

**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION**  
Washington, D.C. 20549

**Amendment No. 2  
to  
FORM S-1  
REGISTRATION STATEMENT  
UNDER  
THE SECURITIES ACT OF 1933**

**Lexeo Therapeutics, Inc.**

(Exact name of registrant as specified in its charter)

Delaware  
(State or other jurisdiction of  
incorporation or organization)

2836  
(Primary Standard Industrial  
Classification Code Number)

85-4012572  
(I.R.S. Employer  
Identification No.)

345 Park Avenue South, Floor 6  
New York, New York 10010  
Tel: (212) 547-9879

(Address, including zip code, and telephone number, including area code, of registrant's principal executive offices)

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**Approximate date of commencement of proposed sale to the public:** As soon as practicable after this Registration Statement is declared effective.

If any of the securities being registered on this Form are to be offered on a delayed or continuous basis pursuant to Rule 415 under the Securities Act of 1933, check the following box.

If this Form is filed to register additional securities for an offering pursuant to Rule 462(b) under the Securities Act, please check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(c) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(d) under the Securities Act, check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, smaller reporting company, or an emerging growth company. See the definitions of "large accelerated filer," "accelerated filer," "smaller reporting company," and "emerging growth company" in Rule 12b-2 of the Exchange Act.

Large accelerated filer   
Non-accelerated filer

Accelerated filer   
Smaller reporting company   
Emerging growth company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 7(a)(2)(B) of the Securities Act.

**The Registrant hereby amends this Registration Statement on such date or dates as may be necessary to delay its effective date until the Registrant shall file a further amendment which specifically states that this Registration Statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act of 1933 or until the Registration Statement shall become effective on such date as the Securities and Exchange Commission, acting pursuant to said Section 8(a), may determine.**

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The information in this preliminary prospectus is not complete and may be changed. These securities may not be sold until the registration statement filed with the Securities and Exchange Commission is effective. This preliminary prospectus is not an offer to sell these securities nor does it seek an offer to buy these securities in any jurisdiction where the offer or sale is not permitted.

Subject to completion, dated \_\_\_\_\_, 2023

**PRELIMINARY PROSPECTUS**



**Common Stock**

This is an initial public offering of shares of common stock of Lexeo Therapeutics, Inc.

We are offering \_\_\_\_\_ shares of our common stock. The initial public offering is expected to be between \$ \_\_\_\_\_ and \$ \_\_\_\_\_ per share.

Prior to this offering, there has been no public market for our common stock. We intend to apply to list our common stock on the Nasdaq Global Market under the trading symbol “LXEO.”

We are an “emerging growth company” and a “smaller reporting company” as defined under U.S. federal securities laws and, as such, will be subject to reduced public company reporting requirements for this prospectus and future filings. See “Prospectus Summary—Implications of Being an Emerging Growth Company and a Smaller Reporting Company.”

**Investing in our common stock involves risks. See “[Risk Factors](#)” beginning on page 15 to read about factors you should consider before buying shares of our common stock.**

**Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities, or passed upon the accuracy or adequacy of this prospectus. Any representation to the contrary is a criminal offense.**

	Per Share	Total
Initial public offering price	\$ _____	\$ _____
Underwriting discounts <sup>(1)</sup>	\$ _____	\$ _____
Proceeds, before expenses, to Lexeo Therapeutics, Inc.	\$ _____	\$ _____

(1) See the section titled “Underwriting” for a description of the compensation payable to the underwriters.

We have granted the underwriters an option for a period of 30 days to purchase up to \_\_\_\_\_ additional shares from us at the public offering price less the underwriting discount.

The underwriters expect to deliver the shares against payment in New York, New York on \_\_\_\_\_, 2023.

**J.P. Morgan**

**SVB Securities**

**Stifel**

**RBC Capital Markets**

**Chardan**

Prospectus dated \_\_\_\_\_, 2023.

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Neither we nor the underwriters have authorized anyone to provide any information or to make any representations other than those contained in this prospectus or in any free writing prospectuses we have prepared. We and the underwriters take no responsibility for, and can provide no assurance as to the reliability of, any other information that others may provide you. We are offering to sell, and seeking offers to buy, shares of common stock only in jurisdictions where offers and sales are permitted. The information contained in this prospectus is accurate only as of the date of this prospectus, regardless of the time of delivery of this prospectus or of any sale of the common stock.

For investors outside of the United States: We have not, and the underwriters have not, done anything that would permit this offering or possession or distribution of this prospectus in any jurisdiction where action for that purpose is required, other than the United States. Persons outside of the United States who come into possession of this prospectus must inform themselves about, and observe any restrictions relating to, the offering of the shares of common stock and the distribution of this prospectus outside of the United States.

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All trademarks, trade names and service marks appearing in this prospectus are the property of their respective owners. Solely for convenience, the trademarks and trade names in this prospectus may be referred to without the ® and ™ symbols, but such references should not be construed as any indicator that their respective owners will not assert their rights thereto.

## Prospectus summary

*This summary highlights, and is qualified in its entirety by, information contained elsewhere in this prospectus. This summary does not contain all of the information you should consider before investing in our common stock. You should read this entire prospectus carefully, especially the sections titled “Risk Factors” and “Management’s Discussion and Analysis of Financial Condition and Results of Operations” and our financial statements and the related notes thereto appearing elsewhere in this prospectus, before making an investment decision. As used in this prospectus, unless the context otherwise requires, references to “we,” “us,” “our,” “the Company,” “Lexeo” and “Lexeo Therapeutics” refer to Lexeo Therapeutics, Inc.*

### Overview

We are a clinical-stage gene therapy company focused on addressing some of the most devastating genetically defined cardiovascular diseases and a genetically defined sub-group of Alzheimer’s disease. We are advancing a deep and diverse pipeline of AAV-based gene therapy candidates for diseases affecting both larger-rare and prevalent patient populations. Utilizing a modular approach that integrates a clinically validated vector, we have a disease area strategy targeting defined patient sub-groups most likely to benefit from our gene therapy candidates, and high-quality, scalable manufacturing, which is designed to overcome many of the challenges facing the field of gene therapy.

Our most advanced cardiovascular product candidate, LX2006 for the treatment of patients with Friedreich’s ataxia, or FA, cardiomyopathy, is currently being evaluated in an ongoing Phase 1/2 clinical trial and we expect to report initial biomarker data from this trial in 2024. Our second most advanced cardiovascular product candidate, LX2020 for the treatment of arrhythmogenic right ventricular cardiomyopathy, or ARVC, caused by mutations in the *PKP2* gene, has completed investigational new drug, or IND, -enabling studies, and we expect to submit an IND application to the U.S. Food and Drug Administration, or FDA, in mid-2023.

Our lead Alzheimer’s disease product candidate, LX1001, for the treatment of *APOE4* homozygous patients with Alzheimer’s disease, is in an ongoing Phase 1/2 trial, and we have initially observed an increase in expression levels of the protective protein, APOE2, in the first and second dose cohorts and a consistent trend towards improvement in core Alzheimer’s disease biomarkers in the first cohort. We expect to report additional biomarker data from the second dose cohort in 2023.

We are targeting diseases that have seen limited penetration of precision medicine, which we define as medications that treat the underlying molecular mechanism of a disease, where we believe there is significant opportunity for gene therapy to play a role as a key therapeutic option. We believe the specific indications we are initially targeting, FA cardiomyopathy, ARVC and *APOE4*-associated Alzheimer’s disease, are highly amenable to gene therapy, where administration of a single dose has the potential to either restore loss-of-function or minimize gain-of-function mutations by treating the underlying genetic cause of the disease. Although few precision medicines are currently approved for the treatment of cardiovascular conditions or Alzheimer’s disease, recent approvals by the FDA suggest a willingness to approve new precision medicines based on biomarkers and functional endpoints. Together with improved diagnostics and increased testing, these developments may offer one of the most substantial opportunities for the uptake of precision medicines in the global pharmaceutical marketplace.

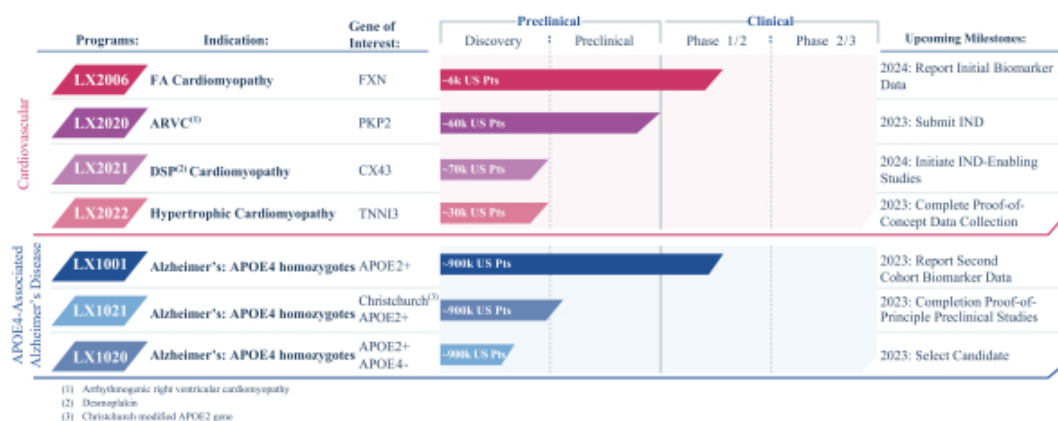
Each of our gene therapy candidates utilizes the vector construct, dose and route of administration that we believe will result in the most favorable biodistribution and safety profile for our product candidate for each disease. Our most advanced cardiovascular and *APOE4*-associated Alzheimer’s disease programs use the

AAVrh10 vector due to its high transduction efficiency in both myocardial cells and neurons, potential for lower toxicity given the opportunity to utilize lower doses compared to other well-established AAV serotypes, and lower pre-existing immunity.

By specifically tailoring our technological approach to each targeted disease, we believe we can optimize our programs to achieve the highest likelihood of having therapeutic impact. We target genetically defined indications in specific sub-groups of patients that offer the potential to demonstrate therapeutic impact through improvement in functional endpoints or biomarkers, have high unmet need and large market opportunities, have established proof-of-concept in relevant preclinical models, and have organized patient advocacy groups and identifiable patient populations. In addition to targeting cardiovascular diseases and APOE4-associated Alzheimer’s disease that we believe can be addressed by our current approach utilizing AAVrh10, we have ongoing discovery efforts to identify next-generation vector technologies with the best potential therapeutic profile. Finally, we continuously seek to bolster our pipeline through relationships with academic institutions, which provide us access to cutting edge gene therapy research that we will utilize in the discovery and development of next generation gene therapy candidates.

## Our pipeline

Utilizing a stepwise, capital-efficient development approach, we are leveraging early proof-of-concept functional and biomarker data to advance a deep and diverse pipeline of cardiovascular and APOE4-associated Alzheimer’s disease programs for larger-rare and prevalent indications. We retain exclusive worldwide development and commercialization rights to all of our product candidates and programs.



### Cardiovascular programs

We are developing a number of disease-modifying gene therapy candidates to treat larger-rare cardiovascular diseases that have significant unmet need and no approved treatments that address the underlying genetic cause of the disease. These programs include:

- LX2006 is an AAVrh10-based gene therapy candidate designed to intravenously deliver a functional frataxin, or *FXN*, gene for the treatment of FA cardiomyopathy. FA cardiomyopathy is the most common cause of mortality in patients with FA and affects approximately 5,600 patients in the United States. LX2006 is designed to promote the expression of the protein frataxin to restore normal mitochondrial function and energy

production in myocardial cells. In preclinical studies, LX2006 demonstrated improvement in cardiac function and survival in a severe *FXN* knockout mouse model. Similarly, in a partial *FXN* knockout mouse model, LX2006 was observed to restore cardiac function and reverse disease abnormalities of FA cardiomyopathy. LX2006 is currently being evaluated in an open-label, dose-escalation Phase 1/2 clinical trial in patients with FA cardiomyopathy, and we expect to report initial biomarker data from this trial in 2024. The FDA has granted Rare Pediatric Disease designation and Orphan Drug designation to LX2006 for the treatment of FA.

- *LX2020* is an AAVrh10-based gene therapy candidate designed to intravenously deliver a fully functional *PKP2* gene to cardiac muscle for the treatment of ARVC caused by mutations in the *PKP2* gene. *PKP2* mutations are associated with approximately 75% of all genetic cases of ARVC, and we estimate they affect approximately 60,000 patients in the United States. *PKP2* mutations can cause replacement of heart muscle with fibrotic tissue and fatty deposits, and severe abnormal heart rhythms, or arrhythmias, that cause cardiac dysfunction and can result in sudden cardiac death. LX2020 is designed to increase desmosomal *PKP2* protein levels, reassemble desmosomes and restore myocardial cell function. In our preclinical studies, using a genetic mouse model of ARVC harboring a *PKP2* mutation that recapitulates the phenotype of ARVC, LX2020 resulted in fewer arrhythmias and increased survival. We intend to submit an IND application for LX2020 in mid-2023.
- *LX2021* is a gene therapy candidate we are developing to intravenously deliver the coding sequence for the functional connexin 43, or Cx43, protein for arrhythmogenic cardiomyopathies. Arrhythmogenic cardiomyopathies, or ACs, are a group of inherited cardiac muscle disorders associated with a high risk of sudden death. We believe restoring the Cx43 protein can potentially treat multiple genetic causes of AC because the cardiac loss of Cx43 is a molecular deficit generally observed in all AC patient populations. Our LX2021 program is initially targeting a distinct form of AC known as Desmoplakin cardiomyopathy, or DSP Cardiomyopathy, with a prevalence that may be as high as 6% of all inherited dilated cardiomyopathies and 7% of all ARVCs, impacting up to approximately 70,000 patients in the United States. We plan to initiate IND-enabling studies for LX2021 by the end of 2024.
- *LX2022* is a gene therapy candidate we are developing to intravenously deliver a functional *TNNI3* gene to myocardial cells to treat a distinct form of hypertrophic cardiomyopathy, or HCM, due to mutations in the *TNNI3* gene. Mutations in *TNNI3* gene often result in left ventricular hypertrophy and restrictive cardiomyopathy, leading to arrhythmias and heart failure. With an estimated prevalence of 1 in 500 people in the United States, HCM is one of the most common forms of genetic cardiomyopathy caused by mutations that affect the cardiac sarcomere in approximately 75% of cases. It is estimated that approximately 30,000 patients in the United States and 36,000 patients in the European Union are affected by HCM caused by mutations in the *TNNI3* gene. We plan to complete proof-of-concept data collection for LX2022 in 2023.

#### *APOE4-associated Alzheimer's disease programs*

We are developing a portfolio of approaches to treat the genetics underlying Alzheimer's disease.

- *LX1001* is an AAVrh10-based gene therapy candidate designed to deliver via one-time administration into the cerebrospinal fluid, or CSF, an *APOE2* gene for the treatment of *APOE4* homozygous patients with Alzheimer's disease. Alzheimer's disease is the leading cause of cognitive decline in late adult life and characterized by complex underlying pathology in the central nervous system, or CNS. Individuals homozygous for *APOE4*, an allele of the gene *APOE*, are approximately 15 times more likely to develop Alzheimer's disease than the general population, and it is estimated that there are 900,000 *APOE4* homozygous patients with Alzheimer's disease in the United States alone. Conversely, individuals homozygous for the *APOE* allele *APOE2* are 40% less likely to develop Alzheimer's disease than the general population. This and other evidence suggest that *APOE2* may play a neuroprotective role. LX1001 is designed

to express the protective *APOE2* gene in the CNS of *APOE4* homozygous patients in order to halt or slow the progression of Alzheimer's disease. LX1001 is being evaluated in an ongoing open-label, dose-escalation Phase 1/2 clinical trial. In the first dose cohort in the trial, we observed a consistent trend towards improvement in Alzheimer's disease CSF biomarkers, such as total tau and phosphorylated tau, that are consistent with improvement in the first three patients with 12 months of data. We have also observed expression of the protective *APOE2* protein in all patients in the first and second dose cohorts with follow-up data. LX1001 has been granted Fast Track designation by the FDA for the treatment of patients with early Alzheimer's disease who are *APOE4* homozygous, to slow disease progression. We expect to report additional biomarker data from the second dose cohort in 2023.

- *LX1021* is a gene therapy candidate we are developing to deliver a Christchurch mutation-modified *APOE2* allele for the treatment of *APOE4* homozygous patients with Alzheimer's disease. The Christchurch mutation has been recognized to protect individuals against Alzheimer's disease even in the presence of significant amyloid pathology. The mechanism of this protection may relate to the fact that *APOE*, in the presence of the Christchurch mutation, binds poorly to heparan sulfate proteoglycans, which are molecules found on the surface of neurons that may inhibit the spread of tau between cells. We believe this approach has the potential to enhance the protective effect of *APOE2* in homozygous *APOE4*-associated Alzheimer's disease. We plan to complete proof-of-principle preclinical studies for LX1021 in 2023.
- *LX1020* is a gene therapy candidate we are developing to deliver both the protective *APOE2* allele and miRNA to suppress *APOE4* for the treatment of *APOE4* homozygous patients. We believe delivery of *APOE2* with concurrent suppression of *APOE4* will achieve a higher degree of conversion to the *APOE4/E2* heterozygous profile, which should lead to greater therapeutic effect. We plan to complete candidate selection in 2023.

## Our approach

Our integrated modular approach enables us to optimize our strategy to pursue larger-rare and prevalent genetically defined indications in specific sub-groups of patients, and is comprised of our technology approach, our precision medicine approach, our disease area strategy, our manufacturing process and academic collaborations.

### *Our technology approach*

We believe that our approach to technology confers the following advantages over existing gene therapy technologies:

- **High Transduction Efficiency and Biodistribution.** The AAVrh10 vector has been shown to be capable of transducing myocardial cells based on preclinical research, and it has been shown to be capable of transducing neurons based on results from preclinical studies and our ongoing clinical trials. We believe these results demonstrate that among currently available, commonly used serotypes, AAVrh10 is an efficient vector for delivery and expression of transgenes for the treatment of the cardiovascular diseases that we are currently targeting and *APOE4*-associated Alzheimer's disease.
- **Reduced Toxicity.** The cardiac tropism of AAVrh10 may allow our gene therapy candidates to be systemically administered at lower doses than many other AAV-based therapies targeting cardiovascular or other systemic diseases.
- **Reduced Pre-Existing Neutralizing Antibodies.** Treatments leveraging vector serotypes to which humans have pre-existing immunity tend to be less effective. Among the naturally occurring and commonly used AAV serotypes, AAVrh10 has been shown in preclinical studies to have among the lowest levels of pre-existing neutralizing antibodies.
- **Optimized Expression.** We are collaborating with our academic partners to develop novel solutions to optimize the potential therapeutic efficacy of our product candidates.



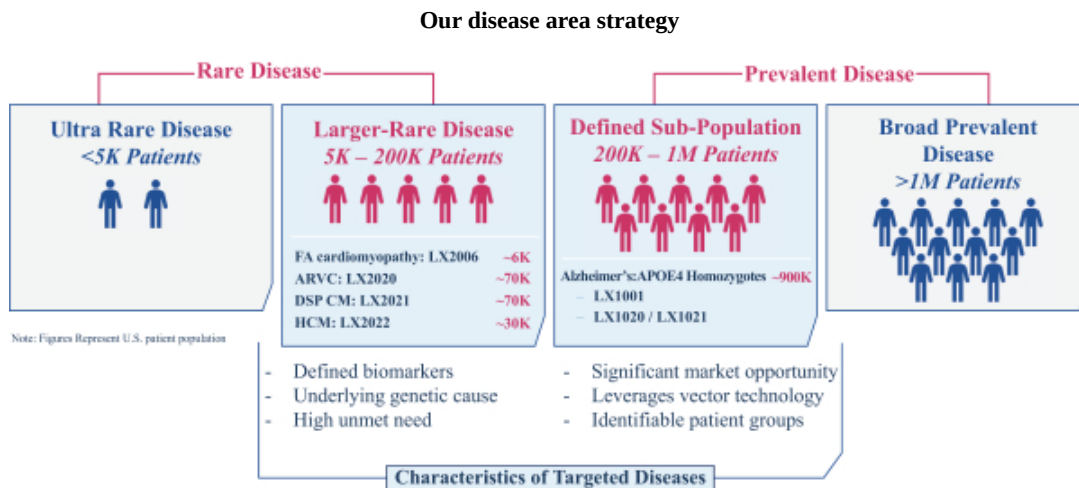
*Our precision medicine approach*

We believe a precision medicine approach for cardiovascular diseases and APOE4-associated Alzheimer’s disease represents a compelling opportunity and the next frontier in expanding the impact of genetic medicines across therapeutic areas:

- *Precision genetic medicines target the underlying cause of the disease.* Next-generation precision based genetic medicines offer the opportunity to directly target the mechanisms underlying cardiovascular disease and Alzheimer’s disease.
- *Significant unmet need in cardiovascular disease and APOE4-associated Alzheimer’s disease.* More than 30 million adults have been diagnosed and approximately 655,000 Americans die yearly from heart disease. Additionally, we believe there are approximately 900,000 APOE4 homozygous patients in the United States, and current treatment options are likely to be less effective for this subpopulation of patients.
- *Improved landscape of regulatory precedents utilizing cardiac and CNS biomarkers.* Biomarkers can help accelerate development timelines, reduce patient attrition, and reduce the overall cost of drug development by providing an early read on potential efficacy and therapeutic impact.
- *Improved diagnostics expand market opportunity.* Implementation of genetic testing has the potential to help identify patients, including those who are asymptomatic or at a higher likelihood of developing the disease being evaluated or who respond less favorably to currently approved treatments.

*Our disease area strategy*

We employ a strategy to select targets in indications that have relatively large patient populations, high unmet need and technically feasible treatment profiles.



Our cardiovascular and APOE4-associated gene therapy programs have the following characteristics:

- *Indications that may be effectively treated by gene therapy.* We select targets that correspond to populations with a specific genetic profile and clearly defined disease phenotype, increasing the homogeneity of our studied patient populations and the potential for the early clinical observation of a beneficial effect.

- **Indications with the potential to demonstrate early evidence of meaningful clinical benefit.** We pursue clearly defined biomarkers and functional endpoints that can provide early proof-of-mechanism and inform clinical development decisions, including the potential to seek accelerated approval pathways.
- **Present opportunity to address high unmet medical need.** We are focused on genetically defined cardiovascular diseases and APOE4-associated Alzheimer's disease where there are no currently approved treatment or where we believe our therapeutic candidates will have a meaningful improvement relative to existing standards of care.
- **Significant market opportunity.** We seek indications with significant commercial opportunities beyond those typically associated with gene therapy companies targeting rare monogenic diseases. Our current focus is on targets that impact larger-rare disease or prevalent disease patient populations.
- **Targets that have established proof-of-concept.** We have leveraged our relationships with academic institutions including Weill Cornell Medical College, or Weill Cornell Medicine, and the University of California, San Diego, or UCSD, to in-license or acquire product candidates with established proof-of-concept in relevant preclinical models that closely resemble the clinical phenotype.
- **Targeted disease areas best treated by optimal delivery technologies.** Our current gene therapy candidates utilize the AAVrh10 vector due to its tropism for the heart and its observed tolerability in clinical trials for CNS disorders. We will pursue the optimal technology to address the diseases of interest for future indications while ensuring sufficient preclinical or clinical validation for any novel approaches.
- **Readily identifiable patient sub-groups.** Our goal is to accelerate patient recruitment for our clinical trials and increase the likelihood of commercial success of our potential products by focusing on diseases with established patient advocacy groups and university researchers who maintain registries of potentially eligible patients. Where possible, we leverage existing natural history studies to better define target patient phenotypes associated with the disease and potentially utilize these natural history datasets as a control group in future studies.

#### *Our manufacturing approach*

We are developing gene therapy candidates for larger-rare and prevalent disease patient populations, which require a high-quality process that can produce vectors in relatively large quantities while utilizing traditional biologics manufacturing infrastructure. We utilize a baculovirus/Sf9 expression system to manufacture our gene therapy candidates. Our manufacturing platform is designed to infect Sf9 cells at high densities in suspension cell culture with both an AAVrh10 and baculovirus containing the transgene. The output is coupled with a chromatography-based purification process which allows for efficient AAV purification, resulting in higher yields and fewer empty AAV capsids than traditional plasmid HEK adherent cell culture approaches. Furthermore, we believe our process has an improved safety profile over traditional adherent HEK manufacturing for the following reasons: we have observed reduced incorporation of non-transgene DNA plasmid DNA impurities, from 15% observed in some HEK systems to 0.2% in our process, our cellular components are non-replicating in mammalian cells; we have eliminated potentially immunogenic or toxic animal-derived proteins; and our cells are grown under serum-free conditions, leading to reduced risk of contamination from animal-derived products.

We believe our manufacturing process enables us to efficiently pursue our goal of targeting larger-rare and prevalent patient populations. Historically, manufacturing challenges largely driven by the quantity of vector required to pursue large commercial opportunities have limited the utility of gene therapy for large patient populations. We believe our proprietary SF9 baculovirus process will allow us to produce vectors at the

necessary scale to support the patient populations we are targeting at a cost-of-goods profile similar to what the pharmaceutical marketplace has seen with biologics. Based on our estimates, we believe our manufacturing process is approximately 10 times more cost efficient than traditional adherent HEK manufacturing, allowing us to potentially produce clinical and commercial material in a more cost efficient manner than peers relying on first-generation manufacturing approaches. We expect that cGMP manufacture of our product candidates using this platform will begin in the second half of 2023.

#### *Academic collaborations*

Our foundational science stems from partnerships and exclusive licenses with leading academic laboratories at Weill Cornell Medicine and UCSD, two preeminent institutions on the cutting edge of gene therapy research. We believe our ongoing work with these institutions, as well as any potential new or expanded collaborations, will continue to be a valuable aspect of our efforts as we seek to further discover and develop novel gene therapies for devastating diseases.

### **Our strategy**

Our vision is a world where gene therapies resolve the burden of disease, transforming the lives of patients. The key elements of our strategy to achieve this vision are to:

- Focus on genetically defined cardiovascular diseases and *APOE4*-associated Alzheimer's disease, two areas of high unmet need and substantial commercial potential.
- Target development efforts at disease pathologies and defined sub-groups of patients most likely to benefit from our therapies.
- Pursue a stepwise and capital-efficient development approach for advancing our programs through clinical development and regulatory approval.
- Utilize a scalable, unified manufacturing platform across all current and future programs.
- Expand our pipeline capabilities through the discovery and development of next-generation gene therapy technologies.
- Leverage and expand upon our partnerships and exclusive licenses with world class academic institutions.
- Build a fully integrated gene therapy company and selectively evaluate strategic opportunities to maximize the impact of our pipeline.

### **Our company and team**

We are led by pioneers and experts with decades of collective experience in genetic medicines, rare disease drug development, manufacturing and commercialization. Our scientific founder and Chief Scientific Advisor, Ronald G. Crystal, M.D., is a world leader in gene therapy research and development and currently serves as Professor and Chairman of Weill Cornell Medicine's Genetic Medicine Department and Director of the Belfer Gene Therapy Core Facility. Our Chief Executive Officer, R. Nolan Townsend, has spent more than a decade working in global biopharmaceutical commercial organizations. Mr. Townsend previously led Pfizer's U.S. rare disease commercial business unit, where he oversaw the successful U.S. launch of blockbuster rare cardiovascular product Vyndaqel/Vyndamax (tafamidis), now a \$2.4 billion product by revenue. Eric Adler, M.D., our Chief Scientific Officer, is a pioneer in cardiovascular gene therapy and Professor of Medicine, head

of the Heart Failure Section, Director of the Strauss Center for Cardiomyopathy, and the Czarina and Humberto S. Lopez Chancellor's Endowed Chair in Cardiology at UCSD. Our Chief Technical Officer, Paul McCormac, Ph.D., has more than 20 years of experience and a proven track record in the field of gene therapy and biologics chemistry, manufacturing, and controls, or CMC. He previously served as Medicinal Sciences Category Lead for Pfizer Rare Disease, where he led gene therapy CMC and vector supply strategy for Pfizer's rare disease business and research units. Our Chief Financial Officer, Micah Zajic, brings nearly 20 years of experience in biotech corporate development, healthcare capital markets and investment banking to our company. Mr. Zajic most recently served as Chief Financial Officer at SQZ Biotechnologies. He began his career at Morgan Stanley in the Global Capital Markets and Healthcare Investment Banking Groups, raising capital and executing strategic transactions for companies across the healthcare industry. Our Chairman, Steven Altschuler, M.D., is currently Managing Director at Ziff Capital Partners and was formerly chairman of the gene therapy pioneer Spark Therapeutics, Inc., which was responsible for the first FDA-approved gene therapy, voretigene neparvovec-rzyl (Luxturna), and was acquired by Roche Holding AG in 2019 for \$4.3 billion.

Since our inception, we have raised \$183.7 million in capital, including from certain funds affiliated with D1 Capital Partners, Eventide Healthcare & Life Sciences Fund, Janus Henderson Investors, Longitude Capital, Lundbeckfonden Ventures, Omega Funds and PBM Capital. Prospective investors should not rely on the investment decisions of our existing investors, as these investors may have different risk tolerances and have received their shares in prior offerings at prices lower than the price offered to the public in this offering. See "Certain Relationships and Related Party Transactions" for more information.

### **Risks associated with our business**

Our business is subject to numerous risks that you should be aware of before making an investment decision. These risks are described more fully in the section titled "Risk Factors" and include, among others:

- We have incurred significant losses since our inception. We expect to incur losses over the next several years and may never achieve or maintain profitability.
- We have a limited operating history and no history of commercializing products, which may make it difficult for an investor to evaluate the success of our business to date and to assess our future viability.
- Even if this offering is successful, we will need substantial additional funding to meet our financial obligations and to pursue our business objectives. If we are unable to raise capital when needed, we could be forced to curtail our planned operations and the pursuit of our growth strategy.
- Our business is dependent on our ability to advance our current and future product candidates through preclinical studies and clinical trials, obtain marketing approval and ultimately commercialize them. If we are unable or experience significant delays in doing so, our business will be materially harmed.
- We are developing novel gene therapy product candidates, which makes it difficult to predict the time, cost and potential success of product candidate development.
- Because gene therapy is novel and of the regulatory landscape that governs any product candidates we may develop (which is rigorous, complex, uncertain and subject to change), we can predict neither the geographic areas in which we could obtain regulatory approval nor the time and cost of obtaining regulatory approval, if we receive it at all, for any product candidates we may develop.
- Preclinical studies and clinical trials are expensive, time-consuming, difficult to design and implement and involve an uncertain outcome. Further, we may encounter substantial delays in completing the development of our product candidates.

- The regulatory approval processes of the FDA, EMA and comparable foreign authorities are lengthy, time consuming and inherently unpredictable. If we are not able to obtain required regulatory approval for our product candidates, our business will be substantially harmed.
- Success in preclinical studies or earlier clinical trials may not be indicative of results in future clinical trials.
- Our preclinical studies and clinical trials may fail to demonstrate the safety and efficacy of our product candidates, or serious adverse or unacceptable side effects may be identified during the development of our product candidates, which could prevent or delay regulatory approval and commercialization, increase our costs or necessitate the abandonment or limitation of the development of some of our product candidates. We may also identify safety and efficacy concerns after the approval of a product candidate which can result in negative consequences to our business and results of operations.
- We and our contract manufacturers are subject to significant regulation with respect to manufacturing our products. The third-party manufacturing facilities on which we rely, and any manufacturing facility that we may have in the future, may have limited capacity or fail to meet the applicable stringent regulatory requirements.
- Gene therapies are novel, complex and difficult to manufacture. We could experience manufacturing problems that result in delays in the development or commercialization of our product candidates or otherwise harm our business.
- We depend on third-party suppliers for materials used in the manufacture of our product candidates, and the loss of these third-party suppliers or their inability to supply us with adequate materials could harm our business.
- If we are unable to obtain or protect intellectual property rights related to any of our product candidates, we may not be able to compete effectively in our market.
- Our future success depends on our ability to retain key executives and advisors and to attract, retain and motivate qualified personnel.

### **Implications of being an emerging growth company and a smaller reporting company**

We qualify as an “emerging growth company” as defined in the Jumpstart Our Business Startups Act of 2012, or the JOBS Act. For so long as we remain an emerging growth company, we may take advantage of relief from certain reporting requirements and other burdens that are otherwise applicable generally to public companies. These provisions include:

- reduced obligations with respect to financial data, including only being required to present two years of audited financial statements, in addition to any required unaudited interim financial statements with correspondingly reduced “Management’s Discussion and Analysis of Financial Condition and Results of Operations” disclosure;
- an exception from compliance with the auditor attestation requirements of Section 404 of the Sarbanes-Oxley Act of 2002, as amended;
- reduced disclosure about our executive compensation arrangements in our periodic reports, proxy statements and registration statements;
- exemptions from the requirements of holding non-binding advisory votes on executive compensation or golden parachute arrangements; and

- an exemption from compliance with the requirements of the Public Company Accounting Oversight Board regarding the communication of critical audit matters in the auditor's report on financial statements.

We may take advantage of these provisions until we no longer qualify as an emerging growth company. We will cease to qualify as an emerging growth company on the date that is the earliest of: (i) the last day of our fiscal year following the fifth anniversary of the date of the completion of this offering, (ii) the last day of the fiscal year in which we have more than \$1.235 billion in total annual gross revenues, (iii) the date on which we are deemed to be a "large accelerated filer" under the rules of the U.S. Securities and Exchange Commission, which means the market value of our common stock that is held by non-affiliates exceeds \$700 million as of the prior June 30, or (iv) the date on which we have issued more than \$1.0 billion of non-convertible debt over the prior three-year period. We may choose to take advantage of some but not all of these reduced reporting burdens. We have taken advantage of certain reduced reporting requirements in this prospectus. Accordingly, the information contained herein may be different than you might obtain from other public companies in which you hold equity interests.

In addition, under the JOBS Act, emerging growth companies can delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have elected to take advantage of the extended transition period to comply with new or revised accounting standards and to adopt certain of the reduced disclosure requirements available to emerging growth companies. As a result of the accounting standards election, we will not be subject to the same implementation timing for new or revised accounting standards as other public companies that are not emerging growth companies, which may make comparison of our financials to those of other public companies more difficult. As a result of these elections, the information that we provide in this prospectus may be different than the information you may receive from other public companies in which you hold equity interests. In addition, it is possible that some investors will find our common stock less attractive as a result of these elections, which may result in a less active trading market for our common stock and higher volatility in our share price.

We are also a "smaller reporting company," meaning that the market value of our shares held by non-affiliates plus the proposed aggregate amount of gross proceeds to us as a result of this offering is less than \$700 million and our annual revenue was less than \$100 million during the most recently completed fiscal year. We may continue to be a smaller reporting company after this offering if either (i) the market value of our shares held by non-affiliates is less than \$250 million or (ii) our annual revenue was less than \$100 million during the most recently completed fiscal year and the market value of our shares held by non-affiliates is less than \$700 million. If we are a smaller reporting company at the time we cease to be an emerging growth company, we may continue to rely on exemptions from certain disclosure requirements that are available to smaller reporting companies. Specifically, as a smaller reporting company, we may choose to present only the two most recent fiscal years of audited financial statements in our Annual Report on Form 10-K and, similar to emerging growth companies, smaller reporting companies have reduced disclosure obligations regarding executive compensation.

## **Corporate information**

In February 2017, we were formed as a Delaware limited liability company under the name LEXEO Therapeutics, LLC. In November 2020, we converted into a Delaware corporation and were renamed Lexeo Therapeutics, Inc. Our principal executive offices are located at 345 Park Avenue South, Floor 6, New York, New York 10010, and our telephone number is (212) 547-9879. Our website address is [www.lexeotx.com](http://www.lexeotx.com). The information contained on, or accessible through, our website is not incorporated by reference into this prospectus.

## The offering

Common stock offered by us	shares
Underwriters' option to purchase additional shares	We have granted the underwriters an option for a period of 30 days from the date of this prospectus to purchase up to additional shares of our common stock.
Common stock to be outstanding immediately after this offering	shares (or shares if the underwriters exercise in full their option to purchase additional shares).
Use of proceeds	<p>We estimate that the net proceeds from this offering will be approximately \$ million, (or approximately \$ million if the underwriters exercise in full their option to purchase up to additional shares of common stock), assuming an initial public offering price of \$ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us.</p> <p>We currently intend to use the net proceeds from this offering, together with our existing cash, to advance the clinical development of LX2006, LX1001 and LX2020, to fund the continued development of our other programs and cardiac discovery efforts, and for working capital and other general corporate purposes. See the section titled "Use of Proceeds" for additional information.</p>
Risk factors	You should read the section titled "Risk Factors" for a discussion of factors you should consider carefully, together with all the other information included in this prospectus, before deciding to invest in our common stock.
Proposed Nasdaq Global Market symbol	"LXEO"
The number of shares of our common stock to be outstanding after this offering is based on	shares of our common stock outstanding as of , 2023 after giving effect to the conversion of all outstanding shares of our convertible preferred stock into an aggregate of shares of common stock in connection with this offering, and excludes:
	<ul style="list-style-type: none"><li>shares of our common stock issuable upon the exercise of options under our 2021 Equity Incentive Plan, or the Existing Plan, granted subsequent to , 2023 at a weighted-average exercise price of \$ per share;</li><li>shares of our common stock reserved for future issuance under the Existing Plan as of , 2023, which shares will cease to be available for issuance at the time our 2023 Equity Incentive Plan, or the 2023 Plan, becomes effective and will be added to, and become available for issuance under, the 2023 Plan;</li></ul>

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- \_\_\_\_\_ shares of our common stock reserved for future issuance under our 2023 Plan, which will become effective in connection with this offering, as well as any automatic increases in the number of shares of common stock reserved for future issuance under the 2023 Plan; and
- \_\_\_\_\_ shares of our common stock reserved for future issuance under our 2023 Employee Stock Purchase Plan, or the 2023 ESPP, which will become effective in connection with this offering, as well as any automatic increases in the number of shares of common stock reserved for future issuance under the 2023 ESPP.

Unless otherwise indicated, all information contained in this prospectus, including the number of shares of common stock that will be outstanding after this offering, assumes or gives effect to:

- the conversion of all outstanding shares of our preferred stock on a one-for-one basis into \_\_\_\_\_ shares of our common stock, which will occur upon the closing of this offering;
- a \_\_\_\_\_-for-\_\_\_\_\_ stock split of our common stock effected on \_\_\_\_\_;
- the filing and effectiveness of our amended and restated certificate of incorporation upon the closing of this offering;
- no exercise of the outstanding options referred to above after \_\_\_\_\_, 2023; and
- no exercise by the underwriters of their option to purchase additional shares of our common stock.



## Summary financial data

The following tables set forth our summary financial data for the three months ended March 31, 2023 and March 31, 2022 and the years ended December 31, 2022 and 2021. We have derived the statement of operations and comprehensive loss data for the three months ended March 31, 2023 and March 31, 2022 and the years ended December 31, 2022 and 2021 and the balance sheet as of March 31, 2023 from our unaudited financial statements included elsewhere in this prospectus. In the opinion of management, the unaudited data reflects all adjustments, consisting only of normal recurring adjustments, considered necessary for a fair presentation of the financial information in those statements.

You should read the following summary data together with our financial statements and the related notes thereto included elsewhere in this prospectus and the section of this prospectus titled “Management’s Discussion and Analysis of Financial Condition and Results of Operations.” The summary financial data in this section are not intended to replace our financial statements and are qualified in their entirety by our financial statements and related notes included elsewhere in this prospectus. Our historical results are not necessarily indicative of the results that should be expected in the future.

	Three Months ended		Year ended December 31	
	2023	March 31 2022	2022	2021
Grant revenue	\$ —	\$ 85,550	\$ 653,956	\$ 1,657,122
<b>Total revenue</b>	<b>—</b>	<b>85,550</b>	<b>653,956</b>	<b>1,657,122</b>
<i>Operating expenses:</i>				
Research and development	16,438,110	10,146,371	49,161,540	45,120,874
General and administrative	2,852,379	2,458,438	12,001,097	7,173,026
<b>Total operating expenses</b>	<b>19,290,489</b>	<b>12,604,809</b>	<b>61,162,637</b>	<b>52,293,900</b>
Loss from operations	(19,290,489)	(12,519,259)	(60,508,681)	(50,636,778)
<i>Other income (expense):</i>				
Other income (expense)	(4,521)	146	(1,852)	(155)
Interest expense	(49,967)	—	(90,874)	—
Interest income	687,495	17,283	1,324,680	15,220
<b>Total other income (expense)</b>	<b>633,007</b>	<b>17,429</b>	<b>1,231,954</b>	<b>15,065</b>
Loss from operations before income taxes	(18,657,482)	(12,501,830)	(59,276,727)	(50,621,713)
Provision for income taxes	—	—	—	—
<b>Net loss and comprehensive loss</b>	<b>\$ (18,657,482)</b>	<b>\$ (12,501,830)</b>	<b>\$ (59,276,727)</b>	<b>\$ (50,621,713)</b>
Net loss per common share, basic and diluted	\$ (1.09)	\$ (0.74)	\$ (3.43)	\$ (3.08)
Weighted average number of shares outstanding used in computation of net loss per common share, basic and diluted	17,069,994	16,863,270	17,272,352	16,449,814
Pro forma net loss per common share, basic and diluted <sup>(1)</sup>				
Pro forma weighted average number of shares outstanding used in computation of net loss per common share, basic and diluted <sup>(1)</sup>				

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- (1) Unaudited pro forma net loss per share, basic and diluted, is calculated giving effect to the automatic conversion of the convertible preferred stock into shares of common stock. Unaudited pro forma net loss per share does not include the shares expected to be sold and related proceeds to be received in this offering. Unaudited pro forma net loss per share for the three months ended March 31, 2023 and the year ended December 31, 2022 was calculated using the weighted-average number of shares of common stock outstanding, including the pro forma effect of the automatic conversion of all outstanding shares of our convertible preferred stock into shares of our common stock, as if the initial public offering had occurred at the beginning of the period or their issuance dates, if later.

	As of March 31, 2023		
	Actual	Pro forma <sup>(1)</sup>	Pro forma as adjusted <sup>(2)</sup>
<b>Balance Sheet Data:</b>			
Cash	\$ 58,853,899	\$	\$
Working capital <sup>(3)</sup>	47,528,254		
Total assets	77,823,873		
Convertible preferred stock	185,033,499		
Accumulated deficit	(134,181,684)		
Total stockholders' equity (deficit)	(131,181,684)		

- (1) The pro forma column reflects the conversion of all of the outstanding preferred shares of our redeemable convertible preferred stock into an aggregate of \_\_\_\_\_ shares of our common stock upon completion of this offering.
- (2) The pro forma as adjusted column reflects the pro forma adjustments set forth above and the sale of \_\_\_\_\_ shares of common stock in this offering at an assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, after deducting estimated underwriting fees and commissions and estimated offering expenses payable by us. This pro forma as adjusted information is illustrative only and will depend on the actual initial public offering price and other terms of this offering determined at pricing. Each \$1.00 increase or decrease in the assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, would increase or decrease the pro forma as adjusted amount of each of cash, working capital, total assets and total stockholders' equity by \$ \_\_\_\_\_ million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting the estimated underwriting discounts and commissions. Each increase or decrease of 1.0 million in the number of shares we are offering would increase or decrease the pro forma as adjusted amount of each of cash, working capital, total assets and total stockholders' equity by \$ \_\_\_\_\_ million, assuming no change in the assumed initial public offering price per share and after deducting the estimated underwriting discounts and commissions.
- (3) We define working capital as current assets less current liabilities. See our financial statements appearing elsewhere in this prospectus.

## Risk factors

*Investing in our common stock involves a high degree of risk. You should consider carefully the risks and uncertainties described below, together with all of the other information in this prospectus, including our financial statements and related notes, before deciding whether to purchase shares of our common stock. The risks and uncertainties described below are not the only ones we face. Additional risks and uncertainties that we are unaware of, or that we currently believe are not material, may also become important factors that affect us. If any of the following risks are realized, our business, financial condition, results of operations and prospects could be materially and adversely affected. In that event, the price of our common stock could decline, and you could lose part or all of your investment.*

### Risks related to our financial position and capital needs

***We have incurred significant losses since our inception. We expect to incur losses over the next several years and may never achieve or maintain profitability.***

Since our inception, we have incurred significant net losses, and we expect to continue to incur significant expenses and operating losses for the foreseeable future. For the fiscal years ended December 31, 2021 and 2022 and the quarter ended March 31, 2023, we incurred net losses of \$50.6 million, \$59.3 million, and \$18.6 million, respectively, and we had an accumulated deficit of \$134.1 million as of March 31, 2023. We have primarily financed our operations with approximately \$183.7 in gross proceeds raised in our private placements of convertible preferred stock in 2020 and 2021. We have no products approved for commercialization and have never generated any revenue from product sales.

We are still in the early clinical stages of development of our product candidates. We expect to continue to incur significant expenses and operating losses over the next several years. Our operating expenses and net losses may fluctuate significantly from quarter to quarter and year to year. We anticipate that our expenses will increase substantially as we:

- continue to advance the preclinical and clinical development of our product candidates and discovery programs;
- initiate and complete additional clinical trials of our current and future product candidates;
- seek regulatory approval for any product candidates that successfully complete clinical trials;
- continue to develop our gene therapy product candidate pipeline;
- scale up our clinical and regulatory capabilities;
- work with our third party manufacturing partners to produce material in accordance with current good manufacturing practices, or cGMP, for clinical trials or potential commercial sales;
- establish, either alone or with a third party, a commercialization infrastructure and scale up manufacturing and distribution capabilities to commercialize any product candidates for which we may obtain regulatory approval;
- adapt our regulatory compliance efforts to incorporate requirements applicable to marketed products;
- maintain, expand and protect our intellectual property portfolio and patent claims;
- hire additional clinical, quality control, regulatory, manufacturing, scientific and administrative personnel;

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- add operational, financial and management information systems and personnel, including personnel to support our product development and planned future commercialization efforts; and
- incur additional legal, accounting and other expenses in operating as a public company.

To date, we have not generated any revenue from the commercialization of our product candidates. To become and remain profitable, we must succeed in developing and eventually commercializing product candidates that generate significant revenue. This will require us to be successful in a range of challenging activities, including completing preclinical testing and clinical trials of our product candidates, obtaining regulatory approval, and manufacturing, marketing and selling any product candidates for which we may obtain regulatory approval, as well as discovering and developing additional product candidates. We are only in the preliminary stages of most of these activities and all of our product candidates are in early clinical trials or preclinical development. We may never succeed in these activities and, even if we do, may never generate any revenue or revenue that is significant enough to achieve profitability.

Even if we achieve profitability, we may not be able to sustain or increase profitability on a quarterly or annual basis. Our failure to become and remain profitable would depress the value of our company and could impair our ability to raise capital, expand our business, maintain our development efforts, obtain product approvals, diversify our offerings or continue our operations. A decline in the value of our company could also cause you to lose all or part of your investment.

***We have a limited operating history and no history of commercializing products, which may make it difficult for an investor to evaluate the success of our business to date and to assess our future viability.***

We are a clinical-stage gene therapy company with a limited operating history. We commenced substantive business operations in 2020, and our operations to date have been largely focused on organizing and staffing our company, business planning, raising capital and entering into collaboration and license agreements for conducting preclinical and clinical research and development activities for our product candidates and gene therapy pipeline. To date, we have not yet demonstrated our ability to successfully complete internally sponsored clinical trials, complete pivotal clinical trials, manufacture a product on a commercial scale, or arrange for a third party to do so on our behalf, or conduct sales and marketing activities necessary for successful commercialization. Consequently, any predictions you make about our future success or viability may not be as accurate as they could be if we had a longer operating history or a history of successfully developing and commercializing products.

***Even if this offering is successful, we will need substantial additional funding to meet our financial obligations and to pursue our business objectives. If we are unable to raise capital when needed, we could be forced to curtail our planned operations and the pursuit of our growth strategy.***

We will require substantial future capital in order to complete planned and future clinical development for our lead product candidates, preclinical development for our other product candidates, and potential commercialization of these product candidates, if any are approved. We expect our spending levels to significantly increase in connection with our planned clinical trials of our lead product candidates. In addition, if we obtain marketing approval for any of our product candidates, we expect to incur significant expenses related to product sales, medical affairs, marketing, manufacturing and distribution. Furthermore, upon the closing of this offering, we expect to incur additional costs associated with operating as a public company. Accordingly, we will need to obtain substantial additional funding in connection with our continuing operations. If we are unable to raise capital when needed or on acceptable terms, we would be forced to delay, reduce or eliminate certain of our clinical trials, our research and development programs or other operations.

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As of March 31, 2023, we had cash and cash equivalents of \$58.9 million. We believe that the anticipated net proceeds from this offering, together with our existing cash, will be sufficient to fund our operating expenses and capital requirements through . This estimate is based on assumptions that may prove to be wrong, and we could use our available capital resources sooner than we expect. Changes may occur beyond our control that would cause us to consume our available capital before that time, including changes in and progress of our development activities, acquisitions of additional product candidates, and changes in regulation. Our future capital requirements will depend on many factors, including:

- the costs of and investment in ongoing and future development of our gene therapy product candidates;
- the scope, progress, costs and results of discovery, preclinical development, laboratory testing and clinical trials for our product candidates;
- the extent to which we develop, in-license or acquire other product candidates and technologies in our gene therapy product candidate pipeline;
- the costs and timing of process development and manufacturing scale-up activities associated with our product candidates and other programs as we advance them through preclinical and clinical development;
- the number of, and development requirements for, product candidates that we may pursue;
- the costs, timing and outcome of regulatory review of our product candidates;
- our headcount growth and associated costs as we expand our research and development capabilities and establish a commercial infrastructure;
- the costs of establishing and maintaining commercial-scale cGMP manufacturing capabilities, either internally or with third parties;
- the costs and timing of future commercialization activities, including product manufacturing, marketing, sales and distribution, for any of our product candidates for which we receive marketing approval;
- the costs and timing of preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending any intellectual property-related claims;
- the revenue, if any, received from commercial sales of our product candidates for which we receive marketing approval;
- our ability to establish and maintain additional collaborations on favorable terms, if at all;
- the success of any collaborations that we may establish and our license agreements;
- the achievement of milestones or occurrence of other developments that trigger payments under our collaboration agreement or any additional collaboration agreements we may enter into; and
- the costs of operating as a public company.

We will require additional capital to achieve our business objectives. While the long-term economic impact of either the COVID-19 pandemic or the conflict between Russia and Ukraine is difficult to assess or predict, each of these events has caused significant disruptions to the global financial markets and contributed to a general global economic slowdown. Furthermore, inflation rates, particularly in the United States, have increased recently to levels not seen in decades. Increased inflation may result in increased operating costs and may affect our operating budgets, specifically with respect to increased labor costs and associated difficulties in recruiting qualified personnel. In addition, the U.S. Federal Reserve has raised, and may further raise, interest rates in response to concerns about inflation. Increases in interest rates, especially if coupled with reduced government

spending and volatility in financial markets, may further increase economic uncertainty and heighten these risks. If the disruptions and slowdown deepen or persist, we may not be able to access additional capital on favorable terms, or at all, which could in the future negatively affect our financial condition and we could be forced to curtail our planned operations and the pursuit of our growth strategy.

***Raising additional capital may cause dilution to our stockholders, restrict our operations or require us to relinquish rights to our technologies or product candidates.***

Until such time, if ever, as we can generate substantial revenue, we may finance our cash needs through a combination of equity offerings, government or private party grants, debt financings and license and collaboration agreements. We do not currently have any committed external source of funds. To the extent that we raise additional capital through the sale of equity or convertible debt securities, your ownership interest will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect your rights as a common stockholder. Debt financing and preferred equity financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making capital expenditures or declaring dividends.

If we raise additional funds through collaborations, strategic alliances or marketing, distribution or licensing arrangements with third parties, we may be required to relinquish valuable rights to our technologies, future revenue streams or product candidates, grant licenses on terms that may not be favorable to us or commit to future payment streams. If we are unable to raise additional funds through equity or debt financings when needed, we may be required to delay, limit, reduce or terminate our product development or future commercialization efforts or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

### **Risks related to the development of our product candidates**

***Our business is dependent on our ability to advance our current and future product candidates through preclinical studies and clinical trials, obtain marketing approval and ultimately commercialize them. If we are unable or experience significant delays in doing so, our business will be materially harmed.***

We have invested a significant portion of our time and financial resources in the development of our product candidates and technology platforms. Our business is dependent on our ability to successfully complete preclinical and clinical development of, obtain regulatory approval for, and, if approved, successfully commercialize LX2006, LX1001, LX2020 and any other product candidates in a timely manner.

Each of our product candidates and programs will require additional preclinical and/or clinical development, regulatory approval and significant marketing efforts, and we will be required to obtain manufacturing supply and expertise and to build a commercial organization or successfully outsource commercialization before we generate any revenue from product sales. We do not have any products that are approved for commercial sale, and we may never be able to develop or commercialize marketable products.

Our ability to generate revenue from our product candidates, which we do not expect to occur for several years, if ever, will depend heavily on the successful development, regulatory approval and eventual commercialization of our product candidates. The success of our lead product candidates, or any other product candidates that we develop or otherwise may acquire will depend on several factors, including:

- timely and successful completion of preclinical studies, including toxicology studies, biodistribution studies and minimally efficacious dose studies in animals, where applicable, under FDA's good laboratory practices, or GLPs;

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- the availability or development of suitable animal disease models for nonclinical studies to enable us to proceed into clinical development or support the submission of a marketing application;
- effective investigational new drug, or IND, applications from the U.S. Food and Drug Administration, or FDA, or comparable foreign applications that allow commencement of our planned clinical trials or future clinical trials for our product candidates;
- sufficiency of our financial and other resources to complete the necessary preclinical studies and clinical trials;
- successful enrollment and completion of clinical trials, including under the FDA's current Good Clinical Practices, or cGCPs;
- establishment of arrangements with third-party manufacturers for our commercial manufacturing processes for any of our product candidates that receive regulatory approval;
- receipt of timely marketing approvals from applicable regulatory authorities;
- launch of commercial sales of products, if approved, whether alone or in collaboration with others;
- acceptance of the benefits and use of our products, including method of administration, if and when approved, by patients, the medical community and third-party payors, for their approved indications;
- the prevalence and severity of adverse events experienced with any of our product candidates;
- the availability, perceived advantages, cost, safety and efficacy of alternative therapies for any diseases for which we are developing our product candidates;
- our ability to produce our product candidates on a commercial scale;
- attainment and maintenance of patent, trademark and trade secret protection and regulatory exclusivity for our product candidates and otherwise protecting our rights in our intellectual property portfolio;
- maintenance of compliance with regulatory requirements such as cGMPs;
- attainment and maintenance of third-party coverage and adequate reimbursement for our product candidates and patients' willingness to pay out-of-pocket in the absence of such coverage and adequate reimbursement; and
- maintenance of a continued acceptable safety, tolerability and efficacy profile of our products following approval.

If we are not successful with respect to one or more of these factors in a timely manner or at all, we could experience significant delays or an inability to successfully commercialize the product candidates we develop, which would materially harm our business. If we do not receive marketing approvals for any product candidate we develop, we may not be able to continue our operations.

### ***We are developing novel gene therapy product candidates, which makes it difficult to predict the time, cost and potential success of product candidate development.***

Our future success depends on the successful development of a novel therapeutic approach. To date, very few products that utilize gene transfer have been approved in the United States or Europe. There have been a limited number of clinical trials using AAVrh10. Although gene therapies have been studied in human clinical trials for over 30 years, there are few AAV-based gene therapy products approved by the FDA: voretigene neparvovec rzyt (Luxturna), for the treatment of a rare ocular condition; onasemnogene abeparvovec xioi (Zolgensma), for

the treatment of a rare neuromuscular condition; etranacogene dezaparvovec (Hemegenix), for the treatment of adults with Hemophilia B who currently use Factor IX prophylaxis therapy; and nadofaragene firadenovec-vnvcg (Adstiladrin), for an unresponsive non-muscle invasive bladder cancer.

We cannot be certain that our AAVrh10-based gene therapy product candidates will successfully complete clinical trials or that any future product candidates utilizing this or other vector constructs will successfully complete preclinical studies or clinical trials. We may not be successful in developing product candidates that avoid triggering toxicities or other side effects in preclinical studies or clinical trials. Our intravenous, intracisternal, intrathecal and intraparenchymal routes of administration may cause unforeseen side effects or present other challenges. Any such results could impact our ability to develop a product candidate, including our ability to enroll patients in our clinical trials. As a result of these factors, it is more difficult for us to predict the time and cost of product candidate development, and we cannot predict whether the application of our approach to gene therapy, or any similar or competitive programs, will result in the identification, development, and regulatory approval of any product candidate, or that other gene therapy programs will not be considered better or more favorable. There can be no assurance that any development problems we experience in the future related to our current gene therapy product candidates or any of our research programs will not cause significant delays or unanticipated costs, or that such development problems can be solved. We may also experience delays and challenges in achieving sustainable, reproducible, and scalable production. Any of these factors may prevent us from completing our preclinical studies or clinical trials or commercializing any product candidates we may develop on a timely or profitable basis, if at all.

***Because gene therapy is novel and of the regulatory landscape that governs any product candidates we may develop (which is rigorous, complex, uncertain and subject to change), we can predict neither the geographic areas in which we could obtain regulatory approval nor the time and cost of obtaining regulatory approval, if we receive it at all, for any product candidates we may develop.***

The regulatory requirements that will govern any novel gene therapy product candidates we develop are not entirely clear and are subject to change. The novel nature of our capsids makes it difficult to further determine how long it will take or how much it will cost to obtain regulatory approvals for our product candidates in the United States, the European Union or other jurisdictions. Within the broader genetic medicine field, very few gene therapy products have received marketing authorization from the FDA or the European Medicines Agency, or EMA. Even with respect to gene therapies, the regulatory landscape is still developing. Regulatory requirements governing gene therapy products have changed frequently and will likely continue to change in the future. Moreover, there is substantial overlap in those responsible for regulation of existing gene therapy products. For example, in 2016, the FDA established the Office of Tissues and Advanced Therapies, or OTAT, within the Center for Biologics Evaluation and Research, or CBER, to consolidate the review of gene therapy and related products, and to advise the CBER on its review. In September 2022, the FDA announced retitling of OTAT to the Office of Therapeutic Products, or OTP, and elevation of OTP to a “Super Office” to meet its growing cell and gene therapy workload.

Our product candidates will need to meet safety and efficacy standards applicable to any new biologic being pursued for a given disease under the regulatory framework administered by the FDA. Although the FDA decides whether individual gene therapy protocols may proceed, the review process and determinations of other reviewing bodies, including Institutional Review Boards, or IRBs, can impede or delay the initiation of a clinical trial.

The same applies in the EU. The EMA’s Committee for Advanced Therapies, or CAT, is responsible for assessing the quality, safety and efficacy of advanced-therapy medicinal products. Advanced-therapy medicinal products include gene therapy medicines, somatic-cell therapy medicines and tissue-engineered medicines. The



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role of the CAT is to prepare a draft opinion on an application for marketing authorization for a gene therapy medicinal candidate that is submitted to the EMA. In the EU, the development and evaluation of a gene therapy product must be considered in the context of the relevant EU guidelines. The EMA may issue new guidelines concerning the development and marketing authorization for gene therapy products and require that we comply with these new guidelines. This could mean that any gene therapy product candidate we may develop in the future could be required to comply with additional and/or more stringent gene therapy guidelines in the EU.

Adverse developments in preclinical studies or clinical trials conducted by others in the field of gene therapy and gene regulation products may cause the FDA, the EMA and other regulatory bodies to revise the requirements for approval of any product candidates we may develop or limit the use of products utilizing gene regulation technologies, either of which could harm our business. In addition, the clinical trial requirements of the FDA, the EMA and other regulatory authorities and the criteria these regulators use to determine the safety and efficacy of a product candidate vary substantially according to the type, complexity, novelty, and intended use and market of the potential products. The regulatory approval process for product candidates such as ours can be more expensive and take longer than for other, better known, or more extensively studied pharmaceutical or other product candidates. Further, as we are developing novel potential treatments for diseases in which, in some cases, there is little clinical experience with potential new endpoints and methodologies, there is heightened risk that the FDA, the EMA or other regulatory bodies may not consider the clinical trial endpoints to provide clinically meaningful results, and the resulting clinical data and results may be more difficult to analyze. In addition, we may not be able to identify or develop appropriate animal disease models to enable or support planned clinical development. Any natural history studies that we may rely upon in our clinical development may not be accepted by the FDA, EMA or other regulatory authorities. Regulatory agencies administering existing or future regulations or legislation may not allow production and marketing of products utilizing gene regulation technology in a timely manner or under technically or commercially feasible conditions. In addition, regulatory action or private litigation could result in expenses, delays, or other impediments to our research programs or the commercialization of resulting products. Further, approvals by one regulatory agency may not be indicative of what other regulatory agencies may require for approval.

The regulatory review committees and advisory groups described above and any new guidelines they promulgate may lengthen the regulatory review process, require us to perform additional preclinical studies or clinical trials, increase our development costs, lead to changes in regulatory positions and interpretations, delay or prevent approval and commercialization of these treatment candidates, or lead to significant post-approval limitations or restrictions. As we advance our research programs and develop future product candidates, we will be required to consult with these regulatory and advisory groups and to comply with applicable guidelines. If we fail to do so, we may be required to delay or discontinue development of any product candidates we identify and develop. These additional processes may result in a review and approval process that is longer than we otherwise would have expected. Delays as a result of an increased or lengthier regulatory approval process or further restrictions on the development of our product candidates can be costly and could negatively impact our ability to complete clinical trials and commercialize our current and future product candidates in a timely manner, if at all.

***Preclinical studies and clinical trials are expensive, time-consuming, difficult to design and implement and involve an uncertain outcome. Further, we may encounter substantial delays in completing the development of our product candidates.***

All of our product candidates are in preclinical or early clinical development, and the risk of failure is high. The preclinical studies, clinical trials and manufacturing of our product candidates are, and the manufacturing and marketing of our products, if approved, will be, subject to extensive and rigorous review and regulation by numerous government authorities in the United States and in other countries where we may test and market our

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product candidates. Before obtaining regulatory approvals for the commercial sale of any of our product candidates, we must demonstrate through lengthy, complex and expensive preclinical testing and clinical trials that our product candidates are both safe and effective for use in each target disease. In particular, because our product candidates are subject to regulation as biologics, we will need to demonstrate that they are safe and of sufficient purity and potency for use in their target diseases. Each product candidate must demonstrate an adequate risk-versus-benefit profile in its intended patient population and for its intended use.

Clinical testing is expensive, can take many years to complete and is subject to uncertainty. We cannot guarantee that any clinical trials will be initiated on schedule, conducted as planned or completed on schedule, if at all. To date, we are sponsoring clinical trials of LX1001 and LX2006, but we have not successfully completed any clinical trial that we have internally sponsored. Failure can occur at any time during the clinical trial process. Even if our ongoing and future clinical trials are completed as planned, we cannot be certain that their results will support the safety and effectiveness of our product candidates for their targeted diseases or support continued clinical development of such product candidates. Our future clinical trial results may not be successful.

In addition, even if such trials are successfully completed, we cannot guarantee that the FDA or foreign regulatory authorities will interpret the results as we do, and more trials could be required before we submit our product candidates for approval. This is particularly true for clinical trials in rare diseases, where the small patient populations make it difficult or impossible to conduct two traditional, adequate and well-controlled trials, and therefore the FDA or comparable foreign regulatory authorities are often required to exercise flexibility in approving therapies for such diseases. Moreover, results acceptable to support approval in one jurisdiction may be deemed inadequate by another regulatory authority to support regulatory approval in that other jurisdiction. To the extent that the results of the trials are not satisfactory to the FDA or foreign regulatory authorities for support of a marketing application, we may be required to expend significant resources, which may not be available to us, to conduct additional trials in support of potential approval of our product candidates.

We may experience delays in initiating and conducting clinical trials of our lead product candidates and we do not know whether our clinical trials will begin on time, need to be redesigned, recruit and enroll patients on time or be completed on schedule, or at all. Events that may prevent successful or timely completion of clinical development include:

- inability to generate sufficient preclinical, toxicology, or other *in vivo* or *in vitro* data to support the initiation of clinical trials;
- delays in sufficiently developing, characterizing or controlling a manufacturing process suitable for advanced clinical trials;
- delays in sourcing or qualifying ancillaries required for administration of our clinical drug product (such as vials, stoppers, or tubing);
- delays in developing suitable assays for screening patients for eligibility for trials with respect to certain product candidates;
- delays in reaching agreement with the FDA, EMA or other regulatory authorities as to the design or implementation of our clinical trials;
- obtaining regulatory approval to commence a clinical trial;
- reaching an agreement on acceptable terms with clinical trial sites or prospective contract research organizations, or CROs, the terms of which can be subject to extensive negotiation and may vary significantly among different clinical trial sites;
- obtaining IRB approval for each clinical trial site;

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- recruiting suitable patients to participate in a clinical trial in a timely manner;
- having patients complete a clinical trial or return for post-treatment follow-up;
- clinical trial sites, CROs or other third parties deviating from trial protocol or dropping out of a trial;
- failure to perform our planned clinical trials in accordance with the FDA's cGCP requirements, or applicable regulatory guidelines in other countries;
- addressing patient-safety concerns that arise during the course of a trial, including occurrence of adverse events associated with the product candidate that are viewed to outweigh its potential benefits;
- adding a sufficient number of clinical trial sites; or
- manufacturing sufficient quantities of a product candidate for use in clinical trials.

We may experience numerous unforeseen events during, or as a result of, clinical trials that could delay or prevent our ability to receive marketing approval or commercialize our product candidates or significantly increase the cost of such trials, including:

- we may experience changes in regulatory requirements or guidance, or receive feedback from regulatory authorities that requires us to modify the design of our clinical trials;
- clinical trials of our product candidates may produce negative or inconclusive results, and we may decide, or regulators may require us, to conduct additional clinical trials or halt development programs;
- the number of patients required for clinical trials of our product candidates may be larger than we anticipate, enrollment in these clinical trials may be slower than we anticipate, or participants may drop out of these clinical trials at a higher rate than we anticipate;
- our third-party contractors may fail to comply with regulatory requirements or meet their contractual obligations to us in a timely manner, or at all;
- we, our investigators or regulators may suspend or terminate clinical trials of our product candidates for various reasons, including non-compliance with regulatory requirements, a finding that our product candidates have undesirable side effects or other unexpected characteristics, or a finding that the participants are being exposed to unacceptable health risks;
- the cost of clinical trials of our product candidates may be greater than we anticipate, and we may not have funds to cover the costs;
- the supply or quality of our product candidates or other materials necessary to conduct clinical trials of our product candidates may be insufficient or inadequate;
- regulators may revise the requirements for approving our product candidates, or such requirements may not be as we anticipate; and
- any future collaborators that conduct clinical trials may face any of the above issues and may conduct clinical trials in ways they view as advantageous to them but that are suboptimal for us.

If we are required to conduct additional clinical trials or other testing of our product candidates beyond those that we currently contemplate, if we are unable to successfully initiate or complete clinical trials of our product candidates or other testing, if the results of these trials or tests are not positive or are only modestly positive or if there are safety concerns, we may:

- incur unplanned costs;

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- be delayed in obtaining marketing approval for our product candidates or not obtain marketing approval at all;
- obtain marketing approval in some countries and not in others;
- obtain marketing approval for diseases or patient populations that are not as broad as intended or desired;
- obtain marketing approval with labeling that includes significant use or distribution restrictions or safety warnings, including boxed warnings or Risk Evaluation and Mitigation Strategies, or REMS;
- be subject to additional post-marketing testing requirements; or
- have the product removed from the market after obtaining marketing approval.

We could encounter delays if a clinical trial is suspended or terminated by us, by the IRBs of the institutions in which such trials are being conducted, by the Data Safety Monitoring Board for such trial or by the FDA, EMA or other regulatory authorities. Such authorities may impose such a suspension or termination due to a number of factors, including failure to conduct the clinical trial in accordance with regulatory requirements or our clinical protocols, inspection of the clinical trial operations or trial site by the FDA, EMA or other regulatory authorities resulting in the imposition of a clinical hold, unforeseen safety issues or adverse side effects, failure to demonstrate a benefit from using a drug, changes in governmental regulations or administrative actions or lack of adequate funding to continue the clinical trial.

All of our product candidates will require extensive clinical testing before we are prepared to submit a biologics license application, or BLA, or marketing authorization application, or MAA, for regulatory approval. We cannot predict with any certainty if or when we might complete the clinical development for our product candidates and submit a BLA or MAA for regulatory approval of any of our product candidates or whether any such BLA or MAA will be approved. We may also seek feedback from the FDA, EMA or other regulatory authorities on our clinical development program, and the FDA, EMA or such regulatory authorities may not provide such feedback on a timely basis, or such feedback may not be favorable, which could further delay our development programs.

We cannot predict with any certainty whether or when we might complete a given clinical trial. If we experience delays in the commencement or completion of our clinical trials, or if we terminate a clinical trial prior to completion, the commercial prospects of our product candidates could be harmed, and our ability to generate revenues from our product candidates may be delayed or lost. In addition, any delays in our clinical trials could increase our costs, slow down the development and approval process and jeopardize our ability to commence product sales and generate revenues. Any of these occurrences may harm our business, financial condition and results of operations. In addition, many of the factors that cause, or lead to, a delay in the commencement or completion of clinical trials may also ultimately lead to the denial of regulatory approval of our product candidates.

***The regulatory approval processes of the FDA, EMA and comparable foreign authorities are lengthy, time-consuming and inherently unpredictable. If we are not able to obtain required regulatory approval for our product candidates, our business will be substantially harmed.***

The time required to obtain approval or other marketing authorizations by the FDA, EMA and comparable foreign authorities is unpredictable, and it typically takes many years following the commencement of clinical trials and depends upon numerous factors, including the substantial discretion of the regulatory authorities. In addition, approval policies, regulations, and the type and amount of clinical data necessary to gain approval may change during the course of a product candidate's clinical development and may vary among jurisdictions. We have not obtained regulatory approval for any product candidate, and it is possible that we may never obtain regulatory approval for any product candidates we may seek to develop in the future. Neither we nor any collaborator is permitted to market any biologic product candidates in the United States until we receive

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regulatory approval of a BLA from the FDA, and we cannot market it in the European Union until we receive approval for an MAA from the EMA, or other required regulatory approval in other countries.

Prior to obtaining approval to commercialize any product candidate in the United States or abroad, we must demonstrate with substantial evidence from well-controlled clinical trials, and to the satisfaction of the FDA or foreign regulatory agencies, that such product candidates are safe, effective and of sufficient purity for their intended uses. Results from preclinical studies and clinical trials can be interpreted in different ways. Even if we believe the preclinical or clinical data for our product candidates are promising, such data may not be sufficient to support approval by the FDA and other regulatory authorities. The FDA may also require us to conduct additional preclinical studies or clinical trials for our product candidates either prior to or after approval, or it may object to elements of our clinical development programs, or require changes to our manufacturing approaches.

Of the large number of products in development, only a small percentage successfully complete the FDA or foreign regulatory approval processes and are commercialized. The lengthy approval and marketing authorization process as well as the unpredictability of future clinical trial results may result in our failing to obtain regulatory approval and marketing authorization to market our product candidates, which would significantly harm our business, financial condition, results of operations and prospects.

We have invested a significant portion of our time and financial resources in the development of our product candidates and technology platforms. Our business is dependent on our ability to successfully complete preclinical and clinical development of, obtain regulatory approval for, and, if approved, successfully commercialize LX2006, LX1001 and our other product candidates in a timely manner.

Even if we eventually complete clinical testing and receive approval of a BLA or foreign marketing application for any of our product candidates, the FDA, EMA or the applicable foreign regulatory agency may grant approval or other marketing authorization contingent on the performance of costly additional clinical trials, including post-marketing clinical trials. The FDA, EMA or the applicable foreign regulatory agency also may approve or authorize for marketing a product candidate for a more limited disease or patient population than we originally request, and the FDA, EMA or applicable foreign regulatory agency may not approve or authorize the labeling that we believe is necessary or desirable for the successful commercialization of a product candidate. Any delay in obtaining, or inability to obtain, applicable regulatory approval or other marketing authorization would delay or prevent commercialization of that product candidate and would materially adversely impact our business and prospects.

In addition, the FDA, EMA and other regulatory authorities may change their policies, issue additional regulations or revise existing regulations, or take other actions, which may prevent or delay approval of our future products under development on a timely basis. Such policy or regulatory changes could impose additional requirements upon us that could delay our ability to obtain approvals, increase the costs of compliance or restrict our ability to maintain any marketing authorizations we may have obtained.

### ***Success in preclinical studies or earlier clinical trials may not be indicative of results in future clinical trials.***

Success in preclinical testing and early clinical trials does not ensure that later clinical trials will generate the same results or otherwise provide adequate data to demonstrate the efficacy and safety of a product candidate. Preclinical tests and Phase 1 and Phase 2 clinical trials are primarily designed to test safety, to study pharmacokinetics and pharmacodynamics and to understand the side effects of product candidates at various doses and schedules. Success in preclinical or animal studies and early clinical trials does not ensure that later large-scale efficacy and safety trials will be successful, nor does it predict final results. For example, we may be unable to identify suitable animal disease models for our product candidates, which could delay or frustrate our

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ability to proceed into clinical trials or obtain marketing approval. In addition, the preclinical studies conducted by Stelios Therapeutics, Inc., or Stelios (an entity that we acquired in 2021), and UCSD for our product candidates LX2021 and LX2022 employed an AAV9-based formulation and studies using this vector may not be predictive of future testing we intend to conduct using an AAVrh10-based formulation.

Our product candidates may fail to show the desired safety and efficacy in clinical development despite positive results in preclinical studies or having successfully advanced through initial clinical trials. Furthermore, our currently ongoing and most future clinical trials involve or will involve a small patient population. Because of the small sample sizes studied in our trials thus far, the results of these trials may not be indicative of results of future clinical trials.

Additionally, some of our ongoing and planned clinical trials utilize, or may utilize, an “open-label” trial design. An “open-label” clinical trial is one where both the patient and investigator know whether the patient is receiving the investigational product candidate or either an existing approved drug or placebo. Most typically, open-label clinical trials test only the investigational product candidate and sometimes may do so at different dose levels. Open-label clinical trials are subject to various limitations that may exaggerate any therapeutic effect as patients in open-label clinical trials are aware when they are receiving treatment. Open-label clinical trials may be subject to a “patient bias” where patients perceive their symptoms to have improved merely due to their awareness of receiving an experimental treatment. In addition, open-label clinical trials may be subject to an “investigator bias” where those assessing and reviewing the physiological outcomes of the clinical trials are aware of which patients have received treatment and may interpret the information of the treated group more favorably given this knowledge. The results from an open-label trial may not be predictive of future clinical trial results with any of our product candidates for which we include an open-label clinical trial when studied in a controlled environment with a placebo or active control.

Many companies in the pharmaceutical and biotechnology industries have suffered significant setbacks in late-stage clinical trials even after achieving promising results in preclinical testing and earlier-stage clinical trials. Data obtained from preclinical and clinical activities are subject to varying interpretations, which may delay, limit or prevent regulatory approval. In addition, we may experience regulatory delays or rejections as a result of many factors, including changes in regulatory policy during the period of our product candidate development. Any such delays could negatively impact our business, financial condition, results of operations and prospects.

***Interim “top-line” and preliminary results from our clinical trials that we announce or publish from time to time may change as more patient data become available and are subject to audit and verification procedures that could result in material changes in the final data.***

From time to time, we may publish interim top-line or preliminary results from our clinical trials. Interim results from clinical trials that we may complete are subject to the risk that one or more of the clinical outcomes may materially change as patient enrollment continues and more patient data become available. Preliminary or top-line results also remain subject to audit and verification procedures that may result in the final data being materially different from the preliminary data we previously published. As a result, interim and preliminary data should be viewed with caution until the final data are available. Differences between preliminary or interim data and final data could significantly harm our business prospects and may cause the trading price of our common stock to fluctuate significantly.

***Our preclinical studies and clinical trials may fail to demonstrate the safety and efficacy of our product candidates, or serious adverse or unacceptable side effects may be identified during the development of our product candidates, which could prevent or delay regulatory approval and commercialization, increase our costs or necessitate the abandonment or limitation of the development of some of our product candidates. We may also identify safety and efficacy concerns after the approval of a product candidate which can result in negative consequences to our business and results of operations.***

Before obtaining regulatory approvals for the commercial sale of our product candidates, we must demonstrate through lengthy, complex and expensive preclinical testing and clinical trials that our product candidates are safe, effective and of sufficient purity for use in each target disease, and failures can occur at any stage of testing. Preclinical studies and clinical trials often fail to demonstrate safety or efficacy of the product candidate studied for the target disease. While we have developed our AAVrh10-mediated gene therapy product candidates to leverage the low seropositivity of AAVrh10, any gene therapy product based on viral vectors carries the risks of immunogenicity, elevated liver enzymes and insertional oncogenesis, which is the process whereby the insertion of a functional gene near a gene that is important in cell growth or division results in uncontrolled cell division, which could potentially enhance the risk of malignant transformation. In one of our preclinical studies of LX2006, we observed one case of hepatocellular carcinoma, or HCC, in a wild-type mouse at 10 months post-treatment. Although data reported by the FDA Cellular, Tissue, and Gene Therapies Advisory Committee in September 2021 suggests that HCC observed in mice after AAV treatment is unlikely to translate to risks for humans, any future instances of HCC in our clinical trials could result in delays or the abandonment of our trials. Health authorities also ask that sponsors closely monitor the risk of elevated liver enzymes and abnormal liver ultrasound on a routine basis in patients participating in gene therapy clinical trials. Patients in our trial of LX1004, our AAVrh10-based gene therapy candidate designed to restore TPP1, the secreted protein that is deficient in CLN2 Batten disease, reported six acute serious adverse events within two weeks of dosing and 15 chronic serious adverse events after two weeks of dosing that were deemed to be related to the study drug or its administration. The six acute serious adverse events consisted of one patient with seizures, two patients with abnormal movements, one patient with emesis, one patient with hematoma, and one patient with pneumocephalus. The 15 chronic serious adverse events consisted of six patients with seizures, one patient with dystonia, one patient with emesis, one patient with hematoma, one patient with hygroma, one patient with bronchospasm, one patient with aspiration, one patient with pneumonia, and two patients with elevated hepatic enzymes.

Possible adverse side effects that could occur with treatment with gene therapy products include an immunologic reaction early after administration, which, while not necessarily adverse to the patient's health, could substantially limit the effectiveness of the treatment. For example, in previous third-party clinical trials involving AAV capsids for gene therapy, some subjects experienced the development of a T-cell antibody response, whereby after the vector is within the target cell types, the cellular immune response system triggers the removal of transduced cell types by activated T-cells. If any of our product candidates demonstrate a similar effect, we may decide or be required to perform additional preclinical studies or to halt or delay further clinical development of our product candidates.

In addition to side effects caused by the product candidate, the administration process or related procedures also can cause adverse side effects. Our APOE-associated Alzheimer's disease product candidates are designed to be delivered via intracisternal administration. While the intracisternal method of administration has been available for some years, its use for gene therapies is new and no gene therapy is currently approved for this method of administration. Intracisternal administration may have greater risk and/or be perceived as having greater risk than more common methods of administration, such as intravenous injection, while parenchymal administration involves neurosurgery and carries all risks attendant with such a procedure. These risks include allergic reaction to anesthesia, bleeding or swelling in the brain, seizures, stroke, coma and infection. Other gene therapy product candidates in clinical development utilizing intracisternal or parenchymal delivery could also generate data that could adversely affect the clinical, regulatory or commercial perception of our product candidates.

If adverse events occur, either as a result of the product candidate or administration process, our clinical trials could be suspended or terminated. If we cannot demonstrate that any adverse events were not caused by the drug or administration process or related procedures, the FDA, EMA or foreign regulatory authorities could order us

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to cease further development of, or deny approval of, our product candidates for any or all targeted diseases. Even if we are able to demonstrate that all future serious adverse events are not product-related, such occurrences could affect patient recruitment or the ability of enrolled patients to complete the trial. Moreover, if we elect, or are required, to not initiate, delay, suspend or terminate any future clinical trial of any of our product candidates, the commercial prospects of such product candidates may be harmed and our ability to generate product revenues from any of these product candidates may be delayed or eliminated. Any of these occurrences may harm our ability to develop other product candidates, and may harm our business, financial condition and prospects significantly. Furthermore, negative results in our development of LX2006, LX1001, or LX2020 could be interpreted as a failure to achieve proof of concept for our technology and result in the abandonment of other development programs.

In addition, gene therapy is still a relatively new approach to disease treatment and additional adverse side effects could develop. There also is the potential risk of delayed adverse events following exposure to gene therapy products due to persistent biologic activity of the genetic material or other components of products used to carry the genetic material. If our product candidates are associated with side effects in clinical trials or have characteristics that are unexpected, we may need to abandon their development or limit development to more narrow uses in which the side effects or other characteristics are less prevalent, less severe or more acceptable from a risk-benefit perspective. The FDA or an IRB may also require that we suspend, discontinue, or limit our clinical trials based on safety information, or that we conduct additional animal or human studies regarding the safety and efficacy of our product candidates which we have not planned or anticipated. Such findings could further result in regulatory authorities failing to provide marketing authorization for our product candidates or limiting the scope of the approved indication, if approved. Many product candidates that initially showed promise in early-stage testing have later been found to cause side effects that prevented further development of the product candidate.

Additionally, if one or more of our product candidates receives marketing approval, and we or others identify undesirable side effects caused by such products or the administration procedure, a number of potentially significant negative consequences could result, including:

- regulatory authorities may withdraw approvals of such product;
- regulatory authorities may require additional warnings on the labels;
- we may be required to create a medication guide outlining the risks of such side effects for distribution to patients or other requirements subject to a REMS;
- we could be sued and held liable for harm caused to patients;
- we may not be able to obtain or maintain third-party payor coverage and adequate reimbursement; and
- our reputation and physician or patient acceptance of our products may suffer.

There can be no assurance that we will resolve any issues related to any product-related adverse events to the satisfaction of the FDA or foreign regulatory agency in a timely manner or at all. Moreover, any of these events could prevent us from achieving or maintaining market acceptance of the particular product candidate, if approved, and could significantly harm our business, results of operations and prospects.

***Some of the diseases we initially seek to treat have low prevalence and it may be difficult to identify and enroll patients with these diseases. If we experience delays or difficulties in the enrollment and/or maintenance of patients in clinical trials, our receipt of necessary regulatory approvals could be delayed or prevented.***

Successful and timely completion of clinical trials will require that we enroll a sufficient number of patients. Patient enrollment, a significant factor in the timing of clinical trials, is affected by many factors, including the size and nature of the patient population and competition for patients with other trials. The rare genetic diseases which some of our product candidates are designed to target have low incidence and prevalence. In particular, because we are focused on patients with specific genetic mutations, our ability to enroll eligible patients may be limited or enrollment may be slower than we anticipate. For example, we estimate that approximately 5,600



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people in the United States have Friedreich's ataxia, or FA, and that approximately 70% of these patients will develop the cardiac manifestation of FA, or FA cardiomyopathy, and accordingly it may be difficult for us to identify and timely recruit a sufficient number of eligible patients to conduct our clinical trials. Even for more prevalent conditions such as Alzheimer's disease, it may be difficult to recruit patients to clinical trials due to the number of approved products, difficulty identifying patients with the specific genotype we are studying, and the number of clinical trials being conducted in this indication.

Our trials may be subject to delays as a result of patient enrollment taking longer than anticipated or patient withdrawal. We may not be able to initiate or continue clinical trials for our product candidates if we are unable to locate and enroll a sufficient number of eligible patients to participate in these trials as required by the FDA, EMA or other foreign regulatory authorities. We cannot predict how successful we will be at enrolling subjects in future clinical trials. Subject enrollment is affected by other factors including:

- the severity of the disease under investigation;
- the eligibility criteria for the trial in question;
- the size of the patient population and process for identifying patients;
- the perceived risks and benefits of the product candidate under study;
- clinicians' and patients' perceptions as to the potential advantages of the product candidate being studied in relation to other available therapies, including any new drugs that may be approved for the diseases we are investigating;
- the availability of competing commercially available therapies and other competing therapeutic candidates' clinical trials;
- the efforts to facilitate timely enrollment in clinical trials;
- the risk that patients enrolled in clinical trials will drop out of the clinical trials before completion of their treatment;
- the patient referral practices of physicians;
- the ability to monitor patients adequately during and after treatment; and
- the proximity and availability of clinical trial sites for prospective patients.

Our inability to enroll a sufficient number of patients for clinical trials would result in significant delays and could require us to abandon one or more clinical trials altogether. Enrollment delays in these clinical trials may result in increased development costs for our product candidates, which would cause the value of our company to decline and limit our ability to obtain additional financing. Furthermore, we rely on CROs and clinical trial sites to help ensure the proper and timely conduct of our clinical trials and we may have limited influence over their performance. For additional information, see the risk factor in this section under the heading "*We intend to continue to rely on third parties to conduct a significant portion of our existing clinical trials and potential future clinical trials for product candidates, and those third parties may not perform satisfactorily, including failing to meet deadlines for the completion of such trials.*"

Furthermore, even if we are able to enroll a sufficient number of patients for our clinical trials, we may have difficulty maintaining enrollment of such patients in our clinical trials.

***We may seek Orphan Drug designation or Rare Pediatric Disease designation for some of our product candidates and we may be unsuccessful, or may be unable to maintain the benefits associated with Orphan Drug designation, including the potential for market exclusivity, for product candidates for which we obtain Orphan Drug designation.***

Regulatory authorities in some jurisdictions, including the United States, may designate drugs or biologics intended to treat relatively small patient populations as orphan drug products. Under the Orphan Drug Act, the FDA may designate a drug or biologic as an orphan drug if it is intended to treat a rare disease or condition, which is generally defined as a patient population of fewer than 200,000 individuals in the United States, or a patient population of 200,000 or more in the United States where there is no reasonable expectation that the cost of developing the drug will be recovered from sales in the United States.

In the United States, Orphan Drug designation entitles a party to potential financial incentives such as tax advantages and user fee waivers. Opportunities for grant funding toward clinical trial costs may also be available for clinical trials of drugs or biologics for rare diseases, regardless of whether the drugs or biologics are designated for the orphan use. In addition, if a drug or biologic with an Orphan Drug designation subsequently receives the first marketing approval for the disease for which it has such designation, the product is entitled to a seven-year period of marketing exclusivity, which precludes the FDA from approving another marketing application for the same drug and disease for that time period, except in limited circumstances. If our competitors are able to obtain orphan drug exclusivity prior to us, for products that constitute the “same drug” and treat the same diseases as our product candidates, we may not be able to have competing products approved by the applicable regulatory authority for a significant period of time.

Similarly, in the European Union, the European Commission, upon the recommendation of the EMA’s Committee for Orphan Medicinal Products, grants Orphan Drug designation to promote the development of drugs that are intended for the diagnosis, prevention or treatment of life-threatening or chronically debilitating conditions and either the prevalence of the condition is not more than 5 in 10,000 persons in the European Union, or, without incentives, it is unlikely that sales of the drug in the European Union would be sufficient to justify the necessary investment in developing the drug. In each case, there must be no satisfactory method of diagnosis, prevention or treatment of the condition that has been authorized, or, if such a method exists, the product in question must be of significant benefit to those affected by such condition. In the European Union, Orphan Drug designation entitles a party to financial incentives such as reduction of fees or fee waivers.

We have obtained from the FDA Orphan Drug designation for LX2006 for treatment of FA. We may seek orphan designation for some or all of our product candidates in orphan indications in which there is a medically plausible basis for the use of these product candidates. However, we may be unsuccessful in obtaining Orphan Drug designation and may be unable to maintain the benefits associated with such designations. Even if we obtain orphan drug exclusivity for any of our product candidates, that exclusivity may not effectively protect those product candidates from competition because different drugs can be approved for the same condition, and orphan drug exclusivity does not prevent the FDA from approving the same or a different drug in another indication. Even after an orphan drug is granted orphan exclusivity and approved, the FDA can subsequently approve a later application for the same drug for the same condition before the expiration of the seven-year exclusivity period if the FDA concludes that the later drug is clinically superior in that it is shown to be safer in a substantial portion of the target populations, more effective or makes a major contribution to patient care. On August 3, 2017, Congress passed the FDA Reauthorization Act of 2017, or FDARA. FDARA, among other things, codified the FDA’s preexisting regulatory interpretation, to require that a drug sponsor demonstrate the clinical superiority of an orphan drug that is otherwise the same as a previously approved drug for the same rare disease in order to receive orphan drug exclusivity. The statute supplants prior precedent holding that the Orphan Drug Act unambiguously requires that the FDA recognize the orphan exclusivity period regardless of a showing of clinical superiority. Moreover, in the Consolidated Appropriations Act of 2021, Congress clarified that the interpretation of orphan drug exclusivity codified in FDARA would apply in cases where the FDA issued an orphan designation before the enactment of FDARA but where product approval came after the enactment of FDARA. In addition, a designated Orphan Drug may not receive orphan drug exclusivity if it is approved for a use that is

broader than the disease for which it received orphan designation. On January 24, 2023, the FDA announced its intention to apply its existing regulations and long-standing approach to grant orphan drug exclusivity based on the indications for which the drug is approved rather than granting the exclusivity for the entire rare disease or condition that was the subject of the orphan drug designation, in response to the U.S. Court of Appeals for the Eleventh Circuit's September 30, 2021, decision in *Catalyst Pharms., Inc. v. Becerra*. The FDA may further reevaluate its regulations and policies under the Orphan Drug Act. We do not know if, when, or how the FDA may change the orphan drug regulations and policies in the future, and it is uncertain how any changes might affect our business. Depending on what changes the FDA may make to its orphan drug regulations and policies, our business could be adversely impacted. Moreover, orphan drug-exclusive marketing rights in the United States may be lost if the FDA later determines that the request for designation was materially defective or if we are unable to manufacture sufficient quantities of the product to meet the needs of patients with the rare disease or condition. Orphan Drug designation neither shortens the development time or regulatory review time of a drug nor gives the drug any advantage in the regulatory review or approval process.

***Fast Track, Breakthrough Therapy, or Regenerative Medicine Advanced Therapy designation by the FDA may not actually lead to a faster development or regulatory review or approval process, and does not assure FDA approval of our product candidates.***

We may seek Fast Track, Breakthrough Therapy or Regenerative Medicine Advanced Therapy, or RMAT, designation from the FDA for some or all of our product candidates, but we may be unable to obtain such designations or to maintain the benefits associated with such designations. The FDA's Fast Track, Breakthrough Therapy, and RMAT designation programs are intended to expedite the development of certain qualifying product candidates intended for the treatment of serious diseases and conditions. If a product candidate is intended for the treatment of a serious or life-threatening condition and preclinical or clinical data demonstrate the product's potential to address an unmet medical need for this condition, the sponsor may apply for FDA Fast Track designation.

A product candidate may be designated as a breakthrough therapy if it is intended, alone or in combination with one or more other drugs or biologics to treat a serious or life-threatening condition and preliminary clinical evidence indicates that the product candidate may demonstrate substantial improvement over existing therapies on one or more clinically significant endpoints. For product candidates that have been designated as breakthrough therapies, interaction and communication between the FDA and the sponsor of the trial can help to identify the most efficient path for clinical development while minimizing the number of patients placed in ineffective control regimens. Drugs and biologics designated as breakthrough therapies by the FDA may also be eligible for accelerated approval.

A product candidate may receive RMAT designation if it is a regenerative medicine therapy that is intended to treat, modify, reverse or cure a serious or life-threatening disease or condition, and preliminary clinical evidence indicates that the product candidate has the potential to address an unmet medical need for such condition. RMAT designation allows companies developing regenerative medicine therapies to work more closely and frequently with the FDA, and RMAT-designated product candidates may be eligible for priority review and accelerated approval. FDA has confirmed that gene therapies, including genetically modified cells, that lead to a sustained effect on cells or tissues may meet the definition of a regenerative medicine therapy. For product candidates that have received an RMAT designation, interaction and communication between the FDA and the sponsor of the trial can help to identify the most efficient path for clinical development while minimizing the number of patients placed in ineffective control regimens.

We have received Fast Track designation for LX1001 for the treatment of patients with early Alzheimer's disease who are *APOE4* homozygous, to slow disease progression. While we may seek Fast Track, Breakthrough Therapy and/or RMAT designation for some or all of our product candidates, there is no guarantee that we will be successful in obtaining any such designation. Even if we do obtain such designation, we may not experience a faster development process, review or approval compared to conventional FDA procedures. A Fast Track,

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Breakthrough Therapy, or RMAT designation does not ensure that the product candidate will receive marketing approval or that approval will be granted within any particular time frame. In addition, the FDA may withdraw Fast Track, Breakthrough Therapy, or RMAT designation if it believes that the designation is no longer supported by data from our clinical development program. Fast Track, Breakthrough Therapy and/or RMAT designation alone does not guarantee qualification for the FDA's priority review procedures.

***We have received Rare Pediatric Disease designation from the FDA for LX2006 for the treatment of FA and we may seek such designation for future product candidates. However, a marketing application for these product candidates, if approved, may not meet the eligibility criteria for a rare pediatric disease priority review voucher.***

We have received Rare Pediatric Disease designation from the FDA for LX2006 for the treatment of FA and LX1004 for the treatment of neuronal ceroid lipofuscinosis type 2 (CLN2) disease and we may seek Rare Pediatric Disease designation for future product candidates. The FDA defines "rare pediatric disease" as a (i) serious or life-threatening disease in which the serious or life-threatening manifestations primarily affect individuals aged from birth to 18 years, including age groups often called neonates, infants, children, and adolescents; and (ii) a rare disease or condition within the meaning of the Orphan Drug Act. Designation of a product candidate as a product for a rare pediatric disease does not guarantee that a marketing application for such product candidate will meet the eligibility criteria for a rare pediatric disease priority review voucher at the time the application is approved. Under the U.S. Federal Food, Drug, and Cosmetic Act, or FDCA, we will need to request a rare pediatric disease priority review voucher in our original marketing application for our product candidates for which we have received Rare Pediatric Disease designation. The FDA may determine that a marketing application for any such product candidates, if approved, does not meet the eligibility criteria for a priority review voucher, including for the following reasons:

- the rare pediatric disease that received such designation no longer meets the definition of a "rare pediatric disease";
- the marketing application contains an active ingredient (including any ester or salt of the active ingredient) that has been previously approved in a marketing application;
- the marketing application is not deemed eligible for priority review;
- the marketing application does not rely on clinical data derived from studies examining a pediatric population and dosages of the product intended for that population (that is, if the marketing application does not contain sufficient clinical data to allow for adequate labeling for use by the full range of affected pediatric patients); or
- the marketing application is approved for a different adult indication than the rare pediatric disease for which our product candidates are designated.

Under the current statutory sunset provisions, after September 30, 2024, the FDA may only award a priority review voucher, or PRV, for an approved rare pediatric disease product application if the sponsor has Rare Pediatric Disease designation for the drug or biologic that is the subject of such application, and that designation was granted by September 30, 2024. After September 30, 2026, the FDA may not award any rare pediatric disease priority review vouchers. However, it is possible the authority for FDA to award rare pediatric disease priority review vouchers will be further extended by Congress. As such, if we do not obtain approval of a marketing application for LX2006 in patients with FA on or before September 30, 2026, and if the priority review voucher program is not extended by Congressional action, we may not receive a priority review voucher.

***Where appropriate, we may seek approval from the FDA, EMA or comparable foreign regulatory authorities through the use of accelerated approval pathways. If we are unable to obtain such approval, we may be required to conduct additional preclinical studies or clinical trials beyond those that we contemplate, which could increase the expense of obtaining, and delay the receipt of, necessary marketing approvals. Even if we receive accelerated approval from the FDA, EMA or comparable regulatory authorities, if our confirmatory trials do not verify clinical benefit, or if we do not comply with rigorous post-marketing requirements, the FDA, EMA or such other regulatory authorities may seek to withdraw accelerated approval.***

Where possible, we may pursue accelerated development strategies in areas of high medical need. We may seek an accelerated approval pathway for one or more of our therapeutic candidates from the FDA, EMA or comparable foreign regulatory authorities. Under the accelerated approval provisions in the FDCA and the FDA's implementing regulations, the FDA may grant accelerated approval to a therapeutic candidate that is designed to treat a serious or life-threatening condition, generally provides a meaningful therapeutic benefit over available therapies, and demonstrates an effect on a surrogate endpoint that is reasonably likely to predict clinical benefit or on a clinical endpoint that can be measured earlier than irreversible morbidity or mortality, or IMM, that is reasonably likely to predict an effect on IMM or other clinical benefit. The FDA considers a clinical benefit to be a positive therapeutic effect that is clinically meaningful in the context of a given disease, such as irreversible morbidity or mortality. For the purposes of accelerated approval, a surrogate endpoint is a marker, such as a laboratory measurement, radiographic image, physical sign, or other measure that is thought to predict clinical benefit, but is not itself a measure of clinical benefit. An intermediate clinical endpoint is a clinical endpoint that can be measured earlier than an effect on irreversible morbidity or mortality that is reasonably likely to predict an effect on irreversible morbidity or mortality or other clinical benefit. The accelerated approval pathway may be used in cases in which the advantage of a new product over available therapy may not be a direct therapeutic advantage, but is a clinically important improvement from a patient and public health perspective. If granted, accelerated approval is usually contingent on the sponsor's agreement to conduct, in a diligent manner, additional post-approval confirmatory studies to verify and describe the product's clinical benefit. If such post-approval studies fail to confirm the product's clinical benefit, the FDA may withdraw its approval of the product. In addition, the FDA currently requires, unless otherwise informed by the agency, pre-approval of promotional materials for products receiving accelerated approval, which could adversely impact the timing of the commercial launch of the product.

Prior to seeking accelerated approval, we would seek feedback from the FDA, EMA or comparable foreign regulatory authorities and would otherwise evaluate our ability to seek and receive such accelerated approval. There can be no assurance that after our evaluation of the feedback and other factors we will decide to pursue or submit a BLA for accelerated approval or any other form of expedited development, review or approval. Similarly, there can be no assurance that after subsequent feedback from the FDA, EMA or comparable foreign regulatory authorities, we will continue to pursue or apply for accelerated approval or any other form of expedited development, review or approval, even if we initially decide to do so. Furthermore, if we decide to submit an application for accelerated approval, there can be no assurance that such application will be accepted or that any approval will be granted on a timely basis, or at all. The FDA, EMA or other comparable foreign regulatory authorities could also require us to conduct further studies prior to considering our application or granting approval of any type, including, for example, if other products are approved via the accelerated pathway and subsequently converted by FDA to full approval. A failure to obtain accelerated approval or any other form of expedited development, review or approval for our therapeutic candidate would result in a longer time period to commercialization of such therapeutic candidate, could increase the cost of development of such therapeutic candidate and could harm our competitive position in the marketplace.

***Priority review designation by the FDA may not lead to a faster regulatory review or approval process and, in any event, does not assure FDA approval of our product candidates.***

If the FDA determines that a product candidate is intended to treat a serious disease or condition and, if approved, would provide a significant improvement in the safety or effectiveness of the treatment, prevention, or diagnosis of such disease or condition, the FDA may designate the product candidate for priority review. A priority review designation means that the goal for the FDA to review a marketing application is six months from filing of the application, rather than the standard review period of ten months. We may request priority review for certain of our product candidates. The FDA has broad discretion with respect to whether or not to grant priority review status to a product candidate, so even if we believe a particular product candidate is eligible for such designation or status, the FDA may disagree and decide not to grant it. Moreover, a priority review designation does not necessarily mean a faster regulatory review process or necessarily confer any advantage with respect to approval compared to conventional FDA procedures. Receiving priority review from the FDA does not guarantee approval within the six-month review cycle or thereafter.

***We may expend our limited resources to pursue a particular product candidate and fail to capitalize on product candidates that may be more profitable or for which there is a greater likelihood of success.***

Because we have limited financial and management resources, we must focus on development programs and product candidates that we identify for specific diseases. As such, currently we are primarily focused on the development of our current pipeline of candidates. As a result, we may forego or delay pursuit of opportunities with other product candidates. For example, we discontinued internal development of our LX1004 program and we are evaluating strategic alternatives to find the appropriate partner to take the program forward. Our resource allocation decisions may cause us to fail to capitalize on viable commercial products or profitable market opportunities. Our spending on current and future development programs and product candidates for specific diseases may not yield any commercially viable products. If we do not accurately evaluate the commercial potential or target market for a particular product candidate, we may relinquish valuable rights to that product candidate through collaboration, licensing or other royalty arrangements in cases in which it would have been more advantageous for us to retain sole development and commercialization rights to such product candidate.

***We may not be successful in our efforts to build a pipeline of additional product candidates.***

Our business model is centered on developing therapies for patients with both larger-rare and prevalent cardiac and central nervous system diseases by establishing focused selection criteria to select, develop and advance product candidates that we believe will have a high probability of technical and regulatory success through development into commercialization. We may not be able to continue to identify and develop new product candidates. Even if we are successful in continuing to build our pipeline, the potential product candidates that we identify may not be suitable for clinical development. For example, they may be shown to have side effects or other characteristics that indicate that they are unlikely to be drugs that will receive marketing approval and achieve market acceptance. If we do not successfully develop and commercialize product candidates based upon our approach, we will not be able to obtain product revenue in future periods, which likely would result in significant harm to our financial position and adversely affect our stock price.

***If we are unable to successfully validate, develop and obtain regulatory approval for companion diagnostic tests for our product candidates that require or would commercially benefit from such tests, or experience significant delays in doing so, we may not realize the full commercial potential of these product candidates.***

In connection with the clinical development of our product candidates for certain indications, we intend to work with collaborators to develop or obtain access to *in vitro* companion diagnostic tests to identify patient subsets

within a disease category who may derive selective and meaningful benefit from our product candidates. To be successful, we or our collaborators will need to address a number of scientific, technical, regulatory and logistical challenges. The FDA and comparable foreign regulatory authorities regulate *in vitro* companion diagnostics as medical devices and, under that regulatory framework, will likely require the conduct of clinical trials to demonstrate the safety and effectiveness of any diagnostics we may develop, which we expect will require separate regulatory clearance or approval prior to commercialization. The FDA generally will require approval or clearance of the diagnostic at the same time that the FDA approves the therapeutic product if the FDA determines that safe and effective use of a therapeutic product depends on an *in vitro* companion diagnostic. The clearance or approval of a companion diagnostic as part of the product label will also limit the use of the product candidate to patients who have met the screening criteria tested for by the companion diagnostic.

We intend to rely on third parties for the design, development and manufacture of companion diagnostic tests for our therapeutic product candidates that may require such tests. If we enter into such collaborative agreements, we will be dependent on the sustained cooperation and effort of our future collaborators in developing and obtaining approval for these companion diagnostics. It may be necessary to resolve issues such as selectivity/specificity, analytical validation, reproducibility, or clinical validation of companion diagnostics during the development and regulatory approval processes. Moreover, even if data from preclinical studies and early clinical trials appear to support development of a companion diagnostic for a product candidate, data generated in later clinical trials may fail to support the analytical and clinical validation of the companion diagnostic. We and our future collaborators may encounter difficulties in developing, obtaining regulatory approval for, manufacturing and commercializing companion diagnostics similar to those we face with respect to our therapeutic candidates themselves, including issues with achieving regulatory clearance or approval, production of sufficient quantities at commercial scale and with appropriate quality standards, and in gaining market acceptance. If we are unable to successfully develop companion diagnostics for these product candidates, or experience delays in doing so, the development of these product candidates may be adversely affected, these product candidates may not obtain marketing approval, and we may not realize the full commercial potential of any of these product candidates that obtain marketing approval. As a result, our business, results of operations and financial condition could be materially harmed. In addition, a diagnostic company with whom we contract may decide to discontinue selling or manufacturing the companion diagnostic test that we anticipate using in connection with development and commercialization of our product candidates or our relationship with such diagnostic company may otherwise terminate. We may not be able to enter into arrangements with another diagnostic company to obtain supplies of an alternative diagnostic test for use in connection with the development and commercialization of our product candidates or do so on commercially reasonable terms, which could adversely affect and/or delay the development or commercialization of our product candidates.

### **Risks related to the manufacturing of our product candidates**

***We and our contract manufacturers are subject to significant regulation with respect to manufacturing our products. The third-party manufacturing facilities on which we rely, and any manufacturing facility that we may have in the future, may have limited capacity or fail to meet the applicable stringent regulatory requirements.***

We currently have relationships with a limited number of suppliers for the manufacturing of all components of our product candidates. However, if we experience slowdowns or problems with our manufacturing partners and are unable to establish or scale our internal manufacturing capabilities, we will need to continue to contract with manufacturers that can produce the preclinical, clinical and commercial supply of our products. Each supplier may require licenses to manufacture such components if such processes are not owned by the supplier or in the public domain and we may be unable to license such intellectual property rights on reasonable commercial terms or to transfer or sublicense the intellectual property rights we may have with respect to such activities.

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All entities involved in the preparation of therapeutics for clinical trials or commercial sale, including our existing contract manufacturers for components of our product candidates, are subject to extensive regulation. Components of a finished therapeutic product approved for commercial sale or used in late-stage clinical trials in the United States and European Union must be manufactured in accordance with cGMP. These regulations govern manufacturing processes and procedures (including recordkeeping) and the implementation and operation of quality systems to control and assure the quality of investigational products and products approved for sale. Poor control of production processes can lead to the introduction of adventitious agents or other contaminants, or to inadvertent changes in the properties or stability of our product candidates that may not be detectable in final product testing. We or our contract manufacturers must supply all necessary documentation in support of a BLA or an MAA on a timely basis. Our potential manufacturing facilities and quality systems and the facilities and quality systems of some or all of our third-party contractors must pass a pre-approval inspection for compliance with the applicable regulations as a condition of regulatory approval of our product candidates or any of our other potential products. In addition, the regulatory authorities may, at any time, audit or inspect a manufacturing facility involved with the preparation of our product candidates or our other potential products or the associated quality systems for compliance with the regulations applicable to the activities being conducted, and they could put a hold on one or more of our clinical trials if the facilities of our contract development and manufacturing organizations, or CDMOs, do not pass such audit or inspections. If these facilities do not pass a pre-approval plant inspection, FDA approval of the products will not be granted.

The regulatory authorities also may, at any time following approval of a product for sale, inspect or audit our manufacturing facilities or those of our third-party contractors. If any such inspection or audit identifies a failure to comply with applicable regulations or if a violation of our product specifications or applicable regulations occurs independent of such an inspection or audit, we or the relevant regulatory authority may require remedial measures that may be costly and/or time-consuming for us or a third party to implement and that may include the temporary or permanent suspension of a clinical trial or commercial sales or the temporary or permanent closure of a facility. Any such remedial measures imposed upon us or third parties with whom we contract could harm our business.

If we or any of our third-party manufacturers fail to maintain regulatory compliance, the FDA can impose regulatory sanctions including, among other things, refusal to approve a pending application for a new drug product or biologic product, or revocation of a pre-existing approval. As a result, our business, financial condition and results of operations may be harmed. Additionally, if supply from one approved manufacturer is interrupted, there could be a significant disruption in commercial supply. An alternative manufacturer would need to be qualified through a BLA and/or an MAA supplement which could result in further delay. The regulatory agencies may also require additional studies if a new manufacturer is relied upon for commercial production. Switching manufacturers may involve substantial costs and is likely to result in a delay in our desired clinical and commercial timelines.

These factors could cause the delay of clinical trials, regulatory submissions, required approvals or commercialization of our product candidates, cause us to incur higher costs and prevent us from commercializing our products successfully, if approved. Furthermore, if our suppliers fail to meet contractual requirements, and we are unable to secure one or more replacement suppliers capable of production at a substantially equivalent cost, our clinical trials may be delayed, or we could lose potential revenue.



***Gene therapies are novel, complex and difficult to manufacture. We could experience manufacturing problems that result in delays in the development or commercialization of our product candidates or otherwise harm our business.***

The manufacture of gene therapy products is technically complex and necessitates substantial expertise and capital investment. Production difficulties caused by unforeseen events may delay the availability of material for our clinical studies.

We rely on third-party manufacturers to manufacture our product candidates for preclinical studies and clinical trials. The manufacturers of pharmaceutical products must comply with strictly enforced cGMP requirements, state and federal regulations, as well as foreign requirements when applicable. Any failure of us or our contract manufacturing organizations, or CMOs, to adhere to or document compliance to such regulatory requirements could lead to a delay or interruption in the availability of our product candidate materials for clinical trials or enforcement action from the FDA, EMA or foreign regulatory authorities. If we or our manufacturers fail to comply with the requirements of the FDA, EMA or other regulatory authority, sanctions could be imposed on us, including clinical holds, fines, injunctions, civil penalties, delays, suspension or withdrawal of approvals, license revocation, seizures or recalls of product candidates or products, operating restrictions and criminal prosecutions, any of which could significantly and adversely affect supplies of our product candidates.

There can be no assurances that our third-party manufacturers will be able to meet our timetable and requirements. If any third party with whom we contract fails to perform its obligations, we may be forced to either manufacture the materials ourselves, for which we may not have the capabilities or resources, or enter into an agreement with a different third-party manufacturer, which we may not be able to do on reasonable terms, if at all. In either scenario, our clinical trials and future commercial supply could be delayed significantly as we establish alternative supply sources. In some cases, the technical skills required to manufacture our product candidates may be unique or proprietary to the original third-party manufacturer and we may have difficulty, or there may be contractual restrictions prohibiting us from, transferring such skills to a back-up or alternate supplier, or we may be unable to transfer such skills at all. In addition, if we are required to change manufacturers for any reason, we will be required to verify that the new manufacturer maintains facilities and procedures that comply with quality standards and with all applicable regulations. We will also need to verify, such as through a manufacturing comparability study, that any new manufacturing process will produce our product candidates according to the specifications previously submitted to the FDA or another regulatory authority. The delays associated with the verification of a new manufacturer could negatively affect our ability to develop product candidates or commercialize our products, if approved, in a timely manner or within budget.

If we are unable to arrange for alternative third-party manufacturing sources on commercially reasonable terms or in a timely manner, we may be delayed in the development of our product candidates. Our dependence upon others for the manufacture of our product candidates may also adversely affect our future profit margins and our ability to commercialize any product candidates that receive regulatory approval on a timely and competitive basis.

Our product candidates require processing steps that are more complex than those required for most chemical pharmaceuticals. Moreover, unlike chemical pharmaceuticals, the physical and chemical properties of a biologic such as our modified virus generally cannot be fully characterized. As a result, assays of the finished product may not be sufficient to ensure that the product will perform in the intended manner. Although we believe that the manufacture of our product candidates may be simplified due to their shared raw materials and other similarities, we cannot be certain that this will be the case and we may be required to develop manufacturing methods that ultimately differ significantly between product candidates, which would require that we invest substantial time and capital to develop suitable manufacturing methods. Our program materials are manufactured using technically complex processes requiring specialized equipment and facilities, highly specific raw materials,

cell types and reagents, and other production constraints. Our production process also requires a number of highly specific raw materials, cell types and reagents with limited suppliers. Even though we aim to have backup supplies of raw materials, cell types and reagents whenever possible, we cannot be certain they will be sufficient if our primary sources are unavailable. A shortage of a critical raw material, cell line, or reagent, or a technical issue during manufacturing may lead to delays in clinical development or commercialization plans. We are particularly susceptible to any shortages, delays or our inability to obtain suitable raw materials for our lead product candidates. Any changes in the manufacturing of components of the raw materials we use could result in unanticipated or unfavorable effects in our manufacturing processes, resulting in delays.

In addition, if any of our product candidates obtain approval, the FDA, EMA and other regulatory authorities may require us to submit samples of any lot of any approved product together with the protocols showing the results of applicable tests at any time. Under some circumstances, the FDA, EMA or other regulatory authorities may require that we not distribute a lot until the agency authorizes its release. Slight deviations in the manufacturing process, including those affecting quality attributes and stability, may result in unacceptable changes in the product that could result in lot failures or product recalls. Lot failures or product recalls could cause us to delay product launches or clinical trials, which could be costly to us and otherwise harm our business, financial condition, results of operations and prospects.

***We depend on third-party suppliers for materials used in the manufacture of our product candidates, and the loss of these third-party suppliers or their inability to supply us with adequate materials could harm our business.***

We rely on third-party suppliers for the materials and components required for the production of our product candidates. Our dependence on these third-party suppliers and the challenges we may face in obtaining adequate supplies of materials involve several risks, including limited control over pricing, availability, and delivery schedules. There is substantial demand and limited supply for certain of the raw materials used to manufacture gene therapy products. As a small company, our negotiation leverage is limited, and we may get lower priority than our competitors that are larger than we are. We cannot be certain that our suppliers will continue to provide us with the quantities of these raw materials within the timelines that we require or satisfy our anticipated specifications and quality requirements. Any supply interruption in limited or sole-sourced raw materials could materially harm our ability to manufacture our product candidates until a new source of supply, if any, could be identified and qualified. We may be unable to find a sufficient alternative supply channel in a reasonable time or on commercially reasonable terms. Any performance failure on the part of our suppliers could delay the development and potential commercialization of our product candidates, including limiting supplies necessary for clinical trials and regulatory approvals, which would have a material adverse effect on our business.

***Any contamination or interruption in our manufacturing process, shortages of raw materials or failure of our suppliers of viruses to deliver necessary components could result in delays in our clinical development or marketing schedules.***

Given the nature of gene therapy manufacturing, there is a risk of contamination occurring during the manufacturing process. Any contamination could adversely affect our ability to produce product candidates on schedule and could, therefore, harm our results of operations and cause reputational damage. Some of the raw materials required in our manufacturing process are derived from biologic sources. Such raw materials are difficult to procure and may be subject to contamination or recall. A material shortage, contamination, recall or restriction on the use of biologically derived substances in the manufacture of our product candidates could adversely impact or disrupt the commercial manufacturing or the production of clinical material, which could adversely affect our development timelines and our business, financial condition, results of operations and prospects.

***Changes in methods of product candidate manufacturing or formulation may result in additional costs or delay.***

As product candidates proceed through preclinical studies to late-stage clinical trials towards potential approval and commercialization, it is common that various aspects of the development program, such as manufacturing methods and formulation, are altered along the way in an effort to optimize processes and product characteristics. Such changes carry the risk that they will not achieve our intended objectives. Any such changes could cause our product candidates to perform differently and affect the results of planned clinical trials or other future clinical trials conducted with the materials manufactured using altered processes. Such changes may also require additional testing, FDA notification or FDA approval. This could delay the initiation and completion of clinical trials, require the conduct of bridging clinical trials or the repetition of one or more clinical trials, increase clinical trial costs, delay approval of our product candidates and jeopardize our ability to commence sales and generate revenue. In addition, we may be required to make significant changes to our upstream and downstream processes across our pipeline, which could delay the development of our future product candidates. Regulatory agencies, and in particular the FDA and EMA, have demonstrated increased caution in their regulation of gene therapies, including increased scrutiny related to chemistry, manufacturing and control, or CMC, issues. This increased regulatory scrutiny around gene therapy CMC may result in us being required to conduct additional preclinical studies or clinical trials with respect to any of our product candidates, which may result in delays and increased costs in the development or commercialization of our product candidates and ultimately could lead to the failure to obtain approval for any gene therapy product.

**Risks related to the commercialization of our product candidates**

***Even if any of our product candidates receive marketing approval, they may fail to achieve the degree of market acceptance by physicians, patients, third-party payors and others in the medical community necessary for commercial success.***

If any of our product candidates receive marketing approval, they may nonetheless fail to gain sufficient market acceptance by physicians, patients, third-party payors and others in the medical community. If our product candidates do not achieve an adequate level of acceptance, we may not generate significant revenue and we may not become profitable. The degree of market acceptance of our product candidates, if approved for commercial sale, will depend on a number of factors, including:

- their efficacy, safety and potential advantages compared to alternative treatments;
- our ability to offer our products for sale at competitive prices;
- their convenience and ease of administration compared to alternative treatments;
- product labeling or product insert requirements of the FDA or foreign regulatory authorities, including any limitations or warnings contained in a product's approved labeling, including any boxed warning or REMS;
- the willingness of the target patient population to try new treatments, such as gene therapy as a novel modality for treatment of our target indications and of physicians to prescribe these treatments;
- our ability to hire and retain a sales force in the United States;
- the strength of marketing and distribution support;
- the availability of coverage and adequate reimbursement for our product candidates, once approved, from third-party payors and government authorities;

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- the prevalence and severity of any side effects; and
- any restrictions on the use of our products together with other medications.

### ***Negative public opinion of gene therapy and increased regulatory scrutiny of gene therapy and genetic research may adversely impact the development or commercial success of our current and future product candidates.***

Our product candidates involve introducing genetic material into a patient's cells via intrathecal and intravenous administration. The clinical and commercial success of our potential products will depend in part on public acceptance of the use of gene therapy and gene regulation for the prevention or treatment of human diseases. Public attitudes may be influenced by claims that gene therapy and gene regulation are unsafe, unethical or immoral, and consequently, our products may not gain the acceptance of the public or the medical community. Adverse public attitudes may adversely impact our ability to enroll clinical trials. Moreover, our success will depend upon physicians prescribing, and their patients being willing to receive, treatments that involve the use of product candidates we may develop in lieu of, or in addition to, existing treatments with which they are already familiar and for which greater clinical data may be available.

More restrictive government regulations or negative public opinion would have a negative effect on our business or financial condition and may delay or impair the development and commercialization of our product candidates or demand for any products once approved. In recent years, sponsors of other clinical trials involving gene therapies have announced imposition of clinical holds by the FDA to evaluate safety issues arising during the trials. Among the risks in any gene therapy product based on viral vectors are the risks of immunogenicity, elevated liver enzymes and insertional oncogenesis. If any of our vectors demonstrate a similar effect, we may decide or be required to halt or delay further clinical development of any product candidates that utilize that vector. Adverse events in our or others' clinical trials, even if not ultimately attributable to our product candidates, and the resulting publicity could result in increased governmental regulation, unfavorable public perception, potential regulatory delays in the testing or approval of our product candidates, stricter labeling requirements for those product candidates that are approved and a decrease in demand for any such product candidates. The risk of cancer remains a concern for gene therapy, and we cannot assure that it will not occur in any of our planned or future clinical trials or in any clinical trials conducted by other companies. In addition, there is the potential risk of delayed adverse events following exposure to gene therapy products due to persistent biological activity of the genetic material or other components of products used to carry the genetic material. In addition, for our regulated gene replacement therapy candidates which require that the expression of a therapeutic transgene be tightly regulated, such as LX1020, we may inadvertently cause overexpression, which could lead to numerous issues, including safety and toxicity concerns. Furthermore, these regulatory gene replacement therapy candidates require the insertion of microRNA, or miRNA, targets into the viral genome, which is a technology that to our knowledge is not present in any approved gene therapy products. If any such adverse events occur, commercialization of our product candidates or further advancement of our clinical trials could be halted or delayed, which would have a negative impact on our business and operations.

### ***The affected populations for our other product candidates may be smaller than we or third parties currently project, which may affect the addressable markets for our product candidates.***

We currently focus our research and product development on several indications that are orphan diseases. However, our projections of the number of people who have the diseases we are seeking to treat, as well as the subset of people with these diseases who have the potential to benefit from treatment with our product candidates, are estimates based on our knowledge and understanding of these diseases. These estimates may prove to be incorrect and new studies may further reduce the estimated incidence or prevalence of this disease.

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The number of patients in the United States, the European Union and elsewhere may turn out to be lower than expected, may not be otherwise amenable to treatment with our product candidate or patients may become increasingly difficult to identify and access, all of which would adversely affect our business, financial condition, results of operations and prospects.

The total addressable market opportunity for our product candidates will ultimately depend upon a number of factors, including the diagnosis and treatment criteria included in the final label, if approved for sale in specified indications, acceptance by the medical community, patient access and product pricing and reimbursement. Incidence and prevalence estimates are frequently based on information and assumptions that are not exact and may not be accurate, and the methodology is forward-looking and potentially speculative. The process we have used in developing an estimated incidence and prevalence range for the indications we are targeting has involved collating limited data from multiple sources. Accordingly, the incidence and prevalence estimates included in this prospectus should be viewed in that context. Further, the data and statistical information used in this prospectus, including estimates derived from them, may differ from information and estimates made by our competitors or from current or future studies conducted by independent sources.

***We face significant competition from other biotechnology and pharmaceutical companies, and our operating results will suffer if we fail to compete effectively.***

Drug development, particularly in the gene therapy field, is highly competitive and subject to rapid and significant technological advancements. As a significant unmet medical need exists in the cardiovascular disease and Alzheimer's disease areas, there are several large and small pharmaceutical companies focused on delivering therapeutics for the treatment of these diseases, including those that we are initially targeting. It is likely that additional drugs will become available in the future for the treatment of our target diseases.

We are aware that our competitors are developing product candidates for the treatment of diseases that our product candidates will target. With respect to LX2006, we are aware of preclinical gene therapy programs in development at Novartis, PTC Therapeutics, Inc., Aavanti Bio, Inc., Lacerta Therapeutics, Inc. and those being developed in collaborations between Voyager Therapeutics, Inc. and Neurocrine Biosciences, Inc. and between Takeda Pharmaceutical Company Limited and StrideBio, Inc. Among other treatment modalities for FA, we are aware that Larimar Therapeutics, Inc. is developing a clinical-stage product candidate, CTI-1601, and that Reata Pharmaceuticals, Inc.'s Omaveloxolone (Skyclarys) was recently approved by the FDA.

With respect to LX2020, both Rocket Pharmaceuticals Inc. and Tenaya Therapeutics Inc. are developing a preclinical AAV-based gene therapy candidate designed to deliver a functional *PKP2* gene to patients with ARVC.

With respect to our portfolio of gene therapy programs for the treatment of homozygous *APOE4*-associated Alzheimer's disease, we are aware that uniQure, N.V. is pursuing AMT-240, a preclinical gene therapy candidate for autosomal dominant Alzheimer's disease intended to silence the toxic variant while expressing the protective variant, and Novartis has a gene therapy candidate for Alzheimer's disease that is in the early preclinical stages of development. Many large and small pharmaceutical companies and academic institutions are developing potential treatments for the condition given the significant unmet need and the large population suffering from Alzheimer's disease. There are multiple FDA-approved treatments for Alzheimer's disease, including donepezil (Aricept), memantine (Namenda), and aducanumab (Aduhelm), the latter of which was recently approved under accelerated approval and will require confirmatory data to verify clinical benefit. In January of 2023, lecanemab was granted accelerated approval by the FDA for the treatment of Alzheimer's disease based on the observed reduction of amyloid beta plaque. In addition, Eli Lilly and Company's product candidate for the treatment of Alzheimer's disease, donanemab, is currently in Phase 3 development. Finally, we are aware that Voyager Therapeutics, Inc. and Taysha Gene Therapies, Inc. are pursuing Alzheimer's disease treatments and have early-stage discovery efforts ongoing based on vectorized antibodies and tau-specific miRNA shuttles, respectively.

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Many of our existing or potential competitors have substantially greater financial, technical and human resources than we do and significantly greater experience in the discovery and development of product candidates, as well as in obtaining regulatory approvals of those product candidates in the United States and in foreign countries. Our current and potential future competitors may also have significantly more experience commercializing drugs, particularly gene therapy and other biologics, that have been approved for marketing. Mergers and acquisitions in the pharmaceutical and biotechnology industries could result in even more resources being concentrated among a small number of our competitors.

We will face competition from other drugs or from other non-drug products currently approved or that will be approved in the future in the cardiac and neurology fields, including for the treatment of diseases and diseases in the therapeutic categories we intend to target. Therefore, our ability to compete successfully will depend largely on our ability to:

- develop and commercialize drugs that are advantageous as compared to other products in the market;
- demonstrate through our clinical trials that our product candidates are differentiated from existing and future therapies;
- attract qualified scientific, product development and commercial personnel;
- obtain patent or other proprietary protection for our medicines;
- obtain required regulatory approvals;
- obtain coverage and adequate reimbursement from, and negotiate competitive pricing with, third-party payors; and
- successfully collaborate with other pharmaceutical companies in the discovery, development and commercialization of new medicines.

The availability of our competitors' products could limit the demand, and the price we are able to charge, for any product candidate we develop. The inability to compete with existing or subsequently introduced drugs would have an adverse impact on our business, financial condition and prospects. In addition, the reimbursement structure of approved gene therapies by other companies could impact the anticipated reimbursement structure of our gene therapies, if approved, and our business, financial condition, results of operations and prospects.

Established pharmaceutical companies may invest heavily to accelerate discovery and development of novel compounds or to in-license novel compounds that could make our product candidates less competitive. In addition, any new product that competes with an approved product must demonstrate compelling advantages in efficacy, convenience, tolerability and safety in order to overcome price competition and to be commercially successful. Accordingly, our competitors may succeed in obtaining patent protection, discovering, developing, receiving regulatory and marketing approval for, or commercializing, drugs before we do, which would have an adverse impact on our business and results of operations.

### ***Any product candidates for which we intend to seek approval as biologic products may face competition sooner than anticipated.***

The Patient Protection and Affordable Care Act, as amended by the Health Care and Education Reconciliation Act, or collectively, the ACA, includes a subtitle called the Biologics Price Competition and Innovation Act of 2009, or BPCIA, which created an abbreviated approval pathway for biologics that are biosimilar to or interchangeable with an FDA-licensed reference biological product. Under the BPCIA, an application for a biosimilar product may not be submitted to the FDA until four years following the date that the reference product

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was first licensed by the FDA. In addition, the approval of a biosimilar product may not be made effective by the FDA until 12 years from the date on which the reference product was first licensed. During this 12-year period of exclusivity, another company may still market a competing version of the reference product if the FDA approves a full BLA for the competing product containing the sponsor's own preclinical data and data from adequate and well-controlled clinical trials to demonstrate the safety, purity and potency of its product. The law is complex and is still being interpreted and implemented by the FDA. As a result, its ultimate impact, implementation and meaning are subject to uncertainty. While it is uncertain when such processes intended to implement BPCIA may be fully adopted by the FDA, any such processes could have an adverse effect on the future commercial prospects for our biologics.

There is a risk that any of our product candidates approved as a biological product under a BLA would not qualify for the 12-year period of exclusivity or that this exclusivity could be shortened due to congressional action or otherwise, or that the FDA will not consider our product candidates to be reference products for competing products, potentially creating the opportunity for generic competition sooner than anticipated. Other aspects of the BPCIA, some of which may impact the BPCIA exclusivity provisions, have also been the subject of recent litigation. Moreover, the extent to which a biosimilar, once approved, will be substituted for any one of our reference products in a way that is similar to traditional generic substitution for non-biologics is not yet clear, and will depend on a number of marketplace and regulatory factors that are still developing. If competitors are able to obtain marketing approval for biosimilars referencing our candidates, if approved, our products may become subject to competition from such biosimilars, with the attendant competitive pressure and potential adverse consequences.

***The success of our product candidates will depend significantly on coverage and adequate reimbursement or the willingness of patients, commercial and government payors to pay for these procedures.***

We believe our success depends on obtaining and maintaining coverage and adequate reimbursement for our product candidates, including LX2006 and LX1001, and the extent to which patients will be willing to pay out-of-pocket for such products, in the absence of reimbursement for all or part of the cost. In the United States and in other countries, patients who are provided medical treatment for their conditions generally rely on third-party payors to reimburse all or part of the costs associated with their treatment. The availability of coverage and adequacy of reimbursement for our products by third-party payors, including government health care programs (e.g., Medicare, Medicaid, TRICARE), managed care providers, private health insurers, health maintenance organizations and other organizations is essential for most patients to be able to afford medical services and pharmaceutical products such as our product candidates. Third-party payors often rely upon Medicare coverage policy and payment limitations in setting their own coverage and reimbursement policies. However, decisions regarding the extent of coverage and amount of reimbursement to be provided are made on a payor-by-payor basis. One payor's determination to provide coverage for a drug product does not assure that other payors will also provide coverage, and adequate reimbursement. For more information, see the section titled "*Business – Government Regulation – Coverage and Reimbursement.*"

The principal decisions about reimbursement for new medicines are typically made by the Centers for Medicare & Medicaid Services, or CMS, an agency within the U.S. Department of Health and Human Services, or HHS. CMS decides whether and to what extent products will be covered and reimbursed under Medicare and private payors tend to follow CMS to a substantial degree.

Third-party payors determine which products and procedures they will cover and establish reimbursement levels. Even if a third-party payor covers a particular product or procedure, the resulting reimbursement payment rates may not be adequate. Patients who are treated in-office for a medical condition generally rely on third-party payors to reimburse all or part of the costs associated with the procedure, including costs associated with

products used during the procedure, and may be unwilling to undergo such procedures in the absence of such coverage and adequate reimbursement. Physicians may be unlikely to offer procedures for such treatment if they are not covered by insurance and may be unlikely to purchase and use our product candidates, if approved, for our stated diseases unless coverage is provided and reimbursement is adequate. In addition, for products administered under the supervision of a physician, obtaining coverage and adequate reimbursement may be particularly difficult because of the higher prices often associated with such drugs. Further, coverage policies and third-party reimbursement rates may change at any time. Therefore, even if favorable coverage and reimbursement status is attained, less favorable coverage policies and reimbursement rates may be implemented in the future. In addition, companion diagnostic tests require coverage and reimbursement separate and apart from the coverage and reimbursement for their companion pharmaceutical or biological products. Similar challenges to obtaining coverage and reimbursement, applicable to pharmaceutical or biological products, will apply to companion diagnostics.

Reimbursement by a third-party payor may depend upon a number of factors, including the third-party payor's determination that a procedure is a covered benefit under its health plan; safe, effective and medically necessary; appropriate for the specific patient; cost-effective; supported by peer-reviewed medical journals; included in clinical practice guidelines; and neither cosmetic, experimental, nor investigational. Further, increasing efforts by third-party payors in the United States and abroad to cap or reduce healthcare costs may cause such organizations to limit both coverage and the level of reimbursement for newly approved products and, as a result, they may not cover or provide adequate payment for our product candidates. In order to secure coverage and reimbursement for any product that might be approved for sale, we may need to conduct expensive pharmacoeconomic studies in order to demonstrate the medical necessity and cost-effectiveness of our products, in addition to the costs required to obtain FDA or comparable regulatory approvals. Additionally, we may also need to provide discounts to purchasers, private health plans or government healthcare programs. Our product candidates may nonetheless not be considered medically necessary or cost-effective. If third-party payors do not consider a product to be cost-effective compared to other available therapies, they may not cover the product after approval as a benefit under their plans or, if they do, the level of payment may not be sufficient to allow a company to sell its products at a profit. We expect to experience pricing pressures from third-party payors in connection with the potential sale of any of our product candidates.

Foreign governments also have their own healthcare reimbursement systems, which vary significantly by country and region, and we cannot be sure that coverage and adequate reimbursement will be made available with respect to the treatments in which our products are used under any foreign reimbursement system.

There can be no assurance that LX2006, LX1001, LX2020 or any other product candidates, if approved for sale in the United States or in other countries, will be considered medically reasonable and necessary, that it will be considered cost-effective by third-party payors, that coverage or an adequate level of reimbursement will be available or that reimbursement policies and practices in the United States and in foreign countries where our products are sold will not adversely affect our ability to sell our product candidates profitably, if they are approved for sale.

***Product liability lawsuits against us could cause us to incur substantial liabilities and to limit commercialization of any products that we may develop.***

We face an inherent risk of product liability exposure related to the testing of our product candidates in human clinical trials and will face an even greater risk if we commercially sell any products that we may develop. If we cannot successfully defend ourselves against claims that our product candidates or drugs caused injuries, we will incur substantial liabilities. Regardless of merit or eventual outcome, liability claims may result in:

- decreased demand for any product candidates or drugs that we may develop;



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- injury to our reputation and significant negative media attention;
- withdrawal of clinical trial participants;
- significant costs to defend the related litigation;
- substantial monetary awards paid to trial participants or patients;
- loss of revenue;
- reduced resources of our management to pursue our business strategy; and
- the inability to commercialize any products that we may develop.

Although we maintain product liability insurance coverage, such insurance may not be adequate to cover all liabilities that we may incur. We may need to increase our insurance coverage as we expand our clinical trials or if we commence commercialization of our product candidates. Insurance coverage is increasingly expensive. We may not be able to maintain insurance coverage at a reasonable cost or in an amount adequate to satisfy any liability that may arise.

### **Risks related to our dependence on third parties**

***Currently, we rely on our collaborations with Cornell University and University of California, San Diego, or UCSD, to conduct research and development for many of our pipeline programs, including conducting preclinical and IND-enabling studies for portions of our near-term future pipeline. Failure or delay of Cornell University or UCSD to fulfill all or part of their respective obligations to us under our agreements, a breakdown in collaboration between the parties or a complete or partial loss of either of these relationships would materially harm our business.***

Our company was founded with a collaboration between us, Cornell University, and Ronald G. Crystal, M.D., Professor and Chairman of Weill Cornell Medicine's Department of Genetic Medicine. Our collaboration with Cornell University is critical to our business and in May 2020, we entered into two separate license agreements with Cornell University for preclinical research and development collaborations and non-exclusive license rights to patents for certain products and technologies. As part of our first license agreement, as amended, we assumed oversight for the conduct of the Phase 1/2 clinical trial of LX1001 that was initiated by Cornell University at the end of 2019. Pursuant to these license agreements, we are obligated to diligently proceed with the development, manufacture, and sale of licensed products. In February 2021, we further expanded our collaboration and entered into a Research Collaboration Agreement with Cornell University, or the Cornell Collaboration Agreement, to enable discovery and preclinical research aimed at expanding our cardiovascular pipeline. If Cornell University delays or fails to perform its obligations under this collaboration agreement, disagrees with our interpretation of the terms of the collaboration or our discovery plan or terminates any of our existing license and collaboration agreements, our pipeline of product candidates would be significantly adversely affected and our prospects may be materially harmed.

Our collaboration with UCSD is also highly important to our business, as we have licensed from UCSD intellectual property rights related to our LX 2021, LX 2022 and LX 2021 programs under three separate license agreements, and we have entered into sponsored research agreements with UCSD for preclinical research and development for these programs. Furthermore, our Chief Scientific Officer, Eric Adler, is the director of the UCSD-affiliated lab at which the preclinical work for the LX2022 program is conducted. If UCSD delays or fails to perform its obligations under either of the sponsored research agreements, disagrees with our interpretation of the terms of the sponsored research or our discovery plan or terminates any of our existing license agreements, our pipeline of product candidates would be significantly adversely affected and our prospects may be materially harmed.

***We intend to continue to rely on third parties to conduct a significant portion of our existing clinical trials and potential future clinical trials for product candidates, and those third parties may not perform satisfactorily, including failing to meet deadlines for the completion of such trials.***

We engage CROs to help conduct our ongoing clinical trials. We expect to continue to rely on third parties, including clinical data management organizations, medical institutions and clinical investigators, to conduct those clinical trials and any future clinical trials. Any of these third parties may terminate their engagements with us, some in the event of an uncured material breach and some at any time for convenience. If any of our relationships with these third parties terminate, we may not be able to timely enter into arrangements with alternative third parties or to do so on commercially reasonable terms, if at all. Switching or adding a CRO involves substantial cost and requires management time and focus. In addition, there is a natural transition period when a new CRO commences work. As a result, delays occur, which can materially impact our ability to meet our desired clinical development timelines. Though we intend to carefully manage our relationships with our CROs, there can be no assurance that we will not encounter challenges or delays in the future or that these delays or challenges will not have a material adverse impact on our business, financial condition and prospects.

In addition, any third parties conducting our clinical trials will not be our employees, and except for remedies available to us under our agreements with such third parties, we cannot control whether or not they devote sufficient time and resources to our clinical programs. If these third parties do not successfully carry out their contractual duties or obligations or meet expected deadlines, if they need to be replaced or if the quality or accuracy of the clinical data they obtain is compromised due to the failure to adhere to our clinical protocols, regulatory requirements or for other reasons, our clinical trials may be extended, delayed or terminated and we may not be able to obtain regulatory approval for or successfully commercialize our product candidates. Consequently, our results of operations and the commercial prospects for our product candidates would be harmed, our costs could increase substantially and our ability to generate revenue could be delayed significantly.

We rely on these parties for execution of our preclinical studies and clinical trials, and generally do not control their activities. Our reliance on these third parties for research and development activities will reduce our control over these activities but will not relieve us of our responsibilities. For example, we will remain responsible for ensuring that each of our clinical trials is conducted in accordance with the general investigational plan and protocols for the trial. Moreover, the FDA requires us to comply with cGCP regulations, for conducting, recording and reporting the results of clinical trials to assure that data and reported results are credible and accurate and that the rights, integrity and confidentiality of trial participants are protected. We also are required to register ongoing clinical trials and post the results of completed clinical trials on a government-sponsored database, ClinicalTrials.gov, within specified time frames. Failure to do so can result in fines, adverse publicity and civil and criminal sanctions. If we or any of our CROs or other third parties, including trial sites, fail to comply with applicable cGCPs, the clinical data generated in our clinical trials may be deemed unreliable and the FDA, EMA or comparable foreign regulatory authorities may require us to perform additional clinical trials before approving our marketing applications. We cannot assure you that upon inspection by a given regulatory authority, such regulatory authority will determine that any of our clinical trials complies with cGCP regulations. In addition, our clinical trials must be conducted with product produced under cGMP conditions. Our failure to comply with these regulations may require us to repeat clinical trials, which would delay the regulatory approval process.

In addition, principal investigators for our clinical trials may serve as scientific advisors or consultants to us from time to time and receive compensation in connection with such services. Under certain circumstances, we may be required to report some of these relationships to the FDA. The FDA may conclude that a financial relationship between us and a principal investigator has created a conflict of interest or otherwise affected interpretation of the trial. The FDA may therefore question the integrity of the data generated at the applicable clinical trial site

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and the utility of the clinical trial itself may be jeopardized. This could result in a delay in approval, or rejection, of our marketing applications by the FDA and may ultimately lead to the denial of marketing approval of LX2006, LX1001, or any other product candidates.

We also expect to rely on other third parties to store and distribute product supplies for our clinical trials. Any performance failure on the part of our distributors could delay clinical development or marketing approval of our product candidates or commercialization of our products, producing additional losses and depriving us of potential revenue.

***We may seek collaborations with non-academic third parties for the development or commercialization of our product candidates. If those collaborations are not successful, we may not be able to capitalize on the market potential of these product candidates.***

We may seek third-party collaborators for the development and commercialization of our product candidates, including for the commercialization of any of our product candidates that are approved for marketing outside the United States. Our likely collaborators for any such arrangements include regional and national pharmaceutical companies and biotechnology companies. If we enter into any additional such arrangements with any third parties, we will likely have limited control over the amount and timing of resources that our collaborators dedicate to the development or commercialization of our product candidates. Our ability to generate revenue from these arrangements will depend on our collaborators' abilities to successfully perform the functions assigned to them in these arrangements. Collaborations involving our product candidates pose the following risks to us:

- collaborators have significant discretion in determining the efforts and resources that they will apply to these collaborations;
- collaborators may not perform their obligations as expected;
- collaborators may not pursue development and commercialization of any product candidates that achieve regulatory approval or may elect not to continue or renew development or commercialization programs based on clinical trial results, changes in the collaborators' strategic focus or available funding, or external factors, such as an acquisition, that divert resources or create competing priorities;
- collaborators may delay clinical trials, provide insufficient funding for a clinical trial program, stop a clinical trial or abandon a product candidate, repeat or conduct new clinical trials or require a new formulation of a product candidate for clinical testing;
- collaborators could independently develop, or develop with third parties, products that compete directly or indirectly with our product candidates if the collaborators believe that competitive products are more likely to be successfully developed or can be commercialized under terms that are more economically attractive than ours;
- we could grant exclusive rights to our collaborators that would prevent us from collaborating with others;
- product candidates discovered in collaboration with us may be viewed by our collaborators as competitive with their own product candidates or drugs, which may cause collaborators to cease to devote resources to the commercialization of our product candidates;
- a collaborator with marketing and distribution rights to one or more of our product candidates that achieve regulatory approval may not commit sufficient resources to the marketing and distribution of such products;
- disagreements with collaborators, including disagreements over proprietary rights, contract interpretation or the preferred course of development, might cause delays or termination of the research, development or

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commercialization of product candidates, might lead to additional responsibilities for us with respect to product candidates, or might result in litigation or arbitration, any of which would be time-consuming and expensive;

- collaborators may not properly maintain or defend our or their intellectual property rights or may use our or their proprietary information in such a way as to invite litigation that could jeopardize or invalidate such intellectual property or proprietary information or expose us to potential litigation;
- collaborators may infringe the intellectual property rights of third parties, which may expose us to litigation and potential liability; and
- collaborations may be terminated for the convenience of the collaborator and, if terminated, we could be required to raise additional capital to pursue further development or commercialization of the applicable product candidates.

Collaboration agreements may not lead to development or commercialization of product candidates in the most efficient manner or at all. If any future collaborator of ours were to be involved in a business combination, the continued pursuit and emphasis on our product development or commercialization program could be delayed, diminished or terminated.

We face significant competition in seeking appropriate collaborators. Whether we reach a definitive agreement for any collaboration will depend, among other things, upon our assessment of the collaborator's resources and expertise, the terms and conditions of the proposed collaboration and the proposed collaborator's evaluation of a number of factors. Those factors may include the design or results of clinical trials, the likelihood of approval by the FDA or similar regulatory authorities outside the United States, the potential market for the subject product candidate, the costs and complexities of manufacturing and delivering such product candidate to patients, the potential of competing products, the existence of uncertainty with respect to our ownership of technology, which can exist if there is a challenge to such ownership without regard to the merits of the challenge and industry and market conditions generally. The collaborator may also consider alternative product candidates or technologies for similar diseases that may be available to collaborate on and whether such a collaboration could be more attractive than the one with us for our product candidate. Collaborations are complex and time-consuming to negotiate and document. In addition, there have been a significant number of recent business combinations among large pharmaceutical companies that have resulted in a reduced number of potential future collaborators.

We may not be able to negotiate additional collaborations on a timely basis, on acceptable terms, or at all. If we are unable to do so, we may have to curtail the development of such product candidate, reduce or delay its development program or one or more of our other development programs, delay its potential commercialization or reduce the scope of any sales or marketing activities, or increase our expenditures and undertake development or commercialization activities at our own expense. If we elect to increase our expenditures to fund development or commercialization activities on our own, we may need to obtain additional capital, which may not be available to us on acceptable terms or at all. If we do not have sufficient funds, we may not be able to further develop our product candidates or bring them to market and generate revenue.

### **Risks related to intellectual property**

*If we are unable to obtain or protect intellectual property rights related to any of our product candidates, we may not be able to compete effectively in our market.*

We rely upon a combination of patents, trade secret protection and confidentiality agreements to protect the intellectual property related to our product candidates, including LX2006, LX2020, LX2021, LX2022, LX1001,

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LX1020, LX1021, and other programs, their respective components, formulations, therapies, methods used to manufacture them and methods treatment. Furthermore, we currently do not have any patents or patent applications covering our LX1004 product candidate.

Our success depends in large part on our ability to obtain and maintain patent and other intellectual property protection in the United States and in other countries with respect to our proprietary technology and product candidates.

We cannot offer any assurances about which of our patent applications will issue, the breadth of any resulting patent or whether any of the issued patents will be found invalid and unenforceable or will be threatened by third parties. We cannot offer any assurances that the breadth of our granted patents will be sufficient to stop a competitor from developing and commercializing a product, including a biosimilar product that would be competitive with one or more of our product candidates. Furthermore, any successful challenge to these patents or any other patents owned by or licensed to us after patent issuance could deprive us of rights necessary for the successful commercialization of any of our product candidates. Further, if we encounter delays in regulatory approvals, the period of time during which we could market a product candidate under patent protection could be reduced.

The patent prosecution process is expensive and time-consuming. We may not be able to prepare, file and prosecute all necessary or desirable patent applications at a commercially reasonable cost or in a timely manner or in all jurisdictions. It is also possible that we may fail to identify patentable aspects of inventions made in the course of development and commercialization activities before it is too late to obtain patent protection on them. We may not be able to obtain or maintain patent applications and patents due to the subject matter claimed in such patent applications and patents being in the public domain. In some cases, the work of certain academic researchers in the gene therapy field has entered the public domain, which may preclude our ability to obtain patent protection for certain inventions relating to such work. Although we enter into nondisclosure and confidentiality agreements with parties who have access to confidential or patentable aspects of our research and development output, such as our employees, corporate collaborators, outside scientific collaborators, contract manufacturers, consultants, advisors and other third parties, any of these parties may breach these agreements and disclose such output before a patent application is filed, thereby jeopardizing our ability to seek patent protection. Consequently, we would not be able to prevent any third party from using any technology that is in the public domain to compete with our product candidates. Moreover, depending on the terms of any future in-licenses to which we may become a party, we may not have the right to control the preparation, filing and prosecution of patent applications, or to maintain the patents, covering technology in-licensed from third parties. Therefore, these patents and patent applications may not be prosecuted and enforced in a manner consistent with the best interests of our business.

The patent position of biotechnology and pharmaceutical companies generally is highly uncertain, involves complex legal and factual questions and has, in recent years, been the subject of much litigation. As a result, the issuance, scope, validity, enforceability and commercial value of any patent rights are highly uncertain. Our owned and licensed pending and future patent applications may not result in issued patents which protect our technology or product candidates, effectively prevent others from commercializing competitive technologies and product candidates or otherwise provide any competitive advantage. In fact, patent applications may not issue as patents at all. Even if patent applications we license or own currently or in the future issue as patents, they may not issue in a form that will provide us with any meaningful protection, prevent competitors or other third parties from competing with us, or otherwise provide us with any competitive advantage. Any patents that we hold or in-license may be challenged, narrowed, circumvented, or invalidated by third parties. Consequently, we do not know whether any of our technologies and product candidates will be protectable or remain protected by valid and enforceable patents. In addition, our existing patents and any future patents we obtain may not be sufficiently broad to prevent others from using our technology or from developing competing technologies and products, and

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the coverage claimed in a patent application can be significantly reduced before the patent is issued, and its scope can be reinterpreted after issuance. Any failure to obtain, maintain or defend our patents and other intellectual property could have a material adverse effect on our business, financial conditions, results of operations and prospects.

We cannot be certain that we are the first to invent the inventions covered by pending patent applications and, if we are not, we may be subject to priority or entitlement disputes. We may be required to disclaim part or all of the term of certain patents or all of the term of certain patent applications. There may be prior art of which we are not aware that may affect the validity or enforceability of a patent claim. There also may be prior art of which we are aware, but which we do not believe affects the validity or enforceability of a claim, which may, nonetheless, ultimately be found to affect the validity or enforceability of a claim. Since patent applications in the United States and other countries are confidential for a period of time after filing, at any moment in time, we cannot be certain that we were in the past or will be in the future the first to file any patent application related to our product candidates. For example, some patent applications in the United States may be maintained in secrecy until the patents are issued. Further, publications in the scientific literature often lag behind actual discoveries. Consequently, we cannot be certain that others have not filed patent applications for technology covered by our owned and in-licensed issued patents or our pending applications, or that we or, if applicable, a licensor, were the first to invent or first to file an application for the technology.

It is possible that defects of form in the preparation or filing of our patents or patent applications may exist, or may arise in the future, for example, with respect to proper priority claims, inventorship, claim scope, or requests for patent term adjustments. If there are material defects in the form, preparation, prosecution, or enforcement of our patents or patent applications, such patents may be invalid and/or unenforceable, and such applications may never result in valid, enforceable patents. Any of these outcomes could impair our ability to prevent competition from third parties, which may have an adverse impact on our business.

In addition to the protection provided by our patent estate, we rely on trade secret protection and confidentiality agreements to protect proprietary know-how that is not amenable to patent protection. Although we generally require all of our employees to assign their inventions to us, and all of our employees, consultants, advisors and any third parties who have access to our proprietary know-how, information, or technology to enter into confidentiality agreements, we cannot provide any assurances that all such agreements have been duly executed, or that our trade secrets and other confidential proprietary information will not be disclosed. In addition, while we have undertaken reasonable efforts to ensure such agreements are enforceable and that employees and third parties comply with their obligations thereunder, these agreements may be found insufficient by a court of law or may be breached, or we may not enter into sufficient agreements with such individuals in the first instance, in either case potentially resulting in the unauthorized use or disclosure of our trade secrets and other intellectual property, including to our competitors, which could cause us to lose any competitive advantage resulting from this intellectual property. Individuals not subject to invention assignment agreements may make adverse ownership claims to our current and future intellectual property. Moreover, our competitors may independently develop knowledge, methods and know-how equivalent to our trade secrets. Competitors could purchase our products, if approved, and replicate some or all of the competitive advantages we derive from our development efforts for technologies on which we do not have patent protection. If any of our trade secrets were to be lawfully obtained or independently developed by a competitor, we would have no right to prevent them, or those to whom they communicate it, from using that technology or information to compete with us. If any of our trade secrets were to be disclosed to or independently developed by a competitor, our competitive position would be harmed. Enforcing a claim that a third-party entity illegally obtained and is using any of our trade secrets is expensive and time-consuming, and the outcome is unpredictable, and we may not be able to obtain adequate remedies for such breaches.

We also seek to preserve the integrity and confidentiality of our data and trade secrets by maintaining physical security of our premises and physical and electronic security of our information technology systems. While we have confidence in these individuals, organizations and systems, our agreements or security measures may be breached, and we may not have adequate remedies for any breach. Also, if the steps taken to maintain our trade secrets are deemed inadequate, we may have insufficient recourse against third parties for misappropriating the trade secret. In addition, others may independently discover our trade secrets and proprietary information. For example, the FDA is considering whether to make additional information publicly available on a routine basis, including information that we may consider to be trade secrets or other proprietary information, and it is not clear at the present time how the FDA's disclosure policies may change in the future. If we are unable to prevent material disclosure of the non-patented intellectual property related to our technologies to third parties, and there is no guarantee that we will have any such enforceable trade secret protection, we may not be able to establish or maintain a competitive advantage in our market, which could materially adversely affect our business, results of operations and financial condition.

***Patent terms may be inadequate to protect our competitive position on our products for an adequate amount of time, and if we do not obtain protection under the Hatch-Waxman Amendments and similar non-United States legislation for extending the term of patents covering each of our product candidates, our business may be materially harmed.***

Given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such candidates might expire before or shortly after such candidates are commercialized. Depending upon the timing, duration and conditions of FDA marketing approval of our product candidates, one or more of our United States patents may be eligible for limited patent term extension under the Drug Price Competition and Patent Term Restoration Act of 1984, referred to as the Hatch-Waxman Amendments, and similar legislation in the European Union. The Hatch-Waxman Amendments permit a patent term extension of up to five years for a patent covering an approved product as compensation for effective patent term lost during product development and the FDA regulatory review process. A patent term extension cannot extend the remaining term of a patent beyond a total of 14 years from the date of product approval. Only one patent may be extended, and only those claims covering the approved drug, a method for using it, or a method for manufacturing it may be extended. However, we may not receive an extension if we fail to apply within applicable deadlines, fail to apply prior to expiration of relevant patents or otherwise fail to satisfy applicable requirements. Moreover, the length of the extension could be less than we request. If we are unable to obtain patent term extension or the term of any such extension is less than we request, the period during which we can enforce our patent rights for that product will be shortened and our competitors may obtain approval to market competing products sooner. As a result, our revenue from applicable products could be reduced and could have a material adverse effect on our business.

***We in-license key intellectual property necessary for the development of each of our current product candidates. If we fail to comply with our obligations in our current and future intellectual property licenses with third parties, resulting in the termination of such licenses, we could lose rights that are important to our business.***

We are heavily reliant upon licenses to certain patent rights and proprietary technology for the development of each of our current product candidates. In particular, we in-license key patents and patent applications from Adverum Biotechnologies, Inc., or Adverum, related to LX2006, we in-license patent applications and know-how from Cornell University related to our LX1001, LX1020 and LX1021 product candidates for the treatment of Alzheimer's disease, and we in-license patent applications and know-how from the Regents of the University of California, San Diego, related to our LX2020, LX2021 and LX2022 product candidates for the

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treatment of cardiomyopathy. Our license agreements impose diligence and milestone and royalty payment obligations on us, and also contain certain development requirements. If we fail to comply with our obligations, our licensors may have the right to terminate our licenses, in which event we will not be able to develop, manufacture or market any product using the intellectual property under any such terminated agreement and may face other penalties. Such an occurrence would materially adversely affect our business prospects.

Certain of our licenses may not provide us with exclusive rights to use the licensed intellectual property and technology, or may not provide us with exclusive rights to use such intellectual property and technology in all relevant fields of use and in all territories in which we may wish to develop or commercialize our technology and product candidates in the future. In addition, the intellectual property rights licensed to us by our licensors, including certain intellectual property licensed by Cornell University, The Regents of the University of California, San Diego, and Adverum, at least in some respects, may be used by such licensors or licensed to third parties, and such third parties may have certain enforcement rights with respect to such intellectual property. Thus, patents licensed to us could be put at risk of being invalidated or interpreted narrowly in litigation filed by or against our licensors or another licensee or in administrative proceedings brought by or against our licensors or another licensee in response to such litigation or for other reasons. As a result, we may not be able to prevent competitors or other third parties from developing and commercializing competitive products, including in territories covered by our licenses.

Licenses to additional third-party technology and materials that may be required for our development programs may not be available in the future or may not be available on commercially reasonable terms, or at all, which could have a material adverse effect on our business and financial condition. In such events, we may be required to expend significant time and resources to redesign our technology, product candidates, or the methods for manufacturing them or to develop or license replacement technology, all of which may not be feasible on a technical or commercial basis. If we are unable to do so, we may be unable to develop or commercialize the affected technology or product candidates. Even if we are able to obtain such additional licenses, they may be non-exclusive thereby giving our competitors and other third parties access to the same technology licensed to us.

If we or our licensors fail to adequately protect our licensed intellectual property, our ability to commercialize our product candidates and technology could suffer. Although we have oversight rights, Cornell University and The Regents of the University of California, San Diego, generally control the prosecution, maintenance and enforcement of our in-licensed patents and patent applications. Therefore, we cannot be certain that the prosecution, maintenance and enforcement of these patent rights will be in a manner consistent with the best interests of our business, or in compliance with applicable laws and regulations, or will result in valid and enforceable patents and other intellectual property rights. It is possible that our licensors' infringement proceedings or defense activities may be less vigorous than had we conducted them ourselves or may not be conducted in accordance with our best interests. If we or our licensors fail to maintain such patents or patent applications, or if we or our licensor lose rights to those patents or patent applications, the rights we have licensed may be reduced or eliminated and our right to develop and commercialize any of our product candidates that are the subject of such licensed rights could be adversely affected. In addition to the foregoing, the risks associated with patent rights that we license from third parties will also apply to patent rights we may own in the future.

Further, if we fail to comply with our development obligations under our license agreements, we may lose our patent rights with respect to such agreement on a territory-by-territory basis, which would affect our patent rights worldwide. In spite of our efforts, our current and future licensors might conclude that we have materially breached our obligations under our license agreements and might therefore terminate such license agreements,



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thereby removing or limiting our ability to develop and commercialize products and technology covered by these license agreements. Disputes may also arise between us and our licensors regarding intellectual property subject to a license agreement, including:

- the scope of rights granted under the license agreement and other interpretation-related issues;
- our financial and other obligations under the license agreement;
- whether and the extent to which our technology and processes infringe on intellectual property of the licensor that is not subject to the licensing agreement;
- our right to sublicense patent and other rights to third parties under collaborative development relationships;
- our diligence obligations with respect to the use of the licensed technology in relation to our development and commercialization of our product candidates and what activities satisfy those diligence obligations;
- the inventorship or ownership of inventions and know-how resulting from the joint creation or use of intellectual property by our licensors and us and our partners; and
- the priority of invention of patented technology.

If disputes over intellectual property that we have licensed prevent or impair our ability to maintain our current licensing arrangements on acceptable terms, we may be unable to successfully develop and commercialize the affected technology or product candidates. In addition, if any such disputes result in the termination of our intellectual property licenses, this could result in the loss of our ability to develop and commercialize our lead product candidates, or we could lose other significant rights, experience significant delays in the development and commercialization of our other product candidates, or incur liability for damages, any of which could have a material adverse effect on our business, financial condition, results of operations and prospects. In addition, we may seek to obtain additional licenses from our licensors and, in connection with obtaining such licenses, we may agree to amend our existing licenses in a manner that may be more favorable to the licensors, including by agreeing to terms that could enable third parties, including our competitors, to receive licenses to a portion of the intellectual property that is subject to our existing licenses and to compete with our product candidates.

Some of our future agreements with certain of our third-party research partners may provide that improvements developed in the course of our relationship may be owned solely by either us or our third-party research partner. If we determine that rights to such improvements owned solely by a third-party research partner or other third party with whom we collaborate are necessary to commercialize our therapeutic candidates or maintain our competitive advantage, we may need to obtain a license from such third party in order to use the improvements and continue developing, manufacturing or marketing our drug candidates. We may not be able to obtain such a license on an exclusive basis, on commercially reasonable terms, or at all, which could prevent us from commercializing our product candidates or allow our competitors or others the chance to access technology that is important to our business.

Termination of our current or any future license agreements would reduce or eliminate our rights under these agreements and may result in our having to negotiate new or reinstated agreements with less favorable terms or cause us to lose our rights under these agreements, including our rights to important intellectual property or technology. Any of the foregoing could prevent us from commercializing our other product candidates, which could have a material adverse effect on our operating results and overall financial condition.

In addition, intellectual property rights that we in-license in the future may be sublicenses under intellectual property owned by third parties, in some cases through multiple tiers. The actions of our licensors may therefore affect our rights to use our sublicensed intellectual property, even if we are in compliance with all of the obligations under our license agreements. Should our licensors or any of the upstream licensors fail to comply

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with their obligations under the agreements pursuant to which they obtain the rights that are sublicensed to us, or should such agreements be terminated or amended, our ability to develop and commercialize our product candidates may be materially harmed.

In addition, a third party may in the future bring claims that our performance under our license agreements, including our sponsoring of clinical trials, interferes with such third party's rights under its agreement with one of our licensors. If any such claim were successful, it may adversely affect our rights and ability to advance our product candidates as clinical candidates or subject us to liability for monetary damages, any of which would have an adverse effect on our business, financial condition, results of operations and prospects.

***We are generally also subject to all of the same risks with respect to protection of intellectual property that we license as we are for intellectual property that we own, which are described above and below. If we or our licensors fail to adequately protect this intellectual property, our ability to commercialize products could suffer. We may not be successful in obtaining or maintaining necessary rights to product components and processes for our development pipeline through acquisitions and in-licenses.***

Presently, we have obtained rights to certain intellectual property rights through licenses from third parties to develop, manufacture and commercialize our lead product candidates and other potential product candidates in our pipeline. Because the commercialization of our product candidates may require the use of additional intellectual property rights held by third parties, the growth of our business likely will depend, in part, on our ability to acquire or license these intellectual property rights. Our product candidates also require specific formulations and manufacturing processes to work effectively and efficiently, and some of these rights are held by others.

We may be unable to acquire or in-license any compositions, methods of use, processes or other third-party intellectual property rights from third parties that we identify as necessary, important or more expedient to further our business operations. In addition, even if we are able to obtain such licenses, we may fail to obtain any of these licenses at a reasonable cost or on reasonable terms, which would harm our business. Were that to happen, we may need to cease use of the product candidates and technologies covered by those third-party intellectual property rights and may need to seek to develop alternative approaches that do not infringe, misappropriate or violate those intellectual property rights, which may entail additional costs and development delays if we are able to develop such alternatives, or which may not be feasible. Even if we are able to obtain a license, it may be non-exclusive, which means that our competitors may also receive access to the same technologies licensed to us. The licensing and acquisition of third-party intellectual property rights is a competitive practice, and companies that may be more established, or have greater resources than we do, may also be pursuing strategies to license or acquire third-party intellectual property rights that we may consider necessary or attractive in order to commercialize our product candidates. More established companies may have a competitive advantage over us due to their larger size and cash resources or greater clinical development and commercialization capabilities. In addition, companies that perceive us to be a competitor may be unwilling to assign or license rights to us. There can be no assurance that we will be able to successfully complete such negotiations and ultimately acquire the rights to the intellectual property surrounding the additional product candidates that we may seek to acquire. We also may be unable to license or acquire third-party intellectual property rights on terms that would allow us to make an appropriate return on our investment.

If we are unable to successfully obtain rights to required third-party intellectual property or maintain the existing intellectual property rights we have licensed, we may be required to expend significant time and resources to redesign our product candidates, or to develop or license replacement technology, all of which may not be feasible on a technical or commercial basis, and we may have to abandon development of our product candidates, any of which could have a material adverse effect on our business, financial condition, results of operations and prospects.

***Our intellectual property licenses with third parties may be subject to disagreements over contract interpretation, which could narrow the scope of our rights to the relevant intellectual property or technology or increase our financial or other obligations to our licensors.***

We currently depend, and will continue to depend, on our license agreements, including the license agreements with Cornell University and Adverum related to LX2006, with Cornell University related to our LX1001, LX1020 and LX1021 product candidates for the treatment of Alzheimer's disease, and with The Regents of the University of California, San Diego, related to LX2020, LX2021 and LX2022 product candidates for the treatment of cardiomyopathy. The agreements under which we currently license intellectual property or technology from third parties are complex, and certain provisions in such agreements may be susceptible to multiple interpretations. The resolution of any contract interpretation disagreement that may arise could narrow what we believe to be the scope of our rights to the relevant intellectual property or technology, or increase what we believe to be our financial or other obligations under the relevant agreement, either of which could have a material adverse effect on our business, financial condition, results of operations and prospects.

If any of our licenses or material relationships or any in-licenses upon which our licenses are based are terminated or breached, we may:

- lose our rights to develop and market our products;
- lose patent protection for our products;
- experience significant delays in the development or commercialization of our products;
- not be able to obtain any other licenses on acceptable terms, if at all; or
- incur liability for damages.

These risks apply to any agreements that we may enter into in the future for our products or for any future product candidates. If we experience any of the foregoing, it could have a material adverse effect on our business, financial condition, results or operations and prospects.

***We cannot be certain that any of our or licensed pending patent applications or our future owned or licensed patent applications will result in issued patent claims covering such aspects of our product candidates.***

Composition-of-matter patents on the active pharmaceutical ingredient, or API, in prescription drug products are generally considered to be the strongest form of intellectual property protection for drug products because those types of patents provide protection without regard to any particular method of use or manufacture or formulation of the API used. Although we intend to file patent applications in the future that cover these product candidates, we cannot be certain that our future owned or licensed patent applications will cover our current or future product candidates.

Method-of-use patents protect the use of a product for the specified method and formulation patents cover formulations of the API. These types of patents do not prevent a competitor or other third party from developing or marketing an identical product for an indication that is outside the scope of the patented method or from developing a different formulation that is outside the scope of the patented formulation. Moreover, with respect to method-of-use patents, even if competitors or other third parties do not actively promote their product for our targeted indications or uses for which we may obtain patents, physicians may recommend that patients use these products off-label, or patients may do so themselves. Although off-label use may infringe or contribute to the infringement of method-of-use patents, the practice is common, and this type of infringement is difficult to prevent or prosecute. In addition, there are numerous publications and other prior art that may be relevant to our owned or in-licensed method-of-use patents and patent applications and may be used to challenge the validity of

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these owned or in-licensed patents and patent applications in litigation or other intellectual property-related proceedings. If these types of challenges are successful, our owned or in-licensed patents and patent applications may be narrowed or found to be invalid, and we may lose valuable intellectual property rights. Any of the foregoing could have a material adverse effect on our business, financial conditions, prospects and results of operations.

The strength of patents in the biotechnology and pharmaceutical field involves complex legal and scientific questions and can be uncertain. The patent applications that we own or in-license may fail to result in issued patents with claims that cover our product candidates or uses thereof in the United States or in other countries. Even if patents do successfully issue, the issuance of a patent is not conclusive as to its inventorship, scope, validity or enforceability and third parties may challenge the validity, enforceability or scope of our owned and licensed patents in courts or patent offices in the United States and abroad, which may result in those patents being narrowed, invalidated or held unenforceable. Furthermore, even if they are unchallenged, our owned and licensed patents and pending patent applications, if issued, may not adequately protect our intellectual property or prevent competitors or others from designing around our patent claims to circumvent our owned or licensed patents by developing similar or alternative technologies or therapeutics in a non-infringing manner. If the breadth or strength of protection provided by the patents and patent applications we own or license with respect to our product candidates is not sufficient to impede such competition or is otherwise threatened, it could dissuade companies from collaborating with us to develop, and threaten our ability to commercialize, our product candidates. Any of the foregoing could have a material adverse effect on our business, financial condition, results of operations and prospects.

### ***We may be involved in lawsuits to protect or enforce our patents, which could be expensive, time-consuming and unsuccessful.***

Competitors may infringe the patents for which we have applied. To counter infringement or unauthorized use, we may be required to file infringement claims, which can be expensive and time-consuming. If we initiate legal proceedings against a third party to enforce a patent covering one of our product candidates, the defendant could counterclaim that the patent covering our product or product candidate is invalid and/or unenforceable. In patent litigation in the United States, counterclaims alleging invalidity and/or unenforceability are common, and there are numerous grounds upon which a third party can assert invalidity or unenforceability of a patent. In an infringement proceeding, a court may decide that the patent claims we are asserting are invalid and/or unenforceable, or may refuse to stop the other party from using the technology at issue on the grounds that our patent claims do not cover the technology in question. Third parties may also raise similar claims before administrative bodies in the United States or abroad, even outside the context of litigation. Such mechanisms include re-examination, post-grant review, inter partes review and equivalent proceedings in foreign jurisdictions (for example, opposition proceedings). Such proceedings could result in revocation of or amendment to our patents in such a way that they no longer cover our product candidates. The outcome following legal assertions of invalidity and unenforceability is unpredictable. With respect to the validity question, for example, we cannot be certain that there is no invalidating prior art, of which we, our patent counsel and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity and/or unenforceability, we would lose at least part, and perhaps all, of the patent protection on our product candidates. An adverse result in any litigation or defense proceedings could put one or more of our patents at risk of being invalidated or interpreted narrowly, could put our patent applications at risk of not issuing and could have a material adverse impact on our business.

Interference proceedings provoked by third parties or brought by us may be necessary to determine the priority of inventions with respect to our patent applications. An unfavorable outcome could require us to cease using the related technology or force us to take a license under the patent rights of the prevailing party, if available.

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Furthermore, our business could be harmed if the prevailing party does not offer us a license on commercially reasonable terms. Our defense of litigation or interference proceedings may fail and, even if successful, may result in substantial costs and distract our management and other employees. We may not be able to prevent misappropriation of our intellectual property rights, particularly in countries where the laws may not protect those rights as fully as in the United States.

Even if we establish infringement of any of our patents by a competitive product, a court may decide not to grant an injunction against further infringing activity, thus allowing the competitive product to continue to be marketed by the competitor. It is difficult to obtain an injunction in U.S. litigation and a court could decide that the competitor should instead pay us a “reasonable royalty” as determined by the court, and/or other monetary damages. A reasonable royalty or other monetary damages may or may not be an adequate remedy. Loss of exclusivity and/or competition from a related product would have a material adverse impact on our business.

For certain of our in-licensed patent rights, such as patent rights in-licensed from Cornell University and Adverum, we may not have the right to file a lawsuit for infringement and may have to rely on a licensor to enforce these rights for us. If we are not able to directly assert our licensed patent rights against infringers or if a licensor does not vigorously prosecute any infringement claims on our behalf, we may have difficulty competing in certain markets where such potential infringers conduct their business, and our commercialization efforts may suffer as a result.

In addition, we or our licensors, as the case may be, may not be able to detect infringement against our owned or in-licensed patents, which may be especially difficult for manufacturing processes or formulation patents. Even if we or our licensors detect infringement by a third party of our owned or in-licensed patents, we or our licensors, as the case may be, may choose not to pursue litigation against or settlement with the third party. If we or our licensors later sue such third party for patent infringement, the third party may have certain legal defenses available to it that otherwise would not be available but for the delay between when the infringement was first detected and when the suit was brought. These legal defenses may make it impossible for us or our licensors to enforce our owned or in-licensed patents, as the case may be, against that third party.

Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during this type of litigation. There could also be public announcements of the results of hearings, motions, or other interim proceedings or developments. If securities analysts or investors perceive these results to be negative, it could have a material adverse effect on the price of our common stock.

### ***Third parties may initiate legal proceedings alleging that we are infringing their intellectual property rights, the outcome of which would be uncertain.***

As our current and future product candidates progress toward commercialization, the possibility of a patent infringement claim against us increases. We cannot provide any assurance that our current and future product candidates do not infringe other parties’ patents or other proprietary rights, and competitors or other parties may assert that we infringe their proprietary rights in any event. We may become party to, or threatened with, adversarial proceedings or litigation regarding intellectual property rights with respect to our current and future product candidates, including interference or derivation proceedings before the USPTO, or oppositions and other proceedings in foreign jurisdictions. We may be exposed to, or threatened with, future litigation by third parties having patent or other intellectual property rights alleging that our product candidates, manufacturing methods, formulations, administration methods and/or proprietary technologies infringe, misappropriate or otherwise violate their intellectual property rights.

Numerous issued patents and pending patent applications that are owned by third parties exist in the fields in which we are developing our product candidates. As the biotechnology and pharmaceutical industries expand and

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more patents are issued, the risk increases that our product candidates may give rise to claims of infringement of the patent rights of others. Moreover, it is not always clear to industry participants, including us, the claim scope that may issue from pending patent applications owned by third parties or which patents cover various types of drugs, products or their methods of use or manufacture. Thus, because of the large number of patents issued and patent applications filed in our fields, there may be a risk that third parties, including our competitors, may allege they have patent rights encompassing our product candidates, technologies or methods and that we are employing their proprietary technology without authorization.

If we were sued for patent infringement, we would need to demonstrate that the relevant product or methods of using the product either do not infringe the patent claims of the relevant patent or that the patent claims are invalid or unenforceable, and we may not be able to do this. In order to successfully challenge the validity of any such United States patent in federal court, we would need to overcome a presumption of validity. As this burden is high and requires us to present clear and convincing evidence as to the invalidity of any such United States patent claim, there is no assurance that a court of competent jurisdiction would agree with us and invalidate the claims of any such United States patent. Moreover, given the vast number of patents in our field of technology, we cannot be certain that we do not infringe existing patents or that we will not infringe patents that may be granted in the future.

While we may decide to initiate proceedings to challenge the validity of these or other patents in the future, we may be unsuccessful, and courts or patent offices in the United States and abroad could uphold the validity of any such patent. Furthermore, because patent applications can take many years to issue and may be confidential for 18 months or more after filing, and because pending patent claims can be revised before issuance, there may be applications now pending which may later result in issued patents that may be infringed by the manufacture, use or sale of our product candidates. Regardless of when filed, we may fail to identify relevant third-party patents or patent applications, or we may incorrectly conclude that a third-party patent is invalid or not infringed by our product candidates or activities. If a patent holder believes that one of our product candidates infringes its patent, the patent holder may sue us even if we have received patent protection for our technology. Moreover, we may face patent infringement claims from non-practicing entities that have no relevant drug revenue and against whom our own patent portfolio may thus have no deterrent effect. If a patent infringement suit were threatened or brought against us, we could be forced to stop or delay research, development, manufacturing or sales of the drug or product candidate that is the subject of the actual or threatened suit.

If any third-party patents are held by a court of competent jurisdiction to be valid and enforceable and to cover any of our technology or product candidates, including the manufacturing process of our product candidates, constructs or molecules used in or formed during the manufacturing process, or any final product itself, we could be forced, including by court order, to cease developing, manufacturing or commercializing the infringing product. Alternatively, we may be required to obtain a license from such third party in order to use the infringing technology and continue developing, manufacturing or marketing the infringing product. If we were required to obtain a license to continue to manufacture or market the affected product, we may be required to pay substantial royalties or grant cross-licenses to our patents. We cannot, however, assure that any such license will be available on acceptable terms, if at all. Ultimately, we could be prevented from commercializing a product, or be forced to cease some aspect of our business operations as a result of claims of patent infringement or violation of other intellectual property rights. Further, the outcome of intellectual property litigation is subject to uncertainties that cannot be adequately quantified in advance, including the demeanor and credibility of witnesses and the identity of any adverse party. This is especially true in intellectual property cases that may turn on the testimony of experts as to technical facts upon which experts may reasonably disagree. Furthermore, we may not be able to obtain any required license on commercially reasonable terms or at all. Even if we were able to obtain a license, it could be non-exclusive, thereby giving our competitors access to the same technologies licensed to us; alternatively, or additionally, it could include terms that impede or destroy our ability to compete successfully in

the commercial marketplace. In addition, we could be found liable for significant monetary damages, including treble damages and attorneys' fees if we are found to have willfully infringed a patent. A finding of infringement could prevent us from commercializing a product or force us to cease some of our business operations, which could harm our business. Claims that we have misappropriated the confidential information or trade secrets of third parties could have a similar negative impact on our business. Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation or administrative proceedings, there is a risk that some of our confidential information could be compromised by disclosure. In addition, any uncertainties resulting from the initiation and continuation of any litigation could have material adverse effect on our ability to raise additional funds or otherwise have a material adverse effect on our business, results of operations, financial condition and prospects.

The cost to us in defending or initiating any litigation or other proceeding relating to patent or other proprietary rights, even if resolved in our favor, could be substantial, and litigation would divert our management's attention. Some of our competitors may be able to sustain the costs of complex patent litigation more effectively than we can because they have substantially greater resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could delay our research and development efforts and limit our ability to continue our operations.

***We may be subject to claims that our employees, consultants, or independent contractors have wrongfully used or disclosed confidential information of third parties.***

We employ individuals who were previously employed at other biotechnology or biopharmaceutical companies. Although we try to ensure that our employees, consultants and advisors do not use the proprietary information or know-how of others in their work for us, we may be subject to claims that we or our employees, consultants, or independent contractors have inadvertently or otherwise used or disclosed confidential information of our employees' former employers or other third parties. We may also be subject to claims that former employers or other third parties have an ownership interest in our future patents. Litigation may be necessary to defend against these claims. There is no guarantee of success in defending these claims, and even if we are successful, litigation could result in substantial cost and be a distraction to our management and other employees. Even if we are successful in defending against these types of claims, litigation or other legal proceedings relating to intellectual property claims may cause us to incur significant expenses and could distract our technical and management personnel from their normal responsibilities. In addition, there could be public announcements of the results of hearings, motions or other interim proceedings or developments, and, if securities analysts or investors perceive these results to be negative, that perception could have a substantial adverse effect on the price of our common stock. This type of litigation or proceeding could substantially increase our operating losses and reduce our resources available for development activities. Some of our competitors may be able to sustain the costs of this type of litigation or proceedings more effectively than we can because of their substantially greater financial resources. Uncertainties resulting from the initiation and continuation of intellectual property litigation or other intellectual property related proceedings could adversely affect our ability to compete in the marketplace.

***We may be subject to claims challenging the inventorship or ownership of our future patents and other intellectual property.***

We may also be subject to claims that former employees, collaborators, or other third parties have an ownership interest in our patent applications, our future patents, or other intellectual property. We may be subject to ownership disputes in the future arising, for example, from conflicting obligations of consultants or others who are involved in developing our product candidates and platform discovery. Although it is our policy to require our employees and contractors who may be involved in the conception or development of intellectual property to execute agreements assigning such intellectual property to us, we may be unsuccessful in executing such an

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agreement with each party who, in fact, conceives or develops intellectual property that we regard as our own, and we cannot be certain that our agreements with such parties will be upheld in the face of a potential challenge, or that they will not be breached, for which we may not have an adequate remedy. The assignment of intellectual property rights may not be self-executing, or the assignment agreements may be breached, and litigation may be necessary to defend against these and other claims challenging inventorship or ownership. If we fail in defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights, such as exclusive ownership of, or right to use, valuable intellectual property. Such an outcome could have a material adverse effect on our business. Even if we are successful in defending against such claims, litigation could result in substantial costs and be a distraction to management and other employees.

***Reliance on third parties requires us to share our trade secrets, which increases the possibility that a competitor will discover them or that our trade secrets will be misappropriated or disclosed.***

If we rely on third parties to manufacture or commercialize our product candidates, or if we collaborate with additional third parties for the development of such product candidates, we must, at times, share trade secrets with them. We may also conduct joint research and development programs that may require us to share trade secrets under the terms of our research and development partnerships or similar agreements. We seek to protect our proprietary technology in part by entering into confidentiality agreements and, if applicable, material transfer agreements, consulting agreements or other similar agreements with our advisors, employees, third-party contractors and consultants prior to beginning research or disclosing proprietary information. These agreements typically limit the rights of the third parties to use or disclose our confidential information, including our trade secrets. Despite the contractual provisions employed when working with third parties, the need to share trade secrets and other confidential information increases the risk that such trade secrets become known by our competitors, are inadvertently incorporated into the technology of others, or are disclosed or used in violation of these agreements. Given that our proprietary position is based, in part, on our know-how and trade secrets, a competitor's discovery of our trade secrets or other unauthorized use or disclosure could have an adverse effect on our business and results of operations.

In addition, these agreements typically restrict the ability of our advisors, employees, third-party contractors and consultants to publish data potentially relating to our trade secrets. Despite our efforts to protect our trade secrets, we may not be able to prevent the unauthorized disclosure or use of our technical know-how or other trade secrets by the parties to these agreements. Moreover, we cannot guarantee that we have entered into such agreements with each party that may have or have had access to our confidential information or proprietary technology and processes. Monitoring unauthorized uses and disclosures is difficult, and we do not know whether the steps we have taken to protect our proprietary technologies will be effective. If any of the collaborators, scientific advisors, employees, contractors and consultants who are parties to these agreements breaches or violates the terms of any of these agreements, we may not have adequate remedies for any such breach or violation, and we could lose our trade secrets as a result. Moreover, if confidential information that is licensed or disclosed to us by our partners, collaborators, or others is inadvertently disclosed or subject to a breach or violation, we may be exposed to liability to the owner of that confidential information. Enforcing a claim that a third party illegally obtained and is using our trade secrets, like patent litigation, is expensive and time consuming, and the outcome is unpredictable. In addition, courts outside the United States are sometimes less willing to protect trade secrets.



***Intellectual property rights do not necessarily address all potential threats to our competitive advantage.***

The degree of future protection afforded by our intellectual property rights is uncertain because intellectual property rights have limitations, and may not adequately protect our business, or permit us to maintain our competitive advantage. The following examples are illustrative:

- others may be able to make or use capsids, nucleic acids and vectors that are similar to the biological compositions of our products that are the same as or similar to our product candidates but that are not covered by the claims of owned or in-licensed patents;
- others may independently develop similar or alternative technologies or duplicate any of our technologies;
- it is possible that others may circumvent our owned or in-licensed patents;
- others, including inventors or developers of our owned or in-licensed patented technologies who may become involved with competitors, may independently develop similar technologies that function as alternatives or replacements for any of our technologies without infringing our intellectual property rights;
- it is possible that our owned or in-licensed patents or patent applications omit individual(s) who should be listed as inventor(s) or include individual(s) who should not be listed as inventor(s), which may cause these patents or patents issuing from these patent applications to be held invalid or unenforceable;
- we or our licensors or our other collaboration partners might not have been the first to conceive and reduce to practice the inventions covered by the patents or patent applications that we own, license or will own or license;
- we or our licensors or our other collaboration partners might not have been the first to file patent applications covering certain of the patents or patent applications that we or they own or have obtained a license, or will own or will have obtained a license;
- we or our licensors may fail to meet obligations to the U.S. government with respect to in-licensed patents and patent applications funded by U.S. government grants, leading to the loss of patent rights;
- it is possible that our pending patent applications will not result in issued patents;
- we may not be able to generate sufficient data to support full patent applications that protect the entire breadth of developments in one or more of our programs;
- no patent protection may be available with regard to formulation or method of use;
- the claims of our owned or in-licensed issued patents or patent applications, if and when issued, may not cover our product candidates;
- it is possible that there are prior public disclosures that could invalidate our or our licensors' patents;
- it is possible that there are unpublished applications or patent applications maintained in secrecy that may later issue with claims covering our products or technology similar to ours;
- issued patents that we own or exclusively license may not provide us with any competitive advantage, or may be held invalid or unenforceable, as a result of legal challenges by our competitors;
- we may not exclusively license our patents and, therefore, may not have a competitive advantage if such patents are licensed to others;
- our competitors might conduct research and development activities in countries where we do not have patent rights, or in countries where research and development safe harbor laws exist, and then use the information learned from such activities to develop competitive products for sale in our major commercial markets;

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- the laws of other countries may not protect our or our licensors', as the case may be, proprietary rights to the same extent as the laws of the United States;
- there may be significant pressure on the U.S. government and international governmental bodies to limit the scope of patent protection both inside and outside the United States for disease treatments that prove successful, as a matter of public policy regarding worldwide health concerns;
- countries other than the United States may, under certain circumstances, force us or our licensors to grant a license under our patents to a competitor, thus allowing the competitor to compete with us in that jurisdiction or forcing us to lower the price of our drug in that jurisdiction;
- we have engaged in scientific collaborations in the past and will continue to do so in the future and our collaborators may develop adjacent or competing products that are outside the scope of our patents;
- we may not successfully commercialize the product candidates, if approved, before our relevant patents expire;
- we may not develop additional proprietary technologies for which we can obtain patent protection;
- it is possible that product candidates or technologies we develop may be covered by third parties' patents or other exclusive rights;
- ownership, validity or enforceability of our or our licensors' patents or patent applications may be challenged by third parties; and
- the patents of third parties or pending or future applications of third parties, if issued, may have an adverse effect on our business.

***We may enjoy only limited geographical protection with respect to certain patents and we may not be able to protect our intellectual property rights throughout the world.***

Filing and prosecuting patent applications and defending patents covering our product candidates in all countries throughout the world would be prohibitively expensive. Competitors may use our technologies in jurisdictions where we have not obtained patent protection to develop their own products and, further, may export otherwise infringing products to territories where we have patent protection, but enforcement rights are not as strong as that in the United States or Europe. These products may compete with our product candidates, and our future patents or other intellectual property rights may not be effective or sufficient to prevent them from competing.

In addition, we may decide to abandon national and regional patent applications before they are granted. The examination of each national or regional patent application is an independent proceeding. As a result, patent applications in the same family may issue as patents in some jurisdictions, such as in the United States, but may issue as patents with claims of different scope or may even be refused in other jurisdictions. It is also quite common that depending on the country, the scope of patent protection may vary for the same product candidate or technology.

While we intend to protect our intellectual property rights in our expected significant markets, we cannot ensure that we will be able to initiate or maintain similar efforts in all jurisdictions in which we may wish to market our product candidates. Accordingly, our efforts to protect our intellectual property rights in such countries may be inadequate, which may have an adverse effect on our ability to successfully commercialize our product candidates in all of our expected significant foreign markets. If we encounter difficulties in protecting, or are otherwise precluded from effectively protecting, the intellectual property rights important for our business in such jurisdictions, the value of these rights may be diminished, and we may face additional competition from others in those jurisdictions.

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The laws of some jurisdictions do not protect intellectual property rights to the same extent as the laws or rules and regulations in the United States and Europe and many companies have encountered significant difficulties in protecting and defending such rights in such jurisdictions. The legal systems of certain countries, particularly certain developing countries, do not favor the enforcement of patents, trade secrets and other intellectual property rights, which could make it difficult for us to stop the infringement of our future patents or marketing of competing products in violation of our proprietary rights generally. Proceedings to enforce our patent rights in other jurisdictions, whether or not successful, could result in substantial costs and divert our efforts and attention from other aspects of our business, could put our future patents at risk of being invalidated or interpreted narrowly and our patent applications at risk of not issuing as patents, and could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate, and the damages or other remedies awarded, if any, may not be commercially meaningful. Accordingly, our efforts to enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we develop or license.

Some countries also have compulsory licensing laws under which a patent owner may be compelled to grant licenses to third parties. In addition, some countries limit the enforceability of patents against government agencies or government contractors. In those countries, the patent owner may have limited remedies, which could materially diminish the value of such patents. If we are forced to grant a license to third parties with respect to any patents relevant to our business, our competitive position may be impaired.

***Obtaining and maintaining our patent protection depends on compliance with various procedural, document submission, fee payment and other requirements imposed by government patent agencies, and our patent protection could be reduced or eliminated for non-compliance with these requirements.***

Periodic maintenance fees, renewal fees, annuity fees and various other government fees on patents and/or applications will be due to be paid to the USPTO and various government patent agencies outside of the United States over the lifetime of our patents and/or applications and any patent rights we may obtain in the future. Furthermore, the USPTO and various non-United States government patent agencies require compliance with several procedural, documentary, fee payment and other similar provisions during the patent application process. In many cases, an inadvertent lapse of a patent or patent application can be cured by payment of a late fee or by other means in accordance with the applicable rules. There are situations, however, in which non-compliance can result in abandonment or lapse of the patents or patent applications, resulting in partial or complete loss of patent rights in the relevant jurisdiction. In such an event, potential competitors might be able to enter the market, which could have a material adverse effect on our business.

***Changes in patent laws or patent jurisprudence could diminish the value of patents in general, thereby impairing our ability to protect our product candidates.***

As is the case with other biotechnology companies, our success is heavily dependent on intellectual property, particularly patents. Obtaining and enforcing patents in the biotechnology and genetic medicine industries involve both technological and legal complexity. Therefore, obtaining and enforcing biotechnology and genetic medicine patents is costly, time-consuming and inherently uncertain. In addition, the Leahy-Smith America Invents Act (AIA), which was passed in September 2011, resulted in significant changes to the U.S. patent system.

An important change introduced by the AIA is that, as of March 16, 2013, the United States transitioned from a “first-to-invent” to a “first-to-file” system for deciding which party should be granted a patent when two or more patent applications are filed by different parties claiming the same invention. Under a “first-to-file” system, assuming the other requirements for patentability are met, the first inventor to file a patent application generally

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will be entitled to a patent on the invention regardless of whether another inventor had made the invention earlier. A third party that files a patent application in the USPTO after that date but before us could therefore be awarded a patent covering an invention of ours even if we made the invention before it was made by the third party. This will require us to be cognizant going forward of the time from invention to filing of a patent application and be diligent in filing patent applications, but circumstances could prevent us from promptly filing patent applications on our inventions.

Among some of the other changes introduced by the AIA are changes that limit where a patentee may file a patent infringement suit and providing opportunities for third parties to challenge any issued patent in the USPTO. This applies to all of our U.S. patents, even those issued before March 16, 2013. Because of a lower evidentiary standard in USPTO proceedings compared to the evidentiary standard in U.S. federal courts necessary to invalidate a patent claim, a third party could potentially provide evidence in a USPTO proceeding sufficient for the USPTO to hold a claim invalid even though the same evidence would be insufficient to invalidate the claim if first presented in a district court action.

Accordingly, a third party may attempt to use the USPTO procedures to invalidate our patent claims that would not have been invalidated if first challenged by the third party as a defendant in a district court action. It is not clear what, if any, impact the AIA will have on the operation of our business. However, the AIA and its implementation could increase the uncertainties and costs surrounding the prosecution of our or our licensors' patent applications and the enforcement or defense of our or our licensors' issued patents.

We may become involved in opposition, interference, derivation, inter partes review or other proceedings challenging our or our licensors' patent rights, and the outcome of any proceedings are highly uncertain. An adverse determination in any such proceeding could reduce the scope of, or invalidate, our owned or in-licensed patent rights, allow third parties to commercialize our technology or products and compete directly with us, without payment to us, or result in our inability to manufacture or commercialize products without infringing third-party patent rights.

In addition, the United States federal government retains certain rights in inventions produced with its financial assistance under the Bayh-Dole Act. The federal government retains a "nonexclusive, nontransferable, irrevocable, paid-up license" for its own benefit. The Bayh-Dole Act also provides federal agencies with "march-in rights." March-in rights allows the government, in specified circumstances, to require the contractor or successors in title to the patent to grant a "nonexclusive, partially exclusive, or exclusive license" to a "responsible applicant or applicants." If the patent owner refuses to do so, the government may grant the license itself. Some of our licensed patents are subject to the provisions of the Bayh-Dole Act. If our licensors fail to comply with the regulations of the Bayh-Dole Act, they could lose title to any patents subject to such regulations, which could affect our license rights under the patents and our ability to stop others from using or commercializing similar or identical technology and products, or limit patent protection for our technology and products.

Additionally, the U.S. Supreme Court has ruled on several patent cases in recent years either narrowing the scope of patent protection available in certain circumstances or weakening the rights of patent owners in certain situations, and there are other open questions under patent law that courts have yet to decisively address. In addition to increasing uncertainty with regard to our ability to obtain patents in the future, this combination of events has created uncertainty with respect to the value of patents, once obtained. Depending on decisions by Congress, the federal courts and the USPTO, the laws and regulations governing patents could change in unpredictable ways and could weaken our ability to obtain new patents or to enforce our existing patents and patents that we might obtain in the future. In addition, the European patent system is relatively stringent in the type of amendments that are allowed during prosecution, but the complexity and uncertainty of European patent laws has also increased in recent years. Complying with these laws and regulations could limit our ability to obtain new patents in the future that may be important for our business.

***If our trademarks and trade names are not adequately protected, then we may not be able to build name recognition in our markets of interest and our business may be adversely affected.***

Our current or future trademarks or trade names may be challenged, infringed, circumvented or declared generic or descriptive or determined to be infringing on other marks. We may not be able to protect our rights to these trademarks and trade names or may be forced to stop using these names, which we need for name recognition by potential partners or customers in our markets of interest. During trademark registration proceedings, we may receive rejections of our applications by the USPTO or in other foreign jurisdictions. Although we would be given an opportunity to respond to those rejections, we may be unable to overcome such rejections. In addition, in the USPTO and in comparable agencies in many foreign jurisdictions, third parties are given an opportunity to oppose pending trademark applications and to seek to cancel registered trademarks. Opposition or cancellation proceedings may be filed against our trademarks, and our trademarks may not survive such proceedings. If we are unable to establish name recognition based on our trademarks and trade names, we may not be able to compete effectively and our business may be adversely affected. We may license our trademarks and trade names to third parties, such as distributors. Although these license agreements may provide guidelines for how our trademarks and trade names may be used, a breach of these agreements or misuse of our trademarks and tradenames by our licensees may jeopardize our rights in or diminish the goodwill associated with our trademarks and trade names.

Moreover, any name we have proposed to use with our product candidate in the United States must be approved by the FDA, regardless of whether we have registered it, or applied to register it, as a trademark. Similar requirements exist in Europe. The FDA typically conducts a review of proposed product names, including an evaluation of potential for confusion with other product names. If the FDA (or an equivalent administrative body in a foreign jurisdiction) objects to any of our proposed proprietary product names, we may be required to expend significant additional resources in an effort to identify a suitable substitute name that would qualify under applicable trademark laws, not infringe the existing rights of third parties and be acceptable to the FDA. Furthermore, in many countries, owning and maintaining a trademark registration may not provide an adequate defense against a subsequent infringement claim asserted by the owner of a senior trademark. At times, competitors or other third parties may adopt trade names or trademarks similar to ours, thereby impeding our ability to build brand identity and possibly leading to market confusion. In addition, there could be potential trade name or trademark infringement claims brought by owners of other registered trademarks or trademarks that incorporate variations of our registered or unregistered trademarks or trade names. If we assert trademark infringement claims, a court may determine that the marks we have asserted are invalid or unenforceable, or that the party against whom we have asserted trademark infringement has superior rights to the marks in question. In this case, we could ultimately be forced to cease use of such trademarks.

### **Risks related to legal and regulatory compliance matters**

***Our current and future relationships with customers, healthcare providers, including physicians, and third-party payors may be subject, directly or indirectly, to federal and state healthcare fraud and abuse laws, false claims laws, health information privacy and security laws and other healthcare laws and regulations. If we are unable to comply, or have not fully complied, with such laws, we could face substantial penalties.***

We are currently or will in the future be subject to healthcare regulation and enforcement by the U.S. federal government and the states in which we will conduct our business once our product candidates are approved by the FDA and commercialized in the United States. In addition to the FDA's restrictions on marketing of pharmaceutical products, the U.S. healthcare laws and regulations that may affect our ability to operate include: the federal fraud and abuse laws, including the federal anti-kickback and false claims laws; federal data privacy and security laws; and federal transparency laws related to payments and/or other transfers of value made to

physicians and other healthcare professionals and teaching hospitals. For more information, see the section titled “*Business – Government Regulation – Other Healthcare Laws and Compliance Requirements.*” Many states have similar laws and regulations that may differ from each other and federal law in significant ways, thus complicating compliance efforts. For example, states have anti-kickback and false claims laws that may be broader in scope than analogous federal laws and may apply regardless of payor. In addition, state data privacy laws that protect the security of health information may differ from each other and may not be preempted by federal law. Moreover, several states have enacted legislation requiring pharmaceutical manufacturers to, among other things, establish marketing compliance programs, file periodic reports with the state, make periodic public disclosures on sales and marketing activities, report information related to drug pricing, require the registration of sales representatives, and prohibit certain other sales and marketing practices. These laws may adversely affect our sales, marketing and other activities with respect to any product candidate for which we receive approval to market in the United States by imposing administrative and compliance burdens on us. It is possible that governmental authorities will conclude that our current or future business practices may not comply with current or future statutes, regulations or case law involving applicable fraud and abuse or other healthcare laws and regulations. If our operations are found to be in violation of any of these laws or any other governmental regulations that may apply to us, we may be subject to significant penalties, including, without limitation, civil, criminal and administrative penalties, damages, fines, disgorgement, individual imprisonment, exclusion from participating in federal and state funded healthcare programs, such as Medicare and Medicaid, additional reporting requirements and oversight if we become subject to a corporate integrity agreement or similar agreement to resolve allegations of non-compliance with these laws, contractual damages, diminished profits and future earnings, reputational harm and the curtailment or restructuring of our operations, any of which could harm our business.

The risk of being found in violation of these laws is increased by the fact that many of them have not been fully interpreted by the regulatory authorities or the courts, and their provisions are open to a variety of interpretations. Efforts to ensure that our business arrangements with third parties will comply with applicable healthcare laws and regulations will involve substantial costs. Any action against us for violation of these laws, even if we successfully defend against it, could cause us to incur significant legal expenses and divert our management’s attention from the operation of our business. The shifting compliance environment and the need to build and maintain robust and expandable systems to comply with multiple jurisdictions with different compliance and/or reporting requirements increases the possibility that a healthcare company may run afoul of one or more of the requirements.

***Even if we obtain FDA or EMA approval for any of our product candidates in the United States or European Union, we may never obtain approval for or commercialize any of them in any other jurisdiction, which would limit our ability to realize their full market potential.***

In order to market any products in any particular jurisdiction, we must establish and comply with numerous and varying regulatory requirements on a country-by-country basis regarding safety and efficacy.

Approval by the FDA in the United States or the EMA in the European Union does not ensure approval by regulatory authorities in other countries or jurisdictions. However, the failure to obtain approval in one jurisdiction may negatively impact our ability to obtain approval elsewhere. In addition, clinical trials conducted in one country may not be accepted by regulatory authorities in other countries, and regulatory approval in one country does not guarantee regulatory approval in any other country.

Approval processes vary among countries and can involve additional product testing and validation and additional administrative review periods. Seeking foreign regulatory approval could result in difficulties and increased costs for us and require additional preclinical studies or clinical trials which could be costly and time

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consuming. Regulatory requirements can vary widely from country to country and could delay or prevent the introduction of our products in those countries. We do not have any product candidates approved for sale in any jurisdiction, including in international markets. If we fail to comply with regulatory requirements in international markets or to obtain and maintain required approvals, or if regulatory approvals in international markets are delayed, our target market will be reduced and our ability to realize the full market potential of any product we develop will be unrealized.

***Even if we receive regulatory approval of our product candidates, we will be subject to ongoing regulatory obligations and continued regulatory review, which may result in significant additional expense, and we may be subject to penalties if we fail to comply with regulatory requirements or experience unanticipated problems with our product candidates.***

Any product candidate for which we obtain marketing approval will be subject to ongoing regulatory requirements for, among other things, manufacturing processes, submission of post-approval clinical data and safety information, labeling, packaging, distribution, adverse event reporting, storage, recordkeeping, export, import, advertising, promotional activities and product tracking and tracing. These requirements include submissions of safety and other post-marketing information and reports, establishment registration and drug listing requirements, applicable tracking and tracing requirements, continued compliance with cGMP requirements relating to manufacturing, quality control, quality assurance and corresponding maintenance of records and documents, requirements regarding the distribution of samples to physicians and recordkeeping and cGCP requirements for any clinical trials that we conduct post-approval.

Any regulatory approvals that we receive for our product candidates or any future product candidates may also be subject to a REMS, limitations on the approved indicated uses for which the product may be marketed or to the conditions of approval, or requirements that we conduct potentially costly post-marketing testing, including Phase 4 trials and surveillance to monitor the quality, safety and efficacy of the product. An unsuccessful post-marketing study or failure to complete such a study could result in the withdrawal of marketing approval. We will further be required to immediately report any serious and unexpected adverse events and certain quality or production problems with our products to regulatory authorities along with other periodic reports.

The FDA and EMA closely regulate the post-approval marketing and promotion of genetic therapy medicines to ensure they are marketed only for the approved indications and in accordance with the provisions of the approved labeling. However, companies may share truthful and not misleading information that is otherwise consistent with a product's FDA approved labeling. The FDA imposes stringent restrictions on manufacturers' communications regarding off-label use and if we market our products for uses beyond their approved diseases, we may be subject to enforcement action for off-label marketing. Violations of the FDCA, relating to the promotion of prescription drugs for unapproved uses may lead to enforcement actions and investigations alleging violations of federal and state health care fraud and abuse laws, as well as state consumer protection laws. The holder of an approved BLA must submit new or supplemental applications and obtain prior approval for certain changes to the approved product, product labeling, or manufacturing process. A company that is found to have improperly promoted off-label uses of their products may be subject to significant civil, criminal and administrative penalties.

In addition, later discovery of previously unknown adverse events or other problems with our products, manufacturers or manufacturing processes, including adverse events of unanticipated severity or frequency, or with our third-party manufacturers or manufacturing processes, or failure to comply with regulatory requirements, may yield various results, including:

- restrictions on manufacturing such products;
- restrictions on the labeling or marketing of a product;

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- restrictions on product distribution or use;
- refusal to allow entry into supply contracts, including government contracts;
- requirements to conduct post-marketing studies or clinical trials;
- warning or untitled letters, or holds on clinical trials;
- withdrawal of the products from the market;
- refusal to approve pending applications or supplements to approved applications that we submit;
- recall of products;
- fines, restitution or disgorgement of profits or revenues;
- suspension or withdrawal of marketing approvals;
- refusal to permit the import or export of our products;
- product seizure or detention; or
- injunctions or the imposition of administrative, civil or criminal penalties or monetary fines.

The FDA's policies, and the policies of foreign regulatory agencies, may change and additional government regulations may be enacted that could prevent, limit or delay regulatory approval of our product candidates.

We also cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative or executive action, either in the United States or abroad. For example, executive orders or other actions could impose significant burdens on, or otherwise materially delay, FDA's ability to engage in routine oversight activities such as implementing statutes through rulemaking, issuance of guidance, and review and approval of marketing applications. If such executive actions were to impose restrictions on FDA's ability to engage in oversight and implementation activities in the normal course, our business could be negatively impacted. If we are slow or unable to adapt to changes in existing requirements or the adoption of new requirements or policies, or if we are not able to maintain regulatory compliance, we may lose any marketing approval that we may have obtained which would adversely affect our business, prospects and ability to achieve or sustain profitability.

Any government investigation of alleged violations of law could require us to expend significant time and resources in response and could generate negative publicity. The occurrence of any event or penalty described above may inhibit our ability to commercialize our current product candidates or any future product candidates and harm our business, financial condition, results of operations and prospects.

***Enacted and future healthcare legislation may increase the difficulty and cost for us to progress our clinical programs and obtain marketing approval of and commercialize our product candidates and may affect the prices we may set.***

In the United States, the European Union and other jurisdictions, there have been, and we expect there will continue to be, a number of legislative and regulatory changes and proposed changes to the healthcare system that could affect our future results of operations. In particular, there have been and continue to be a number of initiatives at the U.S. federal and state levels that seek to reduce healthcare costs and improve the quality of healthcare. For more information, see the below section titled "*Business—Government Regulation—Healthcare Reform.*"



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The continuing efforts of the government, insurance companies, managed care organizations and other payers of healthcare services to contain or reduce costs of healthcare may adversely affect:

- the demand for any of our product candidates, if approved;
- the ability to set a price that we believe is fair for any of our product candidates, if approved;
- our ability to generate revenues and achieve or maintain profitability;
- the level of taxes that we are required to pay; and
- the availability of capital.

Legislative and regulatory proposals have been made to expand post-approval requirements and restrict sales and promotional activities for pharmaceutical and biologic products. We cannot be sure whether additional legislative changes will be enacted, or whether FDA regulations, guidance or interpretations will be changed, or what the impact of such changes on the marketing approvals of our product candidates, if any, may be. In addition, increased scrutiny by Congress of the FDA's approval process may significantly delay or prevent marketing approval, as well as subject us to more stringent product labeling and post-marketing testing and other requirements.

Moreover, payment methodologies may be subject to changes in healthcare legislation and regulatory initiatives. For example, CMS may develop new payment and delivery models, such as bundled payment models. In addition, recently there has been heightened governmental scrutiny over the manner in which manufacturers set prices for their marketed products, which has resulted in several U.S. presidential executive orders, Congressional inquiries and proposed and enacted federal legislation designed to, among other things, bring more transparency to drug pricing, reduce the cost of prescription drugs under Medicare, and review the relationship between pricing and manufacturer patient programs. We expect that additional U.S. federal healthcare reform measures will be adopted in the future, any of which could limit the amounts that the U.S. federal government will pay for healthcare products and services, which could result in reduced demand for our product candidates or additional pricing pressures and could negatively affect our customers and accordingly, our financial operations.

Individual states in the United States have also increasingly passed legislation and implemented regulations designed to control pharmaceutical and biological product pricing, including price or patient reimbursement constraints, discounts, restrictions on certain product access and marketing cost disclosure and transparency measures, and, in some cases, designed to encourage importation from other countries and bulk purchasing. Legally mandated price controls on payment amounts by third-party payors or other restrictions could harm our business, results of operations, financial condition and prospects. In addition, regional healthcare authorities and individual hospitals are increasingly using bidding procedures to determine what pharmaceutical products and which suppliers will be included in their prescription drug and other healthcare programs. This could reduce the ultimate demand for our product candidates or put pressure on our product pricing.

In the European Union, similar political, economic and regulatory developments may affect our ability to profitably commercialize our product candidates, if approved. In addition to continuing pressure on prices and cost containment measures, legislative developments at the European Union or member state level may result in significant additional requirements or obstacles that may increase our operating costs. The delivery of healthcare in the European Union, including the establishment and operation of health services and the pricing and reimbursement of medicines, is almost exclusively a matter for national, rather than European Union, law and policy. National governments and health service providers have different priorities and approaches to the delivery of health care and the pricing and reimbursement of products in that context. In general, however, the healthcare budgetary constraints in most EU member states have resulted in restrictions on the pricing and

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reimbursement of medicines by relevant health service providers. Coupled with ever-increasing European Union and national regulatory burdens on those wishing to develop and market products, this could prevent or delay marketing approval of our product candidates, restrict or regulate post-approval activities and affect our ability to commercialize our product candidates, if approved.

In markets outside of the United States and the European Union, reimbursement and healthcare payment systems vary significantly by country, and many countries have instituted price ceilings on specific products and therapies.

We cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative action in the United States, the European Union or any other jurisdiction. If we or any third parties we may engage are slow or unable to adapt to changes in existing requirements or the adoption of new requirements or policies, or if we or such third parties are not able to maintain regulatory compliance, our product candidates may lose any regulatory approval that may have been obtained and we may not achieve or sustain profitability.

***Changes in funding for the FDA and other government agencies could hinder their ability to hire and retain key leadership and other personnel, or otherwise prevent new products and services from being developed or commercialized in a timely manner, which could negatively impact our business.***

The ability of the FDA to review and approve new products can be affected by a variety of factors, including government budget and funding levels, ability to hire and retain key personnel, accept the payment of user fees, and statutory, regulatory and policy changes. Average review times at the agency have fluctuated in recent years as a result. In addition, government funding of other government agencies that fund research and development activities is subject to the political process, which is inherently fluid and unpredictable.

Disruptions at the FDA and other agencies may also slow the time necessary for new drugs to be reviewed and/or approved by necessary government agencies, which would adversely affect our business. For example, over the last several years, including for 35 days beginning on December 22, 2018, the U.S. government has shut down several times and certain regulatory agencies, such as the FDA, have had to furlough critical FDA employees and stop critical activities. Our business depends upon the ability of the FDA to accept and review our potential regulatory filings. If a prolonged government shutdown occurs, it could significantly impact the ability of the FDA to timely review and process our regulatory submissions, which could have a material adverse effect on our ability to advance clinical development of our product candidates.

***If we are unable to establish sales, marketing and distribution capabilities either on our own or in collaboration with third parties, we may not be successful in commercializing our product candidates or realizing the synergies in the target diseases of our programs, even if they are approved.***

We do not have any infrastructure for the sales, marketing or distribution of our products, and the cost of establishing and maintaining such an organization may exceed the cost-effectiveness of doing so. We expect to build a focused sales, distribution and marketing infrastructure to market our product candidates in the United States and European Union, if approved. There are significant expenses and risks involved with establishing our own sales, marketing and distribution capabilities, including our ability to hire, retain and appropriately incentivize qualified individuals, generate sufficient sales leads, provide adequate training to sales and marketing personnel, and effectively manage a geographically dispersed sales and marketing team. Any failure or delay in the development of our internal sales, marketing and distribution capabilities could delay any product launch, which would adversely impact the commercialization of our product candidates. Additionally, if the commercial launch of our product candidates for which we recruit a sales force and establish marketing capabilities is

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delayed or does not occur for any reason, we would have prematurely or unnecessarily incurred these commercialization expenses. This may be costly, and our investment would be lost if we cannot retain or reposition our sales and marketing personnel.

We may not have the resources in the foreseeable future to allocate to the sales and marketing of our product candidates in certain international markets. Therefore, our future sales in these markets will largely depend on our ability to enter into and maintain collaborative relationships for such capabilities, the collaborator's strategic interest in the product and such collaborator's ability to successfully market and sell the product. We may pursue collaborative arrangements regarding the sale and marketing of LX2006, LX1001 or LX2020, if approved, for certain markets overseas; however, we cannot assure that we will be able to establish or maintain such collaborative arrangements, or if able to do so, that they will have effective sales forces.

If we are unable to build our own sales force or negotiate a collaborative relationship for the commercialization of LX2006, LX1001 or LX2020, or any of our other product candidates, if approved, we may be forced to delay the potential commercialization of LX2006, LX1001 or LX2020 or any of our other product candidates or reduce the scope of our sales or marketing activities for LX2006, LX1001 or LX2020 or any of our other product candidates. If we elect to increase our expenditures to fund commercialization activities internationally, we will need to obtain additional capital, which may not be available to us on acceptable terms, or at all. We could enter into arrangements with collaborative partners at an earlier stage than otherwise would be ideal and we may be required to relinquish rights to LX2006, LX1001 or LX2020 or any of our other product candidates or otherwise agree to terms unfavorable to us, any of which may have an adverse effect on our business, operating results and prospects.

If we are unable to establish adequate sales, marketing and distribution capabilities, either on our own or in collaboration with third parties, we will not be successful in commercializing LX2006, LX1001 or LX2020 or any of our other product candidates, if approved, and may not become profitable and may incur significant additional losses. We will be competing with many companies that currently have extensive and well-funded marketing and sales operations. Without an internal team or the support of a third party to perform marketing and sales functions, we may be unable to compete successfully against these more established companies.

***If we obtain approval to commercialize any products outside of the United States or the European Union, a variety of risks associated with international operations could adversely affect our business.***

If LX2006, LX1001, LX2020 or any of our other product candidates are approved for commercialization, we may seek to enter into agreements with third parties to market them in certain jurisdictions outside the United States and the European Union. We expect that we would be subject to additional risks related to international pharmaceutical operations, including:

- different regulatory requirements for drug and biologic approvals and rules governing drug and biologic commercialization in foreign countries;
- reduced protection for intellectual property rights;
- foreign reimbursement, pricing and insurance regimes;
- unexpected changes in tariffs, trade barriers and regulatory requirements;
- economic weakness, including inflation, or political instability in particular foreign economies and markets;
- foreign currency fluctuations, which could result in increased operating expenses and reduced revenues, and other obligations incident to doing business in another country;

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- business interruptions resulting from geopolitical actions, including war and terrorism or natural disasters including earthquakes, typhoons, floods and fires, or from economic or political instability;
- greater difficulty with enforcing our contracts;
- potential noncompliance with the U.S. Foreign Corrupt Practices Act, the U.K. Bribery Act 2010 and similar anti-bribery and anticorruption laws in other jurisdictions; and
- production shortages resulting from any events affecting raw material supply or manufacturing capabilities abroad.

In addition, there are complex regulatory, tax, labor and other legal requirements imposed by individual countries in Europe with which we will need to comply. If we are unable to successfully manage the challenges of international expansion and operations, our business and operating results could be harmed.

***If we fail to comply with environmental, health and safety laws and regulations, we could become subject to fines or penalties or incur costs that could harm our business.***

We are subject to numerous environmental, health and safety laws and regulations, including those governing laboratory procedures and the handling, use, storage, treatment and disposal of hazardous materials and wastes. From time to time and in the future, our operations may involve the use of hazardous and flammable materials, including chemicals and biological materials, and may also produce hazardous waste products. Even if we contract with third parties for the disposal of these materials and waste products, we cannot completely eliminate the risk of contamination or injury resulting from these materials. In the event of contamination or injury resulting from the use or disposal of our hazardous materials, we could be held liable for any resulting damages, and any liability could exceed our resources. We also could incur significant costs associated with civil or criminal fines and penalties for failure to comply with such laws and regulations.

We maintain workers' compensation insurance to cover us for costs and expenses we may incur due to injuries to our employees, but this insurance may not provide adequate coverage against potential liabilities. However, we do not maintain insurance for environmental liability or toxic tort claims that may be asserted against us.

In addition, we may incur substantial costs in order to comply with current or future environmental, health and safety laws and regulations. Environmental laws and regulations may impair our research, development or production efforts. In addition, failure to comply with these laws and regulations may result in substantial fines, penalties or other sanctions.

## **Risks related to employee matters and managing our growth**

***Our future success depends on our ability to retain key executives and advisors and to attract, retain and motivate qualified personnel.***

We are highly dependent on the management, development, clinical, financial and business development expertise of our executive officers, particularly R. Nolan Townsend, our Chief Executive Officer and a member of our board of directors, Paul McCormac, Ph.D., our Chief Technical Officer, Eric Adler, our Chief Scientific Officer, and Micah Zajic, our Chief Financial Officer, as well as on the scientific expertise of our founder, Ronald G. Crystal, M.D., Professor and Chairman of Weill Cornell Medicine's Department of Genetic Medicine. Each of our executive officers may currently terminate their employment with us at any time and we do not have an employment contract with Dr. Crystal. We do not maintain "key person" insurance for any of our executives or employees.

Recruiting and retaining qualified scientific and clinical personnel and, if we progress the development of our product pipeline toward scaling up for commercialization, manufacturing and sales and marketing personnel, will

also be critical to our success. The loss of the services of our executive officers or other key employees could impede the achievement of our development and commercialization objectives and seriously harm our ability to successfully implement our business strategy. Furthermore, replacing executive officers and key employees may be difficult and may take an extended period of time because of the limited number of individuals in our industry with the breadth of skills and experience required to successfully develop, gain regulatory approval of and commercialize gene therapy products. Competition to hire from this limited pool is intense, and we may be unable to hire, train, retain or motivate these key personnel on acceptable terms given the competition among numerous pharmaceutical and biotechnology companies for similar personnel. We also experience competition for the hiring of scientific and clinical personnel from universities and research institutions. In addition, we rely on consultants and advisors, including scientific and clinical advisors, to assist us in formulating our development and commercialization strategy. Our consultants and advisors may be employed by employers other than us and may have commitments under consulting or advisory contracts with other entities that may limit their availability to us. If we are unable to continue to attract and retain high quality personnel, our ability to pursue our growth strategy will be limited.

***We expect to expand our clinical development, manufacturing and regulatory capabilities and potentially implement sales, marketing and distribution capabilities, and as a result, we may encounter difficulties in managing our growth, which could disrupt our operations.***

As of April 30, 2023, we had 39 full-time employees and one part-time employee. As our development progresses, we expect to experience significant growth in the number of our employees and the scope of our operations, particularly in the areas of clinical product development, regulatory affairs and, if any of our product candidates receives marketing approval, sales, marketing and distribution. To manage our anticipated future growth, we must continue to implement and improve our managerial, operational and financial systems, expand our facilities and continue to recruit and train additional qualified personnel. Due to our limited financial resources and the limited experience of our management team in managing a company with such anticipated growth, we may not be able to effectively manage the expansion of our operations or recruit and train additional qualified personnel. Our choice to focus on multiple therapeutic areas may negatively affect our ability to develop adequately the specialized capability and expertise necessary for operations. The expansion of our operations may lead to significant costs and may divert our management and business development resources. Any inability to manage growth could delay the execution of our business plans or disrupt our operations.

***Our employees, independent contractors, consultants, collaborators, principal investigators, CROs, suppliers and vendors may be improperly classified and may engage in misconduct or other improper activities, including non-compliance with regulatory standards and requirements.***

We endeavor to properly classify our employees as exempt or non-exempt with respect to wage and hour laws (including, but not limited to, for purposes of minimum wage, overtime and applicable meal and rest periods), and we monitor and evaluate such classifications. Although there are no current, pending, or threatened claims or investigations against us asserting that any employees have been incorrectly classified as exempt, the possibility nevertheless exists that certain job roles could be deemed to have been incorrectly classified as exempt. In addition, we endeavor to classify our workforce properly, and we monitor and evaluate such classifications. Although there are no current, pending, or threatened claims or investigations against us asserting that any independent contractors have been incorrectly classified, the possibility nevertheless exists that certain contractors could be deemed to be employees.

We are exposed to the risk that our employees, independent contractors, consultants, collaborators, principal investigators, CROs, suppliers and vendors may engage in fraudulent conduct or other illegal activity. Misconduct by these parties could include intentional, reckless and/or negligent conduct that violates FDA

regulations, including those laws requiring the reporting of true, complete and accurate information to the FDA, manufacturing standards, federal and state healthcare laws and regulations, and laws that require the true, complete and accurate reporting of financial information or data. In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, kickbacks, self-dealing and other abusive practices. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Misconduct by these parties could also involve the improper use of individually identifiable information, including, without limitation, information obtained in the course of clinical trials, which could result in regulatory sanctions and serious harm to our reputation. In connection with this offering, we intend to adopt a code of business conduct and ethics; however, even with such a code of conduct in place, it is not always possible to identify and deter misconduct, and the precautions we take to detect and prevent this activity may not be effective in controlling unknown or unmanaged risks or losses or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to be in compliance with such laws or regulations. If any such actions are instituted against us, and we are not successful in defending ourselves or asserting our rights, those actions could have a significant impact on our business, including the imposition of significant civil, criminal and administrative penalties, including, without limitation, damages, fines, disgorgement, imprisonment, exclusion from participation in government healthcare programs, such as Medicare and Medicaid, additional reporting requirements and oversight if we become subject to a corporate integrity agreement or similar agreement to resolve allegations of non-compliance with these laws, and the curtailment or restructuring of our operations.

***Our business and operations would suffer in the event of system failures, cyberattacks or a deficiency in our or our CMOs', CROs', manufacturers' contractors', consultants' or collaborators' cybersecurity.***

Despite the implementation of security measures, our internal computer systems, as well as those of third parties on which we rely, are vulnerable to damage from, among other things, computer viruses, malware, unauthorized access, natural disasters, terrorism, war telecommunication and electrical failures, system malfunctions, cyberattacks or cyber-intrusions over the Internet, attachments to emails, phishing attacks, persons inside our organization, or persons with access to systems inside our organization. The risk of a security breach or disruption, particularly through cyberattacks or cyber intrusion, including by computer hackers, foreign governments and cyber terrorists, has generally increased as the number, intensity and sophistication of attempted attacks and intrusions from around the world have increased. If such an event were to occur and cause interruptions in our operations, it could lead to the loss, destruction, alteration, prevention of access to, disclosure, dissemination of, or damage or unauthorized access to, our data (including trade secrets or other confidential information, intellectual property, proprietary business information and personal data) or data that is processed or maintained on our behalf, and cause interruptions in our operations, which could result in a material disruption of our product candidate development programs. For example, the loss of preclinical study or clinical trial data from completed, ongoing or planned trials could result in delays in our regulatory approval efforts and significantly increase our costs to recover or reproduce the data. To the extent that any disruption or security breach were to result in a loss of or damage to our data or applications, or inappropriate disclosure of personal, confidential or proprietary information, we could incur liability and the further development of our product candidates could be delayed.

In the ordinary course of our business, we collect and store sensitive data, including intellectual property, clinical trial data, proprietary business information, personal data and personally identifiable information of our clinical trial subjects and employees, in our data centers and on our networks. The secure processing, maintenance and transmission of this information is critical to our operations. Despite our security measures, we cannot ensure that our information technology and infrastructure will prevent breakdowns or breaches in our or their systems or other cybersecurity incidents that cause loss, destruction, unavailability, alteration, dissemination of, or damage

or unauthorized access to, our data, including personal data, assets and other data processed or maintained on our behalf, that could have a material adverse effect upon our reputation, business, operations or financial condition. Although, to our knowledge, we have not experienced any such material security breach to date, any such breach could compromise our networks and the information stored there could be accessed, publicly disclosed, lost or stolen. Any such access, disclosure or other loss of information could result in legal claims or proceedings, liability under laws that protect the privacy of personal information, significant regulatory penalties, and such an event could disrupt our operations, damage our reputation, and cause a loss of confidence in us and our ability to conduct clinical trials, which could adversely affect our reputation and delay clinical development of our product candidates.

To the extent that any disruption or security breach were to result in a loss of or damage to our data or applications, or inappropriate disclosure of confidential or proprietary information or personal data, we could incur material legal claims and liability and damage to our reputation, and the further development of our product candidates could be delayed. Any such event could also compel us to comply with federal and state breach notification laws, and foreign law equivalents, subject us to mandatory corrective action and otherwise subject us to substantial liability under laws, rules, regulations and standards that protect the privacy and security of personal data, which could result in significant legal and financial exposure and reputational damages that could potentially have an adverse effect on our business.

Notifications and follow-up actions related to a data breach or other security incident could impact our reputation and cause us to incur significant costs, including significant legal expenses and remediation costs. We expect to incur significant costs in an effort to detect and prevent security incidents, and we may face increased costs and requirements to expend substantial resources in the event of an actual or perceived security incident. However, we cannot guarantee that we will be able to detect or prevent any such incidents, or that we can remediate any such incidents in an effective or timely manner. Our efforts to improve security and protect data from compromise may also identify previously undiscovered instances of data breaches or other cybersecurity incidents. To the extent that any data breach, disruption or security incident were to result in any loss, destruction, or alteration of, damage, unauthorized access to or inappropriate or unauthorized disclosure or dissemination of, our data, including personal data, or other information that is processed or maintained on our behalf, we could be exposed to litigation and governmental investigations and inquiries, the further development and commercialization of our product candidates could be delayed, and we could be subject to significant fines or penalties for any noncompliance with applicable state, federal and foreign privacy and security laws, rules, regulations and standards.

***We are subject to a variety of privacy and data security laws, rules, regulations, policies, industry standards and contractual obligations, and our failure to comply with them could harm our business.***

We maintain a large quantity of sensitive information, including confidential business and personal information in connection with the conduct of our clinical trials and related to our employees, and we are subject to laws and regulations governing the privacy and security of such information. In the United States, there are numerous federal and state privacy and data security laws and regulations governing the collection, use, disclosure and protection of personal information, including federal and state health information privacy laws, federal and state security breach notification laws and federal and state consumer protection laws. The legislative and regulatory landscape for privacy and data protection continues to evolve, and there has been an increasing focus on privacy and data protection issues, which may affect our business and is expected to increase our compliance costs and exposure to liability. In the United States, numerous federal and state laws and regulations could apply to our operations or the operations of our partners, including state data breach notification laws, state health information privacy laws and federal and state consumer protection laws and regulations, including Section 5 of the Federal Trade Commission Act, that govern the collection, use, disclosure and protection of health-related and other

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personal information. In addition, we may obtain health information from third parties, including research institutions from which we obtain clinical trial data, that are subject to privacy and security requirements under HIPAA, as amended by the Health Information Technology for Economic and Clinical Health Act, and the regulations promulgated thereunder. Depending on the facts and circumstances, we could be subject to significant penalties if we obtain, use or disclose individually identifiable health information in a manner that is not authorized or permitted by HIPAA.

In the European Economic Area, or EEA, and the UK, the collection, use, disclosure, transfer or other processing of personal data, including clinical trial data, of individuals is governed by the General Data Protection Regulation (“EU GDPR”) (with regards to the EEA) and the UK GDPR (with regards to the UK) as well as applicable national data protection legislation and requirements. In this document, “GDPR” refers to both the EU GDPR and the UK GDPR, unless specified otherwise. The GDPR is wide ranging in scope imposes numerous requirements on companies that process personal data, including requirements relating to processing health and other sensitive data, obtaining consent of the individuals to whom the personal data relates, providing information to individuals regarding data processing activities, implementing safeguards to protect the security and confidentiality of personal data, providing notification of data breaches, and taking certain measures when engaging third-party processors. The GDPR imposes substantial fines for breaches and violations (up to the greater of €20 million (£17.5 million for the UK) or 4% of our consolidated annual worldwide gross revenue), and confers a private right of action on data subjects and consumer associations to lodge complaints with supervisory authorities, seek judicial remedies and obtain compensation for damages resulting from violations of the GDPR.

The GDPR also includes restrictions on cross-border data transfers of personal data to countries outside the EEA and the UK that are not considered by the European Commission and UK government as providing “adequate” protection to personal data (“third countries”), including the United States, unless a valid GDPR transfer mechanism (for example, the European Commission approved Standard Contractual Clauses (“SCCs”) and the UK International Data Transfer Agreement/Addendum (“UK IDTA”)) has been put in place. Where relying on the SCCs or UK IDTA for data transfers, we may also be required to carry out transfer impact assessments to assess whether the recipient is subject to local laws which allow public authority access to personal data. The international transfer obligations under the EEA and UK data protection regimes will require significant effort and cost, and may result in us needing to make strategic considerations around where EEA and UK personal data is transferred and which service providers we can utilize for the processing of EEA and UK personal data. Although the UK is regarded as a third country under the EU GDPR, the European Commission has issued a decision recognizing the UK as providing adequate protection under the EU GDPR (“Adequacy Decision”) and, therefore, transfers of personal data originating in the EEA to the UK remain unrestricted.

The UK government has confirmed that personal data transfers from the UK to the EEA remain free flowing. The UK Government has also now introduced a Data Protection and Digital Information Bill (“UK Bill”) into the UK legislative process. The aim of the UK Bill is to reform the UK’s data protection regime following Brexit. If passed, the final version of the UK Bill may have the effect of further altering the similarities between the UK and EEA data protection regime and threaten the UK Adequacy Decision from the European Commission. This may lead to additional compliance costs and could increase our overall risk. The respective provisions and enforcement of the EU GDPR and UK GDPR may further diverge in the future and create additional regulatory challenges and uncertainties.

Compliance with these and any other applicable privacy and data security laws and regulations is a rigorous and time-intensive process, and we may be required to put in place additional mechanisms ensuring compliance with the new data protection rules. If we fail to comply with any such laws or regulations, we may face significant fines and penalties that could adversely affect our business, financial condition and results of operations. Furthermore, the laws are not consistent, and compliance in the event of a widespread data breach is costly. In



addition, states are constantly adopting new laws or amending existing laws, requiring attention to frequently changing regulatory requirements. For example, California enacted the California Consumer Privacy Act, or CCPA, which took effect on January 1, 2020, became enforceable by the California Attorney General on July 1, 2020 and has been dubbed the first “GDPR-like” law in the United States. The CCPA gives California residents expanded rights to access and delete their personal information, opt out of certain personal information sharing and receive detailed information about how their personal information is used by requiring covered companies to provide new disclosures to California consumers (as that term is broadly defined) and provide such consumers new ways to opt out of certain sales of personal information. The CCPA provides for civil penalties for violations, as well as a private right of action for data breaches that is expected to increase data breach litigation. Further, the California Privacy Rights Act, or CPRA, which became effective as of January 1, 2023, amended the CCPA and imposes additional data protection obligations on companies doing business in California, including additional consumer rights processes, limitations on data uses, new audit requirements for higher risk data and opt outs for certain uses of sensitive data. The amendments introduced by the CPRA also created a new California data protection agency authorized to issue substantive regulations, and it is anticipated that this development could result in increased privacy and information security enforcement. Although the CCPA, as amended by the CPRA, currently exempts certain health-related information, including clinical trial data, the CCPA may increase our compliance costs and potential liability if we expand our operations into California. Similar broad consumer privacy laws have been enacted in Colorado, Connecticut, Virginia, Utah, Iowa and Indiana and have been proposed in numerous other states and at the federal level. If passed, these bills may have potentially conflicting requirements that would make compliance challenging.

In addition to these consumer privacy laws, the state of Washington recently a comprehensive privacy bill, called the My Health My Data Act. Effective March 2024, this new law will impose strict requirements on the collection, use and processing of health related information that is not subject to HIPAA. Other states are considering bills with similar requirements. The Washington law and, if passed, the other state bills, will add additional complexity to our existing compliance obligations.

With the GDPR, CCPA and other laws, regulations and other obligations relating to privacy and data protection imposing new and relatively burdensome obligations, and with the substantial uncertainty over the interpretation and application of these and other obligations, we may face challenges in addressing their requirements and making necessary changes to our policies and practices and may incur significant costs and expenses in an effort to do so. We are currently in the process of developing and updating our policies and procedures in accordance with requirements under applicable data privacy and protection laws and regulations. We do not currently have any formal data privacy policies and procedures in place and have not completed formal assessments of whether we are in compliance with all applicable data privacy laws and regulations. Additionally, if third parties with which we work, such as vendors or service providers, violate applicable laws, rules or regulations or our policies, such violations may also put our or our clinical trial and employee data, including personal data, at risk, which could in turn have an adverse effect on our business.

### **Risks related to this offering, ownership of our common stock and our status as a public company**

*An active trading market for our common stock may not develop and you may not be able to resell your shares of our common stock at or above the initial offering price, if at all.*

Prior to this offering, there has been no public market for our common stock. The initial public offering price for our common stock was determined through negotiations with the underwriters and may not be indicative of the price at which our common stock will trade after the closing of this offering. An active trading market for our

shares may never develop or be sustained following this offering. If an active market for our common stock does not develop or is not sustained, it may be difficult for you to sell shares you purchased in this offering at an attractive price or at all.

***The trading price of the shares of our common stock may be volatile, and purchasers of our common stock could incur substantial losses.***

Our stock price may be volatile. The stock market in general and the market for biopharmaceutical companies in particular have experienced extreme volatility that has often been unrelated to the operating performance of particular companies. As a result of this volatility, investors may not be able to sell their common stock at or above the price paid for the shares. The market price for our common stock may be influenced by many factors, including:

- the reporting of unfavorable preclinical and clinical results;
- the commencement, enrollment or results of our clinical trials of LX2006, LX1001, LX2020 or any future clinical trials we may conduct, or changes in the development status of our product candidates;
- any delay in our regulatory filings for LX2006, LX1001, LX2020 or any other product candidate we may develop, and any adverse development or perceived adverse development with respect to the applicable regulatory authority's review of such filings, including without limitation the FDA's issuance of a "refusal to file" letter or a request for additional information;
- adverse results from, delays in or termination of clinical trials;
- adverse regulatory decisions, including failure to receive regulatory approval of our product candidates;
- unanticipated serious safety concerns related to the use of LX2006, LX1001 or LX2020 or any other product candidate;
- changes in financial estimates by us or by any equity research analysts who might cover our stock;
- conditions or trends in our industry;
- changes in the market valuations of similar companies;
- stock market price and volume fluctuations of comparable companies and, in particular, those that operate in the biopharmaceutical industry;
- publication of research reports about us or our industry or positive or negative recommendations or withdrawal of research coverage by securities analysts;
- announcements by us or our competitors of significant acquisitions, strategic partnerships or divestitures;
- our relationships with our collaborators;
- announcements of investigations or regulatory scrutiny of our operations or lawsuits filed against us;
- investors' general perception of our company and our business;
- recruitment or departure of key personnel;
- overall performance of the equity markets;
- trading volume of our common stock;
- disputes or other developments relating to proprietary rights, including patents, litigation matters and our ability to obtain patent protection for our technologies;

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- significant lawsuits, including patent or stockholder litigation or employee or independent contractor litigation;
- changes in the structure of healthcare payment systems;
- general political and economic conditions; and
- other events or factors, many of which are beyond our control.

The global economy, including credit and financial markets and the banking sector, has experienced extreme volatility and disruptions, including, among other things, severely diminished liquidity and credit availability, declines in consumer confidence, declines in economic growth, increases in unemployment rates, supply chain shortages, increases in inflation rates, bank failures, higher interest rates and uncertainty about economic stability. For example, the ongoing war between Russia and Ukraine has created extreme volatility in the global capital markets and may have further global economic consequences, including disruptions of the global supply chain and energy markets. Any such volatility and disruptions may have adverse consequences on us or the third parties on whom we rely. If the equity and credit markets continue to deteriorate, it may make any necessary debt or equity financings more difficult to obtain in a timely manner or on favorable terms, more costly or more dilutive. Increased inflation rates can adversely affect us by increasing our costs, including labor and employee benefit costs. In addition, higher inflation and macro turmoil and uncertainty could also adversely affect our buyers and sellers, which could reduce demand for our products. These factors may negatively affect the market price of our common stock, regardless of our actual operating performance.

In addition, in the past, stockholders have initiated class action lawsuits against pharmaceutical and biotechnology companies following periods of volatility in the market prices of these companies' stock. Such litigation, if instituted against us, could cause us to incur substantial costs and divert management's attention and resources from our business.

***A significant portion of our total outstanding shares are restricted from immediate resale but may be sold into the market in the near future. This could cause the market price of our common stock to drop significantly, even if our business is doing well.***

Sales of a substantial number of shares of our common stock in the public market could occur at any time. These sales, or the perception in the market that the holders of a large number of shares intend to sell shares, could reduce the market price of our common stock. After this offering, we will have \_\_\_\_\_ shares of common stock outstanding based on the number of shares outstanding as of March 31, 2023. This includes the shares that we are selling in this offering, which may be resold in the public market immediately. Following the consummation of this offering, approximately \_\_\_\_\_ % of our outstanding shares will be subject to a 180-day lock-up period provided under lock-up agreements executed in connection with this offering described in "Underwriting" and restricted from immediate resale under the federal securities laws as described in "Shares Eligible for Future Sale." All of these shares will, however, be able to be resold after the expiration of the lock-up period, as well as pursuant to customary exceptions thereto or upon the waiver of the lock-up agreement by or on behalf of the underwriters. We also intend to register shares of common stock that we may issue under our equity compensation plans. Once we register these shares, they can be freely sold in the public market upon issuance, subject to the lock-up agreements. As restrictions on resale end, the market price of our stock could decline if the holders of currently-restricted shares sell them or are perceived by the market as intending to sell them.

***If you purchase shares of common stock in this offering, you will suffer immediate dilution of your investment.***

The initial public offering price of our common stock will be substantially higher than the pro forma as adjusted net tangible book value per share of our common stock. Therefore, if you purchase shares of our common stock

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in this offering, you will pay a price per share that substantially exceeds our pro forma as adjusted net tangible book value per share after this offering. To the extent outstanding options are exercised, you will incur further dilution. Based on an assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the estimated price range set forth on the cover page of this prospectus, you will experience immediate dilution of \$ \_\_\_\_\_ per share, representing the difference between our pro forma as adjusted net tangible book value per share, after giving effect to this offering, and the assumed initial public offering price.

***If equity research analysts do not publish research or reports, or publish unfavorable research or reports, about us, our business or our market, our stock price and trading volume could decline.***

The trading market for our common stock will be influenced by the research and reports that equity research analysts publish about us and our business. As a newly public company, we have only limited research coverage by equity research analysts. Equity research analysts may elect not to provide research coverage of our common stock, and such lack of research coverage may adversely affect the market price of our common stock. In the event we do have equity research analyst coverage, we will not have any control over the analysts or the content and opinions included in their reports. The price of our stock could decline if one or more equity research analysts downgrade our stock or issue other unfavorable commentary or research. If one or more equity research analysts ceases coverage of our company or fails to publish reports on us regularly, demand for our stock could decrease, which in turn could cause our stock price or trading volume to decline.

***Our executive officers, directors and their affiliates, if they choose to act together, have the ability to significantly influence all matters submitted to stockholders for approval and may prevent new investors from influencing significant corporate decisions.***

Upon the closing of this offering, based on the number of common stock outstanding as of \_\_\_\_\_, our executive officers, directors and stockholders who owned more than 5% of our outstanding common stock before this offering and their respective affiliates will, in the aggregate, hold shares representing approximately \_\_\_\_\_% of our outstanding common stock. As a result, if these stockholders choose to act together, they would be able to control or significantly influence all matters submitted to our stockholders for approval, as well as our management and affairs. For example, these persons, if they choose to act together, would control or significantly influence the election of directors, the composition of our management and approval of any merger, consolidation, sale of all or substantially all of our assets or other business combination that other stockholders may desire. Any of these actions could adversely affect the market price of our common stock.

Some of these persons or entities may have interests different than yours. For example, because many of these stockholders purchased their shares at prices substantially below the current market price of our common stock and have held their shares for a longer period, they may be more interested in selling our company to an acquirer than other investors, or they may want us to pursue strategies that deviate from the interests of other stockholders.

***We are an “emerging growth company” and a “smaller reporting company” and, as a result of the reduced disclosure and governance requirements applicable to emerging growth companies and smaller reporting companies, our common stock may be less attractive to investors.***

We are an “emerging growth company” as defined in the Jumpstart Our Business Startups Act of 2012, or JOBS Act, and we intend to take advantage of some of the exemptions from reporting requirements that are applicable to other public companies that are not emerging growth companies, including:

- not being required to comply with the auditor attestation requirements in the assessment of our internal control over financial reporting;

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- not being required to comply with any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation or a supplement to the auditor's report providing additional information about the audit and the financial statements;
- reduced disclosure obligations regarding executive compensation in our periodic reports, proxy statements and registration statements; and
- not being required to hold a nonbinding advisory vote on executive compensation and stockholder approval of any golden parachute payments not previously approved.

We cannot predict if investors will find our common stock less attractive because we will rely on these exemptions. If some investors find our common stock less attractive as a result, there may be a less active trading market for our common stock and our stock price may be more volatile. We may take advantage of these reporting exemptions until we are no longer an emerging growth company. We will remain an emerging growth company until the last day of the fiscal year ending after the fifth anniversary of this offering, or, if earlier, (i) the last day of the fiscal year in which we have total annual gross revenue of at least \$1.235 billion, (ii) the date on which we are deemed to be a large accelerated filer, which means the market value of our common stock that is held by non-affiliates exceeds \$700 million as of the prior June 30, or (iii) the date on which we have issued more than \$1.0 billion in non-convertible debt during the prior three-year period.

In addition, we have elected to take advantage of the extended transition period to comply with new or revised accounting standards and to adopt certain of the reduced disclosure requirements available to emerging growth companies. As a result of the accounting standards election, we will not be subject to the same implementation timing for new or revised accounting standards as other public companies that are not emerging growth companies, which may make comparison of our financials to those of other public companies more difficult. As a result of these elections, the information that we provide in this prospectus may be different than the information investors may receive from other public companies in which they hold equity interests. In addition, it is possible that some investors will find our common stock less attractive as a result of these elections, which may result in a less active trading market for our common stock and higher volatility in our share price.

Even after we no longer qualify as an emerging growth company, we may, under certain circumstances, still qualify as a "smaller reporting company," which would allow us to take advantage of many of the same exemptions from disclosure requirements, including reduced disclosure obligations regarding executive compensation in our periodic reports and proxy statements.

***We will have broad discretion in the use of proceeds from this offering and may invest or spend the proceeds in ways with which you do not agree and in ways that may not increase the value of your investment.***

We will have broad discretion over the use of proceeds from this offering. You may not agree with our decisions, and our use of the proceeds may not yield any return on your investment. We expect to use the net proceeds to us from this offering, together with our existing cash, to advance the clinical development of LX2006, LX1001 and LX2020, to fund the continued development of our other programs and cardiac discovery efforts, and for working capital and other general corporate purposes. See "Use of Proceeds." In addition, we may use a portion of the proceeds from this offering to pursue our strategy to in-license or develop additional product candidates. Our failure to apply the net proceeds from this offering effectively could compromise our ability to pursue our growth strategy and we might not be able to yield a significant return, if any, on our investment of these net proceeds. You will not have the opportunity to influence our decisions on how to use our net proceeds from this offering.

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***Because we do not anticipate paying any cash dividends on our common stock in the foreseeable future, capital appreciation, if any, will be your sole source of gains and you may never receive a return on your investment.***

You should not rely on an investment in our common stock to provide dividend income. We have not declared or paid cash dividends on our common stock to date. We currently intend to retain our future earnings, if any, to fund the development and growth of our business. As a result, capital appreciation, if any, of our common stock will be your sole source of gain for the foreseeable future. Investors seeking cash dividends should not purchase our common stock.

***Our amended and restated certificate of incorporation provides that the Court of Chancery of the State of Delaware and the federal district courts of the United States of America will be the exclusive forums for substantially all disputes between us and our stockholders, which could limit our stockholders' ability to obtain a favorable judicial forum for disputes with us or our directors, officers or employees.***

Our amended and restated certificate of incorporation provides that the Court of Chancery of the State of Delaware is the exclusive forum for the following types of actions or proceedings under Delaware statutory or common law:

- any derivative action or proceeding brought on our behalf;
- any action asserting a breach of fiduciary duty;
- any action asserting a claim against us arising under the Delaware General Corporation Law, our amended and restated certificate of incorporation, or our amended and restated bylaws;
- any claim or cause of action seeking to interpret, apply, enforce or determine the validity of our restated certificate or our amended and restated bylaws;
- any claim or cause of action as to which the Delaware General Corporation Law confers jurisdiction on the Court of Chancery of the state of Delaware; and
- any action asserting a claim against us that is governed by the internal-affairs doctrine.

This provision would not apply to suits brought to enforce a duty or liability created by the Securities Exchange Act of 1934, as amended, or the Exchange Act. Furthermore, Section 22 of the Securities Act creates concurrent jurisdiction for federal and state courts over all such Securities Act actions. Accordingly, both state and federal courts have jurisdiction to entertain such claims. To prevent having to litigate claims in multiple jurisdictions and the threat of inconsistent or contrary rulings by different courts, among other considerations, our amended and restated certificate of incorporation further provides that the federal district courts of the United States of America will be the exclusive forum for resolving any complaint asserting a cause of action arising under the Securities Act. While the Delaware courts have determined that such choice of forum provisions are facially valid, a stockholder may nevertheless seek to bring a claim in a venue other than those designated in the exclusive forum provisions. In such instance, we would expect to vigorously assert the validity and enforceability of the exclusive forum provisions of our amended and restated certificate of incorporation. This may require significant additional costs associated with resolving such action in other jurisdictions and there can be no assurance that the provisions will be enforced by a court in those other jurisdictions.

These exclusive forum provisions may result in increased costs for investors to bring a claim. Further, these exclusive forum provisions may limit a stockholder's ability to bring a claim in a judicial forum that it finds favorable for disputes with us or our directors, officers, or other employees, which may discourage lawsuits against us and our directors, officers and other employees. If a court were to find either exclusive forum

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provision in our amended and restated certificate of incorporation to be inapplicable or unenforceable in an action, we may incur further significant additional costs associated with resolving the dispute in other jurisdictions, all of which could seriously harm our business.

***Provisions in our corporate charter documents and under Delaware law could make an acquisition of us, which may be beneficial to our stockholders, more difficult and may prevent attempts by our stockholders to replace or remove our current management.***

Provisions in our certificate of incorporation and our bylaws that will become effective upon the closing of this offering may discourage, delay, or prevent a merger, acquisition, or other change in control of our company that stockholders may consider favorable, including transactions in which you might otherwise receive a premium for your shares. These provisions could also limit the price that investors might be willing to pay in the future for shares of our common stock, thereby depressing the market price of our common stock. In addition, because our board of directors is responsible for appointing the members of our management team, these provisions may frustrate or prevent any attempts by our stockholders to replace or remove our current management by making it more difficult for stockholders to replace members of our board of directors. Among other things, these provisions:

- establish a classified board of directors such that only a portion of our directors stand for election at any given annual stockholder meeting;
- allow the authorized number of our directors to be changed from time to time by our shareholders or our board of directors;
- limit the manner in which stockholders can remove directors from our board of directors;
- establish requirements for stockholder proposals that can be acted on at stockholder meetings;
- require that stockholder actions must be effected at a duly called stockholder meeting and allow actions by our stockholders by written consent, with certain requirements;
- limit who may call stockholder meetings; and
- authorize our board of directors to issue preferred stock without stockholder approval, which could be used to institute a “poison pill” that would work to dilute the stock ownership of a potential hostile acquirer, effectively preventing acquisitions that have not been approved by our board of directors.

Moreover, because we are incorporated in Delaware, we are governed by the provisions of Section 203 of the Delaware General Corporation Law, or the DGCL, which prohibits a person who owns in excess of 15% of our outstanding voting stock from merging or combining with us for a period of three years after the date of the transaction in which the person acquired in excess of 15% of our outstanding voting stock, unless the merger or combination is approved in a prescribed manner.

***We could be subject to securities class action litigation.***

In the past, securities class action litigation has often been brought against a company following a decline in the market price of its securities. This risk is especially relevant for us because biopharmaceutical companies have experienced significant stock price volatility in recent years. If we face such litigation, it could result in substantial costs and a diversion of management’s attention and resources, which could harm our business.

## General risks

### ***Unstable market and economic conditions may have serious adverse consequences on our business and financial condition.***

The global economy, including credit and financial markets, has experienced extreme volatility and disruptions recently, including, among other things, diminished liquidity and credit availability, declines in consumer confidence, declines in economic growth, supply chain shortages, increases in inflation rates, higher interest rates, and uncertainty about economic stability. For example, the COVID-19 pandemic resulted in widespread unemployment, economic slowdown and extreme volatility in the capital markets. The Federal Reserve has raised interest rates multiple times in response to concerns about inflation and it may raise them again. Higher interest rates, coupled with reduced government spending and volatility in financial markets, may increase economic uncertainty and affect consumer spending. Similarly, the ongoing military conflict between Russia and Ukraine and increasing tensions between China and Taiwan have created extreme volatility in the global capital markets and may have further global economic consequences, including disruptions of the global supply chain. Any such volatility and disruptions may adversely affect our business or the third parties on whom we rely. If the equity and credit markets deteriorate, including as a result of political unrest or war, it may make any necessary debt or equity financing more difficult to complete, more costly, and more dilutive. Failure to secure any necessary financing in a timely manner and on favorable terms could have a material adverse effect on our growth strategy, financial performance and share price and could require us to delay or abandon development or commercialization plans. In addition, there is a risk that one or more of our service providers, manufacturers or other partners would not survive or be able to meet their commitments to us under such circumstances, which could directly affect our ability to attain our operating goals on schedule and on budget. We have experienced and may in the future experience disruptions as a result of such macroeconomic conditions, including delays or difficulties in initiating or expanding clinical trials and manufacturing sufficient quantities of materials. Any one or a combination of these events could have a material and adverse effect on our results of operations and financial condition.

### ***We are subject to U.S. and certain foreign anti-corruption laws and regulations, export and import controls, sanctions and embargoes. We could face liability and other serious consequences for violations.***

We are subject to anti-corruption laws and regulations, including the U.S. Foreign Corrupt Practices Act, or the FCPA, the U.S. domestic bribery statute contained in 18 U.S.C. § 201, the U.S. Travel Act and other state and national anti-bribery laws in the countries in which we may conduct activities in the future. Anti-corruption laws are interpreted broadly and generally prohibit companies and their employees, agents, contractors and other third-party collaborators from offering, promising, giving, or authorizing others to give anything of value, either directly or indirectly through third parties, to any person in the public or private sector to obtain or retain business. The FCPA also requires public companies to make and keep books and records that accurately and fairly reflect the transactions of the corporation and to devise and maintain an adequate system of internal accounting controls.

Our business is heavily regulated and therefore involves significant interaction with public officials, including officials of non-United States governments. Additionally, in many other countries, the healthcare providers who prescribe pharmaceuticals are employed by their government, and therefore will be considered foreign officials for purposes of the FCPA. We also expect to rely on third parties for research, preclinical studies and clinical trials and/or to obtain necessary permits, licenses, patent registrations and other marketing approvals. We can be held liable for the corrupt or other illegal activities of our employees, agents, CROs, contractors and other collaborators and partners, even if we do not explicitly authorize or have actual knowledge of such activities.



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We are also subject to export control and import laws and regulations, including the U.S. Export Administration Regulations, U.S. Customs regulations and various economic and trade sanctions regulations administered by the U.S. Treasury Department's Office of Foreign Assets Controls. Compliance with applicable regulatory requirements regarding the export of our products may create delays in the introduction of our products in international markets or, in some cases, prevent the export of our products to some countries altogether. Furthermore, U.S. export control laws and economic sanctions prohibit the shipment of certain products and services to countries, governments and persons targeted by U.S. sanctions.

There is no certainty that all of our employees, agents, suppliers, manufacturers, contractors or collaborators, or those of our affiliates, will comply with all applicable anti-corruption, export and import control, and sanctions laws and regulations, particularly given the high level of complexity of these laws. Violations of these laws and regulations could result in fines, criminal sanctions against us, our officers, or our employees, the closing down of facilities, including those of our suppliers and manufacturers, requirements to obtain export licenses, cessation of business activities in sanctioned countries, implementation of compliance programs, and prohibitions on the conduct of our business. Any such violations could include prohibitions on our ability to offer our products in one or more countries as well as difficulties in manufacturing or continuing to develop our products, and could materially damage our reputation, our brand, our international expansion efforts, our ability to attract and retain employees, and our business, prospects, operating results and financial condition.

***If we fail to maintain proper and effective internal controls, our ability to produce accurate financial statements on a timely basis could be impaired.***

After the closing of this offering, we will be subject to the reporting requirements of the Exchange Act, the Sarbanes-Oxley Act of 2002, or the Sarbanes-Oxley Act, and the rules and regulations of the stock market on which our common stock is listed. The Sarbanes-Oxley Act requires, among other things, that we maintain effective disclosure controls and procedures and internal control over financial reporting.

Commencing with our fiscal year ending December 31, 2024, we must perform system and process evaluation and testing of our internal control over financial reporting to allow management to report on the effectiveness of our internal control over financial reporting in our Form 10-K filing for that year, as required by Section 404 of the Sarbanes-Oxley Act. This will require that we incur substantial additional professional fees and internal costs to expand our accounting and finance functions and that we expend significant management efforts. Prior to this offering, we have never been required to test our internal control within a specified period, and, as a result, we may experience difficulty in meeting these reporting requirements in a timely manner.

We may identify weaknesses in our system of internal financial and accounting controls and procedures that could result in a material misstatement of our financial statements. Our internal control over financial reporting will not prevent or detect all errors and all fraud. A control system, no matter how well designed and operated, can provide only reasonable, not absolute, assurance that the control system's objectives will be met. Because of the inherent limitations in all control systems, no evaluation of controls can provide absolute assurance that misstatements due to error or fraud will not occur or that all control issues and instances of fraud will be detected. If we fail to remediate our identified material weakness, or identify additional material weaknesses, in our internal control over financial reporting, if we are not able to comply with the requirements of Section 404 of the Sarbanes-Oxley Act in a timely manner, or if we are unable to maintain proper and effective internal controls, we may not be able to produce timely and accurate financial statements. If that were to happen, the market price of our stock could decline and we could be subject to sanctions or investigations by the stock exchange on which our common stock is listed, the Securities and Exchange Commission or other regulatory authorities.

***Our ability to utilize our net operating loss carryforwards and research tax credits to offset future taxable income may be subject to limitations.***

As of December 31, 2022, we had approximately \$48.6 million of U.S. federal net operating loss carryforwards, or NOLs, \$109.3 million of U.S. state and local NOLs, and \$3.9 million of research tax credits. U.S. federal NOLs generated in taxable years beginning after December 31, 2017, do not expire and may be carried forward indefinitely, but the deductibility of such NOLs is limited to no more than 80% of current year taxable income. Our U.S. state and local NOLs begin to expire in 2040 and our federal research credits begin to expire in 2041.

In addition, under Sections 382 and 383 of the Internal Revenue Code of 1986, as amended, or the Code, if a corporation undergoes an “ownership change,” which is generally defined as a greater than 50% change, by value, in its equity ownership by certain stockholders over a rolling three-year period, the corporation’s ability to use its pre-change NOLs and certain other pre-change tax attributes (such as research tax credits) to offset its post-change taxable income or taxes may be limited. We may have undergone an ownership change in the past and may undergo an ownership change in connection with this offering or in the future some of which may be outside of our control. As a result, if we undergo an ownership change, and our ability to use our pre-change NOLs and other pre-change tax attributes (such as research tax credits) to offset our post-change income or taxes is limited, it would harm our future results of operations by effectively increasing our future tax obligations. U.S. state and local NOLs may be similar limited. In addition, at the U.S. state and local level, there may be periods during which the use of NOLs is suspended or otherwise limited, which could accelerate or permanently increase U.S. state and local taxes owed.

Irrespective of the above, our ability to utilize our NOLs and research tax credits to offset future taxable income or taxes is conditioned on our attaining profitability and generating taxable income. We do not know if and when we will generate sufficient taxable income to utilize our NOLs and research tax credits.

***Changes in tax laws or regulations that are applied adversely to us or our customers may materially harm our business.***

New tax laws, statutes, rules, regulations, or ordinances could be enacted at any time. Further, existing tax laws, statutes, rules, regulations, or ordinances could be interpreted differently, changed, repealed, or modified at any time. Any such enactment, interpretation, change, repeal, or modification could adversely affect us, possibly with retroactive effect. The IRA enacts a 15% minimum tax on the adjusted financial statement income of certain large U.S. corporations for taxable years beginning after December 31, 2022, as well as a 1% excise tax on stock repurchases made by public corporations after December 31, 2022. Further, the Tax Cuts and Jobs Act of 2017, or the Tax Act, enacted many significant changes in U.S. federal tax laws, some of which were further modified by the Coronavirus Aid, Relief, and Economic Security Act, or the CARES Act, and may be modified in the future by the current or a future presidential administration. Among other changes, the Tax Act amended the Code to require that certain research and experimental expenditures be capitalized and amortized over five years if incurred in the United States or fifteen years if incurred in foreign jurisdictions for taxable years beginning after December 31, 2021. Although the U.S. Congress has considered legislation that would defer, modify, or repeal the capitalization and amortization requirement, there is no assurance that such changes will be made. If the requirement is not deferred, repealed, or otherwise modified, it may increase our cash taxes and effective tax rate. In addition, it is uncertain if and to what extent various states will conform to the IRA, the Tax Act, the CARES Act, or any future U.S. federal tax laws. Changes in corporate tax rates, the realization of net deferred tax assets relating to our operations, the taxation of foreign earnings, and the deductibility of expenses could have a material impact on the value of our deferred tax assets, result in significant one-time charges, and increase our future U.S. tax expenses.

***We will incur increased costs and demands upon management as a result of being a public company.***

As a public company listed in the United States, we will incur significant additional legal, accounting and other expenses that we did not incur as a private company, including the cost of director and officer liability insurance. These additional costs could negatively affect our financial results. In addition, changing laws, regulations and standards relating to corporate governance and public disclosure, including regulations implemented by the SEC and the Nasdaq Stock Market, may increase legal and financial compliance costs and make some activities more time-consuming. These laws, regulations and standards are subject to varying interpretations and, as a result, their application in practice may evolve over time as new guidance is provided by regulatory and governing bodies. We intend to invest resources to comply with evolving laws, regulations and standards, and this investment may result in increased general and administrative expenses and a diversion of management's time and attention from revenue-generating activities to compliance activities. If notwithstanding our efforts to comply with new laws, regulations and standards, we fail to comply, regulatory authorities may initiate legal proceedings against us and our business may be harmed.

Failure to comply with these rules might also make it more difficult for us to obtain some types of insurance, including director and officer liability insurance, and we might be forced to accept reduced policy limits and coverage or incur substantially higher costs to obtain the same or similar coverage. The impact of these events could also make it more difficult for us to attract and retain qualified persons to serve on our board of directors, on committees of our board of directors or as members of senior management.

***Our disclosure controls and procedures may not prevent or detect all errors or acts of fraud.***

Upon completion of this offering, we will become subject to certain reporting requirements of the Exchange Act. Our disclosure controls and procedures are designed to reasonably assure that information required to be disclosed by us in reports we file or submit under the Exchange Act is accumulated and communicated to management, recorded, processed, summarized, and reported within the time periods specified in the rules and forms of the SEC. We believe that any disclosure controls and procedures or internal controls and procedures, no matter how well conceived and operated, can provide only reasonable, not absolute, assurance that the objectives of the control system are met.

These inherent limitations include the realities that judgments in decision-making can be faulty, and that breakdowns can occur because of simple error or mistake. Additionally, controls can be circumvented by the individual acts of some persons, by collusion of two or more people, or by an unauthorized override of the controls. Accordingly, because of the inherent limitations in our control system, misstatements or insufficient disclosures due to error or fraud may occur and not be detected.

## Special note regarding forward-looking statements

*This prospectus contains forward-looking statements that involve substantial risks and uncertainties. The forward-looking statements are contained principally in the sections titled “Prospectus Summary,” “Risk Factors,” “Management’s Discussion and Analysis of Financial Condition and Results of Operations,” “Business” and elsewhere in this prospectus. In some cases, you can identify forward-looking statements by terms such as “may,” “will,” “should,” “expect,” “plan,” “anticipate,” “could,” “intend,” “target,” “project,” “estimate,” “believe,” “estimate,” “predict,” “potential” or “continue” or the negative of these terms or other similar expressions intended to identify statements about the future. These statements speak only as of the date of this prospectus and involve known and unknown risks, uncertainties and other important factors that may cause our actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements. We have based these forward-looking statements largely on our current expectations and projections about future events and financial trends that we believe may affect our business, financial condition and results of operations. These forward-looking statements include, without limitation, statements about the following:*

- the timing, progress and results of our preclinical studies and clinical trials of our product candidates, including statements regarding the timing of initiation and completion of studies or trials and related preparatory work, the period during which the results of the trials will become available and our research and development programs;
- the timing of our planned IND submissions, initiation of planned clinical trials and timing of expected clinical results for LX2006, LX1001, LX2020 and our other future product candidates;
- the timing of any submission of filings for regulatory approval of, and our ability to obtain and maintain regulatory approvals for LX2006, LX1001, LX2020 and any other product candidates;
- the impact of public health crises (such as COVID-19) and other adverse global economic conditions on our operations and the potential disruption in the operations and business of third-party manufacturers, CROs, other service providers, and collaborators with whom we conduct business;
- our ability to identify patients with the diseases treated by our product candidates, and to enroll patients in trials;
- our expectations regarding the size of the patient populations, market acceptance and opportunity for and clinical utility of our product candidates, if approved for commercial use;
- our manufacturing capabilities and strategy, including the scalability and commercial viability of our manufacturing methods and processes;
- our reliance on third party manufacturing partners to comply with significant regulations with respect to manufacturing our products;
- our expectations regarding the scope of any approved indication for LX2006, LX1001, LX2020 or any other product candidate;
- our ability to successfully commercialize our product candidates, if approved;
- our ability to leverage our platform to identify and develop future product candidates;
- our estimates of our expenses, ongoing losses, future revenue, capital requirements and our need for or ability to obtain additional funding before we can expect to generate any revenue from product sales;

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- our ability to establish or maintain collaborations or strategic relationships;
- our ability to identify, recruit and retain key personnel;
- our reliance upon intellectual property licensed from third parties and our ability to obtain such licenses on commercially reasonable terms or at all;
- our ability to protect and enforce our intellectual property position for our product candidates, and the scope of such protection;
- our financial performance;
- our anticipated use of proceeds from this offering;
- our competitive position and the development of and projections relating to our competitors or our industry;
- our estimates regarding future revenue, expenses and needs for additional financing;
- the impact of laws and regulations; and
- our expectations regarding the time during which we will be an emerging growth company under the JOBS Act.

The foregoing list of risks is not exhaustive. Other sections of this prospectus may include additional factors that could harm our business and financial performance. Moreover, we operate in an evolving environment. New risk factors and uncertainties may emerge from time to time, and it is not possible for management to predict all risk factors and uncertainties. As a result of these factors, we cannot assure you that the forward-looking statements in this prospectus will prove to be accurate. Except as required by applicable law, we do not plan to publicly update or revise any forward-looking statements contained herein, whether as a result of any new information, future events, changed circumstances or otherwise. You should, however, review the factors and risks and other information we describe in the reports we will file from time to time with the SEC after the date of this prospectus.

Because forward-looking statements are inherently subject to risks and uncertainties, some of which cannot be predicted or quantified and some of which are beyond our control, you should not rely on these forward-looking statements as predictions of future events. Although we believe that we have a reasonable basis for each forward-looking statement contained in this prospectus, the events and circumstances reflected in our forward-looking statements may not be achieved or occur and actual results could differ materially from those projected in the forward-looking statements. You should refer to the “Risk Factors” section of this prospectus for a discussion of important factors that may cause our actual results to differ materially from those expressed or implied by our forward-looking statements.

You should read this prospectus and the documents that we reference in this prospectus and have filed as exhibits to the registration statement, of which this prospectus is a part, completely and with the understanding that our actual future results may be materially different from what we expect. We qualify all of our forward-looking statements by these cautionary statements.

## **Market and industry data**

We obtained the industry, statistical and market data in this prospectus from our own internal estimates and research as well as from industry and general publications and research, surveys and studies conducted by third parties. All of the market data used in this prospectus involve a number of assumptions and limitations, and the sources of such data cannot guarantee the accuracy or completeness of such information. While management is responsible for the accuracy of such data and believes that each of these studies and publications is reliable, the industry in which we operate is subject to a high degree of uncertainty and risk due to a variety of important factors, including those described in the section titled “Risk Factors.” These and other factors could cause results to differ materially from those expressed in the estimates made by third parties and by us.

## Use of proceeds

We estimate that the net proceeds from our issuance and sale of \_\_\_\_\_ shares of our common stock in this offering will be approximately \$ \_\_\_\_\_ million (or approximately \$ \_\_\_\_\_ million if the underwriters exercise in full their option to purchase up to \_\_\_\_\_ additional shares), assuming an initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, and after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us.

Each \$1.00 increase (decrease) in the assumed initial public offering price of \$ \_\_\_\_\_ per share would increase (decrease) the net proceeds to us from this offering by approximately \$ \_\_\_\_\_ million, assuming the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us. Each increase (decrease) of 1.0 million in the number of shares we are offering would increase (decrease) the net proceeds to us from this offering, after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us, by approximately \$ \_\_\_\_\_ million, assuming the assumed initial public offering price stays the same. We do not expect that a change in the offering price or the number of shares by these amounts would have a material effect on our intended uses of the net proceeds from this offering, although it may impact the amount of time prior to which we may need to seek additional capital.

As of March 31, 2023, we had a cash and cash equivalents balance of \$58.9 million. We intend to use the net proceeds from this offering, together with our existing cash, as follows:

- approximately \$ \_\_\_\_\_ million to fund our ongoing and planned clinical development of LX2006 for the treatment of FA cardiomyopathy, including \_\_\_\_\_ ;
- approximately \$ \_\_\_\_\_ million to fund our ongoing and planned clinical development of LX1001 for the treatment of Alzheimer’s disease in *APOE4* homozygous patients, including \_\_\_\_\_ ;
- approximately \$ \_\_\_\_\_ million to fund our ongoing and planned clinical development of LX2020 for the treatment of ARVC due to mutations in the *PKP2* gene, including \_\_\_\_\_ ;
- approximately \$ \_\_\_\_\_ million to fund the continued development of our other programs and cardiac discovery efforts, including \_\_\_\_\_ ; and
- the remainder for working capital and other general corporate purposes.

Based on our current operational plans and assumptions, we expect our cash, together with the net proceeds from this offering, will be sufficient to fund our operations through \_\_\_\_\_. However, we do not expect these funds will be sufficient to complete the clinical development of, or commercialize, any of our product candidates or programs. We have based these estimates on assumptions that may prove to be wrong, and we could use our available capital resources sooner than we expect.

We believe opportunities may exist from time to time to expand our current business through acquisitions of complementary companies, products or technologies. While we have no current agreements, commitments or understandings for any specific acquisitions at this time, we may also use a portion of the net proceeds for these purposes.

This anticipated use of our existing cash and net proceeds from this offering represents our intentions based upon our current plans and business conditions, which could change in the future as our plans and business conditions evolve. The amounts and timing of our actual expenditures may vary significantly depending on numerous factors, including the progress of our development, the status of and results from clinical trials and the timing and

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outcome of any regulatory submissions, as well as any collaborations that we may enter into with third parties for our drug candidates, and any unforeseen cash needs. As a result, our management will have broad discretion in the application of the net proceeds from this offering, and investors will be relying on the judgment of our management regarding the application of those net proceeds. Pending our use of the net proceeds from this offering, we intend to invest the net proceeds in a variety of capital preservation investments, including short-term, investment grade, interest-bearing instruments and U.S. government securities.



## **Dividend policy**

We have never declared or paid, and do not anticipate declaring or paying, in the foreseeable future, any cash dividends on our capital stock. We currently intend to retain all available funds and any future earnings to support our operations and finance the growth and development of our business. Any future determination related to our dividend policy will be made at the discretion of our board of directors and will depend upon, among other factors, our results of operations, financial condition, capital requirements, contractual restrictions, business prospects and other factors our board of directors may deem relevant.

## Capitalization

The following table sets forth our cash and our capitalization as of March 31, 2023:

- on an actual basis;
- on a pro forma basis to give effect to: (1) the conversion of all outstanding shares of our preferred stock into an aggregate of \_\_\_\_\_ shares of our common stock; and (2) the filing and effectiveness of our amended and restated certificate of incorporation upon the closing of this offering; and
- on a pro forma as adjusted basis to give effect to: (i) the pro forma adjustments described above; and (ii) our issuance and sale of \_\_\_\_\_ shares of common stock in this offering at an assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, after deducting the estimated underwriting discounts and estimated offering expenses payable by us.

This pro forma as adjusted information is illustrative only and will depend on the actual initial public offering price and other terms of this offering determined at pricing. Our capitalization following the closing of this offering will depend on the actual initial public offering price and other terms of this offering determined at pricing. You should read this information in conjunction with our financial statements and the related notes appearing at the end of this prospectus, the section of this prospectus titled “Management’s Discussion and Analysis of Financial Condition and Results of Operations” and other financial information contained in this prospectus.

	March 31, 2023		
	Actual	Pro Forma	Pro Forma As Adjusted
Series A convertible preferred stock, \$0.0001 par value; 85,495,722 shares authorized, issued, and outstanding as of March 31, 2023; aggregate liquidation value of \$85,495,722 as of March 31, 2023	\$ 85,268,513		
Series B convertible preferred stock, \$0.0001 par value; 58,157,824 shares authorized as of March 31, 2023; 58,157,823 shares issued and outstanding as of March 31, 2023; aggregate liquidation value of \$100,059,953 as of March 31, 2023	99,764,986		
<b>Stockholders’ (deficit) equity:</b>			
Common stock, \$0.0001 par value, 192,020,973 shares authorized as of March 31, 2023; 17,482,991 shares issued and 17,099,158 shares outstanding as of March 31, 2023	1,710		
Additional paid-in capital	2,920,353		
Accumulated deficit	(134,103,747)		
<b>Total stockholders’ (deficit) equity</b>	<b>(131,181,684)</b>		
<b>Total capitalization</b>	<b>\$ 53,851,815</b>	<b>\$</b>	<b>\$</b>

Each \$1.00 increase (decrease) in the assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, would increase (decrease) the pro forma as adjusted amount of each of cash, additional paid in capital, total stockholders’ equity and total capitalization by approximately \$ \_\_\_\_\_ million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting

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discounts and commissions and estimated offering expenses payable by us. Similarly, each increase (decrease) of 1.0 million shares in the number of shares offered by us at the assumed initial public offering price per share of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, would increase (decrease) the pro forma as adjusted amount of each of cash, additional paid in capital, total stockholders' equity and total capitalization by approximately \$ \_\_\_\_\_ million.

The number of shares of our common stock outstanding as of March 31, 2023 in the table above excludes:

- \_\_\_\_\_ shares of our common stock issuable upon the exercise of options under the Existing Plan, granted subsequent to March 31, 2023, at a weighted-average exercise price of \$ \_\_\_\_\_ per share;
- \_\_\_\_\_ shares of our common stock reserved for future issuance under the Existing Plan, which shares will cease to be available for issuance at the time the 2023 Plan becomes effective and will be added to, and become available for issuance under, the 2023 Plan;
- \_\_\_\_\_ shares of our common stock reserved for future issuance under the 2023 Plan, which will become effective in connection with this offering, as well as any automatic increases in the number of shares of common stock reserved for future issuance under the 2023 Plan; and
- \_\_\_\_\_ shares of our common stock reserved for future issuance under the 2023 Employee Stock Purchase Plan, or ESPP, which will become effective in connection with this offering, as well as any automatic increases in the number of shares of common stock reserved for future issuance under the ESPP.

## Dilution

If you invest in our common stock in this offering, your ownership interest will be immediately diluted to the extent of the difference between the initial public offering price per share and the pro forma as adjusted net tangible book value per share of our common stock after this offering.

As of March 31, 2023, we had a historical net tangible book value (deficit) of \$ \_\_\_\_\_ million, or \$ \_\_\_\_\_ per share of common stock. Our historical net tangible book value (deficit) per share represents total tangible assets less total liabilities, divided by the number of shares of our common stock outstanding as of March 31, 2023.

Our pro forma net tangible book value (deficit) as of March 31, 2023 was \$ \_\_\_\_\_ million, or \$ \_\_\_\_\_ per share of our common stock. Pro forma net tangible book value represents the amount of our total tangible assets less our total liabilities, after giving effect to the conversion of all outstanding shares of our convertible preferred stock into an aggregate of \_\_\_\_\_ shares of our common stock, as if such conversion had occurred on March 31, 2023. Pro forma net tangible book value per share represents pro forma net tangible book value divided by the total number of shares outstanding as of March 31, 2023, after giving effect to the pro forma adjustments described above.

After giving further effect to the sale of \_\_\_\_\_ shares of common stock in this offering at an assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, and after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us, our pro forma as adjusted net tangible book value as of March 31, 2023 would have been \$ \_\_\_\_\_ million, or \$ \_\_\_\_\_ per share. This amount represents an immediate increase in pro forma net tangible book value of \$ \_\_\_\_\_ per share to our existing stockholders and immediate dilution of \$ \_\_\_\_\_ per share to new investors in this offering. We determine dilution by subtracting the pro forma as adjusted net tangible book value per share after this offering from the amount of cash that a new investor paid for a share of common stock in this offering.

The following table illustrates this dilution:

Assumed initial public offering price per share	\$
Historical net tangible book value (deficit) per share as of March 31, 2023	\$
Pro forma increase in net tangible book value (deficit) per share attributable to the pro forma transactions described above	_____
Pro forma net tangible book value (deficit) per share as of March 31, 2023	_____
Increase in pro forma as adjusted net tangible book value per share attributable to new investors participating in this offering	_____
Pro forma as adjusted net tangible book value per share after this offering	_____
Dilution per share to new investors participating in this offering	\$

The dilution information discussed above is illustrative only and will change based on the actual initial public offering price and other terms of this offering determined at pricing. Each \$1.00 increase (decrease) in the assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, would increase (decrease) the pro forma as adjusted net tangible book value per share after this offering by approximately \$ \_\_\_\_\_, and dilution in pro forma as adjusted net tangible book value per share to new investors by approximately \$ \_\_\_\_\_, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting

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discounts and commissions and estimated offering expenses payable by us. An increase of 1.0 million shares in the number of shares we are offering would increase the pro forma as adjusted net tangible book value per share after this offering by \$ \_\_\_\_\_ and decrease the dilution per share to new investors participating in this offering by \$ \_\_\_\_\_, assuming no change in the assumed initial public offering price per share and after deducting the estimated underwriting discounts and commissions. A decrease of 1.0 million shares in the number of shares we are offering would decrease the pro forma as adjusted net tangible book value per share after this offering by \$ \_\_\_\_\_ and increase the dilution per share to new investors participating in this offering by \$ \_\_\_\_\_, assuming no change in the assumed initial public offering price per share and after deducting the estimated underwriting discounts and commissions.

If the underwriters exercise their option to purchase additional shares of our common stock in full, the pro forma as adjusted net tangible book value after this offering would be \$ \_\_\_\_\_ per share, the increase in pro forma as adjusted net tangible book value per share would be \$ \_\_\_\_\_ and the dilution per share to new investors would be \$ \_\_\_\_\_ per share, in each case assuming an initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus.

The following table summarizes, as of March 31, 2023, on the pro forma as adjusted basis described above, the differences between the number of shares purchased from us, the total consideration paid to us in cash and the average price per share that existing stockholders and new investors paid for such shares. The calculation below is based on an assumed initial public offering price of \$ \_\_\_\_\_ per share, which is the midpoint of the price range set forth on the cover page of this prospectus, before deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us.

	Total Shares		Total Consideration		Weighted-Average Price Per Share
	Number	Percent	Amount	Percent	
Existing stockholders		%	\$	%	\$
New investors					\$
Total		100%		\$ 100%	

The foregoing tables and calculations are based on the number of shares of our common stock outstanding as of March 31, 2023, after giving effect to the conversion of all outstanding shares of our convertible preferred stock into an aggregate of \_\_\_\_\_ shares of common stock, and excludes:

- \_\_\_\_\_ shares of our common stock issuable upon the exercise of options under the Existing Plan, granted subsequent to March 31, 2023, at a weighted-average exercise price of \$ \_\_\_\_\_ per share;
- \_\_\_\_\_ shares of our common stock reserved for future issuance as of March 31, 2023 under the Existing Plan, which shares will cease to be available for issuance at the time the 2023 Plan becomes effective and will be added to, and become available for issuance under, the 2023 Plan;
- \_\_\_\_\_ shares of our common stock reserved for future issuance under the 2023 Plan, which will become effective in connection with this offering, as well as any automatic increases in the number of shares of common stock reserved for future issuance under the 2023 Plan; and
- \_\_\_\_\_ shares of our common stock reserved for future issuance under the ESPP, which will become effective in connection with this offering, as well as any automatic increases in the number of shares of common stock reserved for future issuance under the 2023 ESPP.

To the extent that stock options are exercised, new stock options are issued under our equity incentive plan or we issue additional shares of common stock in the future, there will be further dilution to investors participating in

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this offering. In addition, we may choose to raise additional capital because of market conditions or strategic considerations, even if we believe that we have sufficient funds for our current or future operating plans. If we raise additional capital through the sale of equity or convertible debt securities, the issuance of these securities could result in further dilution to our stockholders.

## Management's discussion and analysis of financial condition and results of operations

*You should read the following discussion and analysis of our financial condition and results of operations together with our consolidated financial statements and related notes included elsewhere in this prospectus. Some of the information contained in this discussion and analysis or set forth elsewhere in this prospectus, including information with respect to our plans and strategy for our business and related financing includes forward-looking statements that involve risks and uncertainties. Many factors, including those factors set forth in the "Risk Factors" section of this prospectus, may materially and adversely affect our actual results, which may differ materially from the results described in or implied by the forward-looking statements contained in the following discussion and analysis.*

### Overview

We are a clinical-stage gene therapy company focused on addressing some of the most devastating genetically defined cardiovascular diseases and a genetically defined sub-group of Alzheimer's disease. We are advancing a deep and diverse pipeline of AAV-based gene therapy candidates for diseases affecting both larger-rare and prevalent patient populations. Utilizing a modular approach that integrates a clinically validated vector, we have a disease area strategy targeting defined patient sub-groups most likely to benefit from our gene therapy candidates, and high-quality, scalable manufacturing, which is designed to overcome many of the challenges facing the field of gene therapy.

Since our inception, we have devoted substantially all of our efforts to organizing and staffing our company, research and development activities, business planning, raising capital, building our intellectual property portfolio and providing general and administrative support for these operations. To date, we have funded our operations primarily through proceeds from the sale of shares of our convertible preferred stock. As of March 31, 2023, we had \$58.9 million of cash and cash equivalents and we had raised aggregate gross proceeds of \$183.7 million from the sale of convertible equity securities.

We have incurred significant operating losses since the commencement of our operations. Our ability to generate product revenue sufficient to achieve profitability will depend heavily on the successful development and eventual commercialization of one or more of our current gene therapy candidates or any future gene therapy candidates. Our net losses for the three months ended March 31, 2023 and for the year ended December 31, 2022 were \$18.7 million and \$59.3 million, respectively, and our accumulated deficit was \$134.1 million at March 31, 2023. We expect to continue to incur significant losses for the foreseeable future as we advance our current and future product candidates through preclinical and clinical development, continue to build our operations and transition to operating as a public company.

We expect to continue to incur net operating losses for at least the next several years, and we expect our research and development expenses, general and administrative expenses, and capital expenditures to continue to increase. We expect our expenses and capital requirements will increase significantly in connection with our ongoing activities as we:

- continue our ongoing and planned clinical trials as well as research and development of our Friedrich's ataxia cardiomyopathy, or FA cardiomyopathy (LX2006), APOE-associated Alzheimer's disease (LX1001), and arrhythmogenic right ventricular cardiomyopathy, or ARVC (LX2020) programs and other product candidates;
- initiate preclinical studies and clinical trials for any additional product candidates that we may pursue in the future;
- seek to discover and develop additional product candidates and further expand our clinical product pipeline;

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- seek regulatory approvals for any product candidates that successfully complete clinical trials;
- invest in capital equipment in order to expand our research and development and manufacturing activities;
- attract, hire and retain additional clinical, scientific, quality control, regulatory, and manufacturing management and administrative personnel;
- add clinical, operational, financial and management information systems and personnel, including personnel to support our product development;
- develop, maintain, expand, protect and enforce our intellectual property portfolio, including patents, trade secrets and know-how;
- acquire or in-license other product candidates and technologies;
- expand our operations in the United States and to other geographies;
- incur additional legal, accounting, investor relations and other general and administrative expenses associated with operating as a public company; and
- establish a sales, marketing and distribution infrastructure, either ourselves or in partnership with others, to commercialize any product candidates, if approved, and related additional commercial manufacturing costs.

We will not generate revenue from product sales unless and until we successfully complete clinical development and obtain regulatory approval for one or more of our product candidates. If we obtain regulatory approval for any of our product candidates, we expect to incur significant expenses related to product sales, marketing and distribution to the extent that such sales, marketing and distribution are not the responsibility of potential collaborators. We will require substantial additional funding to develop our product candidates and support our continuing operations. We may also require additional capital to pursue in-licenses or acquisitions of other product candidates. We may also require additional capital if we choose to pursue an in-house manufacturing strategy. Further, following the completion of this offering we expect to incur additional costs associated with operating as a public company. We also expect to increase the size of our administrative function to support the growth of our business. Our net losses may fluctuate significantly from quarter-to-quarter and year-to-year, depending on the timing of our clinical trials and our expenses related to other research and development activities.

Until such time that we can generate significant revenue from product sales or other sources, we expect to finance our operations through the sale of equity, debt financings or other capital sources, which could include proceeds from potential collaborations, strategic partnerships or marketing, distribution, licensing or other strategic arrangements with third parties, or from grants. We may be unable to raise additional funds or to enter into such agreements or arrangements on favorable terms, or at all. If we fail to raise capital or enter into such agreements as, and when, needed, we may have to significantly delay, scale back or discontinue the development or commercialization of our product candidates or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

Our failure to obtain sufficient funds with acceptable terms could have a material adverse effect on our business, results of operations or financial condition, including requiring us to have to delay, reduce or eliminate our product development or future commercialization efforts. Because of the numerous risks and uncertainties associated with pharmaceutical product development, we are unable to accurately predict the amount of increased expenses or timing, or if we will be able to achieve or maintain profitability. If we fail to become profitable or are unable to sustain profitability on a continuing basis, then we may be unable to continue our operations at planned levels and be forced to reduce or terminate our operations. We cannot provide assurance that we will ever be profitable or generate positive cash flow from operating activities.



## Summary of key factors impacting the comparability of our performance

### *Stelios Therapeutics Inc. acquisition*

On July 16, 2021, we purchased all of the issued and outstanding capital stock of Stelios Therapeutics, Inc., or Stelios, for initial cash consideration of \$7.0 million, with payments of up to an additional \$20.5 million due upon the achievement of the following milestones: clinical candidate selection for either the ARVC or the TNNI3-associated programs; first patient dosed in a Phase 1 clinical trial for any product in our CX43 program; first patient dosed in a Phase 1 clinical trial for any product in our PKP2 program; or first patient dosed in a Phase 1 clinical trial for any candidate in our TNNI3 program. In the third quarter of 2022, a development milestone associated with a clinical candidate selection was achieved resulting in a \$2.0 million payment to the selling shareholders of Stelios.

We accounted for the acquisition of Stelios as an asset acquisition pursuant to Financial Accounting Standards Board, or FASB, Accounting Standards Codification, or ASC, Section 805, *Business Combinations*. We acquired in-process research and development, or IPR&D, assets from Stelios related to ARVC and TNNI3-associated hypertrophic cardiomyopathy programs. Such programs relate to our development efforts for LX2020, LX2021 and LX2022. The fair value of the IPR&D acquired of \$7.0 million was charged to research and development expense as it had no alternative future use at the time of the acquisition. Stelios was merged into Lexeo Therapeutics, Inc. on December 15, 2022 and ceased to exist.

See Note 3 to the audited consolidated financial statements appearing elsewhere in this prospectus for more information on the Stelios acquisition.

### *Lease agreement*

In February 2022, we entered into a lease agreement for our New York headquarters located at 345 Park Avenue South, 6<sup>th</sup> Floor, New York, NY 10010, pursuant to which we lease a total of approximately 15,839 square feet of office and laboratory space. The lease began in April 2022 and ends in July 2029 with an additional five-year option to extend at the then prevailing effective market rental rate. We accounted for this lease pursuant to FASB ASC Topic 842, *Leases*. Upon commencement of this lease, we recorded operating lease right-of-use assets and operating lease liabilities of approximately \$11.6 million based on the present value of payments over the lease term using an estimated incremental borrowing rate of 8.53%. In connection with this lease, we provided a security deposit to the landlord in the form of a letter of credit totaling \$1.2 million.

See Note 5 to the audited consolidated financial statements appearing elsewhere in this prospectus for more information on this lease.

## Components of our results of operations

### *Revenue*

Our revenue to date has been comprised of grant revenue, which are amounts earned from performing contracted research and development services. These grants generally require us to meet certain research milestones in order to receive the funds. To date, we have not generated any revenue from product sales. If our development efforts for LX2006, LX1001, and LX2020 or any future product candidates, are successful and result in regulatory approval, or if we enter into collaboration or license agreements with third parties, we may generate revenue in the future from product sales, royalties or payments from such collaboration or license agreements, or a combination of product sales and payments from such agreements.

## ***Operating expenses***

### *Research and development*

Research and development expenses consist of costs incurred for our research activities, including our discovery efforts and the preclinical and clinical development of our programs. These expenses include:

- employee-related expenses, including salaries, benefits, and stock-based compensation expense for employees engaged in research and development functions;
- expenses incurred under agreements with third parties, such as consultants, clinical investigators, contractors and clinical research organizations, or CROs, that assist with (i) identification of potential product candidates in discovery platforms and (ii) the preclinical and clinical studies of our product candidates;
- the cost of developing and scaling our manufacturing process and manufacturing product candidates for use in our research, preclinical studies and clinical trials, including under agreements with third parties, such as consultants, contractors and CMOs;
- costs to maintain compliance with FDA and other regulatory requirements;
- laboratory supplies and research materials;
- facilities, depreciation, and other expenses, which include direct and allocated expenses for rent and maintenance of facilities;
- payments made under our licensing agreements with third parties, including milestone payments; and
- other expenses incurred as a result of research and development activities.

We expense research and development costs as incurred. Nonrefundable advance payments for goods and services that will be used in future research and development activities are expensed when the activity has been performed or when the goods have been received rather than when the payment is made. When third-party service providers' billing terms do not coincide with our period-end, we are required to make estimates of our obligations to those third parties incurred in a given accounting period and record accruals at the end of the period. We base these estimates on our knowledge of the research and development programs, services performed for the period, past history for related activities and the expected duration of the third-party service contract, where applicable. If timelines or contracts are modified based upon changes in the scope of work to be performed, we modify our estimates of accrued expenses accordingly on a prospective basis; therefore, actual results could differ from our estimates. Upfront payments under license agreements are expensed upon receipt of the license, and annual maintenance fees under license agreements are expensed in the period in which they are incurred. Milestone payments under license agreements are accrued, with a corresponding expense being recognized, in the period in which the milestone is determined to be probable of achievement and the related amount is reasonably estimable.

Our direct research and development expenses are tracked on a program-by-program basis and consist primarily of external costs, such as fees paid to CROs, CMOs, central laboratories and certain outside consultants in connection with our research and discovery, preclinical development, process development, manufacturing, clinical development, clinical trials, regulatory and quality assurance activities. We do not allocate professional services costs and licensing fees and other similar costs to specific programs because these costs are deployed across multiple programs.

Research and development activities are central to our business model and account for a significant portion of our operating expenses. Product candidates in later stages of clinical development generally have higher

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development costs than those in earlier stages of clinical development, primarily due to the increased size and duration of later-stage clinical trials. As a result, we expect our research and development expenses to increase substantially in absolute dollars for the foreseeable future as we and further advance our LX2006, LX2020, and APOE4-associated Alzheimer's disease programs and any other future product candidates that we may develop, into and through preclinical studies and clinical trials and pursue regulatory approvals. We cannot determine with certainty the timing of initiation, the duration or the completion costs of current or future preclinical studies and clinical trials of our product candidates due to the inherently unpredictable nature of preclinical and clinical development. Clinical and preclinical development timelines, the probability of success and development costs can differ materially from expectations. We anticipate that we will make determinations as to which product candidates to pursue and how much funding to direct to each product candidate on an ongoing basis in response to the results of ongoing and future research, preclinical studies and clinical trials, regulatory developments and our assessments as to each product candidate's commercial potential. In addition, we cannot forecast whether any of our current or future product candidates will be subject to future collaborations, when such arrangements will be secured, if at all, and to what degree such arrangements would affect our development plans and capital requirements. We are also unable to predict when, if ever, we will generate revenue from our product candidates to offset our expenses. Our future expenses may vary significantly each period based on factors such as:

- the timing and progress of preclinical and clinical development activities;
- the number and scope of preclinical and clinical programs we decide to pursue;
- our ability to maintain our current research and development programs and to establish new ones;
- establishing an appropriate safety profile with investigational new IND-enabling toxicology studies;
- successful patient enrollment in, and the initiation and completion of, our ongoing and planned clinical trials;
- the successful completion of clinical trials with safety, tolerability and efficacy profiles that are satisfactory to the FDA or any comparable foreign regulatory authority;
- receipt of regulatory approvals from applicable regulatory authorities;
- the timing, receipt and terms of any marketing approvals from applicable regulatory authorities;
- our ability to establish licensing or collaboration arrangements;
- the performance of our current and potential future collaborators;
- establishing commercial manufacturing capabilities or making arrangements with third-party manufacturers;
- development and timely delivery of commercial-grade therapeutics that can be used in our planned clinical trials and for commercial launch;
- commercializing our product candidates, if approved, whether alone or in collaboration with others;
- obtaining, maintaining, defending and enforcing patent claims and other intellectual property rights;
- maintaining a continued acceptable safety profiles of our therapeutics following approval; and
- hiring and retaining key research and development personnel.

Any changes in the outcome of any of these variables could mean a significant change in the costs and timing associated with the development of our product candidates. We may never obtain regulatory approval for any of our product candidates, and, even if we do, drug commercialization takes several years and involves substantial cost.

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### *General and administrative*

General and administrative expenses consist primarily of personnel expenses, including salaries, benefits and stock-based compensation expense, for personnel in executive, accounting, business development, legal, human resources and administrative functions. General and administrative expenses also include corporate facility costs not otherwise included in research and development expenses, depreciation and other expenses, which include direct or allocated expenses for rent and maintenance of facilities and insurance, not otherwise included in research and development expenses, as well as professional fees for legal, consulting, investor and public relations, accounting and audit services.

We expect that our general and administrative expenses will increase significantly in the near-to-medium term as we incur additional expenses associated with operating as a public company, including increased expenses for insurance premiums and audit, legal, regulatory, and tax-related services associated with maintaining compliance with the rules and regulations of the Securities and Exchange Commission, or SEC, and standards applicable to companies listed on a national securities exchange, investor relations activities and other administrative and professional services. We also plan to increase our headcount to support the continued research and development of our programs and the growth of our business.

### ***Other income (expense)***

Other income (expense) includes net foreign exchange gains and (losses).

### ***Interest expense***

Interest expense is primarily associated with our finance right of use asset equipment leases.

### ***Interest income***

Interest income is related to interest earned from our investments in U.S. government money market funds.

### **Income taxes**

Provision for income taxes consists of U.S. federal and state income taxes in which we conduct business. Since our inception, we have not recorded any income tax benefits for the net losses we have incurred in each year or for our research and development tax credits, as we believe, based upon the weight of available evidence, that it is more likely than not that all of our net operating loss, or NOL, carryforwards and tax credits will not be realized. Accordingly, we have recorded a full valuation allowance against our net deferred tax assets at March 31, 2023, December 31, 2022, March 31, 2022 and December 31, 2021. As of March 31, 2023, December 31, 2022, March 31, 2022 and December 31, 2021, we had no unrecognized tax benefits.

## Results of operations

### Comparison of the three months ended March 31, 2023 and 2022

The following table summarizes our results of operations for the three months ended March 31, 2023 and 2022:

	Three Months Ended March 31,		
	2023	2022	Change
Revenue			
Grant revenue	\$ —	\$ 85,550	\$ (85,550)
Total revenue	—	85,550	(85,550)
Operating expenses			
Research and development	16,438,110	10,146,371	6,291,739
General and administrative	2,852,379	2,458,438	393,941
Total operating expenses	19,290,489	12,604,809	6,685,680
Operating loss	(19,290,489)	(12,519,259)	(6,771,230)
Other income (expense)			
Other income (expense)	(4,521)	146	(4,667)
Interest expense	(49,967)	—	(49,967)
Interest income	687,495	17,283	670,212
Total other income (expense)	633,007	17,429	615,578
Loss from operations before income taxes	(18,657,482)	(12,501,830)	(6,155,652)
Income taxes	—	—	—
Net loss and comprehensive loss	\$ (18,657,482)	\$ (12,501,830)	\$ (6,155,652)

#### Grant revenue

Grant revenue was \$0.1 million for the three months ended March 31, 2022. We did not record any grant revenue for the three months ended March 31, 2023. The decrease of \$0.1 million was due to the wind-down of the underlying grants, which were closed-out by the third quarter of 2022.

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### *Research and development expenses*

The following table summarizes our research and development expenses incurred for the three months ended March 31, 2023 and 2022:

	Three Months Ended March 31,		
	2023	2022	Change
<b>Direct external research and development expenses by program:</b>			
LX2020	\$ 3,613,658	\$ 438,143	\$ 3,175,515
LX1001	2,589,978	1,017,642	1,572,336
LX2006	4,410,066	2,188,310	2,221,756
LX1004	141,278	1,028,204	(886,926)
Other programs	856,804	1,563,781	(706,977)
Total direct external research and development expenses by program	\$ 11,611,784	\$ 6,236,080	\$ 5,375,704
<b>Unallocated research and development expenses:</b>			
Employee compensation and stock-based compensation expense	\$ 2,808,505	\$ 1,944,215	\$ 864,290
Professional fees	581,917	216,455	365,462
Lab-related costs and supplies	285,484	50,760	234,724
License fees	1,000	60,577	(59,577)
Other unallocated	1,149,420	1,638,284	(488,864)
Total unallocated research and development expenses:	4,826,326	3,910,291	916,035
<b>Total research and development expenses</b>	<b>\$ 16,438,110</b>	<b>\$ 10,146,371</b>	<b>\$ 6,291,739</b>

Research and development expenses were \$10.1 million for the three months ended March 31, 2022 compared to \$16.4 million for the three months ended March 31, 2023. The increase of \$6.3 million was primarily due to a net increase of \$5.4 million in direct external research and development program expenses and a \$0.9 million net increase in unallocated research and development expenses.

### *Direct external research and development expenses*

The net increase of \$5.4 million in direct external research and development expenses was primarily due to an increase in ARVC program (LX2020) costs of \$3.2 million during the three months ended March 31, 2023, which consisted mostly of (i) CMC expenses of \$1.7 million related to the development of our manufacturing processes at third-party vendors, and (ii) non-clinical and pre-clinical expenses of \$1.3 million related to the advancement of our LX2020 program into IND-enabling studies. We did not incur significant direct external research and development expenses for our LX2020 program during the three months ended March 31, 2022.

The increase of \$1.6 million in direct research and development expenses for our APOE4-associated Alzheimer's disease product candidate (LX1001) was primarily due to increases in clinical expenses of \$2.1 million related to the ongoing Phase 1/2 clinical trial partially offset by a decrease in CMC expenses of \$0.5 million primarily related to third-party GMP batch costs.

The increase of \$2.2 million in direct external research and development expenses for our FA cardiomyopathy product candidate (LX2006) was primarily due to the achievement of a development milestone in the first quarter of 2023 due to Adverum of \$3.5 million, partially offset by a decrease in CMC expenses of \$1.4 million primarily related to third-party GMP batch costs.

The decrease of \$0.9 million in direct external research and development expenses for our CLN2 Batten disease (product candidate (LX1004) was primarily due to decreases in CMC expenses of \$0.7 million and non-clinical expenses of \$0.2 million.

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Direct external research and development expenses for our other programs decreased by \$0.7 million during the three months ended March 31, 2023, which was primarily attributable to a decrease in pre-clinical and non-clinical expenses for our early stage cardiovascular and central nervous system disease programs.

### *Unallocated research and development expenses*

The net increase of \$0.9 million in unallocated research and development expenses was primarily due to (i) increased employee compensation and stock-based compensation expense of \$0.9 million due to increased headcount, (ii) \$0.4 million related to increased professional fees and consulting expenses, and (iii) \$0.2 million related to lab-related costs and supplies associated with our New York City office and laboratory lease that commenced in April 2022, which were partially offset by a decrease of \$0.5 million related to non-allocable expenses.

### *General and administrative expenses*

General and administrative expenses were \$2.5 million for the three months ended March 31, 2022 compared to \$2.9 million for the three months ended March 31, 2023. The increase of \$0.4 million was primarily due to increases in (i) employee compensation and stock-based compensation expense of \$0.6 million due to increased headcount and (ii) rent and facility expenses of \$0.3 million as our New York City office and laboratory lease commenced in April 2022, which were partially offset by a decrease in accounting consulting expenses of \$0.3 million.

### *Interest income*

We recognized interest income of \$0.7 million for the three months ended March 31, 2023 related to interest earned on our investments in U.S. government money market funds.

**Comparison of the years ended December 31, 2022 and 2021**

The following table summarizes our results of operations for the years ended December 31, 2022 and 2021:

	Years Ended December 31,		
	2022	2021	Change
Revenue			
Grant revenue	\$ 653,956	\$ 1,657,122	\$ (1,003,166)
Total revenue	653,956	1,657,122	(1,003,166)
Operating expenses			
Research and development	49,161,540	45,120,874	4,040,666
General and administrative	12,001,097	7,173,026	4,828,071
Total operating expenses	61,162,637	52,293,900	8,868,737
Operating loss	(60,508,681)	(50,636,778)	(9,871,903)
Other income (expense)			
Other income (expense)	(1,852)	(155)	1,697
Interest expense	(90,874)	—	90,874
Interest income	1,324,680	15,220	1,309,461
Total other income and expense	1,231,954	15,065	1,216,889
Loss from operations before income taxes	(59,276,727)	(50,621,713)	(8,655,014)
Income taxes	—	—	—
Net loss and comprehensive loss	\$ (59,276,727)	\$ (50,621,713)	\$ (8,655,014)

*Grant revenue*

Grant revenue was \$1.7 million for the year ended December 31, 2021 compared to \$0.7 million for the year ended December 31, 2022. The decrease of \$1.0 million was primarily due to the wind-down of the underlying grants, which were closed-out by the third quarter of 2022.



[Table of Contents](#)*Research and development expenses*

The following table summarizes our research and development expenses incurred for the years ended December 31, 2022 and 2021:

	Years Ended December 31,		
	2022	2021	Change
<b>Direct external research and development expenses by program:</b>			
LX2020	\$ 8,551,120	\$ 84,299	\$ 8,466,820
LX1001	7,604,938	4,543,530	3,061,408
LX2006	7,591,077	10,498,841	(2,907,764)
LX1004	1,782,901	4,340,549	(2,557,648)
Other programs	5,225,986	148,661	5,077,325
Total direct external research and development expenses by program	\$ 30,756,022	\$ 19,615,880	\$ 11,140,142
<b>Unallocated research and development expenses:</b>			
Employee compensation and stock-based compensation expense	\$ 10,087,356	\$ 3,547,796	\$ 6,539,560
Professional fees	1,340,928	1,896,551	(555,623)
Lab-related costs and supplies	978,666	21,922	956,744
License fees	451,030	1,305,728	(854,698)
Adverum up-front fee	—	7,500,000	(7,500,000)
Stelios acquisition	—	7,000,000	(7,000,000)
Other unallocated	5,547,538	4,232,997	1,314,541
Total unallocated research and development expenses:	18,405,518	25,504,994	(7,099,476)
Total research and development expenses	\$ 49,161,540	\$ 45,120,874	\$ 4,040,666

Research and development expenses were \$45.1 million for the year ended December 31, 2021 compared to \$49.2 million for the year ended December 31, 2022. The increase of \$4.0 million was primarily due to a net increase of \$11.1 million in direct external research and development program expenses, which was partially offset by a \$7.1 million net decrease in unallocated research and development expenses.

*Direct external research and development expenses*

The net increase of \$11.1 million in direct external research and development expenses was primarily due to an increase in ARVC program (LX2020) costs of \$8.5 million during the year ended December 31, 2022, which consisted mostly of (i) CMC expenses of \$3.8 million related to the development of our manufacturing processes at third-party vendors, (ii) a \$2.0 million milestone payment to the selling shareholders of Stelios, for a development milestone achieved in the third quarter of 2022, and (iii) non-clinical and pre-clinical expenses of \$2.7 million related to the advancement of our LX2020 program into IND-enabling studies. We did not incur significant direct external research and development expenses for our LX2020 program during the year ended December 31, 2021.

The increase of \$3.1 million in direct research and development expenses for our APOE4-associated Alzheimer's disease product candidate (LX1001) was primarily due to increases in CMC expenses of \$2.3 million primarily related to third-party GMP batch costs and clinical expenses of \$0.7 million related to the ongoing Phase 1/2 clinical trial.

The decrease of \$2.9 million in direct external research and development expenses for our FA cardiomyopathy product candidate (LX2006) was primarily due to a decrease in non-clinical expenses of \$6.6 million related to the completion of

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IND-enabling activities in 2021, which was partially offset by increases in clinical expenses of \$2.4 million related to the initiation of the ongoing Phase 1/2 clinical trial and CMC expenses of \$1.2 million primarily related to third-party GMP batch costs.

The decrease of \$2.6 million in direct external research and development expenses for our CLN2 Batten disease product candidate (LX1004) was primarily due to decreases in CMC expenses of \$1.6 million and non-clinical expenses of \$1.1 million.

Direct external research and development expenses for our other programs increased by \$5.1 million during the year ended December 31, 2022, which was primarily attributable to pre-clinical and non-clinical expenses for our early stage cardiovascular and central nervous system disease programs. We did not incur significant direct external research and development expenses for our other programs for the year ended December 31, 2021.

### *Unallocated research and development expenses*

The net decrease of \$7.1 million in unallocated research and development expenses was primarily due to a \$7.5 million upfront fee paid to Adverum to in-license materials and technology in 2021 and \$7.0 million paid for the Stelios acquisition in 2021. These decreases were partially offset by increases in (i) employee compensation and stock-based compensation expense of \$6.5 million due to increased headcount, and (ii) \$1.3 million of other expenses primarily related to rent and facilities costs and lab-related costs and supplies of \$1.0 million, which were associated with our New York City office and laboratory lease that commenced in April 2022.

### *General and administrative expenses*

General and administrative expenses were \$7.2 million for the year ended December 31, 2021 compared to \$12.0 million for the year ended December 31, 2022. The increase of \$4.8 million was primarily due to increases in (i) employee compensation and stock-based compensation expense of \$2.1 million due to increased headcount, (ii) rent and facility expenses of \$1.0 million as our New York City office and laboratory lease commenced in April 2022, and (iii) nonrecurring legal, finance and accounting, consulting and professional fees of \$0.9 million.

### *Interest income*

We recognized interest income of \$1.3 million for the year ended December 31, 2022 related to interest earned on our investments in U.S. government money market funds.

## **Liquidity and capital resources**

### ***Sources of liquidity***

Since our inception, we have not generated any revenue from product sales and have incurred significant operating losses and negative cash flows from our operations. We expect to incur significant expenses and operating losses for the foreseeable future as we advance the clinical development of our product candidates. Since our inception through March 31, 2023, we have funded our operations primarily with net proceeds from sales of our convertible preferred stock and Convertible Securities Agreements of \$183.7 million. As of March 31, 2023, we had cash and cash equivalents of \$58.9 million.

Based on our current operating plan, we expect the net proceeds from this offering, together with our existing cash, will be sufficient to fund our planned operating expenses and capital expenditure requirements into the second quarter of 2024. Our total future capital requirements will depend on many factors and is subject to the risks and uncertainties set forth in the section titled "Risk Factors." Our development plans may be revised, and our estimated use of proceeds may be impacted.

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## Cash flows

The following table summarizes our sources and uses of cash for the three months ended March 31, 2023 and March 31, 2022:

	Three Months Ended March 31,	
	2023	2022
Net cash used in operating activities	\$ (18,375,269)	\$ (15,094,795)
Net cash used in investing activities	—	(173,625)
Net cash (used in) provided by financing activities	(97,440)	255,987
Net decrease in cash	\$ (18,472,709)	\$ (15,012,433)

### Operating activities

During the three months ended March 31, 2022, net cash used in operating activities consisted primarily of our net loss of \$12.5 million and \$2.9 million in net cash used by changes in operating assets and liabilities, which were partially offset by \$0.3 million in stock-based compensation expense.

During the three months ended March 31, 2023, net cash used in operating activities consisted primarily of our net loss of \$18.7 million and \$0.6 million in net cash used by changes in operating assets and liabilities, which were partially offset by \$0.4 million in stock-based compensation expense and \$0.4 million in amortization of our right-of-use assets for our operating and finance leases.

### Investing activities

During the three months ended March 31, 2022, net cash used in investing activities was \$0.2 million and consisted of purchases of lab equipment.

During the three months ended March 31, 2023, there was no net cash used in or provided by investing activities.

### Financing activities

During the three months ended March 31, 2022, net cash provided by financing activities consisted primarily of \$0.2 million of proceeds received from the exercise of stock options.

During the three months ended March 31, 2023, net cash used in financing activities consisted primarily of \$0.1 million of principal payments made on equipment finance leases.

The following table summarizes our sources and uses of cash for the years ended December 31, 2022 and December 31, 2021:

	Years Ended December 31,	
	2022	2021
Net cash used in operating activities	\$ (54,560,593)	\$ (44,182,963)
Net cash used in investing activities	(900,536)	(187,671)
Net cash provided by financing activities	189,288	150,857,485
Net increase (decrease) in cash	\$ (55,271,841)	\$ 106,486,851

### Operating activities

During the year ended December 31, 2021, net cash used in operating activities consisted primarily of our net loss of \$50.6 million, partially offset by (i) \$6.0 million in net cash provided by changes in operating assets and liabilities and (ii) \$0.4 million in stock-based compensation expense.

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During the year ended December 31, 2022, net cash used in operating activities consisted primarily of our net loss of \$59.3 million, partially offset by (i) \$1.8 million in stock-based compensation expense, (ii) \$1.8 million in net cash provided by changes in operating assets and liabilities, and (iii) \$1.0 million in amortization of our right-of-use assets for our operating and finance leases.

### *Investing activities*

During the year ended December 31, 2021, net cash used in investing activities was \$0.2 million and consisted of the purchase of lab equipment.

During the year ended December 31, 2022, net cash used in investing activities was \$0.9 million and consisted of the purchase of lab equipment and internal use software.

### *Financing activities*

During the year ended December 31, 2021, net cash provided by financing activities consisted primarily of \$151.1 million gross proceeds from the issuance of 50,999,997 shares of Series A convertible preferred stock at a purchase price of \$1.00 per share and 58,157,823 shares of Series B convertible preferred stock at a purchase price of \$1.72 per share, as well as proceeds from the exercise of stock options of \$0.2 million, partially offset by \$0.3 million of issuance costs.

During the year ended December 31, 2022, net cash provided by financing activities consisted primarily of proceeds from the exercise of stock options of \$0.3 million, partially offset by \$0.2 million of principal payments made on equipment finance leases.

Our primary uses of cash are to fund our clinical trials, research and development activities related to our discovery programs and our preclinical and clinical product candidates, hiring personnel, raising capital and providing general and administrative support for these operations.

### **Funding requirements**

We expect our expenses and capital requirements to increase significantly in connection with our ongoing activities, particularly as we advance our lead product candidates and other development programs. Further, following the completion of this offering, we expect to incur additional costs associated with operating as a public company. Accordingly, we will require substantial additional funding to support our continuing operations.

The timing and amount of our future operating and capital requirements will largely depend on many factors, including:

- the initiation, scope, progress, timing, results and costs of product discovery, preclinical studies and clinical trials for our product candidates or any future candidates we may develop;
- our ability to maintain our relationships with Weill Cornell Medicine, Adverum, UCSD, and any other key licensors or collaborators;
- the scope, prioritization and number of our research and development programs;
- the costs, timing and outcome of seeking and obtaining regulatory approvals from the FDA and comparable foreign regulatory authorities, including the potential for such authorities to require that we perform more preclinical studies or clinical trials than those that we currently expect or change their requirements on studies that had previously been agreed to;

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- our ability to establish and maintain collaborations on favorable terms, if at all;
- the achievement of milestones or occurrence of other developments that trigger payments under any collaboration agreements we have or may enter into;
- the extent to which we are obligated to reimburse, or entitled to reimbursement of, clinical trial costs under collaboration agreements, if any;
- the costs to establish, maintain, expand, enforce and defend the scope of our intellectual property portfolio, including preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending intellectual property-related claims;
- the extent to which we acquire or in-license other product candidates and technologies;
- the costs of securing manufacturing arrangements for commercial production;
- the costs of establishing or contracting for sales and marketing capabilities if we obtain regulatory approvals to market our product candidates; and
- our need to implement additional internal systems and infrastructure, including financial and reporting systems.

Identifying potential product candidates and conducting preclinical studies and clinical trials is a time-consuming, expensive and uncertain process that takes years to complete, and we may never generate the necessary data or results required to obtain regulatory approval and commercialize our product candidates. In addition, our product candidates, if approved, may not achieve commercial success. Our commercial revenues, if any, will be derived from sales of products that we do not expect to be commercially available for many years, if at all. Accordingly, we will need to continue to rely on additional financing to achieve our business objectives.

Until such time, if ever, as we can generate significant revenue from product sales or other sources, we expect to finance our operations through the sale of equity, debt financings, or other capital sources, which could include proceeds from potential collaborations, strategic partnerships or marketing, distribution, licensing or other similar arrangements with third parties. See the “Business” for a discussion of our material license agreements and sponsored research agreements for further information regarding milestone and royalty payments we may be obligated to make in the future.

We may be unable to raise additional funds or enter into such agreements or arrangements on favorable terms, or at all. Both the COVID-19 pandemic and the Russia-Ukraine war resulted in a significant disruption of global financial markets and contributed to a general global economic slowdown. If the disruption persists and deepens, whether as a result of these events or otherwise, we could experience an inability to access additional capital. To the extent that we raise additional capital through the sale of equity or convertible debt securities, your ownership interest will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect your rights as a common stockholder. Any future debt financing and preferred equity financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, selling or licensing our assets, making capital expenditures, declaring dividends or encumbering our assets to secure future indebtedness. Such restrictions could adversely impact our ability to conduct our operations and execute our business plan.

If we raise funds through collaborations, strategic alliances or licensing arrangements with third parties, we may have to relinquish valuable rights to our technologies, future revenue streams, research programs or product candidates or to grant licenses on terms that may not be favorable to us. If we fail to raise capital or enter into such agreements as, and when, needed, we may have to significantly delay, scale back or discontinue the

development or commercialization of our product candidates or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

### **Known trends, events and uncertainties**

We continue to actively monitor the impact of various macroeconomic trends, such as high rates of inflation, supply chain disruptions and geopolitical instability caused by the Russian-Ukrainian war.

The recent trends towards rising inflation may materially affect our business and corresponding financial position and cash flows. Inflationary factors, such as increases in the cost of our clinical trial materials and supplies, interest rates and overhead costs may adversely affect our operating results. Rising interest rates also present a recent challenge impacting the U.S. economy and could make it more difficult for us to obtain traditional financing on acceptable terms, if at all, in the future. Additionally, the general consensus among economists suggests that we should expect a higher recession risk to continue over the next year, which, together with the foregoing, could result in further economic uncertainty and volatility in the capital markets in the near term, and could negatively affect our operations. We may experience increases in our operating costs, including our labor costs and research and development costs, due to supply chain constraints, consequences associated with COVID-19 and the ongoing conflict between Russia and Ukraine, and employee availability and wage increases, which may result in additional stress on our working capital resources.

In addition, the recent closures of Silicon Valley Bank, Signature Bank, and First Republic Bank have resulted in broader financial institution liquidity risk and concerns, and future adverse developments with respect to specific financial institutions or the broader financial services industry may lead to market-wide liquidity shortages. The failure of any bank in which we deposit our funds could reduce the amount of cash we have available for our operations or corporate development, or delay our ability to access such funds. Any such failure may increase the possibility of a sustained deterioration of financial market liquidity, or illiquidity at clearing, cash management and/or custodial financial institutions. In the event we have a commercial relationship with a bank that has failed or is otherwise distressed, we may experience delays or other issues in meeting our financial obligations. If other banks and financial institutions fail or become insolvent in the future in response to financial conditions affecting the banking system and financial markets, our ability to access our cash and cash equivalents may be threatened and our ability to borrow or raise additional capital when needed to operate our business could be substantially impaired.

### **Critical accounting policies and significant judgements and estimates**

Our management's discussion and analysis of our financial condition and results of operations are based on our consolidated financial statements, which have been prepared in accordance with U.S. generally accepted accounting principles, or U.S. GAAP. The preparation of our consolidated financial statements and related disclosures requires us to make estimates and judgments that affect the reported amounts of assets, liabilities, expenses, and the disclosure of our contingent liabilities in our consolidated financial statements. We base our estimates on historical experience, known trends and events and various other factors that we believe are reasonable under the circumstances, the results of which form the basis for making judgments about the carrying values of assets and liabilities that are not readily apparent from other sources. We evaluate our estimates and assumptions on an ongoing basis. Our actual results may differ from these estimates under different assumptions or conditions.

While our significant accounting policies are described in more detail in Note 2 to our audited consolidated financial statements included elsewhere in this prospectus, we believe that the following accounting policies are those most critical to the judgments and estimates used in the preparation of our audited consolidated financial statements.

### ***Research and development***

As part of the process of preparing our consolidated financial statements, we are required to estimate our accrued research and development expenses. This process involves estimating the level of service performed and the associated cost incurred for the service when we have not yet been invoiced or otherwise notified of actual costs. The majority of our service providers invoice us in arrears for services performed, on a pre-determined schedule or when contractual milestones are met; however, some require advance payments. We make estimates of our accrued expenses as of each balance sheet date in the consolidated financial statements based on facts and circumstances known to us at that time. At each period end, we corroborate the accuracy of these estimates with the service providers and make adjustments, if necessary. Examples of estimated accrued research and development expenses include those related to fees paid to:

- vendors in connection with discovery and preclinical development activities;
- CROs in connection with preclinical studies and testing; and
- CMOs in connection with the process development and scale up activities and the production of materials.

We record the expense and accrual related to contract research and manufacturing based on our estimates of the services received and efforts expended considering a number of factors, including our knowledge of the progress towards completion of the research, development, and manufacturing activities; invoicing to date under contracts; communication from the CROs, CMOs, and other companies of any actual costs incurred during the period that have not yet been invoiced; and the costs included in the contracts and purchase orders. The financial terms of these agreements are subject to negotiation, vary from contract to contract, and may result in uneven payment flows. There may be instances in which payments made to our vendors will exceed the level of services provided and result in a prepayment of the expense. In accruing service fees, we estimate the time period over which services will be performed and the level of effort to be expended in each period. If the actual timing of the performance of services or the level of effort varies from the estimate, we adjust the accrual or the amount of prepaid expense accordingly. Although we do not expect our estimates to be materially different from amounts actually incurred, our understanding of the status and timing of services performed relative to the actual status and timing of services performed may vary and may result in reporting amounts that are too high or too low in any particular period. To date, there have not been any material adjustments to our prior estimates of accrued research and development expenses; however, we cannot guarantee that such adjustments will not be made in the future.

### ***Determination of fair value of common stock***

As there has been no public market for our common stock to date, the estimated fair value of our common stock has been determined by our board of directors as of the date of each option grant, with input from management, considering our most recently available third-party valuations of common stock and our board of directors' assessment of additional objective and subjective factors that it believed were relevant and which may have changed from the date of the most recent valuation through the date of the grant. These third-party valuations were performed in accordance with the guidance outlined in the American Institute of Certified Public Accountants' Accounting and Valuation Guide, Valuation of Privately-Held-Company Equity Securities Issued as Compensation.

Our common stock valuations were prepared using either an option pricing method, or OPM, or a hybrid method of OPM and probability-weighted expected return method, or PWERM. Both the OPM and hybrid methods use market approaches to estimate our enterprise value. These third-party valuations were performed at various dates, which resulted in valuations of our common stock of \$1.38 as of December 17, 2021, \$1.43 as of February 15,

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2022, and \$1.66 as of February 14, 2023. In addition to considering the results of these third-party valuations, our board of directors considered various objective and subjective factors to determine the fair value of our common stock as of each grant date, including:

- the prices at which we sold shares of preferred stock and the superior rights and preferences of the preferred stock relative to our common stock at the time of each grant;
- the progress of our research and development programs, including the status and results of preclinical studies for our product candidates;
- our stage of development and our business strategy;
- external market conditions affecting the biopharmaceutical industry and trends within the biopharmaceutical industry;
- our financial position, including cash on hand, and our historical and forecasted performance and operating results;
- the lack of an active public market for our common stock and our preferred stock;
- significant changes to the key assumptions underlying the factors used could have resulted in different fair values of common stock at each valuation date;
- the likelihood of achieving a liquidity event, such as an initial public offering, or IPO, or sale of our company in light of prevailing market conditions; and
- the analysis of IPOs and the market performance of similar companies in the therapeutics industry.

The assumptions underlying these valuations represented management's best estimate, which involved inherent uncertainties and the application of management's judgment. As a result, if we had used significantly different assumptions or estimates, the fair value of our common stock and our stock-based compensation expense could have been materially different.

Once a public trading market for our common stock has been established in connection with the completion of this offering, it will no longer be necessary for our board of directors to estimate the fair value of our common stock in connection with our accounting for granted stock options and other such awards we may grant, as the fair value of our common stock will be determined based on the quoted market price of our common stock.

### ***Stock option awards***

We recognize share-based compensation expense related to stock option awards granted based on the estimated fair value of the awards on the date of grant using a Black-Scholes option pricing model. The grant date fair value of the share-based awards is recognized on a straight-line basis over the requisite service period, which is the vesting period of the respective awards. Share-based payments to non-employees issued in exchange for services are based upon the fair value of the equity instruments issued. Compensation expense for stock options issued to non-employees is calculated using the Black-Scholes option pricing model and is recorded over the requisite service performance period.

The Black-Scholes option-pricing model requires the use of highly subjective assumptions, which are used to determine the fair value stock option awards granted. These assumptions include:

*Expected Term.* The expected term represents the period that our stock option awards granted are expected to be outstanding and is determined using the simplified method (based on the mid-point between the vesting date and the end of the contractual term).



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*Expected Volatility.* Because we do not have any trading history for our shares of common stock, the expected volatility is estimated using averages of the historical volatility of our peer group of companies for a period equal to the expected life of the stock options granted. Our peer group of publicly traded companies was chosen based on their similar size, stage in the life cycle or area of specialty.

*Risk-Free Interest Rate.* The risk-free interest rate is based on the interest rates paid on securities issued by the U.S. Treasury with a term approximating the expected life of stock options granted.

*Expected Dividend.* We have never paid, and do not anticipate paying, cash dividends on our shares of common stock. Therefore, the expected dividend yield was assumed to be zero.

The following table summarizes by grant date the number of shares subject to options granted since December 1, 2021, through March 31, 2023, the per share exercise price of the options, the per share fair value of our common stock on each grant date and the per share estimated fair value of the options:

<b>Grant Date</b>	<b>Number of Shares Subject to Options Granted</b>	<b>Per Share Exercise Price of Options</b>	<b>Per Share Fair Value of Common Stock on Grant Date</b>	<b>Per Share Estimated Fair Value of Options</b>
December 1, 2021	7,500	\$ 0.46	\$ 0.46	\$ 0.30
December 6, 2021	28,295	\$ 0.46	\$ 0.46	\$ 0.30
January 26, 2022	449,060	\$ 1.38	\$ 1.38	\$ 0.89
March 7, 2022	15,000	\$ 1.43	\$ 1.43	\$ 0.92
May 6, 2022	850,786	\$ 1.43	\$ 1.43	\$ 0.95
July 20, 2022	388,542	\$ 1.43	\$ 1.43	\$ 0.97
December 2, 2022	25,250	\$ 1.43	\$ 1.43	\$ 0.98
March 14, 2023	4,922,272	\$ 1.66	\$ 1.66	\$ 1.03

### **Emerging growth company status**

The JOBS Act permits an “emerging growth company” such as us to take advantage of an extended transition period to comply with new or revised accounting standards applicable to public companies until those standards would otherwise apply to private companies. We have elected not to “opt out” of such extended transition period, which means that when a standard is issued or revised and it has different application dates for public or private companies, we will adopt the new or revised standard at the time private companies adopt the new or revised standard and will do so until such time that we either (i) irrevocably elect to “opt out” of such extended transition period or (ii) no longer qualify as an emerging growth company. There are other exemptions and reduced reporting requirements provided by the JOBS Act that we are currently evaluating. For example, as an “emerging growth company,” we are exempt from Sections 14A(a) and (b) of the Securities Exchange Act of 1934, as amended, which would otherwise require us to (1) submit certain executive compensation matters to shareholder advisory votes, such as “say-on-pay,” “say-on-frequency,” and “golden parachutes;” and (2) disclose certain executive compensation related items such as the correlation between executive compensation and performance and comparisons of our chief executive officer’s compensation to our median employee compensation. We also intend to rely on an exemption from the rule requiring us to provide an auditor’s attestation report on our internal controls over financial reporting pursuant to Section 404(b) of the Sarbanes-Oxley Act. We will continue to remain an “emerging growth company” until the earliest of the following: (1) the last day of the fiscal year following the fifth anniversary of the date of the completion of this offering; (2) the last day of the fiscal year in

which our total annual gross revenue is equal to or more than \$1.235 billion; (3) the date on which we have issued more than \$1.0 billion in nonconvertible debt during the previous three years; or (4) the date on which we are deemed to be a large accelerated filer under the rules of the SEC.

### **Recently issued accounting pronouncements**

A description of recently issued accounting pronouncements that may potentially impact our financial position, results of operations and cash flows is disclosed in Note 2 to our audited consolidated financial statements included elsewhere in this prospectus.

### **Qualitative and quantitative disclosures about market risk**

#### *Interest rate risk*

As of March 31, 2023, we had cash and cash equivalents of \$58.9 million. Interest income is sensitive to changes in the general level of interest rates. Our surplus cash has been invested in money market fund accounts and interest-bearing savings accounts. We have not entered into investments for trading or speculative purposes.

Due to the conservative nature of our investment portfolio, which is predicated on capital preservation of investments with short-term maturities, we do not believe an immediate one percentage point change in interest rates would have a material effect on the fair market value of our portfolio, and therefore we do not expect our operating results or cash flows to be significantly affected by changes in market interest rates. As of December 31, 2021, we had no debt outstanding, and as of December 31, 2022 and March 31, 2023, we had approximately \$2.2 million and \$2.1 million, respectively, of total finance lease liabilities that bear interest on a fixed basis. Therefore, we are not significantly exposed to interest rate risk with respect to debt.

All of our employees and our operations are currently located in the United States. We have, from time to time, engaged with contractors or other vendors in a currency other than the U.S. dollar. To date, we have not been exposed to significant market risk related to changes in foreign currency exchange rates. Our operations may be subject to fluctuations in foreign currency exchange rates in the future.

We do not believe that inflation has had a material effect on our business, financial condition or results of operations. Our operations may be subject to inflation in the future.

## Business

### Overview

We are a clinical-stage gene therapy company focused on addressing some of the most devastating genetically defined cardiovascular diseases and a genetically defined sub-group of Alzheimer's disease. We are advancing a deep and diverse pipeline of AAV-based gene therapy candidates for diseases affecting both larger-rare and prevalent patient populations. Utilizing a modular approach that integrates a clinically validated vector, we have a disease area strategy targeting defined patient sub-groups most likely to benefit from our gene therapy candidates and high-quality, scalable manufacturing, which is designed to overcome many of the challenges facing the field of gene therapy.

Our most advanced cardiovascular product candidate, LX2006 for the treatment of patients with Friedreich's ataxia, or FA, cardiomyopathy, is currently being evaluated in an ongoing Phase 1/2 clinical trial, and we expect to report initial biomarker data from this trial in 2024. Our second most advanced cardiovascular product candidate, LX2020 for the treatment of arrhythmogenic right ventricular cardiomyopathy, or ARVC, caused by mutations in the PKP2 gene, has completed investigational new drug, or IND, -enabling studies, and we expect to submit an IND application to the U.S. Food and Drug Administration, or FDA, in mid-2023.

Our lead Alzheimer's disease product candidate, LX1001 for the treatment of *APOE4* homozygous patients with Alzheimer's disease, is in an ongoing Phase 1/2 trial, and we have initially observed an increase in expression levels of the protective protein, *APOE2*, in the first and second dose cohorts and a consistent trend towards improvement in core Alzheimer's disease biomarkers in the first dose cohort. We expect to report additional biomarker data from the second dose cohort in 2023.

We are targeting diseases that have seen limited penetration of precision medicine, which we define as medications that treat the underlying molecular mechanism of a disease, where we believe there is a significant opportunity for gene therapy to play a role as a key therapeutic option. We believe the specific indications we are initially targeting, FA cardiomyopathy, ARVC and *APOE4*-associated Alzheimer's disease, are highly amenable to gene therapy, where administration of a single dose has the potential to either restore loss-of-function or minimize gain-of-function mutations by treating the underlying genetic cause of the disease. Although few precision medicines are currently approved for the treatment of cardiovascular conditions or Alzheimer's disease, recent approvals by the FDA suggest a willingness to approve new precision medicines based on biomarkers and functional endpoints. Together with improved diagnostics and increased testing, these developments may offer one of the most substantial opportunities for the uptake of precision medicines in the global pharmaceutical marketplace.

Each of our gene therapy candidates utilizes the vector construct, dose and route of administration that we believe will result in the most favorable biodistribution and safety profile for our product candidate for each disease. Our most advanced cardiovascular and *APOE4*-associated Alzheimer's disease programs use the AAVrh10 vector due to its high transduction efficiency in both myocardial cells and neurons, potential for lower toxicity given the opportunity to utilize lower doses compared to other well-established AAV serotypes, and lower pre-existing immunity.

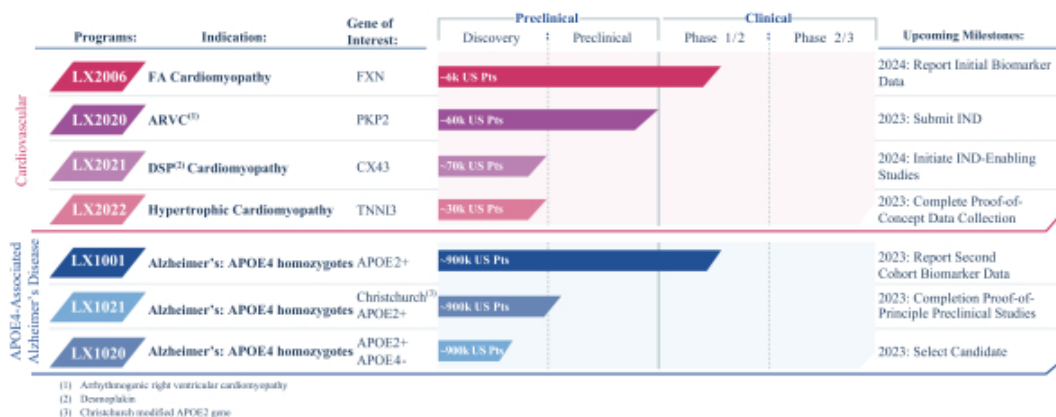
By specifically tailoring our technological approach to each targeted disease, we believe we can optimize our programs to achieve the highest likelihood of having therapeutic impact. We target genetically defined indications in specific sub-groups of patients that offer the potential to demonstrate therapeutic impact through improvement in functional endpoints or biomarkers, have high unmet need and large market opportunities, have established proof-of-concept in relevant preclinical models, and have organized patient advocacy groups and identifiable patient populations. In addition to targeting cardiovascular diseases and *APOE4*-associated Alzheimer's disease that we believe can be addressed by our current approach utilizing AAVrh10, we have ongoing discovery efforts to identify next-generation vector technologies with the best potential therapeutic

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profile. Finally, we continuously seek to bolster our pipeline through relationships with academic institutions, which provide us access to cutting edge gene therapy research that we will utilize in the discovery and development of next-generation gene therapy candidates.

### Our pipeline

Utilizing a stepwise, capital-efficient development approach, we are leveraging early proof-of-concept functional and biomarker data to advance a deep and diverse pipeline of cardiovascular and APOE4-associated Alzheimer's disease programs for larger-rare and prevalent indications. We retain exclusive worldwide development and commercialization rights to all of our product candidates and programs.



### Cardiovascular programs

We are developing a number of disease-modifying gene therapy candidates to treat larger-rare cardiovascular diseases that have significant unmet need and no approved treatments that address the underlying genetic cause of the disease. These programs include:

- **LX2006** is an AAVrh10-based gene therapy candidate designed to intravenously deliver a functional frataxin, or *FXN*, gene for the treatment of FA cardiomyopathy. FA cardiomyopathy is the most common cause of mortality in patients with Friedreich's ataxia and affects approximately 5,600 patients in the United States. LX2006 is designed to promote the expression of the protein frataxin to restore normal mitochondrial function and energy production in myocardial cells. In preclinical studies, LX2006 demonstrated improvement in cardiac function and survival in a severe *FXN* knockout mouse model. Similarly, in a partial *FXN* knockout mouse model, LX2006 was observed to restore cardiac function and reverse disease abnormalities of FA cardiomyopathy. LX2006 is currently being evaluated in an open-label, dose-escalation Phase 1/2 clinical trial in patients with FA cardiomyopathy, and we expect to report initial biomarker data from this trial in 2024. The FDA has granted Rare Pediatric Disease designation and Orphan Drug designation to LX2006 for the treatment of FA.
- **LX2020** is an AAVrh10-based gene therapy candidate designed to intravenously deliver a fully functional *PKP2* gene to cardiac muscle for the treatment of ARVC caused by mutations in the *PKP2* gene. *PKP2* mutations are associated with approximately 75% of all genetic cases of ARVC, and we estimate they affect approximately 60,000 patients in the United States. *PKP2* mutations can cause replacement of heart muscle with fibrotic tissue and fatty deposits and severe abnormal heart rhythms, or arrhythmias, that cause cardiac

dysfunction and can result in sudden cardiac death. LX2020 is designed to increase desmosomal PKP2 protein levels, reassemble desmosomes and restore myocardial cell function. In our preclinical studies using a genetic mouse model of ARVC harboring a *PKP2* mutation that recapitulates the phenotype of ARVC, LX2020 resulted in fewer arrhythmias and increased survival. We intend to submit an IND application for LX2020 in mid-2023.

- *LX2021* is a gene therapy candidate we are developing to intravenously deliver the coding sequence for the functional Cx43 protein for arrhythmogenic cardiomyopathies, or ACs. ACs are a group of inherited cardiac muscle disorders associated with a high risk of sudden death. We believe restoring the Cx43 protein can potentially treat multiple genetic causes of AC because the cardiac loss of Cx43 is a molecular deficit generally observed in all AC patient populations. Our LX2021 program is initially targeting a distinct form of AC called Desmoplakin cardiomyopathy, or DSP Cardiomyopathy, with a prevalence that may be as high as 6% of all inherited dilated cardiomyopathies and 7% of all ARVCs, impacting up to approximately 70,000 patients in the United States. We plan to initiate IND-enabling studies for LX2021 by the end of 2024.
- *LX2022* is a gene therapy candidate we are developing to intravenously deliver a functional *TNNI3* gene to myocardial cells to treat a distinct form of hypertrophic cardiomyopathy, or HCM, due to mutations in the *TNNI3* gene. Mutations in the *TNNI3* gene often result in left ventricular hypertrophy and restrictive cardiomyopathy, leading to arrhythmias and heart failure. With an estimated prevalence of 1 in 500 people in the United States, HCM is one of the most common forms of genetic cardiomyopathy caused by mutations that affect the cardiac sarcomere in approximately 75% of cases. It is estimated that approximately 30,000 patients in the United States and 36,000 patients in the European Union are affected by HCM caused by mutations in the *TNNI3* gene. We plan to complete proof-of-concept data collection for LX2022 in 2023.

### *APOE4-associated Alzheimer's programs*

We are developing a portfolio of approaches to treat the genetics underlying Alzheimer's disease.

- *LX1001* is an AAVrh10-based gene therapy candidate designed to deliver via one-time administration into the cerebrospinal fluid, or CSF, an *APOE2* gene for the treatment of *APOE4* homozygous patients with Alzheimer's disease. Alzheimer's disease is the leading cause of cognitive decline in late adult life and characterized by complex underlying pathology in the central nervous system, or CNS. Individuals homozygous for *APOE4*, an allele of the gene *APOE*, are approximately 15 times more likely to develop Alzheimer's disease than the general population, and it is estimated that there are 900,000 *APOE4* homozygous patients with Alzheimer's disease in the United States alone. Conversely, individuals homozygous for the *APOE* allele *APOE2* are 40% less likely to develop Alzheimer's disease than the general population. This and other evidence suggest that *APOE2* may play a neuroprotective role. *LX1001* is designed to express the protective *APOE2* gene in the CNS of *APOE4* homozygous patients in order to halt or slow the progression of Alzheimer's disease. *LX1001* is being evaluated in an ongoing open-label, dose-escalation Phase 1/2 clinical trial. In the first dose cohort in the trial, we observed a consistent trend towards improvement in Alzheimer's disease CSF biomarkers, such as total tau and phosphorylated tau, that are consistent with improvement in the first three patients with 12 months of data. We have also observed expression of the protective *APOE2* protein in all patients in the first and second dose cohorts with follow-up data. *LX1001* has been granted Fast Track designation by the FDA for the treatment of patients with early Alzheimer's disease who are *APOE4* homozygous, to slow disease progression. We expect to report additional biomarker data from the second dose cohort in 2023.
- *LX1021* is a gene therapy candidate we are developing to deliver a Christchurch mutation-modified *APOE2* allele for the treatment of *APOE4* homozygous patients with Alzheimer's disease. The Christchurch mutation has been recognized to protect individuals against Alzheimer's disease even in the presence of significant amyloid pathology. The mechanism of this protection may relate to the fact that *APOE*, in the presence of the Christchurch mutation, binds poorly to heparan sulfate proteoglycans, which are molecules found on the surface of neurons that may inhibit the spread of tau between cells. We believe this approach has the potential to enhance the protective effect of *APOE2* in homozygous *APOE4*-associated Alzheimer's disease. We plan to complete proof-of-principle preclinical studies in 2023.

- *LX1020* is a gene therapy candidate we are developing to deliver both the protective *APOE2* allele and miRNA to suppress *APOE4* for the treatment of *APOE4* homozygous patients. We believe delivery of *APOE2* with concurrent suppression of *APOE4* will achieve a higher degree of conversion to the *APOE4/E2* heterozygous profile, which should lead to greater therapeutic effect. We plan to complete candidate selection in 2023.

### **Our company and team**

We were founded through a collaboration between the world-class gene therapy research at Weill Cornell Medicine and a team of pioneering scientists, clinicians, and business leaders with deep expertise in gene therapy. We continue to leverage our long-standing relationship with Weill Cornell Medicine through sponsored research and development efforts to progress our cardiovascular platform technology as well as our *APOE4-associated Alzheimer's disease* product candidates. Through our acquisition of Stelios Therapeutics, Inc., we have an ongoing collaboration with the University of California, San Diego, or UCSD, through which we sponsor research and development designed to advance our earlier stage cardiovascular pipeline. We work with highly experienced teams at both Weill Cornell Medicine and UCSD that have deep expertise in the underlying biology of cardiovascular and *APOE4-associated Alzheimer's disease* and the patient populations that they treat. We believe our ongoing work with these preeminent institutions, as well as any potential new or expanded collaborations, will continue to be a valuable aspect of our efforts as we seek to further discover and develop novel gene therapies for devastating diseases.

We are led by pioneers and experts with decades of collective experience in genetic medicines, rare disease drug development, manufacturing and commercialization. Our scientific founder and Chief Scientific Advisor, Ronald G. Crystal, M.D., is a world leader in gene therapy research and development and currently serves as Professor and Chairman of Weill Cornell Medicine's Genetic Medicine Department and Director of the Belfer Gene Therapy Core Facility. Dr. Crystal has sponsored 14 cleared gene therapy IND applications and has published more than 300 papers on gene therapy. He has more than thirty years of experience with AAVs and vector design and expertise in CNS, cardiac, pulmonary and liver-mediated diseases. Our Chief Executive Officer, R. Nolan Townsend, has spent more than a decade working in global biopharmaceutical commercial organizations, including leading Pfizer's \$1.6 billion international rare disease commercial business unit. Mr. Townsend also led Pfizer's U.S. rare disease commercial business unit, where he oversaw the successful U.S. launch of blockbuster rare cardiovascular product Vyndaqel/Vyndamax (tafamidis), now a \$2.4 billion product by revenue. He has overseen other successful rare disease product launches and been involved in business and commercial development efforts for several gene therapy programs. Eric Adler, M.D., our Chief Scientific Officer, is a pioneer in cardiovascular gene therapy, and Professor of Medicine, head of the Heart Failure Section, Director of the Strauss Center for Cardiomyopathy, and the Czarina and Humberto S. Lopez Chancellor's Endowed Chair in Cardiology at UCSD. Our Chief Technical Officer, Paul McCormac, Ph.D., has more than 20 years of experience and a proven track record in the field of gene therapy and biologics chemistry, manufacturing, and controls, or CMC. He previously served as Medicinal Sciences Category Lead for Pfizer Rare Disease, where he led gene therapy CMC and vector supply strategy for Pfizer's rare disease business and research units. Our Chief Financial Officer, Micah Zajic, brings nearly 20 years of experience in biotech corporate development, healthcare capital markets and investment banking to our company. Mr. Zajic most recently served as Chief Financial Officer at SQZ Biotechnologies. He began his career at Morgan Stanley in the Global Capital Markets and Healthcare Investment Banking Groups, raising capital and executing strategic transactions for companies across the healthcare industry. Our Chairman, Steven Altschuler, M.D., is currently Managing Director at Ziff Capital Partners and was formerly Chairman of the gene therapy pioneer Spark Therapeutics, Inc., which was responsible for the first FDA-approved gene therapy, voretigene neparvovec-rzyl (Luxturna), and was acquired by Roche Holding AG in 2019 for \$4.3 billion. Our senior leadership is supported by the rest of our management team, which has decades of experience across research, development,

manufacturing and commercialization in gene therapy and rare disease both from prominent biotechnology companies and global biopharmaceutical companies that are active in the space.

Since our inception, we have raised \$183.7 million in capital, including from certain funds affiliated with D1 Capital Partners, Eventide Healthcare & Life Sciences Fund, Janus Henderson Investors, Longitude Capital, Lundbeckfonden Ventures, Omega Funds and PBM Capital. Prospective investors should not rely on the investment decisions of our existing investors, as these investors may have different risk tolerances and have received their shares in prior offerings at prices lower than the price offered to the public in this offering. See “Business—Certain Relationships and Related Party Transactions” for more information.

## Our strategy

We are building a fully integrated company with the potential to significantly advance gene therapy’s impact on many of society’s most challenging genetically defined cardiovascular diseases and *APOE4*-associated Alzheimer’s disease. Our vision is a world where gene therapies resolve the burden of disease and transform the lives of patients. The key elements of our strategy to achieve this vision are to:

- *Focus on genetically defined cardiovascular diseases and APOE4-associated Alzheimer’s disease, two areas of high unmet need and substantial commercial potential.* We are focused on genetically defined cardiovascular diseases and *APOE4*-associated Alzheimer’s disease where there are no currently approved treatment or where we believe our therapies will provide a meaningful improvement relative to existing standards of care. Furthermore, we seek indications with commercial opportunities beyond those typically associated with gene therapy companies targeting rare monogenic diseases. Among our cardiovascular programs, LX2006 has the potential to address a prevalent population of approximately 5,600 FA cardiomyopathy patients in the United States, and LX2020 has the potential to address approximately 60,000 patients in the United States with ARVC caused by *PKP2* mutations. We believe that LX1001 has the potential to address up to approximately 900,000 patients in the United States with homozygous *APOE4*-associated Alzheimer’s disease.
- *Target development efforts at disease pathologies and defined sub-groups of patients most likely to benefit from our therapies.* Our programs are designed to target either the aspect of the disease that is most amenable to gene therapy or the genetic sub-groups that we believe are most likely to respond to gene therapy. For example, in the case of LX2006, we are targeting the cardiac manifestation of the disease because we believe it is more amenable to gene therapy than the neurologic manifestation. For LX2020, we are targeting the *PKP2* sub-group that represents approximately 75% of the genetic ARVC cases and presents a clinical phenotype amenable to treatment. For LX1001, we are targeting homozygous *APOE4*-associated Alzheimer’s disease patients, the highest risk population to develop Alzheimer’s disease and the sub-group most likely to demonstrate a treatment effect from the expression of the protective *APOE2* protein and where currently approved treatments have demonstrated clinical benefit less favorable compared to other *APOE* genotypes.
- *Pursue a stepwise and capital-efficient development approach for advancing our programs through clinic development and regulatory approval.* We are advancing our pipeline of AAV-based gene therapies for cardiovascular and CNS indications. We expect to report initial biomarker data from our Phase 1/2 clinical trial of LX2006 for the treatment of FA cardiomyopathy in 2024. We expect to report initial biomarker data from the second dose cohort in our Phase 1/2 clinical trial of LX1001 for the treatment of *APOE4*-associated Alzheimer’s disease in 2023. We plan to leverage key learnings from our lead programs in cardiovascular and *APOE4*-associated Alzheimer’s disease to guide the development of our next-generation candidates and follow-on indications. Given that there are several targets in myocardial cells that are dysregulated in cardiomyopathy, we believe there will be read-through to similar indications following proof-of-concept from our initial FA cardiomyopathy and ARVC programs. Similarly, we believe that positive clinical data from our

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*APOE4*-associated Alzheimer's disease program has the potential to validate our novel approach to treating the *APOE4* homozygous sub-group and can help inform the development plan of our next-generation candidates. Furthermore, we plan to leverage biomarkers for each of our development programs, potentially enabling us to rapidly validate proof-of-mechanism and support further development efforts. We believe our strategy could facilitate shorter clinical trial timelines, which, coupled with expedited global regulatory approval pathways we plan to pursue, could potentially accelerate clinical development and reduce overall development costs.

- *Utilize a scalable, unified manufacturing platform across all current and future programs.* We plan to use a unified manufacturing process across all current and future programs to manufacture vector for clinical and commercial use via a proprietary platform suspension baculovirus process using Sf9 cells. This production platform has demonstrated the capability to produce high-quality, high-yield, and high-potency vectors that can accommodate demand for both larger rare and prevalent patient populations. We are currently working with third parties for our manufacturing capabilities; however, we will evaluate building out our own current good manufacturing practices, or cGMP, -compliant manufacturing facility as, and if, we achieve further clinical success or any of our product candidates ultimately receive marketing authorization.
- *Expand our pipeline capabilities through the discovery and development of next-generation gene therapy technologies.* We have combined our robust expertise in vector design, regulatory and clinical strategy, and disease area knowledge to establish a deep cardiovascular gene therapy pipeline. We have chosen to use AAVrh10 across our initial programs and will continue to evaluate the latest scientific understanding of capsid technology for each of our future programs. We will continue to pursue the development of novel capsids, promoter technologies, routes of delivery and other next-generation capabilities to optimize the potential therapeutic efficacy of our programs through our own efforts and in collaboration with others.
- *Leverage and expand upon our partnerships and exclusive licenses with world-class academic institutions.* Our foundational science stems from partnerships and exclusive licenses with leading academic laboratories at Weill Cornell Medicine and UCSD, two preeminent institutions on the cutting edge of gene therapy research. Of note, our Chief Scientific Advisor, Ronald G. Crystal, M.D., has sponsored 14 cleared gene therapy IND applications across multiple disease areas, and researchers at UCSD led the discovery efforts of a cardiovascular gene therapy program that is expected to move into late-stage clinical trials. We will continue to draw on the scientific expertise provided by these partnerships while evaluating new opportunities for complementary research and development with other academic collaborators.
- *Build a fully integrated gene therapy company and selectively evaluate strategic opportunities to maximize the impact of our pipeline.* We aim to discover, develop, manufacture, and eventually commercialize our gene therapy candidates. Despite the larger populations in most of our target indications, we believe genetically defined patient populations enable us to pursue a rare disease commercialization strategy characterized by targeted patient identification and facilitating an efficient patient journey through the diagnostic process, engagement with patient advocacy groups, the potential for value-based market access efforts, and sales, account management, and field reimbursement teams focused on the small number of centers that typically diagnose and treat genetic diseases. We may seek strategic collaborations where we believe the resources and expertise of third-party pharmaceutical or biotechnology companies could accelerate the clinical development or maximize the market potential of our product candidates, or where such collaborations could expand our internal capabilities and platform technology.



## Our approach

### *Background and successes of gene therapy*

Gene therapy is one of the most important emerging modalities, given its potential, in a single administration, to treat or cure life threatening diseases. Other non-genetic medicine approaches have alleviated certain symptoms and conditions associated with genetic diseases, but they do not directly address the underlying genetic cause of the disease.

Gene therapies are designed to deliver transgenes, which are functional versions of the genes that are mutated or are the cause of deficient proteins that manifest in genetic diseases. These transgenes are utilized by the body's cellular machinery to naturally produce functional proteins that were deficient or non-functional prior to treatment. The production of these functional proteins is intended to provide consistent and durable therapeutic benefit. Gene therapies are typically comprised of three key components: a vector—a vehicle that delivers a transgene to cells in the body; a transgene—a gene intended to produce a functional protein; and a promoter—a specialized DNA sequence that directs cells to initiate transcription. AAV-based gene therapy has been shown to be highly effective in targeting many organ systems and has been studied in more than 3,300 patients worldwide.

Although gene therapies have been studied in human clinical trials for over 30 years, there are few AAV-based gene therapy products approved by the FDA: voretigene neparvovec rzy1 (Luxturna), for the treatment of a rare ocular condition; onasemnogene abeparvovec xioi (Zolgensma), for the treatment of a rare neuromuscular condition; etranacogene dezaparvovec (Hemegenix), for the treatment of adults with Hemophilia B who currently use Factor IX prophylaxis therapy; and nadofarogene firadenovec-vncg (Adstiladrin), for an unresponsive non-muscle invasive bladder cancer. Less than four years after its 2019 FDA approval, Zolgensma generated \$1.4 billion of net product sales in 2022, highlighting the commercial viability of novel gene therapies.

### *Challenges associated with gene therapies*

Although substantial advances have been made in the gene therapy field, numerous challenges remain that are associated with safety, efficacy and manufacturing, including:

- *High doses of systemic gene therapy resulting in toxicity.* Like many other drug modalities, higher doses of gene therapies have been associated with toxicity events. Thus far, systemic gene therapies that are intended to treat larger tissues or organ systems (such as in neuromuscular diseases) have typically required higher doses to demonstrate a therapeutic effect ( $1 \times 10^{14}$  vg/kg and higher). Adverse effects such as liver toxicity and microvascular thrombosis may arise from higher doses of gene therapy, leading to direct injury to hepatocytes and activation of aversive immune responses.
- *Delivery and biodistribution challenges in CNS gene therapy.* Gene therapies for CNS indications may cause dorsal root ganglia pathology due to a preferential expression profile in dorsal root ganglia if administered to the CNS. In addition, gene therapies administered to the CNS must overcome the challenge of insufficient biodistribution to neurons and other cells of the CNS in the case of therapies that target intracellular proteins, diminishing their potential efficacy.
- *Lack of scalable, high-quality manufacturing.* Early-stage manufacture of AAV gene therapy has traditionally relied on the use of HEK adherent and suspension cell culture using plasmid transfection and subsequent ultracentrifugation-based purification. This approach involves the use of expensive plasmids and results in a long production cycle. Additional issues include low yield and quality of the material produced, including incorporation of non-transgene DNA impurities, ratio of empty-to-full capsid and inefficient removal of process residuals. It has been reported that empty capsids and DNA impurities can cause innate and adaptive immune responses to AAV and even inhibit transduction by filled capsids. Furthermore, the HEK adherent and suspension cell culture approach has proven difficult to scale up.

## ***Our gene therapy approach***

Our integrated modular approach enables us to optimize our strategy to pursue larger-rare and prevalent genetically defined indications in specific sub-groups of patients. Our gene therapy candidates utilize the vector construct, dose and route of administration that we believe will result in the most favorable biodistribution profile for each disease. By specifically tailoring our technological approach to each targeted disease, we believe we can optimize treatments to achieve the highest likelihood of having potential therapeutic impact. Our disease area strategy is focused on defined sub-groups within selected cardiovascular diseases and *APOE4*-associated Alzheimer's disease that we believe are most amenable to gene therapy. Across our portfolio, we are utilizing a scalable baculovirus/Sf9 expression system that has demonstrated the capability to produce high-quality, high-yield, and high-potency vectors that can accommodate demand for both rare and prevalent patient populations. Finally, we continuously seek to bolster our development pipeline with relationships with academic institutions, which provide us access to cutting edge gene therapy research that we will utilize in the discovery and development of next generation gene therapy candidates.

## ***Our technology approach***

Our most advanced cardiac and *APOE4*-associated Alzheimer's disease programs use the AAVrh10 vector due to its high transduction efficiency in both myocardial cells and neurons, higher ratio of cardiac to liver biodistribution, potentially lower toxicity given the ability to utilize lower doses, compared to other well-established AAV serotypes, and lower pre-existing immunity. We believe that our approach to technology has the potential to confer the following advantages over existing gene therapy technologies:

- ***High Transduction Efficiency and Biodistribution.*** The AAVrh10 vector has been shown to be capable of transducing myocardial cells based on preclinical research, and it has been shown to be capable of transducing neurons based on results from preclinical studies and our ongoing clinical trials. We believe these results demonstrate that among currently available, commonly used serotypes, AAVrh10 is an efficient vector for delivery and expression of transgenes for the treatment of the cardiovascular diseases that we are currently targeting and *APOE4*-associated Alzheimer's disease. We have observed organ-specific biodistribution advantages for our AAVrh10 vector compared to AAV9, a well-known serotype that has been evaluated in several clinical trials, including one cardiovascular clinical trial. For example, we have reported vector distribution of AAVrh10 in cardiac tissue of non-human primates, or NHPs, that is two times greater than for AAV9. Additionally, our preliminary studies in Yucatan minipigs have shown that AAVrh10 has approximately 1.5 times greater biodistribution in the heart than AAV9. Preclinical data from a study using a murine model conducted by other researchers has demonstrated that more AAVrh10 particles transduced a single brain cell than AAV9. In addition, preclinical studies conducted by Weill Cornell Medicine have demonstrated that systemic administration of AAVrh10 promotes a ratio of cardiac-to-liver biodistribution that is more favorable than what is known of other commonly used vector serotypes.
- ***Reduced Toxicity.*** The cardiac tropism of AAVrh10 may allow our gene therapy candidates to be systemically administered at lower doses than many other AAV-based therapies targeting cardiovascular or other systemic diseases. We have selected target indications that we believe will be responsive to AAVrh10-based therapies administered at comparatively lower doses, which will potentially reduce toxicities that have been reported in clinical trials of higher-dose gene therapies. For example, compared to doses used in other systemic gene therapy programs which can exceed  $1 \times 10^{14}$  genome copies per kilogram, we intend to use doses of low to mid  $\times 10^{11}$  genome copies per kilogram in our Phase 1/2 clinical trial of LX2006 for the treatment of FA cardiomyopathy. Similarly, early data suggest a lower dorsal root ganglia toxicity profile than other commonly used vectors, making it amenable for treating *APOE4*-associated Alzheimer's disease at potentially higher doses than other vectors.

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- *Reduced Pre-Existing Neutralizing Antibodies.* Treatments leveraging vector serotypes to which humans have pre-existing immunity tend to be less effective. Among the naturally occurring and commonly used AAV serotypes, AAVrh10 has been shown in preclinical studies to have among the lowest levels of pre-existing neutralizing antibodies.
- *Optimized Expression.* We are collaborating with our academic partners to develop tissue-specific promoters and enhancers, tissue-specific codon optimization and tissue-specific, miRNA-based knockdown which can be utilized to limit transgene expression in non-target organs. We are also exploring additional genetic medicine approaches to increase tissue tropism using novel capsids and developing next generation expression systems to optimize the potential therapeutic efficacy of our product candidates.

### *Our precision medicine approach*

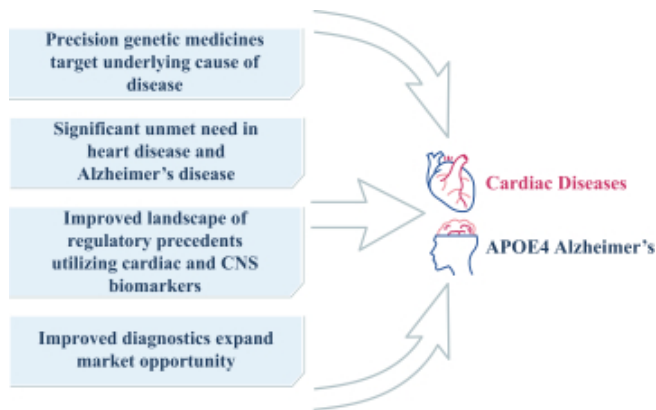
We are focused on disease areas that have seen limited penetration of precision medicines, which we define as medications that treat the underlying molecular mechanism of a disease. The development of precision medicines has transformed treatment paradigms in certain therapeutic areas. For example, the top five precision medicines approved in oncology between 2016 and 2021 have a combined estimated U.S. annual peak sales of approximately \$25 billion. Conversely, relatively few precision medicines have been approved for treating cardiac or central nervous system conditions; this is despite a rapid advancement in the understanding of the genetic underpinning of cardiovascular and neurologic diseases and the improved ability of viral vectors to transduce the heart and brain safely.

Cardiovascular disease remains the most significant cause of morbidity and mortality in the western world and is rapidly becoming a primary cause of death worldwide. We believe the current lack of precision medicines across the cardiovascular landscape is due to historical regulatory precedents focused on “hard endpoints” and delivery technology that has been insufficiently cardiotropic. However, recent approvals by the FDA suggest a willingness to approve new therapies based on biomarkers and functional endpoints. Furthermore, modern AAV delivery technology generally targets the heart and mediates the delivery of genetic payloads. These developments should foster the introduction of genetic medicines in the cardiovascular treatment landscape, offering one of the most substantial opportunities for the uptake of precision medicines in the global pharmaceutical marketplace. There is significant potential for cardiovascular precision medicine. For example, approximately 30% of cases of dilated cardiomyopathy, a common cause of heart failure, stem from genetic mutations in which more than 50 distinct genes have been implicated. Advancements in genome sequencing have led to increasing identification of these genetic mutations over the last two decades. In the majority of genetic cardiomyopathies, there is no existing disease-modifying therapeutic option. In cases where a disease-modifying treatment option is available, the current standard of care only forestalls disease progression. We believe our cardiovascular precision medicine approach offers the opportunity to treat the underlying root cause of the disease and more effectively reverse disease progression with a single dose.

Precision medicine has been more relevant across CNS disorders, in particular gene therapies for ophthalmologic and neuromuscular degeneration. However, there has been limited advancement in developing precision medicines for Alzheimer’s disease, the most significant commercial opportunity within the CNS landscape. *APOE* is the leading genetic risk factor for Alzheimer’s disease, and, despite differences in treatment effect and safety seen between subpopulations in recent clinical trials, currently-approved, disease-modifying treatments for Alzheimer’s disease utilize a “one-size-fits-all” approach, regardless of *APOE* subpopulation. We believe this highlights the unmet need for precision medicines as the Alzheimer’s disease treatment paradigm continues to shift.

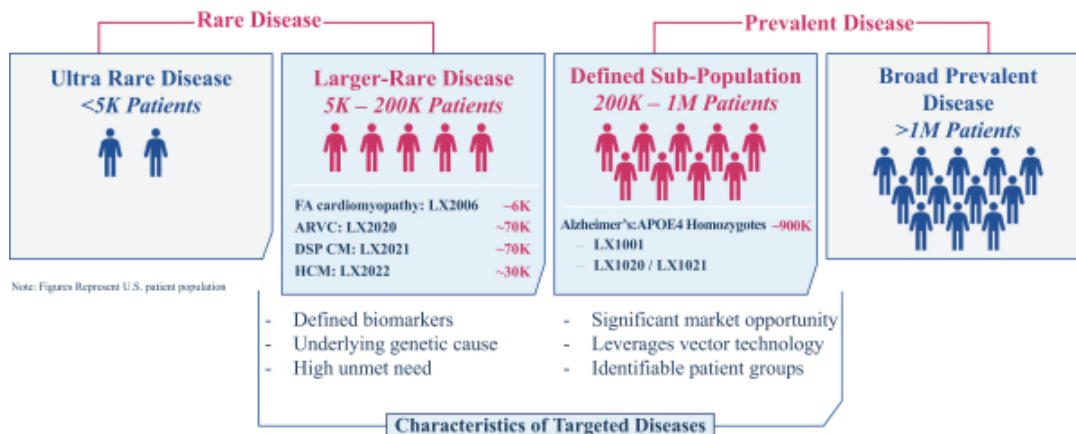
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We believe a precision medicine approach for cardiovascular diseases and APOE4-associated Alzheimer’s disease represents a compelling opportunity and the next frontier in expanding the impact of genetic medicines across therapeutic areas:



- *Precision genetic medicines target the underlying cause of the disease:* next-generation, precision-based genetic medicines offer the opportunity to directly target the mechanisms underlying cardiovascular disease and Alzheimer’s disease.
- *Significant unmet need in cardiovascular disease and APOE4-associated Alzheimer’s disease:* more than 30 million adults have been diagnosed and approximately 655,000 Americans die yearly from heart disease. Additionally, we believe there are approximately 900,000 APOE4 homozygous patients in the United States, and current treatment options are likely to be less effective for this subpopulation of patients.
- *Improved landscape of regulatory precedents utilizing cardiac and CNS biomarkers:* biomarkers can help accelerate development timelines, reduce patient attrition, and reduce the overall cost of drug development by providing an early read on potential efficacy and therapeutic impact.
- *Improved diagnostics expand market opportunity:* implementation of genetic testing has the potential to help identify patients, including those who are asymptomatic or at a higher likelihood of developing the disease being evaluated or who respond less favorably to currently approved treatments.

*Our disease area strategy*



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Our cardiovascular and *APOE4*-associated gene therapy programs are designed to have the following characteristics:

- *Indications that may be effectively treated by gene therapy.* We select targets that correspond to populations with a specific genetic profile and clearly defined disease phenotype, increasing the homogeneity of our studied patient populations and the potential for the early clinical observation of a beneficial effect.
- *Indications with the potential to demonstrate early evidence of meaningful clinical benefit.* We pursue clearly defined biomarkers and functional endpoints that can provide early proof-of-mechanism and inform clinical development decisions, including the potential to seek accelerated approval pathways. Early data regarding functional endpoints and biomarkers allows us to pursue an efficient capital allocation strategy. Certain of our clinical trials are designed to provide initial biomarker data as early as three months from the treatment date. In the case of LX2006 targeting FA cardiomyopathy, we are also pursuing functional and histological endpoints that may demonstrate signs of meaningful clinical benefit as early as three months. We expect these early signals will enable us to design late-stage clinical studies with endpoints that best mirror therapeutic effects that should cure the disease. In addition, we believe these endpoints should allow us to perform trials smaller than traditional cardiac trials that can be completed in a reasonable time frame.
- *Present opportunity to address high unmet medical need.* We are focused on genetically defined cardiovascular diseases and *APOE4*-associated Alzheimer's disease where there are no currently approved treatment or where we believe our therapeutic candidates will have a meaningful improvement relative to existing standards of care.
- *Significant market opportunity.* We seek indications with significant commercial opportunities beyond those typically associated with gene therapy companies targeting rare monogenic diseases. Our current focus is on targets that impact larger-rare disease populations, ranging from approximately 5,600 U.S. patients, in the case of FA cardiomyopathy, to approximately 60,000 U.S patients, in the case of ARVC due to *PKP2* mutations, or prevalent diseases such as homozygous *APOE4*-associated Alzheimer's disease, which has a patient population of up to approximately 900,000 in the United States.
- *Targets that have established proof-of-concept.* We have leveraged our relationships with academic institutions including Weill Cornell Medicine and UCSD to in-license or acquire product candidates with established proof-of-concept in relevant preclinical models that closely resemble the clinical phenotype. We will continue to seek additional similar opportunities via collaborations with academia and the industry.
- *Targeted disease areas best treated by optimal delivery technologies.* Our current gene therapy candidates utilize the AAVrh10 vector due to its tropism for the heart and its observed tolerability in clinical trials for CNS disorders. We will pursue the optimal technology to address the diseases of interest for future indications while ensuring sufficient preclinical or clinical validation for any novel approaches.
- *Readily identifiable patient sub-groups.* Our goal is to accelerate patient recruitment for our clinical trials and increase the likelihood of commercial success of our potential products by focusing on diseases with established patient advocacy groups and university researchers who maintain registries of potentially eligible patients. Where possible, we leverage existing natural history studies to better define target patient phenotypes associated with the disease and may utilize these natural history datasets as a control group in future studies.

### *Our manufacturing approach*

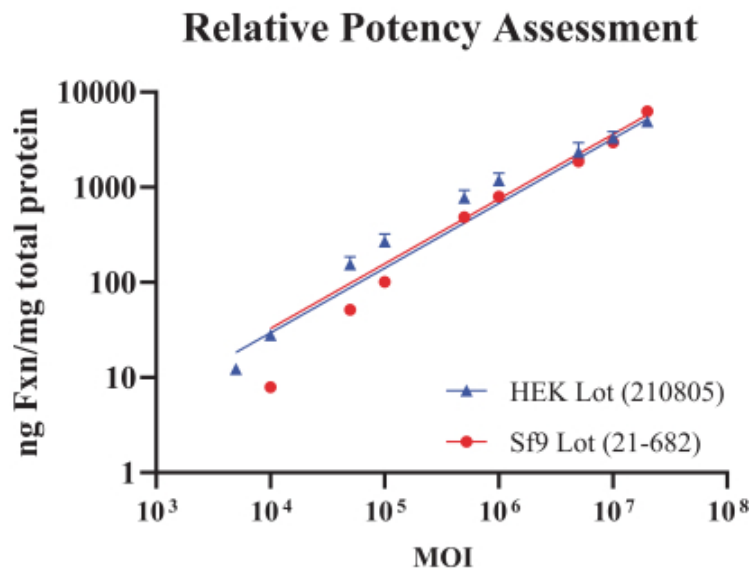
We are developing gene therapy candidates for larger-rare and prevalent disease patient populations, which require a high-quality process that can produce vectors in relatively large quantities while utilizing traditional biologics manufacturing infrastructure. We utilize a baculovirus/Sf9 expression system to manufacture our gene therapy candidates. Our manufacturing platform is designed to infect Sf9 cells at high densities in suspension cell culture with both an AAVrh10 and baculovirus containing the transgene. The output is coupled with a

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chromatography-based purification process which allows for efficient AAV purification, resulting in higher yields and fewer empty AAV capsids than traditional plasmid HEK adherent cell culture approaches. Furthermore, we believe our process has an improved safety profile over traditional adherent HEK manufacturing for the following reasons: we have observed reduced incorporation of non-transgene DNA plasmid DNA impurities, from 15% observed in some HEK systems to 0.2% in our process; our cellular components are non-replicating in mammalian cells; we have eliminated potentially immunogenic or toxic animal-derived proteins; and our cells are grown under serum-free conditions, leading to reduced risk of contamination from animal-derived products.

We believe our manufacturing process enables us to efficiently pursue our goal of targeting larger-rare and prevalent patient populations. Historically, manufacturing challenges, largely driven by the quantity of vector required to pursue large commercial opportunities have limited the utility of gene therapy for large patient populations. We believe our proprietary SF9 baculovirus process will allow us to produce vectors at the necessary scale to support the patient populations we are targeting at a cost-of-goods profile similar to what the pharmaceutical marketplace has seen with biologics. Based on our estimates, we believe our manufacturing process is approximately 10 times more cost efficient than traditional adherent HEK manufacturing, allowing us to potentially produce clinical and commercial material in a more cost efficient manner than peers relying on first-generation manufacturing approaches. We expect that cGMP manufacture of our product candidates using this platform will begin in the second half of 2023.

First generation SF9 baculovirus platforms have historically been less infectious than traditional HEK systems, which is associated with lower levels of the capsid viral protein 1, or VP1, one of the three viral proteins that make up the AAV capsid. However, we have implemented the Virovek “Bac to AAV” technology which enables wild type levels of VP1 allowing us to achieve potency levels largely in-line with a HEK approach. This is shown in the graphic below from an *in-vitro* potency assessment of two lots of LX2006 comparing HEK and SF9 which resulted in an equivalent potency between manufacturing methods.



As representative examples of our process, we have been able to achieve purified yield levels of greater than  $1 \times 10^{14}$  vector genomes per liter of cell culture in a 50 Liter run manufactured for our LX2020 IND-enabling murine efficacy and NHP safety studies with approximately 70% full capsid levels. We believe these levels demonstrate the feasibility of our process and our ability to provide late-stage clinical and commercial material in a cost-efficient manner.

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Based on our approach, we expect that our existing partnerships can supply material for all of our currently ongoing and planned clinical trials as well as potential commercial production of some of our programs. We have secured relationships with Virovek, Inc. for its “Bac-to-AAV” system, Millipore Corporation for its RVN Sf9 cell line, and Fujifilm Diosynth Biotechnologies U.S.A., Inc. for viral vector process development and cGMP manufacture of viral vectors. We will own the intellectual property created by our manufacturing process development activities or have the ability to license it and will maintain the option to transfer the process to other CDMOs in the future and/or to our own potential facility to ensure ongoing redundancy and reliability.

### *Academic collaborations*

To support our integrated modular approach, we have partnered with leading academic institutions who are on the cutting edge of gene therapy research. Our relationships with Weill Cornell Medicine and UCSD have provided us with access to the latest gene therapy research. Our collaboration efforts with Weill Cornell Medicine currently focus on the discovery of second- and third generation cardiac vector technology and novel cardiac transcriptional promoters, all of which have the potential to be deployed in our early-stage research efforts. Our collaboration with UCSD focuses on preclinical candidate selection and translational research in several larger-rare cardiovascular disease indications. Both institutions are recognized leaders in gene therapy, and the collaborations are designed to best leverage their respective skill sets. We may seek additional academic collaborations where such collaborations could expand our internal capabilities and platform technology.

## **Our cardiovascular gene therapy programs**

We have combined our robust expertise in cardiac-targeted vector design, regulatory and clinical strategy, and disease area knowledge to establish a deep cardiovascular gene therapy pipeline. We are developing a number of disease-modifying gene therapy candidates to treat larger-rare cardiovascular diseases that have significant unmet need and no approved disease-modifying treatments. Our most advanced program, LX2006, is an AAVrh10-based gene therapy candidate for the treatment of FA cardiomyopathy caused by mutations in the *FXN* gene. We are also advancing several other AAV-based gene therapy programs to treat additional genetically defined cardiac diseases, including LX2020 to treat ARVC caused by mutations in the *PKP2* gene, LX2021 to treat DSP Cardiomyopathy associated with Cx43 deficiency, and LX2022 to treat HCM associated with mutations in the *TNNI3* gene. We plan to leverage our experience and learnings from LX2006 and apply them to our earlier-stage cardiac programs.

### ***LX2006 for the treatment of FA cardiomyopathy***

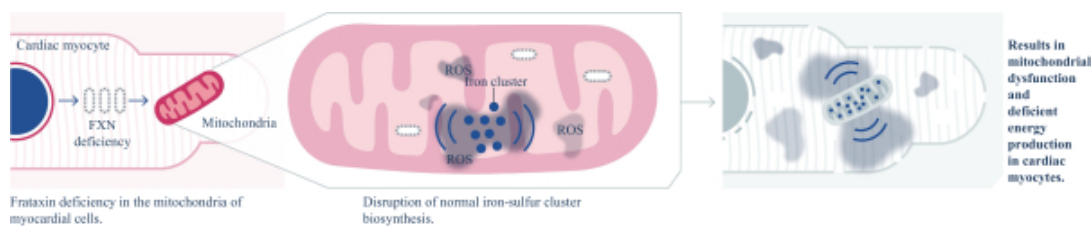
We are developing LX2006, an AAVrh10-based gene therapy candidate for the treatment of FA cardiomyopathy that is designed to deliver functional *FXN* intravenously to promote the expression of frataxin in order to restore normal mitochondrial function in myocardial cells. Cardiomyopathy is the most common cause of mortality in patients with FA. The FDA has granted Rare Pediatric Disease designation and Orphan Drug designation to LX2006 for the treatment of FA cardiomyopathy. LX2006 is currently being evaluated in an open-label, dose-escalation Phase 1/2 clinical trial in patients with FA cardiomyopathy with biomarker data expected in 2024.

### *Overview of Friedreich’s ataxia*

Friedreich’s ataxia is a genetic, progressive, and degenerative multi-system disorder with a prevalence of 1:40,000 or approximately 8,000 people in the United States. It is estimated that approximately 70%, or 5,600 of these patients, will develop FA cardiomyopathy. In the European Union, FA affects approximately 11,000 people, with approximately 8,000 of these cases projected to include FA cardiomyopathy. FA is caused by a mutation in the *FXN* gene that disrupts the normal production of the protein frataxin, which is critical to the

function of mitochondria in a cell and to the maintenance of cardiac function. FA patients with classically presenting disease have peripheral, non-cardiac tissue frataxin protein levels that range from 2% to 30% of normal levels. Carriers who do not develop symptoms of FA generally have peripheral tissue frataxin protein levels that range from 30% to 80% of normal levels. This frataxin deficiency in the mitochondria of myocardial cells causes disruption of normal iron-sulfur cluster biosynthesis leading to mitochondrial dysfunction and deficient energy production, as shown below.

### Overview of FA cardiomyopathy disease mechanism

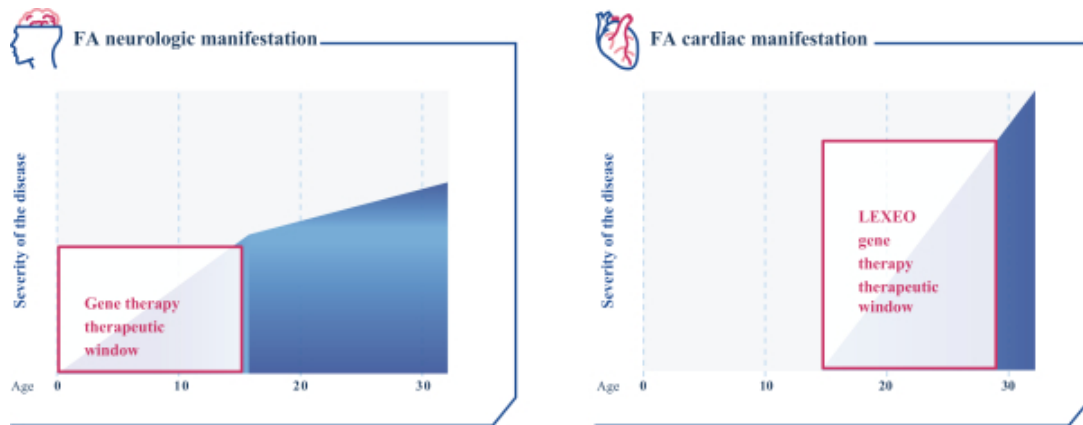


The neurologic disease and cardiac disease are two distinct manifestations of FA. The disease is inherited in an autosomal recessive manner, where both inherited genes are abnormal, and symptoms usually begin in childhood. Absence of fully functional frataxin leads to damage to peripheral nerves and the parts of the brain that controls movement and balance, leading to neurological symptoms that include impaired muscle coordination, or ataxia, that worsen over time. Initial symptoms may include unsteady posture, frequent falling, and progressive difficulty in walking due to impaired ability to coordinate voluntary movements. Affected individuals often develop slurred speech, hearing loss, scoliosis, diabetes, characteristic foot deformities, and an irregular curvature of the spine. The typical age of onset of neurological symptoms is five to 15 years old. As the disease progresses, patients typically experience various heart conditions, including thickening of the heart muscle, or hypertrophic cardiomyopathy, and arrhythmias. Hypertrophic cardiomyopathy, fibrosis, heart failure and arrhythmias are the cause of death in approximately two-thirds of FA patients. Typical onset of the cardiac disease is 15 to 30 years old.

By the time the cardiac disease of FA emerges, the neurologic disease is generally significantly advanced and may not be amenable to gene therapy. The optimal patient age for treatment of the neurologic disease is between five and 15 years of age while the optimal patient age for demonstrating treatment effect of the cardiac disease is 15 to 30 years of age. There are currently no approved treatments for the cardiac manifestations of FA. *Skyclarys*, the only approved treatment for FA, is an Nrf2 activator which only targets the CNS component of the disease. As a result, patients with FA have significant unmet need due to cardiovascular complications associated with the disease.



**Neurological disease and cardiac disease are distinct manifestations of Friedreich's ataxia with different potential therapeutic treatment windows**

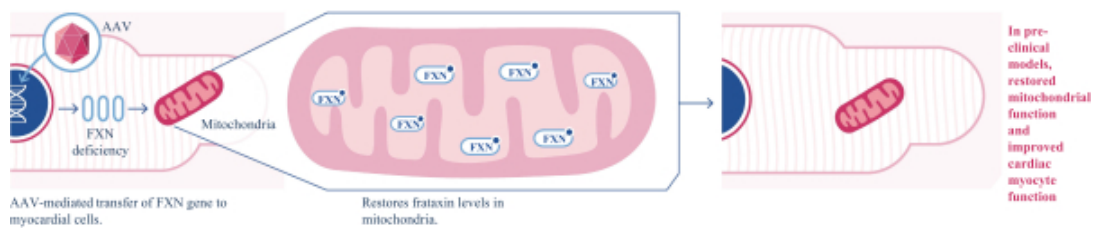


*Our solution: LX2006*

We are developing LX2006 as an AAVrh10-based gene therapy delivered intravenously for the treatment of FA cardiomyopathy. LX2006 is designed to deliver the *FXN* gene under the transcriptional control of the CAG promoter, a strong synthetic promoter frequently used in viral vectors. LX2006 utilizes AAVrh10 based on its favorable cardiac affinity and vector distribution profile observed in preclinical studies, as compared to AAV9. In preclinical studies, vector distribution in cardiac tissue with AAVrh10 has been observed to be between 1.5 to 2 times the distribution associated with the use of AAV9. In addition, there is a lower rate of pre-existing antibodies with AAVrh10 when compared to other commonly used vectors, such as AAV9 and AAV2.

LX2006 is designed to transfer the *FXN* gene to myocardial cells and increase frataxin levels in the mitochondria. The increase in frataxin levels in the mitochondria restores mitochondrial function and energy production in cardiac myocytes, as shown below.

**LX2006 is designed to deliver functional frataxin in order to restore normal mitochondrial function**

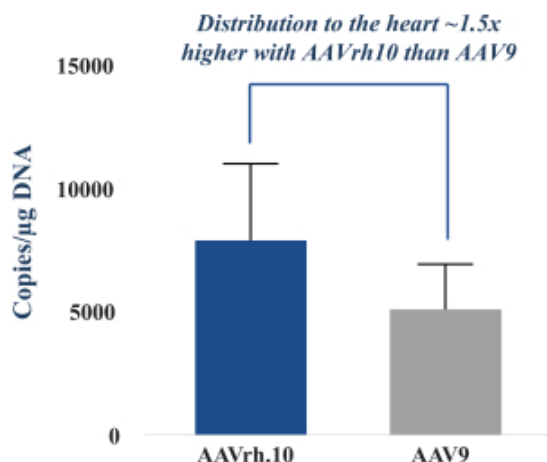


*AAVrh10 demonstrates cardiac tropism*

We have conducted preliminary studies in Yucatan minipigs which have shown that AAVrh10 has a greater biodistribution in the heart compared to AAV9 in this model. The porcine heart is very similar in size, anatomy, and function to the human heart, and we believe it represents an excellent animal model to compare biodistribution across vectors in the heart. A reporter protein, green fluorescent protein, or GFP, is the most commonly used tool to evaluate the biodistribution and expression of a vector in a preclinical study and has well established assays to track the protein. By making GFP constructs in AAVrh10 and AAV9, we are able to

evaluate the biodistribution and transgene expression throughout the different regions of the heart for each vector serotype. AAVrh10-GFP and AAV9-GFP (approximately  $1 \times 10^{13}$  gc/kg) were intravenously administered to three minipigs per group and evaluated two weeks after injection. Vector biodistribution in various regions of the heart were evaluated with qPCR using GFP-specific primers. As shown in the graphic below, AAVrh10 demonstrated higher distribution to the heart compared to AAV9.

**Biodistribution of AAVrh10 and AAV9 in Yucatan minipigs**



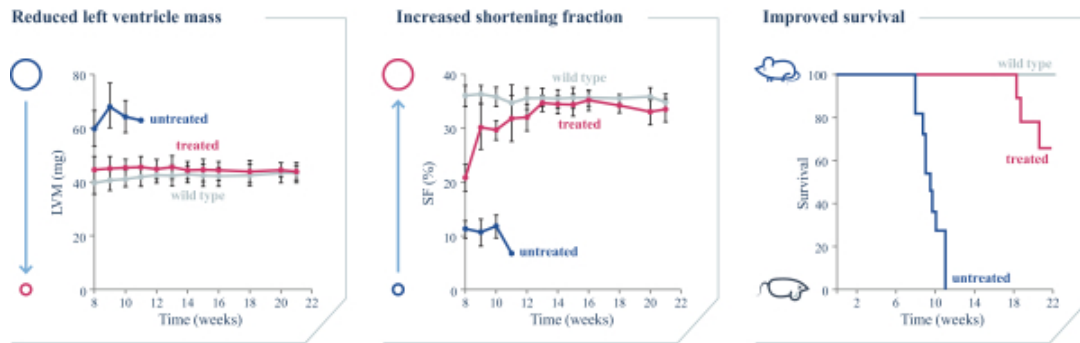
Previous preclinical studies in NHPs conducted by Weill Cornell Medicine demonstrated that biodistribution to the heart was approximately two times higher with AAVrh10 than AAV9. We believe findings from NHP models, the Yucatan minipig model, and murine models conducted by UCSD support our belief that AAVrh10 is a better vector than AAV9 for the cardiovascular diseases we are targeting.

*Preclinical studies in FA mouse models*

LX2006 has demonstrated the ability to significantly reverse the cardiac phenotype in preclinical studies. The ability to restore FXN expression levels using an AAVrh10 vector has been established in numerous preclinical studies of FA conducted internally and by third parties. The most frequently used preclinical mouse models for FA are the MCK model which represents the most severe phenotype of FA cardiomyopathy, and the Myosin heavy chain, isoform, or aMyHC, model, which represents a milder phenotype.

In the preclinical studies conducted by Weill Cornell Medicine utilizing the MCK model, LX2006 reduced left ventricle mass, increased shortening fraction, or pumping ability of the heart, and increased survival, as shown in the graphic below. The MCK model completely lacks the myocardial FXN protein, and untreated mice have a lifespan of approximately 11 weeks of age due to heart failure. The untreated mice cohort developed left ventricular hypertrophy starting at four to five weeks of age, which is associated with a rapid and progressive geometric remodeling of the heart. This cohort exhibited decreased systolic function at eight weeks of age, which led to a severe decrease in resting cardiac output. When administered to the MCK model at three weeks, LX2006 prevented the onset of the disease. In addition, when administered at seven weeks, LX2006 reduced hypertrophy and normalized heart function, based on shortening fraction, within five weeks, and improved survival, as shown in the graphic below. Based on these results, we believe LX2006 has the potential to improve cardiac function and reverse the disease abnormalities in FA cardiomyopathy patients.

**LX2006 improved cardiac function and survival in severe FXN knockout model**

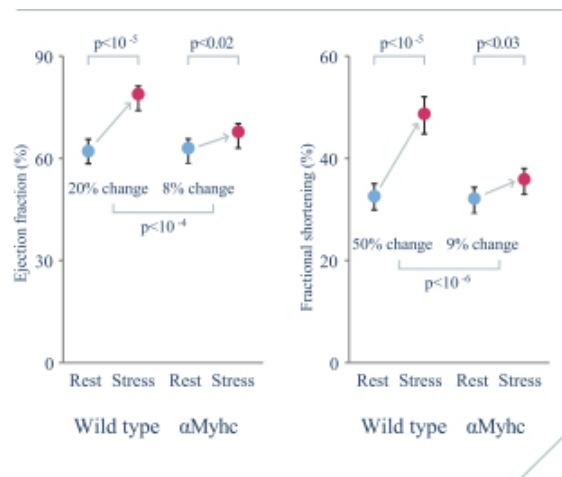


Wild type mice, n=10; untreated MCK mice, n=9; and treated MCK mice, n=9. The treated MCK mice were given a dose of  $5.4 \times 10^{13}$  vg/kg at seven weeks of age.

Additionally, we have completed preclinical studies of LX2006 using an aMyHC mouse model. The aMyHC model has myocardial frataxin protein levels that are less than 50% of normal frataxin expression levels and present with a milder cardiac phenotype. When the mouse is given dobutamine to induce cardiac stress, the cardiac disease becomes apparent, similar to the mild phenotype in FA cardiomyopathy patients at an early stage of their cardiac disease.

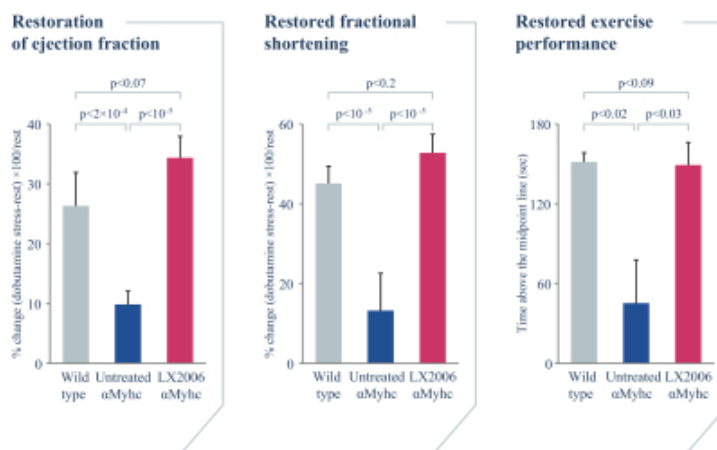
The figure below shows that aMyHC mouse models displayed ejection fraction and fractional shortening levels similar to the wild type phenotype when at rest. However, when both wild type and aMyHC animals are stressed by the administration of dobutamine, the model demonstrated that ejection fraction and fractional shortening levels for the aMyHC mice were below wild type levels.

**aMyHC mouse model demonstrated diminished cardiac function in response to induced stress**



In the aMyHC mouse model, as demonstrated in the graphic below, intravenous administration of LX2006 at a dose of  $4 \times 10^{12}$  vg/kg resulted in improvements in the cardiac phenotype as indicated by correction of dobutamine stress-induced ejection fraction and fractional shortening as well as improvement in treadmill performance. These results show that LX2006 improved key cardiac function measures to wild type levels, helping reverse the pathology of FA cardiomyopathy and suggesting that it could potentially improve cardiac function in FA patients.

**LX206 reversed  $\alpha$ MyHC cardiac phenotype in key cardiac functional measures in preclinical models**



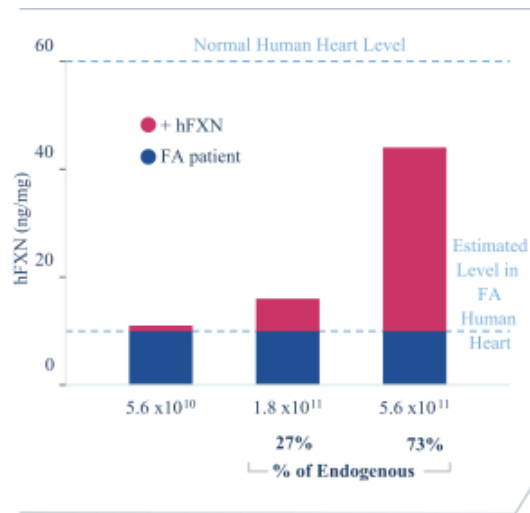
*Preclinical dose range finding study*

Differences between FA patients and phenotypically normal carriers of FA mutations suggest that achieving approximately 30% of normal endogenous frataxin levels would have a potential clinical beneficial effect in FA patients.

Seven-week-old MCK mice were administered increasing doses of 5.6 x10<sup>10</sup> vg/kg, 1.8 x10<sup>11</sup> vg/kg and 5.6 x10<sup>11</sup> vg/kg of LX206 and evaluated for efficacy as a measure of body weight improvement, survival and cardiac function, and associated myocardial FXN expression at one month post dose. MCK mice showed a dose-dependent improvement in efficacy measures with LX206. In these mice, the dose of 1.8 x10<sup>11</sup> vg/kg resulted in trends of improvement in body weight and cardiac function survival, and the level of myocardial frataxin was 27% of normal human endogenous levels. The dose of 5.6 x10<sup>11</sup> vg/kg resulted in statistically significant improvements in body weight, cardiac function, and survival, and the level of myocardial frataxin expression was 73% of normal human endogenous levels.

To establish the endogenous levels in the myocardial cells of non-FA subjects, cardiac samples were obtained during autopsy from five different individuals who did not have FA. The average level of myocardial frataxin measured in non-FA patient samples was approximately 60 ng/mg protein (represented by the upper light blue dashed line in the figure below). In FA patients, an average of 16% of normal endogenous levels would lead to an estimate of 10 ng/mg protein in myocardial cells (represented by the lower light blue dashed line and the blue bars in the figure below).

**Dose-dependent FXN expression in MCK mouse model**

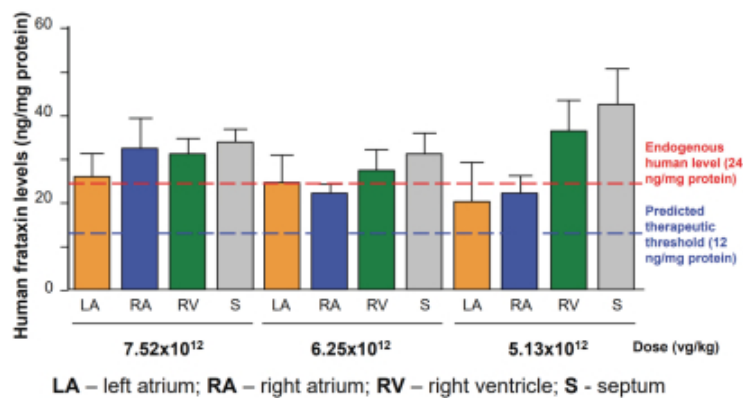


Since the myocardial FXN expression in response to LX2006 treatment in a FA patient’s stressed heart is expected to be similar to the high level of expression in MCK mice observed upon administration of LX2006, dose selection for first-in-human study was based on the MCK mouse data. The  $1.8 \times 10^{11}$  vg/kg dose, which showed 27% of normal myocardial FXN expression, was identified as the minimally effective dose, and the  $5.6 \times 10^{11}$  vg/kg dose, which showed 73% of myocardial FXN expression, was identified as the significantly effective dose.

*Preclinical expression study*

In a separate NHP study conducted at a third-party lab in cynomolgus monkeys that were administered in increasing doses ranging from  $5.13 \times 10^{12}$  to  $7.52 \times 10^{12}$  vg/kg of LX2006, the frataxin expression in myocardial tissue at three weeks following administration exceeded the predicted therapeutic threshold and, in some cases, was higher than the endogenous FXN level in humans, in the atrium, septum and right ventricle, as shown in the figure below. This data was presented at the 2021 American Society of Gene and Cell Therapy annual meeting.

**LX2006 led to frataxin levels in NHP Cardiac Tissues that exceed predicted therapeutic levels**



*Preclinical safety studies*

In a safety study of African green monkeys, increasing doses of LX2006  $1.2 \times 10^{12}$  vg/kg and  $3.9 \times 10^{12}$  vg/kg resulted in dose-dependent increases in myocardial frataxin expression three months post dose. The dose of  $1.2 \times 10^{12}$  vg/kg resulted in FXN expression which is 22% of normal human FXN, or hFXN, levels, while the dose of  $3.9 \times 10^{12}$  vg/kg resulted in FXN expression which is 45% of normal hFXN levels. Both doses demonstrated favorable safety profiles, including no effects on cardiac pathology, physiology and echocardiography. The liver showed minimal to moderate vacuoles, or membrane-enclosed spaces in cells, not considered clinically significant, with no evidence of liver injury. No dorsal root ganglia toxicity was observed.

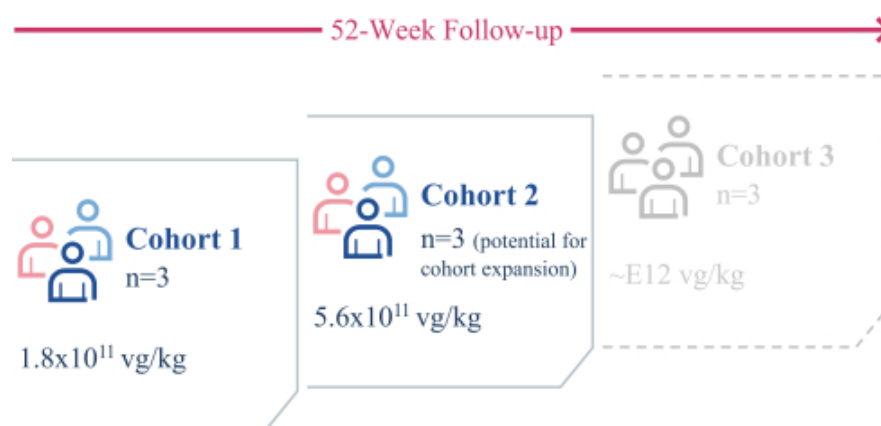
The safety of LX2006 was also tested in seven-week-old wild type mice administered  $1.2 \times 10^{12}$ ,  $3.7 \times 10^{12}$ , and  $1.2 \times 10^{13}$  vg/kg for one and three months, and  $5.6 \times 10^{11}$ ,  $1.8 \times 10^{12}$ , and  $5.6 \times 10^{12}$  vg/kg for 10 months. Animals in the one- and three-month cohorts survived to their scheduled terminations, while those in the 10-month cohort had body weight loss at two time periods that necessitated early euthanasia. The first timepoint was approximately six weeks for four of 12 males in the  $5.6 \times 10^{12}$  vg/kg 10-month cohort; the weight loss was attributed to liver toxicity, which may have been related to high levels of liver hFXN, although hFXN levels were not measured in these mice. No adverse effects occurred in females up to three months post-dose. At 30 weeks post dose, the remaining animals in the high dose ( $5.6 \times 10^{12}$  vg/kg) group and six of 12 female mice in the mid-dose ( $1.8 \times 10^{12}$  vg/kg) group had minimal to moderate hepatic changes and weight loss necessitating early euthanasia. It is possible that both hepatic and possibly cardiac frataxin levels may have contributed to the weight loss. No dorsal root ganglia toxicity was observed at any doses in any animals.

The dose of  $5.6 \times 10^{11}$  vg/kg in the 10-month cohort was generally well tolerated with no treatment-related clinical signs, body weight changes, hematologic or clinical chemistry findings, nor histopathology findings, except for hepatocellular carcinoma, or HCC, which was observed in one of six males at  $5.6 \times 10^{11}$  vg/kg and in three of six males at  $1.8 \times 10^{12}$  vg/kg. While this finding was considered adverse in the context of the study, data reported by the FDA Cellular, Tissue, and Gene Therapies Advisory Committee in September 2021 suggests that HCC observed in mice after AAV treatment is unlikely to translate to risks for humans, as it has not been observed in higher species or humans. The incidence of HCC has been seen in multiple long-term mouse studies with systemic administration of other AAVs.

*LEXEO-sponsored LX2006 clinical development and trial design*

LX2006 is currently being evaluated in an ongoing Phase 1/2 clinical trial in patients with FA cardiomyopathy. The Phase 1/2 study is a first-in-human, 52-week, dose-ascending, and open-label trial of LX2006 in patients who have FA cardiomyopathy. LX2006 is administered as a one-time intravenous infusion to patients in at least two dose-ascending cohorts. The trial consists of at least three patients in each cohort, with the potential for cohort expansion. A potential third cohort will receive a dose that will be selected based on the dose-response relationships seen in cohort 1 and cohort 2. There will be a long-term follow-up for patients who receive LX2006 to monitor ongoing safety for a total of five years, per FDA requirement. We plan to seek endorsement from the data safety monitoring board, or DSMB, to advance into the second cohort in mid-2023.

### Design of Phase 1/2 clinical trial of LX2006



Key patient inclusion criteria for the study include genotyping, left ventricular hypertrophy and echocardiography, or ECHO, changes, and ejection fraction  $\geq 40\%$ . The primary endpoint of the Phase 1/2 trial is to assess the safety and tolerability of one-time administration of LX2006 for the purpose of selecting the appropriate dose for further clinical development. Exploratory endpoints assess biomarkers, including cardiac frataxin expression levels, and preliminary functional efficacy assessments relevant in evaluating the ability of LX2006 to stop progression and improve FA cardiomyopathy. The following biomarker, functional and imaging efficacy assessments will be conducted:

- **CPET:** Peak VO<sub>2</sub> and other functional measures will be performed at regular intervals.
- **Cardiac MRI:** All patients undergo MRI scans of the heart to measure left ventricular mass index, left ventricular ejection fraction, stroke volume, strain, and fibrosis.
- **Cardiac ECHO:** All patients undergo cardiac ECHO to measure left ventricular mass index, left ventricular ejection fraction and strain.
- **Cardiac biopsy:** Biopsies will measure, among other things, frataxin protein expression.
- **Biomarkers:** Serum cardiac biomarkers, including troponin.

#### *Additional ongoing clinical trial of AAVrh10.hFXN (LX2006 at LEXEO)*

AAVrh10.hFXN, which we refer to as LX2006, is currently being evaluated in an additional ongoing clinical trial conducted by Weill Cornell Medicine (NCT05302271). This is an investigator-sponsored, single-site study, and our trial (NCT05445323) is an industry-sponsored multi-site trial. Both clinical studies utilize drug product manufactured at Weill Cornell utilizing the same process with identical doses in cohort 1 and cohort 2. Safety information is being shared across both clinical studies. We expect to use the combined safety data from patients across both studies to seek DSMB endorsement to advance to the second dose-cohort in our trial.

#### ***LX2020 for the treatment of ARVC caused by PKP2 mutations***

We are developing LX2020 as an AAVrh10-based gene therapy for the treatment of ARVC caused by mutations in the *PKP2* gene. LX2020 is designed to provide a fully functional *PKP2* gene to increase desmosomal PKP2 protein levels, reassemble desmosomes, and prevent cardiac arrhythmias and prevent or treat cardiac dysfunction.

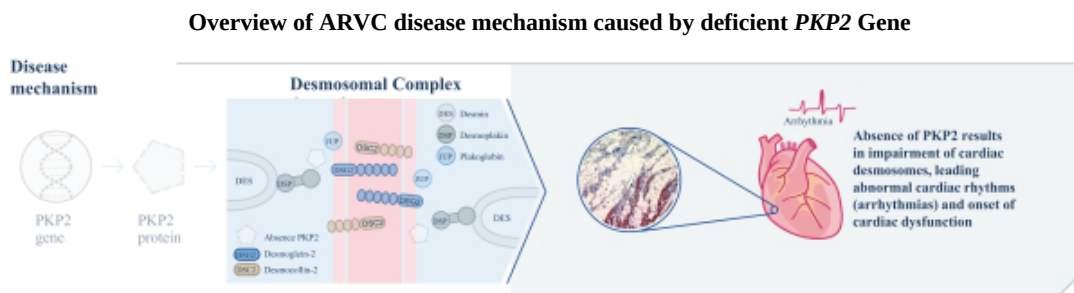
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*PKP2* is the most frequently mutated desmosomal gene, and we believe there are more than 60,000 ARVC patients affected by the *PKP2* mutation in the United States. We intend to submit an IND application to the FDA for LX2020 in mid-2023.

### Overview of ARVC

ARVC is a genetic heart disease primarily characterized by myocardial cell loss and the replacement of heart muscle with fibrotic tissue and fatty deposits. ARVC can result from mutations in several desmosomal genes. These genetic mutations impair the structure and function of cardiac desmosomes, which are membrane protein complexes engaged in cell-to-cell adhesion and the structural integrity of the ventricular myocardium. Lack of functioning cardiac desmosomes can lead to myocardial cell death and fibrosis, heart dysfunction, rhythm abnormalities, and sudden death. The disease is associated with high mortality as more than 40% of ARVC patients die or have heart transplantation within 10 years of diagnosis.

The figure below shows the role of the *PKP2* protein in the formation of the desmosome.



We estimate that ARVC has a prevalence between 66,000 and 165,000 patients in the United States and estimate that over half of all ARVC patients have a genetic form of the disease. Five desmosomal genes account for nearly all of the identified genetic causes of ARVC. We have initially targeted the *PKP2* gene because mutations in this gene are the most commonly known genetic cause of ARVC. We believe that mutations in the *PKP2* gene are associated with approximately 75% of all genetic cases of ARVC, resulting in approximately 60,000 and 75,000 ARVC patients affected by the *PKP2* mutation in the United States and European Union, respectively. Most familial cases of the disease have an autosomal dominant pattern of inheritance, meaning one copy of an altered gene in each cell is sufficient to cause the disorder. Since having only one functioning copy of the *PKP2* gene is insufficient to produce the wild-type phenotype, this results in a phenomenon in genetics known as haploinsufficiency.

Symptoms of the disease can include palpitations, dizziness, heart failure and sudden death. Abnormal function of the right ventricle, fatty or fibrotic infiltrates in the myocardium, abnormal electrocardiogram, or ECG, arrhythmias, or a family history of ARVC can all lead physicians to diagnose the disease. No effective treatments or cures for ARVC exist.

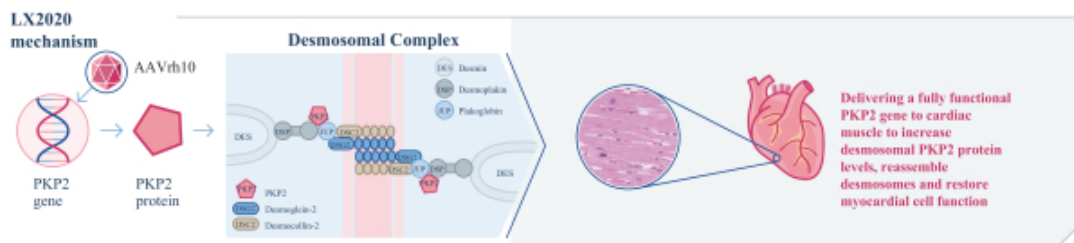
### Our solution: LX2020

We are developing LX2020 as an AAVrh10-based gene therapy candidate for the treatment of ARVC caused by mutations in the *PKP2* gene. LX2020 is designed to intravenously deliver a fully functional *PKP2* gene to cardiac muscle to increase desmosomal *PKP2* protein levels and restore myocardial cell function. We believe that by delivering a fully functional *PKP2* gene, LX2020 has the potential to address the underlying cause of ARVC



for many patients and have a significant effect on lifespan by reassembling the cardiac desmosomes, preventing cardiac arrhythmias and preventing or treating cardiac dysfunction.

**LX2020 is designed to deliver functional PKP2 to restore normal desmosomal function**



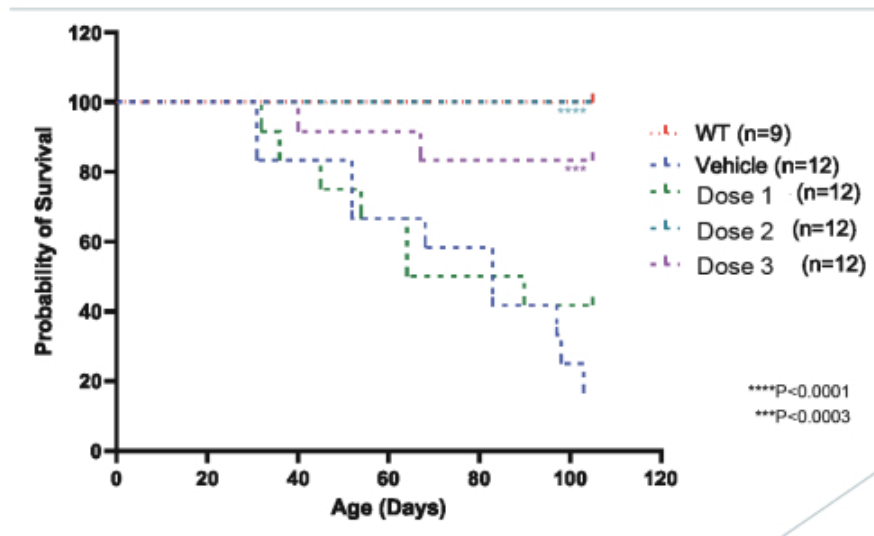
**Preclinical studies**

Our preclinical mice studies have demonstrated that delivery of LX2020 (*AAVRh10-hPK2*) improves cardiac structure, function, and the survival of mice harboring a pathologic human mutation in PKP2. The genetic mouse model we utilize in our preclinical studies is a CRISPR Cas9-edited model constitutively expressing a mutation in the *PKP2* gene found in humans with ARVC. *PKP2* RNA splicing and protein levels are impacted in this model, which recapitulates all classic ARVC disease features. We have also demonstrated that LX2020 has a favorable safety profile in a study of non-human primates with no safety signals.

**Preclinical efficacy studies**

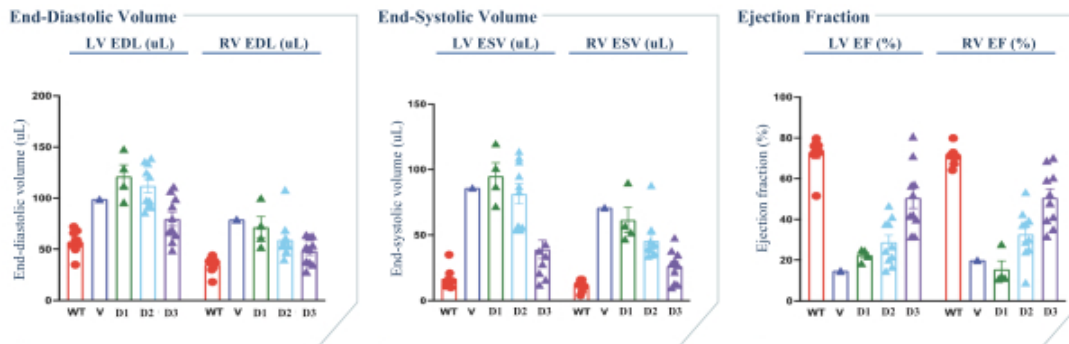
We have completed preclinical studies demonstrating that delivery of LX2020 led to the reassembling of the cardiac desmosome, prevented arrhythmias and cardiac dysfunction, and increased survival in a genetic murine model of ARVC harboring *PKP2* patient genetics. Three dose levels were administered to approximately three-week-old homozygous *PKP2* mice (a more severe disease murine model) with necropsy at twelve weeks post LX2020 treatment and eight-week-old *PKP2* heterozygous mice (a less severe murine model) with necropsy at eight weeks post LX2020 treatment. In the *PKP2* homozygous mouse study, 12 mice (six males and six females) in each group were treated with one of three dose cohorts (Dose 1 gc/kg, Dose 2 gc/kg, or Dose 3 gc/kg, which varied by approximately a half-log increase between each dose). As shown in the graphic below, within twelve weeks, approximately 90% of mice that received vehicle and 58% of mice that received Dose 1 gc/kg dose of LX2020 died. Conversely, 100% of mice that received the Dose 2 gc/kg dose and 83% of the mice that received the Dose 3 gc/kg dose survived until the 12-week necropsy time period. No PKP2 protein expression was found in the mice that died in the Dose 3 gc/kg dose cohort, whereas all mice that survived in this cohort expressed PKP2, indicating a potential for technical dosing error. Further investigation is ongoing to validate this potential error.

**LX2020 significantly extended survival in severe murine model**



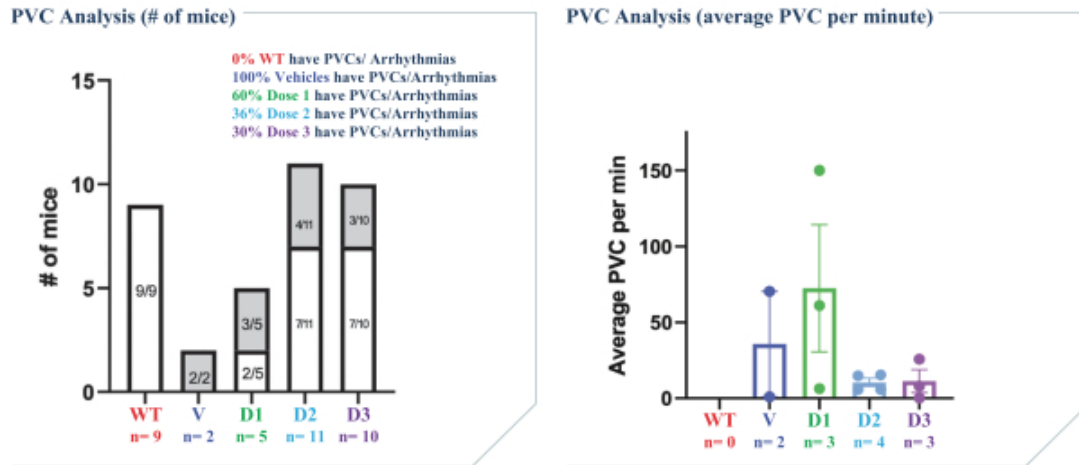
We have also performed quantitative MRI analysis which showed dose-dependent improvement in cardiac function (end-diastolic volume, or EDV, end-systolic volume, or ESV, and ejection fraction, or EF) in *PKP2* homozygous mice, as shown in the graphic below.

**Quantitative MRI analysis showed improvement in cardiac function in homozygous murine model**



As shown in the graphic below, we also completed surface ECG analyses which showed a reduction in arrhythmias as measured by premature ventricular contractions in *PKP2* homozygous murine models.

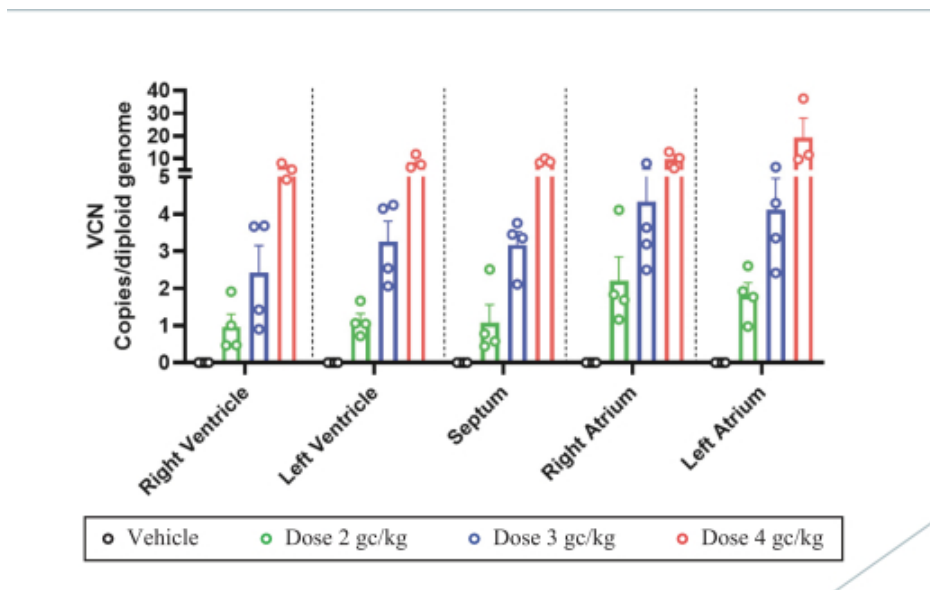
**LX2020 treatment reduced premature ventricular contractions**



**Preclinical safety studies**

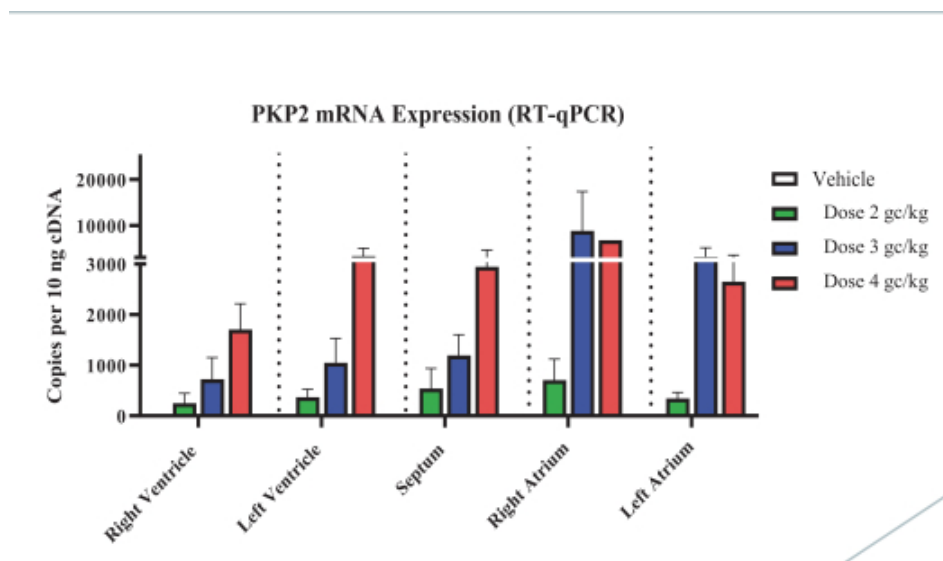
In a twelve week safety study of NHPs, three doses of LX2020 (Dose 2 gc/kg, Dose 3 gc/kg, or Dose 4 gc/kg, which vary by approximately a half-log increase between each dose) were administered to four monkeys per group (two males and two females) and resulted in dose-dependent increases in LX2020 biodistribution (reported as vector copy number, or VCN) in various regions of the heart as shown below. The biodistribution assay utilized qPCR technique with LX2020 transgene-specific primers.

**VCN in various heart regions in NHPs**



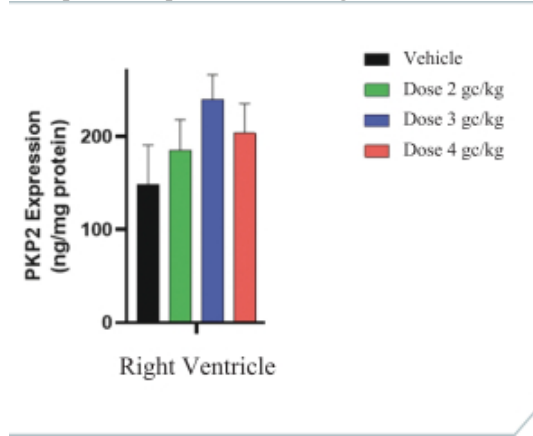
Additionally, in the NHP study, mRNA expression following LX2020 administration was assessed using an RT-qPCR assay with human PKP2 transgene-specific mRNA primers. Data is reported as copies per nanogram of cDNA input, as shown below, and demonstrates that administration of LX2020 showed a dose-dependent increase of human PKP2 mRNA in various regions of the heart.

**PKP2 mRNA expression (RT-qPCR) in NHPs**



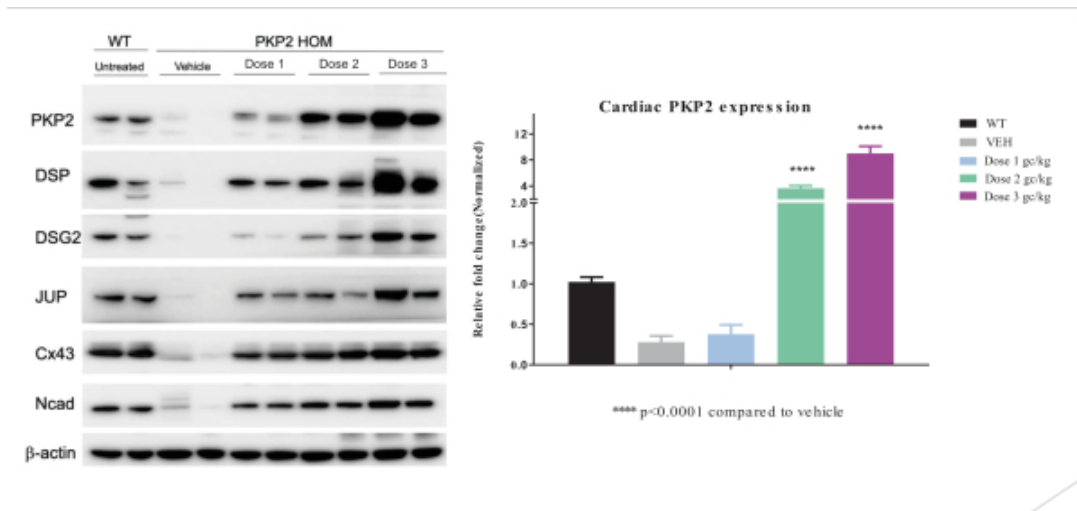
Human PKP2 protein expression was assessed in the NHP hearts using an electrochemiluminescence, or ECL, immunoassay. Notably, unlike the VCN and mRNA assays shown above, where LX2020-specific primers were synthesized to eliminate the detection of endogenous NHP DNA and mRNA, the commercially available antibodies used to detect human PKP2 protein expressed by LX2020 also detect endogenous NHP PKP2. Given the high level of amino acid sequence similarity between the human and NHP PKP2 proteins, generating anti-PKP2 antibodies specific to one of the two species is impossible. Thus, the PKP2 levels measured by the ECL assay represent the sum of endogenous NHP PKP2 and exogenous human PKP2 encoded by LX2020. Data are reported as ng of PKP2 protein per mg of total protein.

**PKP2 protein expression in the right ventricle in NHPs**



Whereas a dose-response level in DNA and RNA was observed in most heart regions in the NHPs treated with LX2020, protein levels did not correlate directly with dose. However, a dose-dependent increase in human PKP2 protein expression was seen in the PKP2 homozygous mouse model as shown in the graphic below that translated well to cardiac activity. Three-week-old adult, PKP2 homozygous mice were administered a single dose of LX2020 (Dose 1 gc/kg, Dose 2 gc/kg, and Dose 3 gc/kg) or formulation buffer (vehicle) intravenously and sacrificed twelve weeks post dose. LX2020 administration resulted in a dose-dependent increase in hPKP2 expression which was sufficient to prevent the cardiac cell-cell junction deficits of the classic proteins that make up the desmosome, including DSP, desmoglein-2, or DSG2, plakoglobin, or JUP, and N-cadherin, or Ncad, that have all been implicated in human ARVC along with the gap junction protein connexin 43 (Cx43).

**Cardiac PKP2 expression in homozygous mouse model**

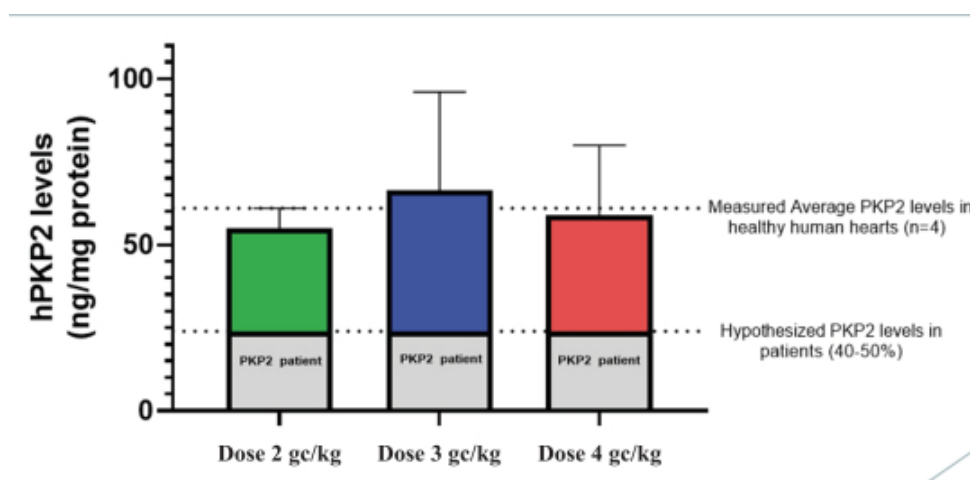


Possible explanations for the differences seen in mice and primate models include the following:

- *Inhibited Translation/Intracellular Protein Degradation:* Desmosomal proteins such as PKP2 are tightly regulated within striated muscle to maintain intracellular homeostasis. Therefore, excessive protein levels, such as those in wild-type primates, will likely result in decreased RNA translation and PKP2 protein degradation. This has been observed in previous studies and may explain why this effect was only observed in the wild-type primates but not in the murine model of PKP2 deficiency.
- *Interference from NHP endogenous PKP2 levels:* PKP2 protein is abundant in NHP cardiac muscle and can fluctuate considerably. Furthermore, for the most part, antibodies cannot distinguish between human and primate PKP2. Hence, the different protein levels in primates may be due to fluctuations in basal primate PKP2, not human PKP2 transcribed from LX2020.

As shown in the graphic below, despite the NHP PKP2 background, with LX2020 administration, the human PKP2 levels reached either close to or higher than the healthy human endogenous PKP2 levels in ventricles. For this purpose, the endogenous human PKP2 levels in the myocardium were measured from autopsy samples of four different individuals who did not have ARVC. The average myocardial PKP2 measured in non-PKP2 patient samples was approximately 61 ng/mg protein. Previous studies have shown that disease-causing mutations in PKP2 result in loss of function mutations, leading to haploinsufficiency. Given that one of the two copies of the gene is mutated, patients are expected to have approximately up to 50% of wild-type protein levels. However, this may decrease as the disease progresses over time. We have assumed that PKP2 patients express 40-50% wild-type protein, and have estimated PKP2 patients to have approximately 24 ng/mg protein. As such, we believe that we achieved at or above 90% of endogenous healthy human PKP2 levels in the ventricles at all three doses evaluated in the NHP study.

**Hypothetical model: human PKP2 in ventricles (NHP PKP2 background subtracted)**

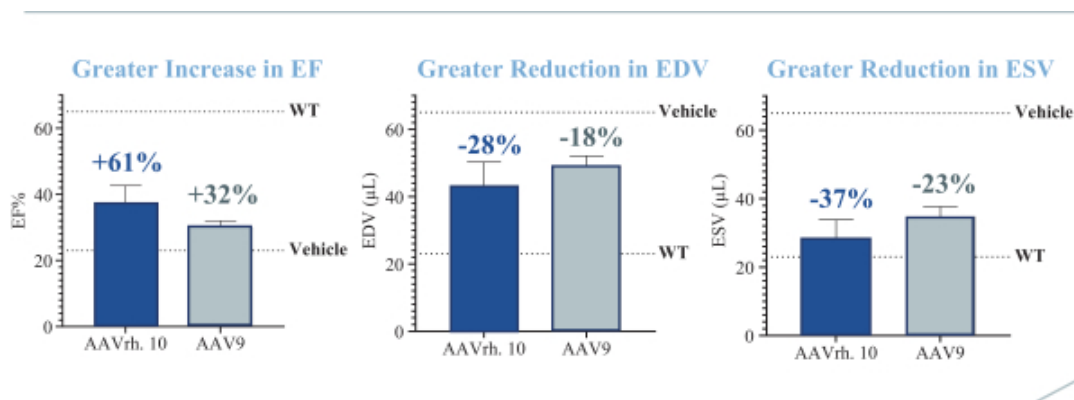


Importantly, in the NHP safety study, all LX2020 doses (Dose 2 gc/kg, Dose 3 gc/kg, and Dose 4 gc/kg) demonstrated favorable safety profiles, including no abnormal effects on clinical signs, body weight, clinical pathology, hematology, histopathology, ECHOs, cardiac biomarkers, or ECG. Collectively these data indicate LX2020 was generally well tolerated during the twelve-week duration of the study in NHPs. The maximum dose of Dose 4 gc/kg is thus recognized as the NOAEL (no observed adverse event level) in NHPs.

### Preclinical Vector Comparability Studies

Additionally, we have completed preclinical studies comparing AAVrh10 and AAV9 in head-to-head studies measuring cardiac function in a PKP2 homozygous mouse model administered with human PKP2. Human PKP2 was administered before disease onset on day two (neonatal administration) to five mice in each group, and cardiac function was measured by MRI. As shown in the graphic below, AAVrh10 demonstrated a trend of superior results compared to AAV9 as measured by improvements in ejection fraction, end-diastolic volume, and end-systolic volume.

#### AAVrh10 showed greater trends of cardiac function improvement versus AAV9 in PKP2 mouse models



### LX2020 clinical development and trial design

We expect to submit an IND application to the FDA for LX2020 in mid-2023 for the treatment of ARVC caused by mutations in the PKP2 gene. If cleared, we expect to initiate a Phase 1/2 study as a first-in-human, 52-week, dose-ascending, and open-label trial of LX2020 in patients with ARVC caused by mutations in the PKP2 gene in 2023. LX2020 will be administered as a one-time intravenous infusion to patients in at least two dose-ascending cohorts. The trial will consist of at least three patients in each cohort, with the potential for cohort expansion. A potential third cohort will receive a dose that will be selected based on the dose-response relationships seen in the first and second cohorts. There will be a long-term follow-up for patients who receive LX2020 to monitor ongoing safety for a total of five years, per FDA requirement.

### Additional cardiovascular gene therapy programs: LX2021 and LX2022

We believe there are additional targets in genetically defined cardiac disease that have the potential to be addressed through AAV-based gene therapies. We plan to continue to innovate with novel capsids, promoters, and delivery methods to optimize our early-stage assets.

### LX2021 for the treatment of DSP Cardiomyopathy

#### Overview

LX2021 is a gene therapy candidate we are developing to intravenously deliver the coding sequence for the functional Cx43 protein for arrhythmogenic cardiomyopathies, or ACs. ACs are a group of inherited cardiac muscle disorders associated with a high risk of sudden death. They are associated with fibrofatty replacement of myocardium, typically due to abnormalities in cardiac desmosomes, structures that attach cells to each other.

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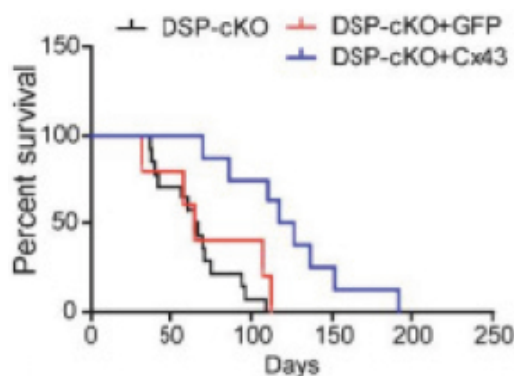
Cx43 is a structural component of gap junctions, protein complexes that allow small molecules and ions to flow directly between cells. In patients with heart disease, including heart failure, Cx43 is often relocalized in the lateral walls of myocardial cells. As a result, it is significantly reduced at cardiac muscle cell junctions, especially in AC populations. Restoring the Cx43 protein can potentially treat multiple genetic causes of AC because the cardiac loss of Cx43 is a molecular deficit generally observed in all AC patient populations. Desmoplakin, or DSP, is a structural protein critical for force transmission in heart muscle. Mutations in DSP cause a distinct form of arrhythmogenic cardiomyopathy (aka DSP Cardiomyopathy). Unlike many forms of AC that predominantly affect the right side of the heart, DSP Cardiomyopathy frequently affects the left side of the heart. The prevalence of DSP Cardiomyopathy is unknown but may be as high as 6% of all inherited dilated cardiomyopathies and 7% of arrhythmogenic right ventricular cardiomyopathies.

We have conducted a preclinical study in a cardiac-specific desmoplakin loss-of-function mouse model, or DSP-cKO, a genetic model of AC. DSP-cKO mice develop histologic and physiologic features of AC and ultimately die prematurely from heart failure. In this study, we observed that severely diseased mice treated with AAV-based therapy expressing Cx43 display fewer arrhythmias, have improved cardiac function, and experience almost twice the survival duration of untreated DSP-cKO adult mice. We plan to initiate IND-enabling studies for LX2021 by the end of 2024.

### *Preclinical studies*

Our preclinical studies have demonstrated that mice harboring a DSP mutation experience nearly twice the survival duration when treated with an AAV-based therapy candidate expressing Cx43 compared to controls. A single dose of an AAV-based gene therapy candidate representing Cx43 was administered intravenously to the cardiac-specific desmoplakin loss-of-function mouse model, or DSP-cKO, at four to six weeks of age. The DSP-cKO mice are a genetic model of ARVC harboring severe structural disease and loss of Cx43. RNA analysis of the treated mice revealed that cardiac-specific Cx43 gene delivery increased Cx43 RNA levels. Additionally, the Cx43-treated DSP-cKO mice showed a re-expression of mechanical junction proteins in their hearts compared to controls. Essential basement membrane proteins, including N-cadherin, PKP2, and junction plakoglobin, were restored to cell-cell junctions in Cx43-treated DSP-cKO hearts compared to end-stage untreated DSP-cKO hearts, which had limited localization of junctional proteins at cell-cell junctions. These findings further validate the ability of Cx43 to resurrect the cardiac mechanical muscle junction complex in diseased hearts.

### **Administration of AAV-based therapy expressing Cx43 improved survival in severe ARVC mouse model**





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These findings demonstrate that restoration of Cx43 has the potential to positively impact cardiomyocyte physiology and lifespan in the DSP-cKO mice and provide direction for our ongoing development efforts, including the evaluation of other vector serotypes. We plan to initiate IND-enabling studies for LX2021 by the end of 2024.

### *LX2022 for the treatment of HCM caused by TNNI3 mutations*

#### *Overview*

LX2022 is a gene therapy candidate we are developing to intravenously deliver a functional *TNNI3* gene to myocardial cells to treat a distinct form of HCM due to mutations in the *TNNI3* gene. With an estimated prevalence of 1 in 500 people in the United States, HCM is one of the most common forms of genetic cardiomyopathy caused by mutations that affect the cardiac sarcomere in approximately 75% of cases. It is estimated that approximately 30,000 patients in the United States and 36,000 patients in the European Union are affected by HCM caused by mutations in the gene *TNNI3*. The *TNNI3* gene encodes troponin I, an essential protein in the thin filament of the sarcomere, which is involved in cardiac contraction and relaxation. Mutations in the gene often result in left ventricular hypertrophy and restrictive cardiomyopathy, leading to arrhythmias and heart failure. We plan to complete proof-of-concept data collection for LX2022 in 2023.

#### *Overview of HCM caused by TNNI3 Mutations*

HCM is a genetic disorder characterized by left ventricular hypertrophy and one of the most common forms of genetic cardiomyopathy. Mutations in over a dozen genes encoding the thick and thin filaments of the cardiac sarcomere have been linked to HCM. *TNNI3*, a critical component of the thin filament of the sarcomere, accounts for approximately 5% of all HCM cases. Genetic mutations may affect any portion of the sarcomere, including thin and thick filaments, ultimately leading to cardiac muscle dysfunction. While thick filament mutations are the most common and best characterized, thin filament mutations remain clinically relevant. Some evidence suggests thin filament mutations are associated with more arrhythmias and earlier progression to advanced heart failure due to a disease mechanism distinct from other forms of inherited cardiomyopathy. In addition, *TNNI3* mutations are less likely to cause cardiac obstruction and hence less likely to benefit from surgical (myectomy) and medical interventions (myosin inhibitors) targeting obstruction.

*Preclinical Studies*

Our preclinical data from an A157V TNNT3 murine model, a model which results in cardiac dysfunction, was generated to validate the delivery of an AAV-based therapy candidate expressing human TNNT3 regulated by a cardiac-specific promoter. At four weeks post injection, the level and specificity of TNNT3 gene expression were evaluated in the liver, skeletal muscle, and heart tissue. Endogenous TNNT2 protein expression indicates cardiomyocyte-specific tissue, whereas ACTN2 indicates expression in only myocytes. HSPA8-1 and HSPA8-2 were blotted as housekeeping gene controls. As TNNT3 was regulated by a cardiac-specific promoter, expression of TNNT3 was only detected in the heart tissue, as shown in the figure below.

**TNNT3 Expression in Liver, Skeletal Muscle, and Heart Tissue in A157V Mouse Model**



Additionally, retro-orbital injection of AAV9-hTNNT3 expressing A157V mutation was performed on three-month-old mice euthanized four weeks later. The hearts were dissected and immunostained. Immunostaining revealed robust expression of human TNNT3 in the cardiomyocyte after treatment with AAV-based TNNT3.

Additionally, high magnification imaging demonstrated the incorporation of TNNT3 within the cardiac sarcomere. These preclinical findings demonstrate that AAV-based therapy expressing TNNT3 can be administered to achieve transgene expression in heart tissue and guide our ongoing development efforts, including evaluation of other vector serotypes. We have also developed a novel CRISPR CAS9-edited novel porcine model of HCM that resembles the human phenotype. This model will allow us to perform preclinical testing in a system to resemble the human condition and perform physiological assays to evaluate efficacy.

**APOE4-associated Alzheimer’s disease programs**

Alzheimer’s disease is a progressive neurodegenerative disorder that is the leading cause of cognitive decline in late adult life. The Alzheimer’s Association estimates that there were 6.2 million patients living in the United States with Alzheimer’s disease in 2021, with costs to the nation exceeding \$350 billion. Aging of the population is expected to significantly increase the socioeconomic burden of this disease in the coming decades, and the Alzheimer’s Association further estimates that as many as 12.4 million patients in the United States could have the disease by 2050.

## Our Alzheimer's disease franchise

In a similar theme to the cardiovascular treatment landscape, we see an opportunity of a comparable magnitude in precision therapies treating Alzheimer's disease. There are two currently approved disease-modifying Alzheimer's disease therapies, each indicated to treat the entirety of the 6.2 million patients in the United States suffering from Alzheimer's disease. However, within this prevalent population there are number of subpopulations: importantly, APOE4 patients who comprise approximately 2-3% of the U.S. population but comprise approximately 15% of Alzheimer's disease patients. We believe they represent a unique subpopulation within the broader disease space. In clinical trials of lecanemab, a different side effect profile was observed for APOE4 homozygous patients compared to patients to other patient genotypes. We believe this points to the unmet need within the APOE4 population and also raises questions of whether there is a unique disease pathology that APOE4 genetics confer to Alzheimer's disease patients. Because of this differentiated treatment effect, we believe precision therapies, particularly those focused on the underlying genetics of Alzheimer's disease, may have a substantial impact on this treatment landscape.

We are building a portfolio of approaches aimed at treating the genetics underlying Alzheimer's disease. In our lead Alzheimer's disease program, LX1001, we are initially targeting homozygous *APOE4*-associated Alzheimer's disease patients by administering AAVrh10 containing the *APOE2* gene. Our approach to treating Alzheimer's disease is predicated on the belief that expressing the protective APOE2 in the CNS of *APOE4* homozygous patients will halt or slow the progression of Alzheimer's disease. We believe these patients represent an ideal target for gene therapy because the *APOE4* homozygous profile is the most common genetic driver of Alzheimer's disease. We believe LX1001 is the only clinical-stage gene therapy candidate designed to treat this genetic cause of Alzheimer's disease. We plan to leverage biomarkers for LX1001 to potentially enable us to rapidly validate proof-of-mechanism and support further development efforts. We believe that positive clinical data from our ongoing Phase 1/2 trial has the potential to validate our novel approach to treating the *APOE4* sub-group and can help inform the development plan of our next generation candidates LX1021 and LX1020 which we believe could demonstrate an even more dramatic effect to slow or halt the progression of Alzheimer's disease. Importantly, we believe gene therapy allows for a unique approach to treating the genetics of Alzheimer's disease by delivering a therapeutic which acts upstream of both the amyloid- $\beta$  and tau-driven pathology of Alzheimer's disease. This treatment strategy is designed to impact multiple pathways, as opposed to most other treatments in development which target a single mechanism of Alzheimer's disease.

Our current gene therapy programs under development for the treatment of Alzheimer's disease are:

- *LX1001*: an AAVrh10-based gene therapy candidate that is designed to express the protective APOE2 protein in the CNS of *APOE4* homozygous patients. In our ongoing Phase 1/2 clinical trial of LX1001, three patients from the first dose cohort have demonstrated a consistent trend towards improvement in CSF tau biomarkers at the 12-month visit. We have also observed expression of the protective APOE2 protein in all patients in the first and second dose cohorts with follow-up data.
- *LX1021*: an AAVrh10-based gene therapy candidate that is designed to express the Christchurch-modified APOE2 protein in the CNS of *APOE4* homozygous patients. The Christchurch mutation has been observed to protect patients against Alzheimer's disease even in the presence of significant amyloid pathology.
- *LX1020:1*: an AAVrh10-based gene therapy candidate that is designed to express the protective APOE2 protein in the CNS of *APOE4* homozygous patients, while concurrently delivering miRNA to suppress the expression of the APOE4 protein.

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We are developing LX1001 to target homozygous *APOE4*-associated Alzheimer's disease patients, the highest risk population to develop Alzheimer's disease and the sub-group we believe will be the most likely to demonstrate a potential treatment effect from *APOE2*-based gene therapy. We anticipate reporting additional biomarker data from the second dose cohort in 2023.

### Background of Alzheimer's disease

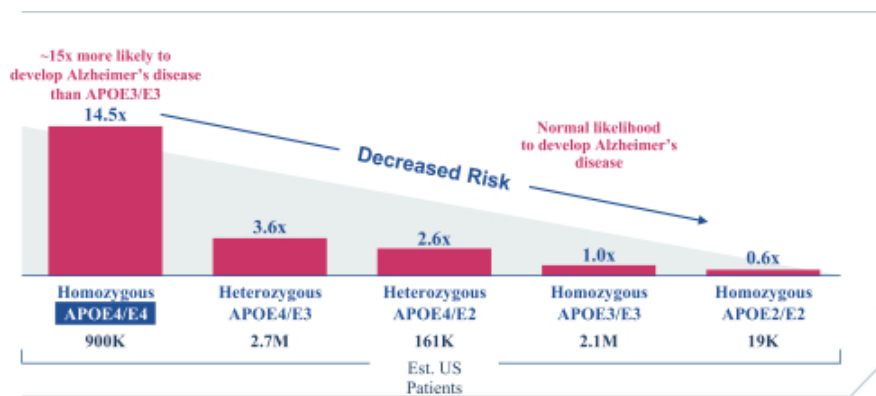
Alzheimer's disease is characterized by a complex underlying pathology in the CNS, including accumulation of A $\beta$  plaques, abnormal phosphorylation of tau, development of tau tangles, inflammation, and progressive loss of neurons, all of which combine to precipitate a progressive decline in cognitive function. APOE, a lipid transport protein, is the major transporter of cholesterol in the brain and is involved in synaptic integrity and plasticity, glucose metabolism, and cerebrovascular function.

Extracellular amyloid beta plaques and tau neurofibrillary tangles are the principle pathological hallmarks of Alzheimer's disease. Therapeutics that modulate plaques and tangles historically have been the focus of substantial research and development efforts; however, there is currently no treatment for Alzheimer's disease that is approved as disease modifying. The current standard of care for Alzheimer's disease is symptomatic therapies, such as donepezil (Aricept) and memantine (Namenda), that are viewed to have limited benefit and do not address the underlying cause of the disease. The FDA granted accelerated approval to the anti-amyloid antibody intravenous infusion therapies aducanumab (Aduhelm) and lecanemab (Leqembi), for the treatment of patients with either the mild cognitive impairment or mild dementia stage of Alzheimer's disease. These recent approvals demonstrate that the FDA has utilized an accelerated approval pathway for Alzheimer's disease therapies.

### Background on homozygous *APOE4*-associated Alzheimer's disease

Presence of *APOE4* is the most common genetic risk factor for Alzheimer's disease. The prevalent *APOE* alleles are *APOE4*, *APOE3* and *APOE2*, with the *E4* allele increasing risk and reducing the age of onset and the *E2* allele decreasing risk and markedly delaying the age of onset. *APOE4* homozygous patients, individuals who have two copies of the *E4* allele, are at the highest risk and are approximately fifteen times more likely to develop Alzheimer's disease than the general population. It is estimated that approximately 60% of Alzheimer's disease patients carry at least one *APOE4* allele as compared with approximately 25% of age-matched and healthy controls. The *APOE3* allele is believed to have a neutral impact on disease progression.

Alzheimer's disease risk by *APOE* genotype (by odds ratio)



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Third-party genetic epidemiology data from humans suggest that *APOE4* and *APOE2* are co-dominant, or both expressed, such that *APOE2/E4* heterozygous individuals have a substantially reduced risk of developing the disease as compared to *APOE4/E4* homozygous individuals. Therefore, the expression of *APOE2* appears to significantly offset the deleterious effects of the *APOE4* allele.

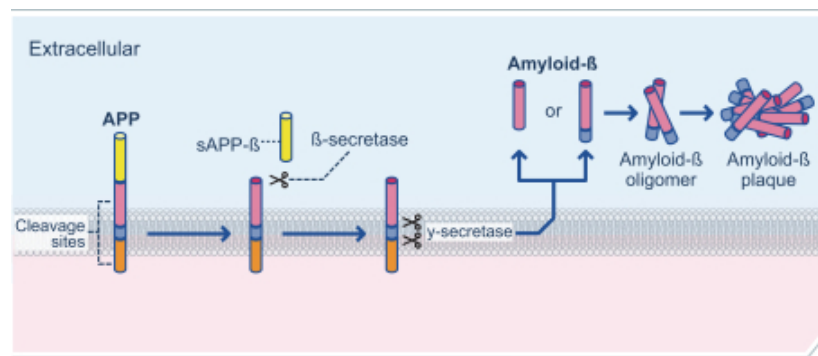
We believe that administering the *APOE2* gene to *APOE4* homozygous patients is a promising approach because it has the potential to address several pathways that are involved in the progression of the disease.

### Role of amyloid in pathogenesis of Alzheimer's disease and APP processing

*APOE4* is associated with decreased clearance of Ab and increased b-amyloid (Ab) production and plaque aggregation. Amyloid-b (Ab) is produced through the sequential cleavage of the amyloid-b precursor protein (APP) by the b-secretase beta-site APP cleaving enzyme 1 (BACE 1) and g-secretase in neurons and other cell types.

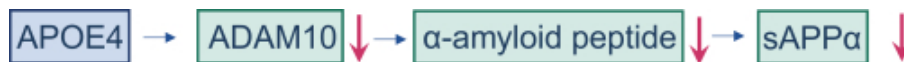
As illustrated in graphic below, cleavage of APP via this pathway results in production of Ab<sub>42</sub> and Ab<sub>40</sub>, and Ab peptides undergo oligomerization and subsequent aggregation into insoluble plaques.

#### APP cleavage resulting in amyloid beta production



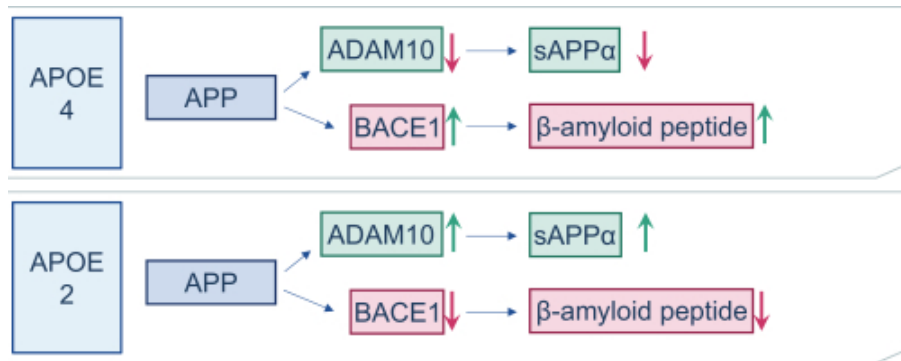
Note: Nature. 2018.

As an alternative to cleavage by BACE1, APP can also be cleaved by α-secretases such as A Disintegrin and Metalloproteinase Domain-containing Protein 10 (ADAM10). Cleavage of APP by ADAM10 results in the formation of soluble sAPPα (sAPPα), which has been demonstrated to improve synaptic function and promote neurite growth.



The α-secretase pathway results in neurotropic sAPPα, whereas the β-secretase pathway results in AD-associated Ab peptides. We believe *APOE4*-associated reduced ADAM10 activity and increased BACE1 activity would result in decreased sAPPα and elevated Ab levels; whereas presence of *APOE2* would result in the opposite effect.

**APOE and APP processing**

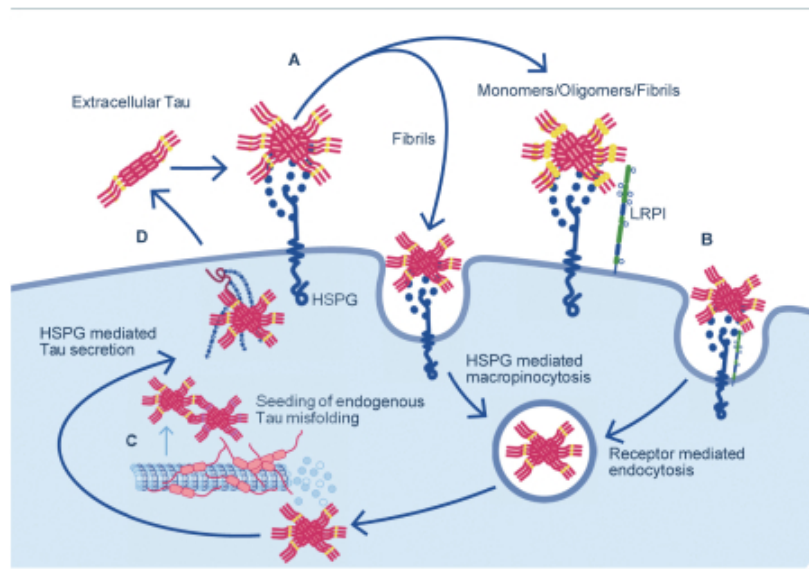


**Role of Tau in Pathogenesis of Alzheimer’s Disease**

Tau proteins are abundant in neurons and maintain the stability of microtubules in axons. However, excessive/abnormal phosphorylation of tau results in microtubular destabilization and transformation of monomers into paired-helical-filament, or PHF, tau and neurofibrillary tangles, or NFTs. This buildup of tau lesions throughout the brain increases with time, correlating with neurodegeneration and Alzheimer’s disease symptomatology. APOE4 effects tau indirectly via its effect on amyloid and the downstream effect of the amyloid pathway on tau. APOE4 also directly affects tau pathology as it is associated with increased tau phosphorylation and the accumulation and aggregation of neurofibrillary tangles. Importantly, we believe therapeutic intervention at the symptomatic stage could slow progression of tau pathology and neurodegeneration.

Heparin sulfate proteoglycans, or HSPGs, have been extensively characterized as a major mediator of tau uptake and have been shown to mediate transcellular tau propagation. HSPGs bind to tau and colocalize with tau-based lesions in Alzheimer’s disease as shown in the figure below.

**Role of HSPGs in tau pathology**

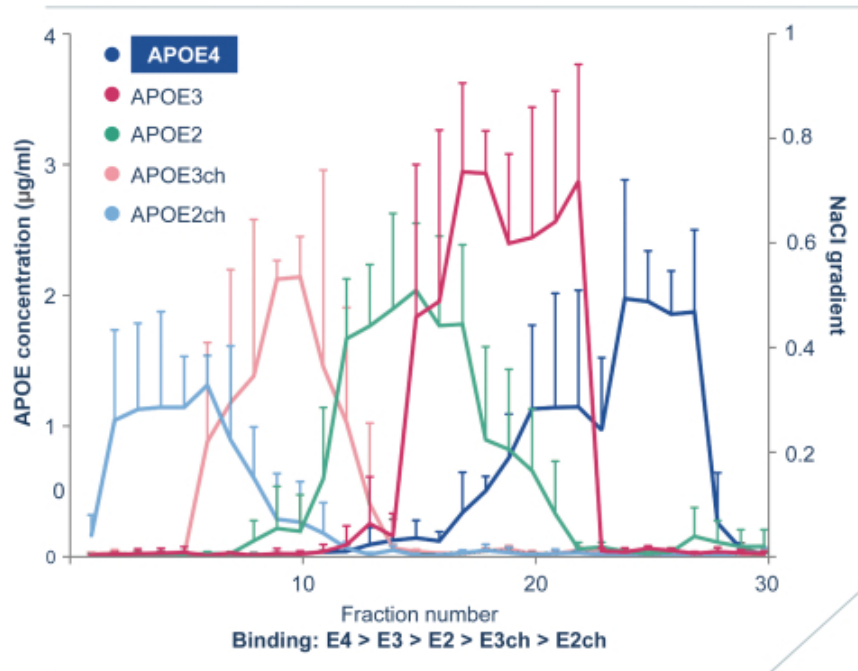


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An *in vitro* study using heparinase III, which removes polysaccharides on the cell surface, has demonstrated the ability to impair HSPGs. In that study, heparinase treatment led to a 50% impairment of tau oligomer uptake in general and reversed the increased accumulation of tau oligomers caused by APOE4. We believe this data demonstrates that HSPGs can mediate increased accumulation of tau observed under the APOE4 condition.

Additionally, Cornell, through our sponsored research agreement with them, has generated *in vitro* data showing APOE isoforms differ in their HSPG binding characteristics, with APOE4 binding strongly to HSPGs and APOE2 with a lower binding affinity. As illustrated in the figure below, the APOE Christchurch mutation has the lowest HSPG binding affinity and is associated with protection against the development of tau pathology and Alzheimer's disease. We believe these data demonstrate that the low HSPG binding affinity of APOE2 is associated with protective effect on tau pathology and neurodegeneration.

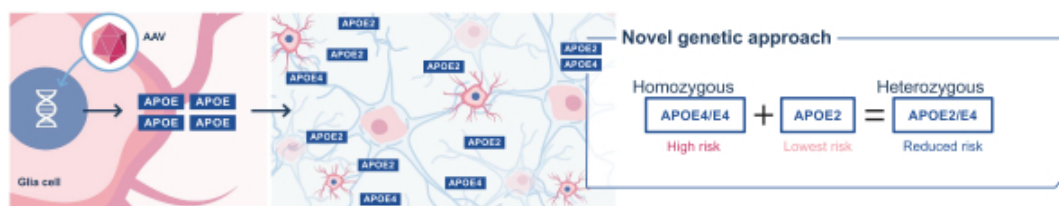
**APOE isoforms had differential effects on heparin binding**



### Our solution: LX1001

LX1001 is an AAVrh10-based gene therapy candidate for the treatment of APO4 homozygous patients. It is designed to deliver the human APOE2 gene into the CNS via one-time administration to the CSF to halt or slow the progression of Alzheimer's disease.

**LX1001 is designed to address the genetic driver of Alzheimer’s disease by delivering APOE2 gene into CNS of APOE4 homozygous patients**



Based on what is known regarding gene therapies delivered to the CNS, we believe LX1001 will only require a single dose because neurons are post-mitotic, or incapable of further cell division, so there will be no dilution of extra-chromosomal LX1001 caused by cell division. APOE is a secreted protein; thus, only a fraction of neurons needs to be transduced with LX1001 to secrete what we anticipate to be potentially efficacious levels of the protective isoform. We have designed LX1001 so that the protective isoform has the potential to be widely available to cells in the brain and able to compensate for non-transduced cells.

The FDA granted Fast Track designation to LX1001 for the treatment of APOE4 homozygous patients with Alzheimer’s disease to slow disease progression. We plan to seek other regulatory designations and consider accelerated clinical development pathways based on biomarker data that may establish early proof-of-mechanism and is expected to predict clinical benefit.

*Preclinical studies*

Weill Cornell Medicine, independently and through our sponsored research agreement with them, has conducted preclinical studies in large and small animals that have demonstrated the potential for LX1001 to induce APOE2 expression levels that we believe will be sufficient to halt or slow the progression of Alzheimer’s disease in APOE4 homozygous patients. The preclinical results discussed below are from those preclinical studies unless otherwise noted.

Preclinical murine studies of LX1001 were conducted in a widely used model of brain amyloidosis (PDAPP mice) as well as a mouse model that develops APOE4-dependent amyloid deposition (APP.PS1/TRE4 mice). APP.PS1/TRE4 triple transgenic mice develop early and robust Aβ and amyloid deposition, which is dramatically reduced in the absence of APOE4. Intracerebral delivery of LX1001 led to the widespread brain expression of APOE2. Studies in the APP.PS1/TRE4 model showed an approximately 80% reduction in insoluble Aβ1-42 in the hippocampus eight weeks following intrahippocampal administration of LX1001.

An additional preclinical study in NHPs was conducted to determine the most active and well-tolerated route of administration for LX1001 while achieving broad CNS biodistribution and therapeutic levels of the APOE2 proteins. In these studies, we utilized an intracisternal delivery route to deliver the gene therapy vector to the CSF in order to achieve widespread biodistribution of LX1001. The intracisternal route has been shown in multiple studies in large animals to result in transgene expression throughout the CNS and involves a minimally invasive procedure to infuse into the CSF.

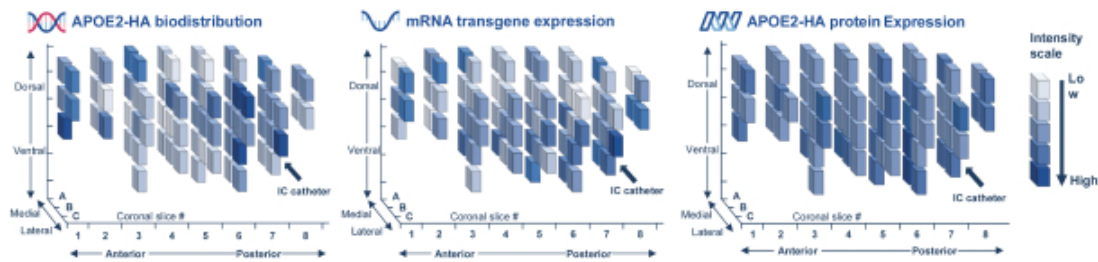
To evaluate this route of delivery, at eight weeks following intracisternal administration of total vector dose of 5 x10<sup>13</sup> vg/ml CSF of LX1001, with the APOE transgene tagged with the common antibody epitope tag hemagglutinin, or APOE-HA, three NHP brains in one study were evaluated for the extent of vector APOE2-HA biodistribution, mRNA transgene expression, and APOE2-HA protein expression. To visualize the extent of vector biodistribution and transgene and protein expression, a 3D representation of one-centimeter cubes of the



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brain was used, as shown below in the example of one NHP. As expected for a secreted protein like APOE2, diffuse expression was observed throughout the brain. Greater-than-baseline levels of all three targets were observed in most cubes throughout the CNS, as well as in the hippocampal regions, the area of the brain most relevant in Alzheimer's disease.

### Biodistribution of APOE2-HA vector genomic copies, transgene mRNA, and protein levels in brain of an NHP following intracisternal administration of LX1001



The results from this study demonstrate that *LX1001* was consistently expressed through the brain and achieved broad distribution in the CNS of the NHPs. In addition, administration of the HA tagged version of *LX1001* directly to the CNS of NHPs exhibited a favorable safety profile.

## Clinical development

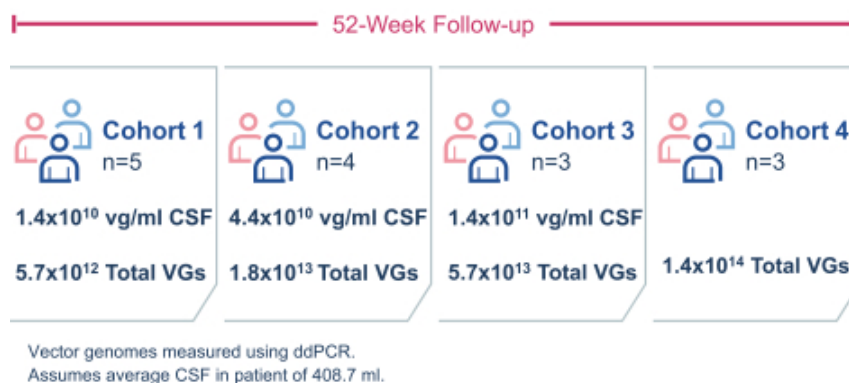
### Trial design

We are evaluating *LX1001* in an open-label, dose-ranging Phase 1/2 clinical trial in patients who are *APOE4* homozygous patients with clinical diagnoses ranging from mild cognitive impairment to mild or moderate dementia due to Alzheimer's disease. All patients have evidence of amyloid plaque by PET scan and CSF biomarkers consistent with Alzheimer's disease.

The primary objective of the trial is to evaluate the safety of *LX1001* administered to the CNS via injection between cervical vertebrae 1 and 2, or intracisternal injection, and to establish the maximum tolerated dose. The trial is also designed to evaluate the conversion of the CSF from the *APOE4* homozygous profile to an *APOE4/E2* profile. Additional secondary endpoints include CSF biomarkers, including A $\beta$ 42, total tau, and phosphorylated tau, amyloid PET scan, structural MRI imaging and cognitive tests.

As shown below, the clinical trial is a dose-ranging trial of *LX1001* in four ascending dose cohorts ( $1.4 \times 10^{10}$ vg/ml,  $4.4 \times 10^{10}$ vg/ml,  $1.4 \times 10^{11}$ vg/ml and  $1.4 \times 10^{14}$  total genomes), with the dose for each patient in the first three cohorts determined based on CSF volume measured by MRI and patients in the fourth cohort receiving a flat dose. Each dose cohort will consist of approximately three to five patients for a total of approximately 15 patients in the trial. Key patient enrollment criteria include *APOE4* homozygous genetic profile, patient age of 50 years or older, positive amyloid PET scan, evidence of CSF biomarkers consistent with Alzheimer's disease, and diagnosis that ranges from mild cognitive impairment to mild or moderate dementia due to Alzheimer's disease.

**Design of Phase 1/2 clinical trial of LX1001**



*Preliminary trial results*

Preliminary data from cohort 1 and cohort 2 showed trends towards improvement in CSF APOE4 levels, in all patients with follow-up data at three months or longer and remained largely steady through the 12-month visit. These results, shown below, demonstrate that the APOE2 transgene is being expressed in the CNS in all patients treated with LX1001 and signal the potential for treatment to convert the APOE4 homozygous profile to a combined APOE2/APOE4 profile.

**Interim APOE2 CSF expression levels (fmol/mL)<sup>(1)</sup>**

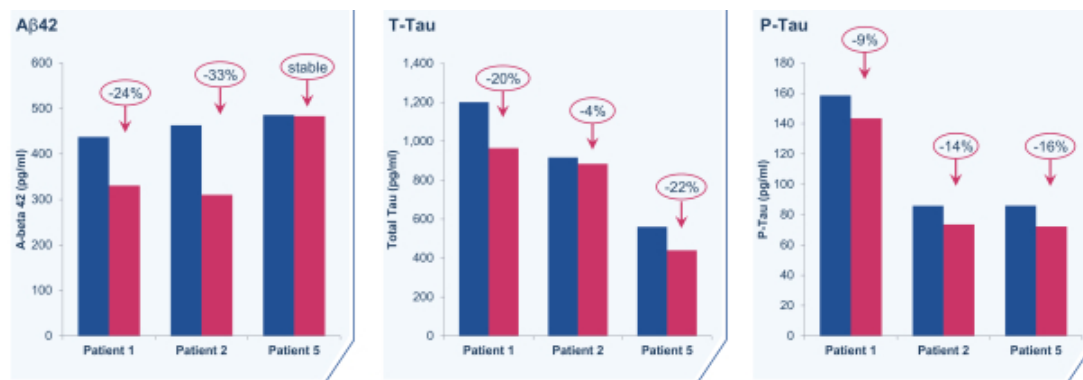
	Patient No.	Dose (Total gc) <sup>(2)</sup>	Month 0	Month 3	Month 6	Month 12
Cohort 1	101-1001	6.7E+12	0.0	0.3	0.4	0.4
	101-1002	7.9E+12	0.0	--	--	0.1
	101-1004	5.0E+12	0.0	0.8	--	--
	101-1005	6.8E+12	0.0	0.3	0.3	0.3
Cohort 2	102-1003	2.3E+13	0.0	2.1	1.6	0.7
	102-1005	1.6E+13	0.0	0.7	0.6	Ongoing
	102-1017	1.7E+13	0.0	1.8	1.2	Ongoing
	102-1013	1.8E+13	0.0	0.7	Ongoing	Ongoing

(1) APOE2 quantified by liquid chromatography mass spectroscopy. Research grade assay; preliminary results represent the average of all runs for each sample (1-5 runs / sample).  
 (2) Cohort 1 & 2 patient doses were administered using a nominal LX1001 titer based on qPCR. Values represent actual total dose administered using LX1001 titer by ddPCR  
 Note: Patient 101-1003 transitioned to a long-term care-facility prior to Month 3 and unlikely to have follow-up efficacy data. Patient 101-1004 was unable to travel to the study site for the month 6 and 12 visits.

Additionally, in the three patients from cohort 1 with data available at 12-months post treatment, we have seen consistent trends towards improvement in the core CSF biomarkers T-tau and P-tau that are associated with neurodegeneration and Alzheimer’s disease specifically and we have also seen declines or stabilization of Ab42, the beta-amyloid peptide associated with Alzheimer’s disease as shown below. We believe this interim data reflects

evidence of proof-of-mechanism and supports our unique approach to treating the genetics of Alzheimer’s disease by delivering a therapeutic which acts upstream of both the amyloid- $\beta$  and tau-driven pathology of Alzheimer’s disease.

**Interim CSF core biomarker data in Cohort 1 patients with 12-month data**



If 2 pre-therapy values were done, the average was used to calculate the change at 12 months  
Note: Patient 3 transitioned to a long-term care facility prior to Month 3 and unlikely to have follow-up efficacy data. Patient 4 was unable to travel to the study site for the month 6 and 12 visits.

Natural history data for T-tau and P-tau would suggest that on average these biomarkers should increase over the course of the year, versus the decline that we have seen following treatment. Among all patients in cohort 1 and cohort 2, treatment with LX1001 has been well-tolerated with no serious adverse events reported as of March 2023. The DSMB meeting identified no safety concerns and recommended the study proceed to cohort 3 which is currently enrolling patients. We expect to report the additional biomarker data from the second dose cohort in 2023.

**Next-generation gene therapy solutions for Alzheimer’s disease: LX1021 and LX1020**

We believe that positive clinical data from our ongoing Phase 1/2 clinical trial of LX1001 has the potential to validate our novel approach to treating the APOE4 sub-group and can help inform the development plan of our next-generation candidates, LX1021 and LX1020.

*LX1021 for the treatment of APOE4 homozygotes*

**LX1021:** We are designing LX1021 to treat APOE4 homozygous patients by adding a Christchurch mutation-modified APOE2 to the CNS. The Christchurch mutation has been recognized to protect individuals against Alzheimer’s disease even in the presence of significant amyloid pathology. The mechanism of this protection may relate to the fact that APOE, in the presence of the Christchurch mutation, binds poorly to heparan sulfate proteoglycans (HSPGs), molecules found on the surface of neurons, which may inhibit the spread of tau pathology between cells. We believe this approach has the potential to enhance the protective effect of APOE2 in homozygous APOE4-associated Alzheimer’s disease. We plan to complete proof-of-principle studies for LX1021 in 2023.

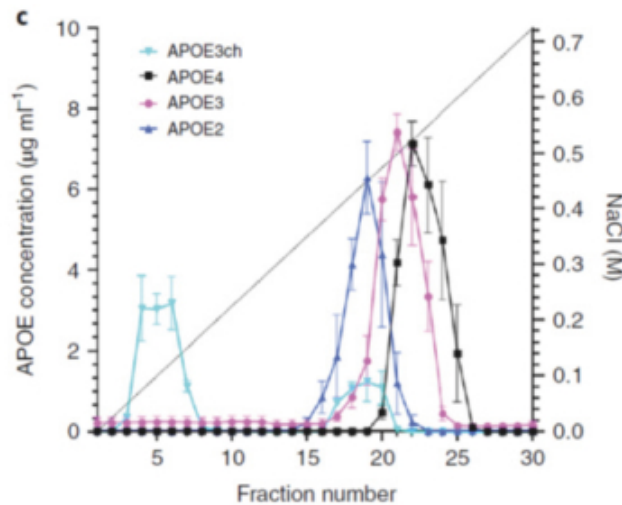
LX1021 is designed to add a Christchurch mutation-modified *APOE2* gene to the CNS of *APOE4* homozygous patients



*Preclinical Studies*

*In vitro* experiments conducted by other researchers and replicated at Weill Cornell Medicine have demonstrated that Christchurch mutation-modified *APOE3* behaves similarly to the *APOE2* allele in that both isoforms bind poorly to heparin sulfate proteoglycans, or HSPGs. As shown in the figure below, differences in one to two amino acids drive the major physiological differences between the three *APOE* isoforms, including differences in protein structure and regulation of amyloid- $\beta$  aggregation, in addition to binding affinities to HSPGs. This increased binding to HSPGs is implicated in the propagation of tau tangles and uptake of toxic forms of tau and amyloid- $\beta$  aggregation.

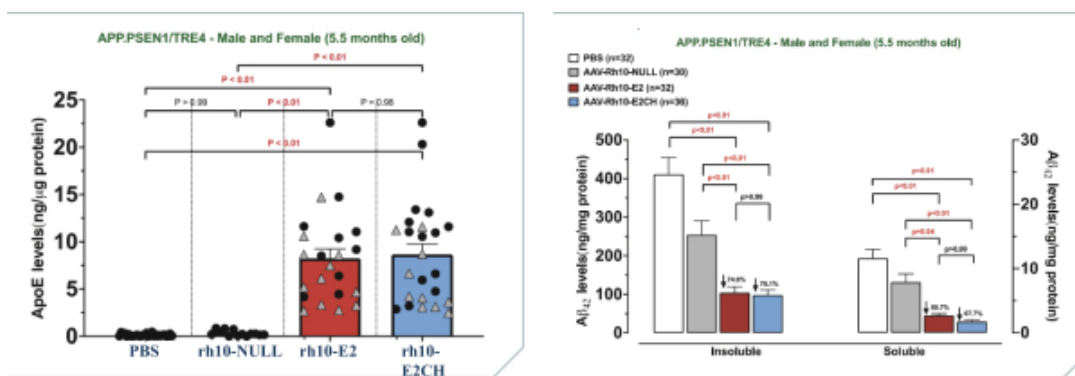
**Binding affinity of different *APOE* alleles and Christchurch mutation-modified *APOE3* and *APOE2***



Preclinical studies have demonstrated that *APOE4*-driven amyloid pathology can be suppressed by AAV-mediated delivery of Christchurch-modified *APOE2* to the CNS in a well-established murine Alzheimer's disease mouse models (amyloid mice, APP.PS1/TRE4).

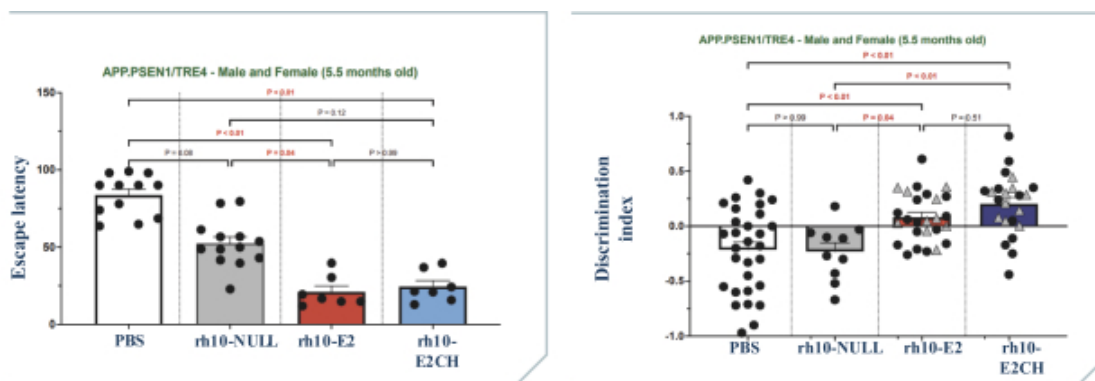
In the APP.PS1/TRE4 mouse model, 2.5-month old mice were administered  $2 \times 10^{10}$  total genome dose of LX1001 and LX1021 via intrahippocampal injection and evaluated for *APOE* expression and AB42 levels. As shown in the graphic below, the interim data from this study show that *APOE* levels were increased compared to vehicle in both groups, and we saw a statistically significant reduction in AB42 in both treated groups versus the PBS-treated group.

**APOE expression and Ab42 reduction in amyloid mouse model**



Additionally, behavioral tests have been conducted in the amyloid APP.PS1/TRE4 mouse model, including novel objection recognition and Barnes maze tests evaluated the impact of treatment. Novel object tests are an indicator of working memory, with a higher discrimination index indicating beneficial results from better working memory. The Barnes maze is a hippocampal-dependent spatial memory test based on the desire to find an escape hole from a platform, lower escape latency demonstrates positive impact of treatment. As shown in the graphic below, both LX1001 and LX1021 demonstrated improvements versus the PBS-treated group.

**Barnes maze (left) and novel objection recognition (right) in amyloid mouse model**

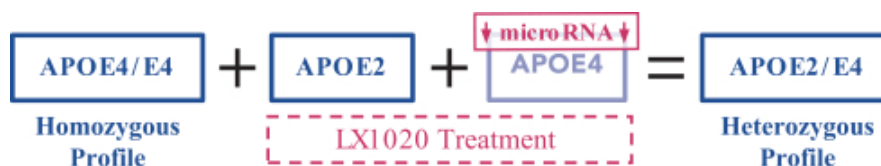


We believe that the preclinical data in Amyloid mice (APP.PS1/TRE4) demonstrated that LX1021 has similar therapeutic potential as LX1001 as both vectors suppress the amyloid pathology in the APP.PS1/TRE4 mice to a similar extent and also showed similar behavioral improvements. We plan to continue preclinical development efforts, including completing additional preclinical studies in a P301S/TRE4 mouse model (tau mice) in 2023 where we believe LX1021 will function even more effectively than LX1001 in suppressing tau pathology.

*LX1020 for the treatment of APOE4 homozygotes*

**LX1020:** We are designing LX1020 to treat APOE4 homozygous patients by delivering both the protective APOE2 allele and miRNA to suppress APOE4. We believe delivery of APOE2 with concurrent suppression of APOE4 will achieve a higher degree of conversion to the APOE4/E2 heterozygous profile, which should lead to greater therapeutic effect. We plan to complete candidate selection for LX1020 in 2023.

**LX1020 is designed to add APOE2 gene while suppressing APOE4**



*Preclinical Studies*

In preclinical studies, AAV9 vectors expressing APOE-targeting miRNAs (AAV-mirAPOE) was administered to the hippocampus of TRE4 mice, a mouse model with murine *APOE* gene deleted and replaced with human *APOE4*. When compared with an AAV9-mCherry (a control vehicle), the AAV-mirAPOE downregulated hippocampal *APOE4* mRNA levels by 2.8-fold, suggesting that an AAV-mediated gene therapy incorporating APOE-targeting miRNA into the expression cassette may potentially reduce human *APOE4* in the CNS. These findings demonstrate that AAV-based delivery of APOE-targeting miRNA may be able to serve as part of a therapeutic approach for treating *APOE4* homozygous Alzheimer’s patients and provide direction for our ongoing development efforts.

We plan to complete candidate selection for LX1020 in 2023.

**Additional Program Outside of Core Focus Area**

***LX1004 for the treatment of CLN2 Batten disease***

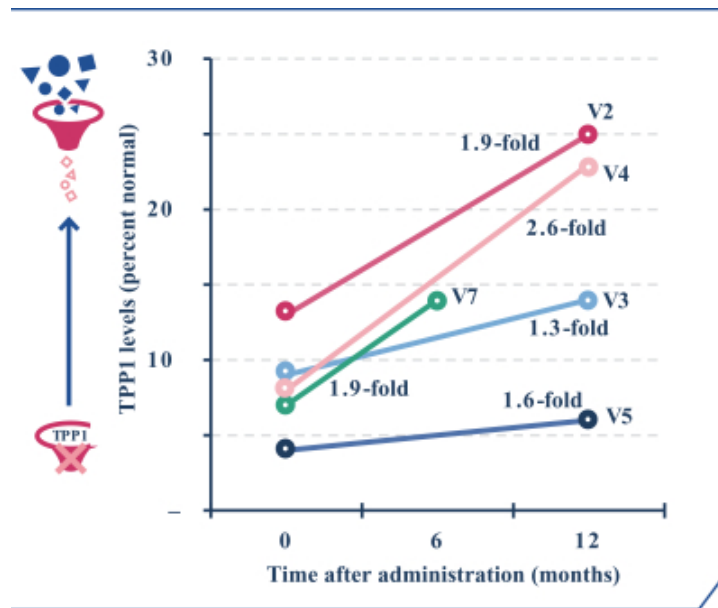
LX1004 is an AAVrh10-based gene therapy candidate designed to deliver a fully functional *CLN2* gene to the CNS via intracisternal injection in order to restore TPP1, the secreted protein that is deficient in CLN2 Batten disease. Batten disease is the common name for a broad class of rare, fatal, inherited disorders of the nervous system also known as neuronal ceroid lipofuscinoses. CLN2 Batten disease is caused by a defect in the *CLN2* gene found on chromosome 11 and, based on our calculations, we believe CLN2 Batten disease affects up to approximately 900 patients in the United States and European Union. The FDA has granted Rare Pediatric Disease designation and Orphan Drug designation to LX1004 for the treatment of CLN2 disease.

In a Phase 1/2 trial that was conducted by Weill Cornell Medicine under an IND that was subsequently transferred to us, LX1004 demonstrated an increase in TPP1 levels leading to a slower decline of motor and language function in treated patients compared to natural history controls, at 18 months post treatment. In November 2022, we held a Type C meeting with the FDA to align on the development path for LX1004 and are currently evaluating strategic alternatives to find the appropriate partner to take the program forward.

*Phase 1/2 clinical trial results*

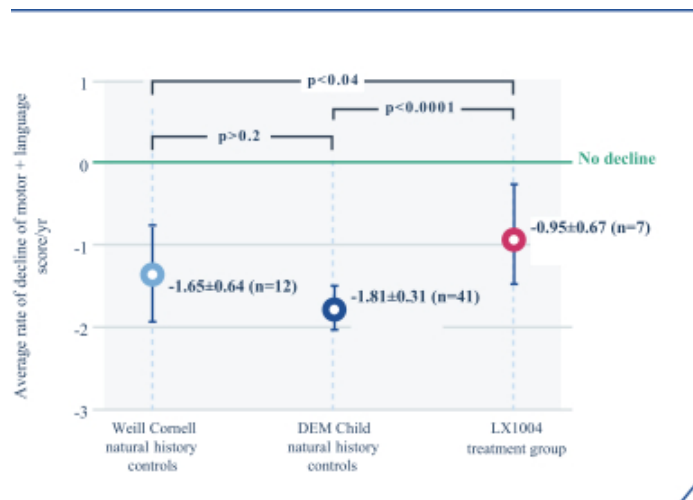
LX1004 was administered intraparenchymally to the CNS of eight children with CLN2 Batten disease. Seven children had follow-up visits during the 18-month duration of the study, and of these, five children had CSF samples pre and post treatment. All five of those patients had increased TPP1 levels in their CSF up to 12 months, indicating substantial expression of the protein and durability of the therapy.

**LX1004 demonstrated increases in TPP1 expression levels up to 12 months**



Additionally, as demonstrated in the below graphic, the treated patients had a statistically significant reduction in the rate of disease progression compared to natural history controls. The trial used the established CLN2 Clinical Rating Scale to assess efficacy. This scale was developed to assess the loss of function in CLN2 Batten disease patients that occurs over the course of the disease. It is a four-part assessment of language, motor function, visual function, and seizures. The rate of decline in the motor + language subscores of the treatment group was compared to two historical control groups of untreated patients: the Weill Cornell Medicine natural history cohort and the European DEM Child natural history cohort. The annual rate of decline was similar in the two control groups. In contrast, the rate of decline in the treated patients was slower than that in both control groups.

**LX1004 demonstrated a slower rate of decline relative to natural history controls**



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Furthermore, patients in the trial reported six acute serious adverse events within two weeks of dosing and 15 chronic serious adverse events after two weeks of dosing that were deemed to be related to the study drug or its administration. The six acute serious adverse events consisted of one patient with seizures, two patients with episodes of increased abnormal movements, one patient with emesis, one patient with hematoma, and one patient with pneumocephalus. The 15 chronic serious adverse events consisted of six patients with seizures, one patient with dystonia, one patient with emesis, one patient with hematoma, one patient with hygroma, one patient with bronchospasm, one patient with aspiration, one patient with pneumonia, and two patients with elevated hepatic enzymes.

## Competition

The biotechnology and pharmaceutical industries are characterized by rapidly changing technologies, significant competition and a strong emphasis on intellectual property. This is also true for the development and commercialization of treatments for cardiovascular and neurodegenerative diseases such as Friedreich's ataxia, Alzheimer's disease, CLN2 Batten disease and broadly across gene therapies. While we believe that our management and scientific team's deep expertise in gene therapy provides us with competitive advantages, we face competition from several sources, including large and small biopharmaceutical companies, government agencies and academic and private research institutions. Not only must we compete with other companies that are focused on gene therapy technology, but any product candidates that we successfully develop and commercialize will compete with existing therapies, to the extent applicable, and new therapies that may become available in the future.

Drug development, particularly in the gene therapy field, is highly competitive and subject to rapid and significant technological advancements. A significant unmet medical need exists in each of the indications that we are targeting, and it is likely that additional drugs will become available in the future for the treatment of these diseases.

We are aware that our competitors are developing product candidates for the treatment of diseases that our product candidates will target. With respect to LX2006, we are aware of preclinical gene therapy programs in development at Novartis, PTC Therapeutics, Inc., Aavanti Bio, Inc., Lacerta Therapeutics, Inc. and those being developed in collaborations between Voyager Therapeutics, Inc. and Neurocrine Biosciences, Inc. and between Takeda Pharmaceutical Company Limited and StrideBio, Inc. Among other treatment modalities for FA, we are aware that Larimar Therapeutics, Inc. is developing a clinical stage product candidate, CTI-1601, and that Reata Pharmaceuticals, Inc.'s Omaveloxolone (Skyclarys) was recently approved by the FDA.

With respect to LX2020, both Rocket Pharmaceuticals Inc. and Tenaya Therapeutics Inc. are developing a preclinical AAV-based gene therapy candidate designed to deliver a functional *PKP2* gene to patients with ARVC.

With respect to our portfolio of gene therapy programs for the treatment of homozygous *APOE4*-associated Alzheimer's disease, we are aware that uniQure, N.V. is pursuing AMT-240, a preclinical gene therapy candidate for autosomal dominant Alzheimer's disease intended to silence the toxic variant while expressing the protective variant and Novartis has a gene therapy candidate for Alzheimer's disease which is in the early preclinical stages of development. Many large and small pharmaceutical companies and academic institutions are developing potential treatments for the condition given the significant unmet need and the large population suffering from Alzheimer's disease. There are multiple FDA-approved treatments for Alzheimer's disease, including donepezil (Aricept), memantine (Namenda), and aducanumab (Aduhelm), the latter of which was recently approved under accelerated approval and will require confirmatory data to verify clinical benefit. In January of 2023, lecanemab was granted accelerated approval by the FDA for the treatment of Alzheimer's disease based on the observed reduction of amyloid beta plaque. In addition, Eli Lilly and Company's product candidate for the treatment of Alzheimer's disease, donanemab, is currently in Phase 3 development. Finally, we are aware that Voyager Therapeutics, Inc. and Taysha Gene Therapies, Inc. are pursuing Alzheimer's disease treatments and have early-stage discovery efforts ongoing based on vectorized antibodies and tau-specific miRNA shuttles, respectively.



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Many of our existing or potential competitors have substantially greater financial, technical and human resources than we do and significantly greater experience in the discovery and development of product candidates, as well as in obtaining regulatory approvals of those product candidates in the United States and in foreign countries.

These competitors also compete with us in recruiting and retaining qualified scientific and management personnel and establishing clinical trial sites and patient registration for clinical trials, as well as in acquiring technologies complementary to, or necessary for, our programs. Mergers and acquisitions in the pharmaceutical and biotechnology industries may result in even more resources being concentrated among a smaller number of our competitors. Smaller or early-stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large and established companies.

We will face competition from other drugs or from other non-drug products and treatments currently approved or that will be approved in the future in the cardiovascular and neurology field, including for the treatment of diseases and diseases in the therapeutic categories we intend to target. Therefore, our ability to compete successfully will depend largely on our ability to:

- develop, manufacture and commercialize drugs that are superior to other products in the market;
- demonstrate through our clinical trials that our product candidates are differentiated from existing and future therapies;
- attract qualified scientific, product development and commercial personnel;
- obtain patent or other proprietary protection for our medicines;
- obtain required regulatory approvals;
- obtain coverage and adequate reimbursement from, and negotiate competitive pricing with, third-party payors; and
- successfully collaborate with pharmaceutical companies in the discovery, development and commercialization of new medicines.

The availability of our competitors' products could limit the demand, and the price we are able to charge, for any product candidate we develop. The inability to compete with existing or subsequently introduced drugs would have an adverse impact on our business, financial condition and prospects. In addition, the reimbursement structure of currently approved or future gene therapies by other companies could impact the anticipated reimbursement structure of our gene therapies, if approved, and our business, financial condition, results of operations and prospects.

Established pharmaceutical companies may invest heavily to accelerate discovery and development of novel compounds or to in-license novel compounds that could make our product candidates less competitive. In addition, any new product that competes *with* an approved product must demonstrate compelling advantages in efficacy, convenience, tolerability and safety in order to overcome price competition and to be commercially successful. Accordingly, our competitors may succeed in obtaining patent protection, discovering, developing, receiving regulatory and marketing approval for, or commercializing, drugs before we do, which would have an adverse impact on our business and results of operations.

## **Intellectual property**

We actively seek to protect our proprietary technology, inventions, and other intellectual property that is commercially important to the development of our business by a variety of means, such as seeking, maintaining,

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and defending patent rights, whether developed internally or licensed from third parties. We also may rely on trade secrets and know-how relating to our proprietary technology platform, on continuing technological innovation and on in-licensing opportunities to develop, strengthen and maintain the strength of our position in the field of gene therapy that may be important for the development of our business. We also intend to seek patent protection or rely upon trade secret rights to protect other technologies that may be used to discover and validate targets, and that may be used to manufacture and develop novel gene therapy products. We are a party to license agreements that give us rights to use specific technologies in our gene therapy candidates and in manufacturing our products. Additional regulatory protection may also be afforded through data exclusivity, market exclusivity and patent term extensions where available.

As of March 1, 2023, we in-licensed two U.S. patents, nine pending U.S. non-provisional patent applications, three pending U.S. provisional applications, two pending PCT applications, three foreign patents and 45 pending foreign applications. We do not own or in-license any composition of matter patents or patent applications in the United States or any other jurisdiction with respect to our LX1004 product candidate.

In regard to our LX2006 product candidate, we in-license from Cornell University and Adverum Biotechnologies, Inc. two U.S. patents, two pending U.S. non-provisional patent applications, three foreign patents granted in India, Mexico, and New Zealand and 9 pending foreign applications pending in such jurisdictions as Australia, Brazil, China, Eurasia, Europe, Israel, Mexico and South Africa. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, these patents, if granted, will expire in 2033. These patents and pending patent applications disclose and/or contain composition-of-matter claims to an AAV vector encoding *FXN*, or a fragment thereof, and disclose and/or contain claims to methods of producing and methods of treatment using the AAV *FXN* vector. Cornell University co-owns these patents and patent applications with Institut National de la Santé et de la Recherche Médicale (INSERM), Centre National de la Recherche Scientifique (CNRS), Université de Strasbourg, Université Paris-Saclay and Assistance Publique-Hôpitaux de Paris (APHP).

We co-own with Cornell University one pending PCT application that relates to our LX2006 cardiac FA program. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, patent applications claiming benefit of this PCT application, will expire in 2043. The PCT application discloses and/or contains composition-of-matter claims to a pharmaceutical dosage form of viral vectors encoding *FXN*, and discloses and/or contains claims to methods of producing and methods of treatment using the pharmaceutical dosage form.

In regard to our LX2020 product candidate, we in-license from The Regents of the University of California one pending PCT application. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, patent applications claiming the benefit of this PCT application, will expire in 2042. This PCT application discloses and/or contains composition-of-matter claims to a vector encoding human plakophilin-2 (*PKP2*), and discloses and/or contains claims to methods of producing and methods of treatment using the *PKP2* vector.

In regard to our LX2021 product candidate, we in-license from The Regents of the University of California two pending U.S. non-provisional patent application and six pending foreign applications pending in Australia, Canada, China, Europe, Israel, and Japan. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, these patent applications will expire in 2038. These pending patent applications disclose and/or contain composition-of-matter claims to a vector encoding a human connexin 43 (*CX43*) polypeptide, and disclose and/or contain claims to methods of producing and methods of treatment using the *CX43* vector.

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In regard to our LX2022 product candidate, we in-license from The Regents of the University of California one pending U.S. non-provisional patent application. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, patent applications claiming the priority to this patent application, will expire in 2040. This pending U.S. non-provisional patent application discloses and/or contains composition-of-matter claims to a vector encoding a human *TNNI3* gene, and discloses and/or contains claims to methods of producing and methods of treatment using the *TNNI3* vector.

In regard to our LX1020 product candidate, we in-license from Cornell University one pending U.S. non-provisional application and 13 pending foreign applications pending in Australia, Brazil, Canada, China, Colombia, Europe, Hong Kong, Israel, Japan, Korea, Mexico, New Zealand, Singapore and South Africa. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, these patent applications will expire in 2040. The pending patent applications disclose and/or contain composition-of-matter claims to a vector encoding human APOE2 protein and encoding one or more RNAi nucleic acid sequences for inhibition of APOE4 mRNA, and disclose and/or contain claims to methods of producing and methods of treatment using the APOE2+/APOE4- vector.

In regard to our LX1021 product candidate, we in-license from Cornell University one pending U.S. non-provisional application and 13 pending foreign applications pending in Australia, Brazil, Canada, China, Europe, Hong Kong, Israel, Japan, Korea, Mexico, New Zealand, Singapore and South Africa. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, these patent applications will expire in 2040. The pending patent applications disclose and/or contain composition-of-matter claims to a vector encoding a mutated human apolipoprotein E protein, and disclose and/or contain claims to methods of producing and methods of treatment using the mutated human apolipoprotein E vector.

In regard to our LX1001 product candidate, we co-own with Cornell University two pending U.S. provisional applications. Not accounting for any patent term adjustment and assuming that all annuity and/or maintenance fees are paid timely, if granted, patent applications claiming the priority to these U.S. provisional applications, will expire in 2043. These pending patent applications disclose and/or contain composition-of-matter claims to a pharmaceutical dosage form of viral vectors encoding human APOE2, and disclose and/or contain claims to methods of producing and methods of treatment using the pharmaceutical dosage form.

### ***License, research collaboration and sponsored research agreements***

#### *First license agreement with Cornell University*

In May 2020, we entered into a license agreement with Cornell University, or the Cornell First License Agreement, pursuant to which we obtained a sublicensable, worldwide license under certain patents to make, use, and sell products that are covered by a valid claim of a licensed patent, are based on the transferred IND from Cornell University, use any licensed materials, or are produced using a licensed method, and to practice certain licensed technology, in all cases, for all human and non-human prophylactic and therapeutic uses. In July 2022, we entered into Amendment No. 1 to the Cornell First License Agreement that updated fees and royalties payable by us under the Cornell First License Agreement. In September 2022, we entered into Amendment No. 2 to the Cornell First License Agreement that clarified patent prosecution and maintenance obligations and added certain inventions and patents to the list of inventions and patent rights covered under the Cornell First License Agreement. The technology under the license includes portfolios for APOE, Alzheimer's disease, and Anti-Tau, although our license is not restricted by such indications. The license is exclusive with respect to certain patents and non-exclusive with respect to other patents. Additionally, under the Cornell First License Agreement, Cornell University assigned to us an IND for the use of AAVrh10.hAPOE2 vector to treat *APOE4* homozygous patients

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who are at risk of having or have Alzheimer's disease. Cornell University reserved the rights to publish and disseminate information about inventions included in the licensed technology and licensed patents, and to use, and allow other nonprofit institutions to use, the patents and technology for educational and research purposes.

We are obligated to diligently proceed with the development, manufacture, and sale of licensed products, to raise certain amounts within specified time frames, to achieve certain development milestones within specified time frames, to meet agreed minimum-spend requirements, and to meet other diligence obligations. If we fail to perform these obligations, Cornell University could terminate the Cornell First License Agreement in full or in part in specified circumstances, subject to certain rights for us to extend these time frames.

Under the Cornell First License Agreement, we paid Cornell University an upfront payment of approximately \$0.15 million and entered into a purchase agreement for approximately \$0.6 million of convertible preferred securities. We are obligated to pay Cornell University an annual license maintenance fee ranging from the low four digits to the mid five digits, increasing annually until such time that we are commercially selling a Product. Under the Cornell First License Agreement, we are obligated to pay Cornell University up to \$8.4 million for each portfolio upon the achievement of specific clinical and regulatory milestones. We are obligated to pay Cornell University a flat royalty in the mid-single-digits based on net sales of a licensed product in a country, which royalty rate increases by one percent if the licensed product is an orphan drug, subject to a reduction upon the expiration of valid claims in licensed patents and certain reductions for third-party licenses. In addition, in certain specific instances where sales are made by a sublicensee, the royalty rate increases by an amount in the low to mid-single digits. If the royalties are below certain agreed amounts, we are required to pay Cornell University minimum annual royalties ranging from low-six digits to low-seven digits. The royalty term continues for each licensed product on a country-by-country basis beginning on the first commercial sale of such licensed product and ending on the latest of (a) expiration or invalidation of the last valid claim in the licensed patent, (b) the expiration of regulatory exclusivity, and (c) the month of the first commercial launch of a generic equivalent in such country. We are also obligated to pay Cornell University a percentage of sublicensing fees in the low-double digits to mid-double digits.

We may terminate the Cornell First License Agreement or any portfolio thereunder at any time upon ninety (90) days' advance written notice to Cornell University. Cornell University may terminate the Cornell First License Agreement if we commit a material breach and fail to cure such breach within a specified cure period after written notice or if we challenge the validity of a licensed patent. Upon expiration of the royalty term of a given licensed product in a country, the license becomes non-exclusive and royalty-free. Upon termination of the Cornell First License Agreement, all licenses and rights granted by either party will terminate, although we will have a period of time to sell off any remaining licensed product.

### *Second license agreement with Cornell University*

In May 2020, we entered into a second license agreement with Cornell University, or the Cornell Second License Agreement, pursuant to which we obtained a sublicensable, worldwide license under certain patents to make, use, and sell products that are covered by a valid claim of a licensed patent, are based on the transferred IND from Cornell University, use any licensed materials, or are produced using a licensed method, and to practice certain licensed technology, in all cases, for all human and non-human prophylactic and therapeutic uses. In January 2022, we entered into Amendment No. 1 to the Cornell Second License Agreement, whereby we paid Cornell a material transfer fee and a license fee totaling \$30,000, and Cornell added certain materials to the list of original materials subject to the Cornell Second License Agreement. In July 2022, we entered into Amendment No. 2 to the Cornell Second License Agreement that updated fees and royalties payable by us under the Cornell Second License Agreement. The technology under the license includes portfolios for CLN2 (Batten disease) and FA, although our license is not restricted by such indications. The license is exclusive with respect to certain patents

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and non-exclusive with respect to other patents. Additionally, under the Cornell Second License Agreement, Cornell University assigned to us an IND for the use of AAVrh.10cUhCLN2 to treat children with infantile neuronal ceroid lipofuscinosis (also called CLN2 or Batten disease). Cornell University reserved the rights to publish and disseminate information about inventions included in the licensed technology and licensed patents, and to use, and allow other nonprofit institutions to use, the patents and technology for educational and research purposes.

We are obligated to diligently proceed with the development, manufacture, and sale of licensed products, to raise certain amounts within specified time frames, to achieve certain development milestones within specified time frames, to meet agreed minimum-spend requirements, and to meet other diligence obligations. If we fail to perform these obligations, Cornell University could terminate the Cornell Second License Agreement in full or in part in specified circumstances, subject to certain rights for us to extend these time frames.

Under the Cornell Second License Agreement, we paid Cornell University an upfront payment of approximately \$0.15 million and entered into a purchase agreement for approximately \$0.6 million of convertible preferred securities. We are obligated to pay Cornell University an annual license maintenance fee ranging from the low four digits to the mid five digits, increasing annually until such time that we are commercially selling a licensed product. Under the Cornell Second License Agreement, we are obligated to pay Cornell University up to \$4.3 million for two portfolios and up to \$0.6 million for a third portfolio upon the achievement of specific clinical and regulatory milestones. Upon submitting our IND application for LX2006 to the FDA in the first quarter of 2022, we achieved the first clinical milestone under the Cornell Second License Agreement, and we paid \$0.1 million to Cornell University in the second quarter of 2022 in connection with this milestone. We are obligated to pay Cornell University a flat royalty in the mid-single-digits based net sales of a licensed product in a country, which royalty rate increases by one percent if the licensed product is an orphan drug, subject to a reduction upon the expiration of valid claims in licensed patents and certain reductions for third-party licenses. In addition, in certain specific instances where sales are made by a sublicensee, the royalty rate increases by an amount in the low to mid-single digits. If the royalties are below certain agreed amounts, we are required to pay Cornell University minimum annual royalties ranging from low six digits to low seven digits. The royalty term continues for each licensed product on a country-by-country basis beginning on the first commercial sale of such licensed product and ending on the latest of (a) expiration or invalidation of the last valid claim in the licensed patent, (b) the expiration of regulatory exclusivity, and (c) the month of the first commercial launch of a generic equivalent in such country. We are also obligated to pay Cornell University a percentage of sublicensing fees ranging in the low double digits.

We may terminate the Cornell Second License Agreement or any portfolio thereunder at any time upon ninety (90) days' advance written notice to Cornell University. Cornell University may terminate the Cornell Second License Agreement if we commit a material breach and fail to cure such breach within a specified cure period after written notice or if we challenge the validity of a licensed patent. Upon expiration of the royalty term of a given licensed product in a country, the license becomes non-exclusive and royalty-free. Upon termination of the Cornell Second License Agreement, all licenses and rights granted by either party will terminate, although we will have a period of time to sell off any remaining licensed product.

### *Research collaboration agreement with Weill Cornell Medicine*

In February 2021, we entered into a Research Collaboration Agreement with Cornell University on behalf of Joan & Sanford I. Weill Medical College, or Weill Cornell Medicine, or the Cornell Collaboration Agreement, in connection with the Cornell First License Agreement and the Cornell Second License Agreement entered into in May 2020, collectively, the Cornell License Agreements. We committed to fund scientific research at Weill Cornell Medicine related to the technology licensed to us pursuant to the Cornell License Agreements. Cornell

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University reserved the rights to publish and disseminate information about the results, excluding any inventions, generated from Cornell University's investigator's conduct of the research.

Under the Cornell License Agreements, we committed to provide funding for research projects at Cornell University for each rolling three-year period during the term of the last to expire of the Cornell License Agreements, with a minimum annual funding commitment of \$1.0 million. As such, under the terms of the Cornell Collaboration Agreement, we will pay Weill Cornell Medicine at least \$3.0 million over the course of three years. With respect to each Weill Cornell Medicine invention and joint invention and related joint results that is either an improvement to, or is dominated by, the patent rights under the Cornell License Agreements or is specifically designed for a licensed product under the Cornell License Agreements, or Improvement, and for which we have made an election to obtain such inventions, the Cornell License Agreements will be amended to include license grants to such Improvements following payment of the relevant amendment fee in the low five digits. With respect to each Weill Cornell Medicine invention and joint invention, and related joint results that are not Improvements, we have the first option to negotiate for a royalty-bearing, worldwide license under such intellectual property to develop, make, have made, use, offer for sale, sell, have sold and import products on commercially reasonable terms.

The Cornell Collaboration Agreement will expire, unless earlier terminated, on the expiration of the agreed upon three-year term. Cornell University may terminate the Cornell Collaboration Agreement if we commit a material breach and fail to cure such breach within a specified cure period after written notice. In addition, the Cornell Collaboration Agreement may be terminated by us in the event that the Weill Cornell Medicine Principal Investigator, currently designated as Dr. Crystal, ceases to supervise the research and Weill Cornell Medicine is unable to or declines to find a substitute.

### *License agreement with Adverum*

In January 2021, we entered into a license agreement, or the Adverum Agreement, with Adverum Biotechnologies, Inc., or Adverum, pursuant to which we obtained an exclusive, sublicensable, worldwide license under certain patents, know-how, and other intellectual property relating to viral vector technology for gene therapy applications for the treatment of FA cardiomyopathy.

We are responsible for the development, manufacture, and commercialization of gene therapy products that consist of a specific nucleic acid sequence that is delivered by a specific gene therapy, or the Products. We are obligated to use commercially reasonable efforts to develop, obtain regulatory approval for, and commercialize the Product.

Under the Adverum Agreement, we paid Adverum a \$7.5 million upfront payment. We are obligated to pay Adverum up to \$17.5 million upon the achievement of specified development and regulatory milestones, including a \$3.5 million development milestone that was achieved in the first quarter of 2023, and up to \$49 million in commercialization and sales milestones for the Products. We are obligated to pay Adverum tiered royalties ranging from 8% to 10% based on annual aggregate worldwide net sales of Products, subject to reductions upon the expiration of valid claims in licensed patents and third-party licenses. The royalty term continues for each Product on a country-by-country basis beginning on the first commercial sale of such Product and ending on the latest of (a) expiration of the last valid claim in the licensed patent that covers the manufacture, use, or sale of the Product in such country, (b) the expiration of all regulatory and data exclusivity in such country, and (c) ten years after the first commercial sale of such Product in such country.

The Adverum Agreement will expire, unless earlier terminated, on the expiration of the last royalty term for a Product in a particular country. We have the right to terminate the Adverum Agreement at any time upon one-hundred twenty (120) days' advance written notice to Adverum. In addition, subject to certain conditions,

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either we or Adverum may terminate the Adverum Agreement upon the insolvency of the other or if the other party commits a material breach of the agreement and fails to cure such breach within a specified cure period after written notice is provided. Additionally, Adverum may terminate the Adverum Agreement if we challenge the validity of any licensed patents. Upon expiration of the Adverum Agreement, the license becomes royalty-free, irrevocable and perpetual. Upon termination of the Adverum Agreement, all licenses, sublicenses, and rights granted by either party will terminate.

### *First license agreement with the Regents of University of California, San Diego*

Stelios, which we acquired in August 2021, is successor-in-interest to ARVC Therapeutics, Inc., or ARVC Therapeutics, under a worldwide license agreement entered into by ARVC Therapeutics and The Regents of The University of California on April 23, 2020, or the UCSD First License Agreement, pursuant to which they obtained a license to certain intellectual property related to gene therapies for ARVC. The UCSD First License Agreement relates to our development efforts for our LX2021 program. Pursuant to the UCSD First License Agreement, we obtained an exclusive, sublicensable, worldwide license under certain patents to make, use, sell, offer for sale and import services, methods, composition and products that incorporate or are developed using any licensed invention or licensed methods, or is covered by a valid claim of a licensed patent, or a Product, in all cases for all diagnostic and therapeutic uses. UCSD reserved the right to practice the relevant invention and licensed patents for educational and research purposes, to publish and disseminate information about the relevant invention and licensed patents, and to allow other nonprofit institutions to use, publish, or disseminate information about the relevant inventions and licensed patents for educational and research purposes.

Under the UCSD First License Agreement, we are obligated to achieve certain development milestones within specified time frames, to use commercially reasonable efforts to diligently develop, manufacture, and sell Products, and to meet agreed minimum-spend requirements. If we fail to perform any of these obligations, UCSD could either terminate the UCSD First License Agreement or convert the exclusive license to a nonexclusive license, subject to certain rights for us to extend these time frames.

The UCSD First License Agreement required Stelios to pay one-time up-front non-refundable cash fees of \$20,000. We are obligated to pay annual license maintenance fees in the mid-four digits to low-five digits, increasing until such time that we are commercially selling a Product. We are also obligated to pay up to \$4.75 million upon the achievement of specific development and commercialization milestones for the first Product and low- to mid-single digit royalties based on aggregate net sales, subject to certain reductions for third-party licenses. The royalty term continues until the expiration of the UCSD First License Agreement. Under the UCSD First License Agreement, if the royalties are below certain agreed amounts, we are required to pay UCSD minimum annual royalties ranging from low- to mid-five digits. We are also obligated to pay UCSD a percentage of sublicensing fees ranging in the low double digits. In the event that we assign the UCSD First License Agreement, we will be obligated to pay an assignment fee that will be determined based on certain aspects of the assignment.

The UCSD First License Agreement will expire, unless earlier terminated, on the expiration of the last claim of any licensed patent or patent applications in a particular country. We have the right to terminate the UCSD First License Agreement at any time upon sixty (60)-days' written notice to UCSD. UCSD may terminate the UCSD First License Agreement if we commit a material breach, if we fail to meet certain specified milestones within the prescribed time periods, if we are delinquent on any report or payment or provide an intentionally misleading report, if we fail to diligently develop and commercialize the licensed products, or if we challenge the validity of a licensed patent, in each case only if we fail to cure such problem within a specified cure period after written notice. Upon termination of the UCSD First License Agreement, all licenses and rights granted by UCSD to us will terminate.

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### *Second license agreement with the Regents of University of California, San Diego*

Stelios, which we acquired in August 2021, entered into a worldwide license agreement in August 2020 with The Regents of The University of California, or the UCSD Second License Agreement, pursuant to which they obtained a license to certain intellectual property and materials related to gene therapies for HCM. The UCSD Second License Agreement relates to our development efforts for our LX2022 programs. Pursuant to the UCSD Second License Agreement, we obtained an exclusive, sublicensable, worldwide license under certain patents to make, use, sell, offer for sale and import services, methods, composition and products that incorporate or are developed using any licensed invention or licensed methods or is covered by a valid claim of a licensed patent, or a Product, and a non-exclusive license to use nonpublic technical information, or Technology, in all cases for all diagnostic and therapeutic uses. UCSD reserved the rights to practice the relevant invention, Technology and licensed patents for educational and research purposes, to publish and disseminate information about the relevant invention, Technology and licensed patents, and to allow other nonprofit institutions to use, publish, or disseminate information about the relevant invention, Technology and licensed patents for educational and research purposes.

Under the UCSD Second License Agreement, we are obligated to achieve certain development milestones within specified time frames and to use commercially reasonable efforts to diligently develop, manufacture, and sell Products. If we fail to perform any of these obligations, UCSD could either terminate the UCSD Second License Agreement or convert the exclusive license to a nonexclusive license subject to certain rights for us to extend these time frames.

The UCSD Second License Agreement required Stelios to pay one-time up-front non-refundable cash fees of \$20,000. We are obligated to pay annual license maintenance fees in the mid-four digits to low-five digits, increasing until such time that we are commercially selling a Product. We are also obligated to pay up to \$2.4 million upon the achievement of certain development and commercialization milestones for the first Product and low-single digit royalties based on aggregate net sales, subject to certain reductions for third-party licenses. The royalty term continues until the expiration of the UCSD Second License Agreement. We are also obligated to pay UCSD a percentage of sublicensing fees ranging in the low double digits.

The UCSD Second License Agreement will expire, unless earlier terminated, on the expiration of the last claim of any licensed patent or patent applications in a particular country. We have the right to terminate the UCSD Second License Agreement at any time upon sixty (60)-days' written notice to UCSD. UCSD may terminate the UCSD Second License Agreement if we commit a breach, if we fail to meet certain specified milestones within the prescribed time periods, if we are delinquent on any report or payment, if we are not diligently developing or commercializing Products in accordance with our diligence obligations, if we provide any intentionally false report, or if we challenge the validity of a licensed patent, in each case only if we fail to cure such problem within a specified cure period after written notice. In the event that we assign the UCSD Second License Agreement, we will be obligated to pay an assignment fee that will be determined based on the acquisition price. Upon termination of the UCSD Second License Agreement, all licenses and rights granted by UCSD to us will terminate.

### *Third license agreement with the Regents of University of California, San Diego*

In October 2021, we entered into a worldwide license agreement with UCSD, or the UCSD Third License Agreement, and collectively with the UCSD First License Agreement and the UCSD Second License Agreement, The UCSD License Agreements, pursuant to which we obtained a license to materials and intellectual property related to a gene therapy for ARVC. The UCSD Third License Agreement relates to our development efforts for our LX2020 program. Pursuant to the Third UCSD Agreement, we obtained an exclusive, sublicensable, worldwide license under certain patents to make, use, and sell, offer for sale and import services, methods,



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composition and products that incorporate or are developed using any licensed invention or licensed methods, or is covered by a valid claim of a licensed patent, or a Product, and an exclusive license to use nonpublic technical information, or Technology, in all cases for all diagnostic and therapeutic uses. UCSD reserved the rights to practice the relevant invention, Technology and licensed patents for educational and research purposes, to publish and disseminate information about the relevant invention, Technology and licensed patents, and to allow other nonprofit institutions to use, publish, or disseminate information about the relevant invention, Technology and licensed patents for educational and research purposes.

Under the UCSD Third License Agreement, we are obligated to achieve certain development milestones within specified time frames, to use commercially reasonable efforts to diligently develop, manufacture, and sell licensed products, and to meet agreed minimum-spend requirements. If we fail to perform any of these obligations, UCSD could either terminate the UCSD Third License Agreement or convert the exclusive license to a nonexclusive license subject to certain rights for us to extend these time frames.

The UCSD Third License Agreement required us to pay a one-time up-front non-refundable cash fee of \$20,000. We are obligated to pay annual license maintenance fees in the mid-four digits to low-five digits, increasing until such time that we are commercially selling a Product. We are also obligated to pay up to \$4.0 million upon the achievement of specified development and commercialization milestones for the first Product and low- to mid-single digit royalties based on aggregate net sales, subject to certain reductions for third-party licenses. The royalty term continues until the expiration of the UCSD Third License Agreement. If the royalties are below certain agreed amounts, we are required to pay UCSD minimum annual royalties ranging from low- to mid-five digits. We are also obligated to pay UCSD a percentage of sublicensing fees ranging in the low double digits.

The UCSD Third License Agreement will expire, unless earlier terminated, on the expiration of the last claim of any licensed patent or patent applications in a particular country. We have the right to terminate the UCSD Third License Agreement at any time upon sixty (60)-days' written notice to the UCSD. UCSD may terminate the UCSD Third License Agreement if we commit a material breach, if we fail to meet certain specified milestones within the prescribed time periods, if we are delinquent on any report or payment, if we are not diligently developing or commercializing Products in accordance with our diligence obligations, if we provide any intentionally false report, or if we challenge the validity of a licensed patent, in each case only if we fail to cure such problem within a specified cure period after written notice. In the event that we assign the UCSD Third License Agreement or we undergo a change of control, we will be obligated to pay a flat low-six digit fee or a fee that will be determined based on the acquisition price. Upon termination of the UCSD Third License Agreement, all licenses and rights granted by UCSD to us will terminate.

### *Sponsored research agreements with the Regents of University of California, San Diego*

In connection with the UCSD Agreements, on December 3, 2021, we entered into a sponsored research agreement with the Regents of UCSD for our LX2020 and LX2021 programs, and another sponsored research agreement with the Regents of UCSD for our LX2002 program, which was subsequently amended on April 19, 2023. We refer to the agreement and the as-amended agreement as the UCSD SRAs. UCSD reserved the rights to publish and disseminate information about the results generated from UCSD's investigator's conduct of the research. The total costs to be invoiced to us under the terms of the UCSD SRAs are approximately \$5.0 million. Under the terms of the UCSD SRAs, which have two-year terms, we have the first rights to obtain non-exclusive or exclusive, sublicensable, royalty-bearing, perpetual and transferable worldwide licenses to any inventions generated by UCSD or UCSD's interest in any inventions generated jointly by UCSD and us, and we retain the rights to any inventions generated solely by us.

The UCSD SRAs will expire, unless earlier terminated, on the expiration of the two-year term. We have the right to terminate the UCSD SRAs at any time upon the giving of thirty (30) days' written notice to UCSD.

## **Government regulation**

Government authorities in the United States at federal, state and local levels, as well as in foreign countries, extensively regulate, among other things, the research, development, testing, manufacture, quality control, import, export, safety, effectiveness, labeling, packaging, storage, distribution, record keeping, approval, advertising, promotion, marketing, post-approval monitoring and post-approval reporting of biologics, including gene therapies, such as those we are developing. Generally, before a new biologic can be marketed, considerable data demonstrating its quality, safety and efficacy must be obtained, organized into a format specific for each regulatory authority, submitted for review and approved, authorized, or cleared by the applicable regulatory authority. We, along with third-party contractors, will be required to navigate the various preclinical, clinical and commercial approval requirements of the governing regulatory agencies of the countries in which we wish to conduct studies or seek approval or licensure of our product candidates.

### ***U.S. biologics regulation***

In the United States, biological products are subject to regulation under the U.S. Federal Food, Drug, and Cosmetic Act, or FDCA, and the Public Health Service Act, or PHSA, and their implementing regulations and other federal, state, local and foreign statutes and regulations. Failure to comply with the applicable United States requirements at any time during the product development process, approval process or following approval may subject an applicant to administrative actions or judicial sanctions. These actions and sanctions could include, among other actions, the FDA's refusal to approve pending applications, withdrawal of an approval, license revocation, a clinical hold, untitled or warning letters, voluntary or mandatory product recalls or market withdrawals, product seizures, total or partial suspension of production or distribution, injunctions, fines, refusals of government contracts, restitution, disgorgement and civil or criminal fines or penalties.

Our product candidates must be approved by the FDA through the Biologics License Application, or BLA, the process which is required by the FDA before biological product candidates may be marketed in the United States and generally involves the following:

- completion of extensive preclinical laboratory tests and animal studies performed in accordance with applicable regulations, including the GLP regulations;
- submission to the FDA of an IND application, which must become effective before human clinical trials may begin;
- approval by an IRB or ethics committee at each clinical site before the trial is commenced;
- performance of adequate and well-controlled human clinical trials in accordance with applicable IND regulations, cGCPs and other clinical trial-related regulations to establish the safety, purity and potency of the proposed biological product candidate for its intended purpose;
- preparation of and submission to the FDA of a BLA that contains sufficient data to demonstrate substantial evidence of effectiveness;
- a determination by the FDA within 60 days of its receipt of a BLA to file the application for review;
- payment of user fees for FDA review of the BLA;

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- satisfactory completion of an FDA pre-license inspection of the manufacturing facility or facilities at which the proposed product will be produced to assess compliance with cGMPs and to assure that the facilities, methods and controls are adequate to ensure and preserve the biological product's identity, strength, quality and purity, and of selected clinical investigation sites to assess compliance with the cGCPs;
- satisfactory completion of an FDA Advisory Committee review, if applicable;
- FDA review and approval, or licensure, of a BLA to permit commercial marketing of the product for particular indications for use in the United States.

### ***Preclinical and clinical development***

Prior to beginning the first clinical trial with a product candidate, the product candidate must undergo rigorous preclinical testing. Preclinical studies include laboratory evaluation of product chemistry and formulation, as well as *in vitro* and animal studies to assess safety and in some cases to establish a rationale for therapeutic use. The conduct of preclinical studies is subject to federal and state regulations and requirements, including GLP regulations for safety/toxicology studies.

An IND sponsor must submit the results of the preclinical tests, together with manufacturing information, analytical data, any available clinical data or literature and plans for clinical trials, among other things, to the FDA as part of an IND application to the FDA. An IND application is a request for authorization from the FDA to administer an investigational new drug product to humans. The central focus of an IND submission is on the general investigational plan and the protocol or protocols for preclinical studies and clinical trials. The IND application also includes results of animal and *in vitro* studies assessing the toxicology, pharmacokinetics, pharmacology and pharmacodynamic characteristics of the product, chemistry, manufacturing and controls information, and any available human data or literature to support the use of the investigational product. An IND application must become effective before human clinical trials may begin. The IND application automatically becomes effective 30 days after receipt by the FDA, unless the FDA, within the 30-day period, raises concerns or questions about the proposed clinical trial. In such a case, the IND application may be placed on clinical hold and the IND sponsor and the FDA must resolve any outstanding concerns or questions before the clinical trial can begin. Submission of an IND application therefore may or may not result in FDA authorization to begin a clinical trial. Additionally, the review of information in an IND submission may prompt FDA to, among other things, scrutinize existing INDs and could generate requests for information or clinical holds on other product candidates or programs.

Clinical trials involve the administration of the investigational product to human subjects under the supervision of qualified investigators, generally physicians not employed by or under the sponsor's control, in accordance with cGCPs, which include the requirement that all research subjects provide their informed consent for their participation in any clinical study. Clinical trials are conducted under protocols detailing, among other things, the objectives of the trial, dosing procedures, subject selection and exclusion criteria and the parameters to be used in monitoring safety and the effectiveness criteria to be evaluated. A separate submission to the existing IND application must be made for each successive clinical protocol conducted during product development and for any subsequent protocol amendments. Furthermore, an independent IRB for each site proposing to conduct the clinical trial must review and approve the plan for any clinical trial and its informed consent form before the clinical trial begins at that site, and must monitor the study until completed to ensure that the risks to individuals participating in the clinical trial are minimized and are reasonable in relation to anticipated benefits.

A sponsor who wishes to conduct a clinical trial outside of the United States may, but need not, obtain FDA authorization to conduct the clinical trial under an IND application. If a foreign clinical trial is not conducted under an IND application, the sponsor may submit data from the clinical trial to the FDA in support of a BLA.

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The FDA will accept a well-designed and well-conducted foreign clinical trial not conducted under an IND application if the foreign data are applicable to the United States population and medical practice, the trial was performed by clinical investigators of recognized competence, the trial was conducted in accordance with cGCP requirements, and the data may be considered valid without the need for an on-site inspection by the FDA or the FDA is able to validate the data through an onsite inspection if deemed necessary.

For purposes of BLA approval of a product candidate, human clinical trials are typically conducted in three sequential phases that may overlap or be combined:

- *Phase 1.* For gene therapies in general, the investigational product is initially introduced into patients with the target disease or condition. These studies are designed to test the safety, dosage tolerance, absorption, metabolism and distribution of the investigational product in humans, the side effects associated with increasing doses, and, if possible, to gain early evidence on effectiveness.
- *Phase 2.* The investigational product is administered to a limited patient population to evaluate the preliminary efficacy, optimal dosages and dosing schedule and to identify possible adverse side effects and safety risks.
- *Phase 3.* The investigational product is administered to an expanded patient population to further evaluate dosage, to provide statistically significant evidence of clinical efficacy and to further test for safety, generally at multiple geographically dispersed clinical trial sites. These clinical trials are intended to establish the overall risk/benefit ratio of the investigational product and to provide an adequate basis for product approval.

When these phases overlap or are combined, the trials may be referred to as Phase 1/2 or Phase 2/3.

In some cases, the FDA may require, or companies may voluntarily pursue, additional clinical trials after a product is approved to gain more information about the product. These so-called Phase 4 studies may be made a condition to approval of the BLA. Concurrent with clinical trials, companies may complete additional animal studies and develop additional information about the characteristics of the product candidate, and must finalize a process for manufacturing the product in commercial quantities in accordance with cGMP requirements. The manufacturing process must be capable of consistently producing quality batches of the product candidate and, among other things, must develop methods for testing the identity, strength, potency, quality and purity of the final product. Additionally, appropriate packaging must be selected and tested and stability studies must be conducted to demonstrate that the product candidate does not undergo unacceptable deterioration over its shelf life.

During all phases of clinical development, regulatory agencies require extensive monitoring and auditing of all clinical activities, clinical data and clinical study investigators. Written IND safety reports must be promptly submitted to the FDA and the investigators for serious, unexpected and suspected adverse events, any findings from other studies, tests in laboratory animals or in vitro testing that suggest a significant risk for human subjects, or any clinically important increase in the rate of a serious, and unexpected suspected adverse reaction over that listed in the protocol or investigator brochure. The sponsor must submit an IND safety report within 15 calendar days after the sponsor determines that the information qualifies for reporting. The sponsor also must notify the FDA of any unexpected fatal or life-threatening suspected adverse reaction within seven calendar days after the sponsor's initial receipt of the information. The FDA or the sponsor may suspend a clinical study at any time on various grounds, including a finding that the research patients or patients are being exposed to an unacceptable health risk. Similarly, an IRB can suspend or terminate approval of a clinical study at its institution if the clinical study is not being conducted in accordance with the IRB's requirements or if the biological product candidate has been associated with unexpected serious harm to patients. Some studies also include oversight by an independent group of qualified experts organized by the clinical study sponsor, known as a data safety monitoring board, which provides authorization for whether or not a study may move forward at designated

checkpoints based on access to certain data from the study and may recommend halting the clinical trial if it determines that there is an unacceptable safety risk for subjects or other grounds, such as no demonstration of efficacy. There are also requirements governing the reporting of ongoing clinical trials and completed clinical trial results to public registries. Sponsors of clinical trials of FDA-regulated products, including biologics, are required to register and disclose certain clinical trial information, which is publicly available at [www.clinicaltrials.gov](http://www.clinicaltrials.gov).

### ***BLA submission and review***

Assuming successful completion of all required testing in accordance with all applicable regulatory requirements, the results of product development, preclinical studies and clinical trials are submitted to the FDA as part of a BLA requesting approval to market the product for one or more indications. FDA approval of a BLA must be obtained before a biologic may be marketed in the United States. The BLA must include all relevant data available from pertinent preclinical studies and clinical trials, including negative or ambiguous results as well as positive findings, together with detailed information relating to the product's chemistry, manufacturing, controls and proposed labeling, among other things. The submission of a BLA requires payment of a substantial application user fee to the FDA, unless a waiver or exemption applies. No user fees are assessed on BLAs for products designated as orphan drugs, unless the product also includes a non-orphan indication. The FDA reviews all submitted BLAs before it accepts them for filing, and may request additional information rather than accepting the BLA for filing. The FDA has sixty days from the applicant's submission of a BLA to either issue a refusal to file letter or accept the BLA for filing, indicating that it is sufficiently complete to permit substantive review.

Once a BLA has been accepted for filing, the FDA's goal is to review standard applications within ten months after it accepts the application for filing, or, if the application qualifies for priority review, six months after the FDA accepts the application for filing. In both standard and priority reviews, the review process can be significantly extended by FDA requests for additional information or clarification. The FDA reviews a BLA to determine, among other things, whether a product is safe, pure and potent for its intended use, and whether the facility in which it is manufactured, processed, packed or held meets standards designed to ensure and preserve the product's identity, safety, strength, quality, potency and purity. The FDA may convene an advisory committee, typically a panel that includes clinicians and other experts, to provide clinical insight on applications for novel products or products which present difficult questions of safety or efficacy. The advisory committee will provide a recommendation as to whether the application should be approved and under what conditions, if any. The FDA is not bound by the recommendations of an advisory committee, but it considers such recommendations when making decisions on approval. Before approving a BLA, the FDA will typically inspect the facility or facilities where the product is manufactured. The FDA will not approve an application unless it determines that the manufacturing processes and facilities are in compliance with cGMP requirements and adequate to assure consistent production of the product within required specifications. Additionally, before approving a BLA, the FDA will typically inspect one or more clinical sites to assure compliance with cGCPs. If the FDA determines that the application, manufacturing process or manufacturing facilities are not acceptable, it will outline the deficiencies in the submission and often will request additional testing or information. Notwithstanding the submission of any requested additional information, the FDA ultimately may decide that the application does not satisfy the regulatory criteria for approval.

After the FDA evaluates a BLA and conducts inspections of manufacturing facilities where the investigational product and/or its drug substance will be manufactured, the FDA may issue an approval letter or a Complete Response Letter. An approval letter authorizes commercial marketing of the product with specific prescribing information for specific indications. A Complete Response Letter indicates that the review cycle of the application is complete, and the application will not be approved in its present form. A Complete Response

Letter will usually describe all of the deficiencies that the FDA has identified in the BLA, except that where the FDA determines that the data supporting the application are inadequate to support approval, the FDA may issue the Complete Response Letter without first conducting required inspections, testing submitted product lots and/or reviewing proposed labeling. In issuing the Complete Response Letter, the FDA may recommend actions that the applicant might take to place the BLA in condition for approval, including requests for additional information or clarification, which may include the potential requirement for additional preclinical studies or clinical trials or additional manufacturing activities. If a Complete Response Letter is issued, the applicant may either resubmit the BLA, addressing all of the deficiencies identified in the letter, or withdraw the application or request an opportunity for a hearing. The FDA may delay or refuse approval of a BLA if applicable regulatory criteria are not satisfied, require additional testing or information and/or require post-marketing testing and surveillance to monitor the safety or efficacy of a product.

If regulatory approval of a product is granted, such approval will be granted for particular indications and may entail limitations on the indicated uses for which such product may be marketed. For example, the FDA may approve the BLA with a Risk Evaluation and Mitigation Strategy, or REMS, to ensure the benefits of the product outweigh its risks. A REMS is a safety strategy to manage a known or potential serious risk associated with a product and to enable patients to have continued access to such medicines by managing their safe use, and could include medication guides, physician communication plans, or elements to assure safe use, such as restricted distribution methods, patient registries and other risk minimization tools. The FDA also may condition approval on, among other things, changes to proposed labeling or the development of adequate controls and specifications. Once approved, the FDA may withdraw the product approval if compliance with pre- and post-marketing requirements is not maintained or if problems occur after the product reaches the marketplace. The FDA may require one or more Phase 4 post-market studies and surveillance to further assess and monitor the product's safety and effectiveness after commercialization, and may limit further marketing of the product based on the results of these post-marketing studies.

### ***Expedited development and review programs***

The FDA offers a number of expedited development and reviews programs for qualifying product candidates. The Fast Track program is intended to expedite or facilitate the process for developing new products that meet certain criteria. Specifically, product candidates are eligible for Fast Track designation if they are intended to treat a serious or life-threatening disease or condition and demonstrate the potential to address unmet medical needs for the disease or condition. Fast Track designation applies to the combination of the product candidate and the specific indication for which it is being studied. The sponsor of a Fast Track designated product candidate has opportunities for frequent interactions with the FDA review team during product development and, once a BLA is submitted, the product candidate may be eligible for priority review. A Fast Track designated product candidate may also be eligible for rolling review, where the FDA may consider for review sections of the BLA on a rolling basis before the complete application is submitted, if the sponsor provides a schedule for the submission of the sections of the BLA, the FDA agrees to accept sections of the BLA and determines that the schedule is acceptable, and the sponsor pays any required user fees upon submission of the first section of the BLA. The sponsor can request the FDA to designate the product candidate for Fast Track status any time before receiving BLA approval, but ideally no later than the pre-BLA meeting.

A product candidate intended to treat a serious or life-threatening disease or condition may also be eligible for Breakthrough Therapy designation to expedite its development and review. A product candidate may receive Breakthrough Therapy designation if preliminary clinical evidence indicates that the product candidate, alone or in combination with one or more other drugs or biologics, may demonstrate substantial improvement over available therapies on one or more clinically significant endpoints, such as substantial treatment effects observed early in clinical development. The designation includes all of the Fast Track program features, as well as more

intensive FDA interaction and guidance beginning as early as Phase 1 and an organizational commitment to expedite the development and review of the product candidate, including involvement of senior managers.

As part of the 21st Century Cures Act, Congress amended the FDCA to facilitate an efficient development program for, and expedite review of regenerative medicine therapies, which include cell therapies, therapeutic tissue engineering products, human cell and tissue products, and combination products using any such therapies or products. Gene therapies, including genetically modified cells that lead to a durable modification of cells or tissues may meet the definition of a “regenerative medicine therapy.” A product candidate may be eligible for regenerative medicine therapy, or RMAT, designation if it meets the following criteria: (1) it is a regenerative medicine therapy; (2) it is intended to treat, modify, reverse, or cure a serious or life-threatening disease or condition; and (3) preliminary clinical evidence indicates that it has the potential to address unmet medical needs for such a disease or condition. A sponsor may request that the FDA designate a product candidate as an RMAT concurrently with or at any time after the submission of an IND. A BLA for a product candidate that has received RMAT designation may be eligible for priority review or accelerated approval through (1) surrogate or intermediate endpoints reasonably likely to predict long-term clinical benefit or (2) reliance upon data obtained from a meaningful number of sites. Benefits of such designation also include early interactions with FDA to discuss any potential surrogate or intermediate endpoint to be used to support accelerated approval. A product candidate with RMAT designation that is granted accelerated approval and is subject to post-approval requirements may fulfill such requirements through the submission of clinical evidence, clinical studies, patient registries, or other sources of real world evidence, such as electronic health records; the collection of larger confirmatory data sets; or post-approval monitoring of all patients treated with such therapy prior to its approval.

Any marketing application for a biologic submitted to the FDA for approval, including a product with a Fast Track designation and/or Breakthrough Therapy designation, may be eligible for other types of FDA programs intended to expedite the FDA review and approval process, such as priority review. A product is eligible for priority review if it is designed to treat a serious or life threatening disease or condition and, if approved, would provide a significant improvement in safety and effectiveness compared to available therapies. For original BLAs, priority review designation means the FDA’s goal is to take action on the marketing application within six months of the 60-day filing date.

Additionally, a product candidate may be eligible for accelerated approval if it is designed to treat a serious or life-threatening disease or condition and demonstrates an effect on a surrogate endpoint that is reasonably likely to predict clinical benefit, or on a clinical endpoint that can be measured earlier than irreversible morbidity or mortality, that is reasonably likely to predict an effect on irreversible morbidity or mortality or other clinical benefit, taking into account the severity, rarity, or prevalence of the condition and the availability or lack of alternative treatments. As a condition of accelerated approval, the FDA will generally require the sponsor to perform adequate and well-controlled post-marketing clinical studies with due diligence to verify and describe the anticipated effect on irreversible morbidity or mortality or other clinical benefit. The FDA may withdraw approval of a product or indication approved under accelerated approval if, for example, the confirmatory trial fails to verify the predicted clinical benefit of the product. In addition, for products being considered for accelerated approval, the FDA generally requires, unless otherwise informed by the agency, pre-approval of all advertising and promotional materials intended for dissemination or publication, which could adversely impact the timing of the commercial launch of the product.

Fast Track designation, Breakthrough Therapy designation, RMAT designation, and priority review do not change the standards for approval but may expedite the development or approval process. Even if a product qualifies for one or more of these programs, the FDA may later decide that the product no longer meets the conditions for qualification or decide that the time period for FDA review or approval will not be shortened.

### ***Orphan Drug designation***

Under the Orphan Drug Act, the FDA may grant Orphan Drug designation to a product candidate intended to treat a rare disease or condition, which is generally a disease or condition that affects fewer than 200,000 individuals in the United States, or 200,000 or more individuals in the United States for which there is no reasonable expectation that the cost of developing and making available in the United States a drug or biologic for this type of disease or condition will be recovered from sales in the United States for that product candidate. Orphan Drug designation must be requested before submitting a BLA. After the FDA grants Orphan Drug designation, the identity of the therapeutic agent and its potential orphan use are disclosed publicly by the FDA. The Orphan Drug designation does not convey any advantage in, or shorten the duration of, the regulatory review or approval process.

If a product that has Orphan Drug designation subsequently receives the first FDA approval for the disease or condition for which it has such designation, the product is entitled to orphan drug exclusive approval (or exclusivity), which means that the FDA may not approve any other applications, including a full BLA, to market the same product for the same indication for seven years, except in limited circumstances, such as a showing of clinical superiority to the product with orphan drug exclusivity by means of greater effectiveness, greater safety or providing a major contribution to patient care or if the holder of the orphan drug exclusivity cannot assure the availability of sufficient quantities of the orphan drug to meet the needs of patients with the disease or condition for which the product was designated. Orphan drug exclusivity does not prevent the FDA from approving a different drug or biologic for the same disease or condition, or the same drug or biologic for a different disease or condition. Among the other benefits of Orphan Drug designation are tax credits for certain research and a waiver of the BLA application fee.

A designated orphan drug may not receive orphan drug exclusivity if it is approved for a use that is broader than the indication for which it received Orphan Drug designation. In addition, exclusive marketing rights in the United States may be lost if the FDA later determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantities of the product to meet the needs of patients with the rare disease or condition.

### ***Rare Pediatric Disease designation and priority review vouchers***

Under the FDCA, as amended, the FDA incentivizes the development of product candidates that meet the definition of a “rare pediatric disease,” defined to mean a serious or life-threatening disease in which the serious or life-threatening manifestations primarily affect individuals aged from birth to 18 years and the disease affects fewer than 200,000 individuals in the United States or affects 200,000 or more in the United States and for which there is no reasonable expectation that the cost of developing and making in the United States a drug for such disease or condition will be recovered from sales in the United States of such drug. The sponsor of a product candidate for a rare pediatric disease may be eligible for a voucher that can be used to obtain a priority review for a subsequent human drug or biologic application after the date of approval of the rare pediatric disease drug product. A sponsor may request Rare Pediatric Disease designation from the FDA prior to the submission of its BLA. A Rare Pediatric Disease designation does not guarantee that a sponsor will receive a priority review voucher, or PRV, upon approval of its BLA. Moreover, a sponsor who chooses not to submit a Rare Pediatric Disease designation request may nonetheless receive a PRV upon approval of their marketing application if they request such a voucher in their original marketing application and meet all of the eligibility criteria. If a PRV is received, it may be sold or transferred an unlimited number of times. Congress has extended the PRV program until September 30, 2024, with the potential for PRVs to be granted until September 30, 2026.



***Pediatric information and pediatric exclusivity***

Under the Pediatric Research Equity Act, or PREA, certain BLAs and certain supplements to a BLA must contain data to assess the safety and efficacy of the product for the claimed indications in all relevant pediatric subpopulations and to support dosing and administration for each pediatric subpopulation for which the product is safe and effective. The FDA may grant deferrals for the submission of pediatric data or full or partial waivers. The Food and Drug Administration Safety and Innovation Act, or FDASIA, amended the FDCA to require that a sponsor who is planning to submit a marketing application for a product that includes a new active ingredient, new indication, new dosage form, new dosing regimen or new route of administration submit an initial Pediatric Study Plan, or PSP, within 60 days of an end-of-Phase 2 meeting or, if there is no such meeting, as early as practicable before the initiation of the Phase 3 or Phase 2/3 study. The initial PSP must include an outline of the pediatric study or studies that the sponsor plans to conduct, including study objectives and design, age groups, relevant endpoints and statistical approach, or a justification for not including such detailed information, and any request for a deferral of pediatric assessments or a full or partial waiver of the requirement to provide data from pediatric studies along with supporting information. The FDA and the sponsor must reach an agreement on the PSP. A sponsor can submit amendments to an agreed-upon initial PSP at any time if changes to the pediatric plan need to be considered based on data collected from preclinical studies, early phase clinical trials and/or other clinical development programs. A biological product can also obtain pediatric market exclusivity in the United States. Pediatric exclusivity, if granted, adds six months to existing exclusivity periods. This six-month exclusivity, which runs from the end of other exclusivity protection, may be granted based on the voluntary completion of a pediatric study in accordance with an FDA-issued “Written Request” for such a study.

***Post-approval requirements***

Any products manufactured or distributed by us pursuant to FDA approvals are subject to pervasive and continuing regulation by the FDA, including, among other things, requirements relating to record-keeping, reporting of adverse experiences, periodic reporting, product sampling and distribution, and advertising and promotion of the product. As part of the manufacturing process, the manufacturer is required to perform certain tests on each lot of the product before it is released for distribution. After a BLA is approved for a biological product, the product also may be subject to official lot release. If the product is subject to official release by the FDA, the manufacturer submits samples of each lot of product to the FDA together with a release protocol showing a summary of the history of manufacture of the lot and the results of all of the manufacturer’s tests performed on the lot. The FDA also may perform certain confirmatory tests on lots of some products before releasing the lots for distribution by the manufacturer. In addition, the FDA conducts laboratory research related to the regulatory standards on the safety, purity, potency and effectiveness of biologics. After approval, most changes to the approved product, such as adding new indications or other labeling claims, are subject to prior FDA review and approval. There also are continuing user fee requirements, under which the FDA assesses an annual program fee for each product identified in an approved BLA. Biologic manufacturers and their subcontractors are required to register their establishments with the FDA and certain state agencies, and are subject to periodic unannounced inspections by the FDA and certain state agencies for compliance with cGMPs, which impose certain procedural and documentation requirements upon us and our third-party manufacturers. Changes to the manufacturing process are strictly regulated, and, depending on the significance of the change, may require prior FDA approval before being implemented. FDA regulations also require investigation and correction of any deviations from cGMPs and impose reporting requirements upon us and any third-party manufacturers that we may decide to use. Manufacturers and other parties involved in the drug supply chain for prescription drug products must also comply with product tracking and tracing requirements and for notifying the FDA of counterfeit, diverted, stolen and intentionally adulterated products or products that are otherwise unfit for distribution in the United States. Accordingly, manufacturers must continue to expend time, money and effort in the area of production and quality control to maintain compliance with cGMPs and other aspects of regulatory compliance.

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The FDA may withdraw approval if compliance with regulatory requirements and standards is not maintained or if problems occur after the product reaches the market. Later discovery of previously unknown problems with a product, including adverse events of unanticipated severity or frequency, or with manufacturing processes, or failure to comply with regulatory requirements, may result in revisions to the approved labeling to add new safety information; imposition of post-market studies or clinical studies to assess new safety risks; or imposition of distribution restrictions or other restrictions under a REMS program. Other potential consequences include, among other things:

- restrictions on the marketing or manufacturing of a product, complete withdrawal of the product from the market or product recalls;
- fines, warning or untitled letters or holds on post-approval clinical studies;
- refusal of the FDA to approve pending applications or supplements to approved applications, or suspension or revocation of existing product approvals;
- product seizure or detention, or refusal of the FDA to permit the import or export of products;
- consent decrees, corporate integrity agreements, debarment or exclusion from federal healthcare programs;
- mandated modification of promotional materials and labeling and the issuance of corrective information;
- the issuance of safety alerts, Dear Healthcare Provider letters, press releases and other communications containing warnings or other safety information about the product; or
- injunctions or the imposition of civil or criminal penalties.

The FDA closely regulates the marketing, labeling, advertising and promotion of biologics. A company can make only those claims relating to safety and efficacy, purity and potency that are approved by the FDA and in accordance with the provisions of the approved label. However, companies may share truthful and not misleading information that is otherwise consistent with a product's FDA-approved labeling. The FDA and other agencies actively enforce the laws and regulations prohibiting the promotion of off-label uses. Failure to comply with these requirements can result in, among other things, adverse publicity, warning letters, corrective advertising and potential civil and criminal penalties. Physicians may prescribe legally available products for uses that are not described in the product's labeling and that differ from those tested by us and approved by the FDA. Such off-label uses are common across medical specialties. Physicians may believe that such off-label uses are the best treatment for many patients in varied circumstances. The FDA does not regulate the behavior of physicians in their choice of treatments. The FDA does, however, restrict manufacturer's communications on the subject of off-label use of their products.

### ***Biosimilars and reference product exclusivity***

The ACA includes a subtitle called the Biologics Price Competition and Innovation Act, or BPCIA, which created an abbreviated approval pathway for biologics that are biosimilar to or interchangeable with an FDA-approved reference biological product. To date, a number of biosimilars have been licensed under the BPCIA, and numerous biosimilars have been approved in Europe. The FDA has issued several guidance documents outlining an approach to review and approval of biosimilars.

Biosimilarity, which requires that there be no clinically meaningful differences between the biological product and the reference product in terms of safety, purity and potency, can be shown through analytical studies, animal studies and a clinical study or studies. Interchangeability requires that a product is biosimilar to the reference product and the product must demonstrate that it can be expected to produce the same clinical results as the

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reference product in any given patient and, for products that are administered multiple times to an individual, the biologic and the reference biologic may be alternated or switched after one has been previously administered without increasing safety risks or risks of diminished efficacy relative to exclusive use of the reference biologic. Complexities associated with the larger, and often more complex, structures of biologics, as well as the processes by which such products are manufactured, pose significant hurdles to the implementation of the abbreviated approval pathway that are still being worked out by the FDA.

Under the BPCIA, an application for a biosimilar product may not be submitted to the FDA until four years following the date that the reference product was first licensed by the FDA. In addition, the approval of a biosimilar product may not be made effective by the FDA until 12 years from the date on which the reference product was first licensed. During this 12-year period of exclusivity, another company may still market a competing version of the reference product if the FDA approves a full BLA for the competing product containing that applicant's own preclinical data and data from adequate and well-controlled clinical trials to demonstrate the safety, purity and potency of its product. The BPCIA also created certain exclusivity periods for biosimilars approved as interchangeable products. At this juncture, it is unclear whether products deemed "interchangeable" by the FDA will, in fact, be readily substituted by pharmacies, which are governed by state pharmacy law.

The BPCIA is complex and continues to be interpreted and implemented by the FDA. In addition, government proposals have sought to reduce the 12-year reference product exclusivity period. Other aspects of the BPCIA, some of which may impact the BPCIA exclusivity provisions, have also been the subject of recent litigation. As a result, the ultimate impact and implementation of the BPCIA are subject to significant uncertainty.

### ***Regulation of companion diagnostics***

If the safe and effective use of a therapeutic depends on an *in vitro* diagnostic, then the FDA generally will require approval or clearance of that diagnostic, known as a companion diagnostic, at the same time that the FDA approves the therapeutic product. In August 2014, the FDA issued final guidance clarifying the requirements that will apply to approval of therapeutic products and *in vitro* companion diagnostics. According to the guidance, for novel candidates, a companion diagnostic device and its corresponding therapeutic should be approved or cleared contemporaneously by the FDA for the use indicated in the therapeutic product's labeling. Once cleared or approved, the companion diagnostic must adhere to post-marketing requirements including the requirements of FDA's quality system regulation, medical device reporting, recalls and corrections along with product marketing requirements and limitations. Companion diagnostic manufacturers are subject to unannounced FDA inspections at any time during which the FDA will conduct an audit of the product(s) and the company's facilities for compliance with its authorities.

### ***Foreign regulation***

In order to market any product outside of the United States, we would need to comply with numerous and varying regulatory requirements of other countries and jurisdictions regarding quality, safety, and efficacy and governing, among other things, clinical trials, marketing authorization, commercial sales and distribution of our products. Whether or not we obtain FDA approval for a product, we would need to obtain the necessary approvals by the comparable foreign regulatory authorities before we can commence clinical trials or marketing of the product in foreign countries and jurisdictions. Although many of the issues discussed above with respect to the United States apply similarly in the context of the European Union, the approval process varies between countries and jurisdictions and can involve additional product testing and additional administrative review periods. The time required to obtain approval in other countries and jurisdictions might differ from and be longer than that required to obtain FDA approval. Regulatory approval in one country or jurisdiction does not ensure regulatory approval in another, but a failure or delay in obtaining regulatory approval in one country or jurisdiction may negatively impact the regulatory process in others.

***Other healthcare laws and compliance requirements***

Pharmaceutical companies are subject to additional healthcare regulation and enforcement by the federal government and by authorities in the states and foreign jurisdictions in which they conduct their business, particularly once they have a commercialized product that is reimbursable by third-party payor programs. Such laws include, without limitation: the U.S. federal Anti-Kickback Statute, the civil False Claims Act, the federal Health Insurance Portability and Accountability Act of 1996, or HIPAA, and similar foreign, federal and state fraud and abuse, transparency and privacy laws.

- The U.S. federal Anti-Kickback Statute prohibits, among other things, persons and entities from knowingly and willfully soliciting, receiving, offering or paying remuneration, to induce, or in return for, either the referral of an individual, or the purchase or recommendation of an item or service for which payment may be made under any federal healthcare program. The term remuneration has been interpreted broadly to include anything of value, including stock options. There are a number of statutory exceptions and regulatory safe harbors protecting some common activities from prosecution, but they are drawn narrowly, and practices that involve remuneration, such as consulting agreements, that may be alleged to be intended to induce prescribing, purchasing or recommending may be subject to scrutiny if they do not qualify for an exception or safe harbor. Our practices may not in all cases meet all of the criteria for protection under a statutory exception or regulatory safe harbor. A person or entity does not need to have actual knowledge of the statute or specific intent to violate it in order to have committed a violation. Violations are subject to civil and criminal fines and penalties for each violation, plus up to three times the remuneration involved, imprisonment, and exclusion from government healthcare programs. In addition, a claim including items or services resulting from a violation of the U.S. federal Anti-Kickback Statute constitutes a false or fraudulent claim for purposes of the civil False Claims Act.
- Civil and criminal false claims laws, and civil monetary penalty laws, including the civil False Claims Act, which can be enforced through civil whistleblower or qui tam actions, prohibit, among other things, individuals or entities from knowingly presenting, or causing to be presented, claims for payment to the federal government, including federal healthcare programs, that are false or fraudulent. For example, the civil False Claims Act prohibits any person or entity from knowingly presenting, or causing to be presented, a false claim for payment to the federal government or knowingly making, using or causing to be made or used a false record or statement material to a false or fraudulent claim to the federal government.
- HIPAA created additional federal civil and criminal liability for, among other things, knowingly and willfully executing a scheme to defraud any healthcare benefit program, or obtain, by means of false or fraudulent pretenses, representations, or promises, any of the money or property owned by, or under the custody or control of, any healthcare benefit program, regardless of the payor (e.g., public or private) and knowingly and willfully falsifying, concealing, or covering up by any trick or device a material fact or making any materially false statements in connection with the delivery of, or payment for, healthcare benefits, items or services relating to healthcare matters. Similar to the U.S. federal Anti-Kickback Statute, a person or entity can be found guilty of violating HIPAA without actual knowledge of the statute or specific intent to violate it.
- HIPAA, as amended by the Health Information Technology for Economic and Clinical Health Act of 2009, or HITECH, and their respective implementing regulations, impose certain requirements on HIPAA covered entities, which include certain healthcare providers, healthcare clearing houses and health plans, and individuals and entities that provide services on their behalf that involve individually identifiable health information, known as business associates, relating to the privacy, security and transmission of individually identifiable health information, as well as their covered subcontractors. HITECH also created new tiers of civil monetary penalties, amended HIPAA to make civil and criminal penalties directly applicable to business

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associates, and gave state attorneys general new authority to file civil actions for damages or injunctions in federal courts to enforce the federal HIPAA laws and seek attorneys' fees and costs associated with pursuing federal civil actions.

- The U.S. federal Physician Payments Sunshine Act requires certain manufacturers of drugs, devices, biologics and medical supplies for which payment is available under Medicare, Medicaid or the Children's Health Insurance Program, with specific exceptions, to annually report to the Centers for Medicare and Medicaid Services, or CMS, information related to payments or other transfers of value made to physicians (defined to include doctors, dentists, optometrists, podiatrists and chiropractors), certain other licensed healthcare professionals (such as physician assistants and nurse practitioners), and teaching hospitals, as well as ownership and investment interests held by physicians and their immediate family members.

We may also be subject to additional similar U.S. state and foreign law equivalents of each of the above federal laws, such as anti-kickback and false claims laws which may apply to sales or marketing arrangements and claims involving healthcare items or services reimbursed by non-governmental third-party payors, including private insurers, or that apply regardless of payor, state laws which require pharmaceutical companies to comply with the pharmaceutical industry's voluntary compliance guidelines and the relevant compliance guidance promulgated by the federal government, state and local laws which require pharmaceutical companies to report information related to payments and other transfers of value to physicians and other healthcare providers or marketing expenditures, state laws which require the reporting of information related to product pricing, state and local laws requiring the registration of pharmaceutical sales representatives, and state and foreign laws governing the privacy and security of health information which, in some cases, differ from each other in significant ways, and may not have the same effect, thus complicating compliance efforts. If our operations are found to be in violation of any of such laws or any other governmental regulations that apply, we may be subject to penalties, including, without limitation, significant civil, criminal and administrative penalties, damages, fines, exclusion from government-funded healthcare programs, such as Medicare and Medicaid or similar programs in other countries or jurisdictions, integrity oversight and reporting obligations to resolve allegations of non-compliance, disgorgement, imprisonment, contractual damages, reputational harm, diminished profits and the curtailment or restructuring of our operations.

### ***Coverage and reimbursement***

Significant uncertainty exists as to the coverage and reimbursement status of any pharmaceutical or biological product for which we obtain regulatory approval in the future. Sales of any product, if approved, depend, in part, on the extent to which such product will be covered by third-party payors, such as federal, state and foreign government healthcare programs, commercial insurance and managed healthcare organizations, and the level of reimbursement, if any, for such product by third-party payors. Decisions regarding whether to cover any of our product candidates, if approved, the extent of coverage and amount of reimbursement to be provided are made on a plan-by-plan basis. Further, no uniform policy for coverage and reimbursement exists in the United States, and coverage and reimbursement can differ significantly from payor to payor.

Third-party payors often rely upon Medicare coverage policy and payment limitations in setting their own reimbursement rates, but also have their own methods and approval process apart from Medicare determinations. As a result, the coverage determination process is often a time-consuming and costly process that will require us to provide scientific and clinical support for the use of our product candidates to each payor separately, with no assurance that coverage and adequate reimbursement will be applied consistently or obtained in the first instance. For example, the FDA recently approved a monoclonal antibody treatment for Alzheimer's disease, aducanumab (Aduhelm), which is the first Alzheimer's disease drug to be approved in nearly 20 years. On April 7, 2022, CMS released a national coverage determination for FDA-approved monoclonal antibodies directed against amyloid

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for the treatment of Alzheimer's disease, including aducanumab (Aduhelm). Further, on January 6, 2023, CMS announced that this coverage determination also applies to lecanemab (Leqembi). Nevertheless, it is possible that CMS may not approve Medicare coverage and reimbursement for any of our product candidates for treatment of Alzheimer's disease, once approved.

Furthermore, it is possible that one or more of our product candidates may not be considered medically necessary or cost effective. A decision by a third-party payor not to cover any product candidates we may develop could reduce physician utilization of such product candidates once approved and have a material adverse effect on our sales, results of operations and financial condition.

For products administered under the supervision of a physician, obtaining coverage and adequate reimbursement may be particularly difficult because of the higher prices often associated with such drugs. Additionally, separate reimbursement for the product itself or the treatment or procedure in which the product is used may not be available, which may impact physician utilization.

In addition, the U.S. government, state legislatures and foreign governments have continued implementing cost-containment programs, including price controls, restrictions on coverage and reimbursement and requirements for substitution of generic products. Third-party payors are increasingly challenging the prices charged for medical products and services, examining the medical necessity and reviewing the cost-effectiveness of pharmaceutical or biologics, medical devices and medical services, in addition to questioning safety and efficacy. Adoption of price controls and cost-containment measures, and adoption of more restrictive policies in jurisdictions with existing controls and measures, could further limit sales of any product that receives approval. Decreases in third-party reimbursement for any product or a decision by a third party not to cover a product could reduce physician usage and patient demand for the product. In addition, companion diagnostic tests require coverage and reimbursement separate and apart from the coverage and reimbursement for their companion pharmaceutical or biological products. Similar challenges to obtaining coverage and reimbursement, applicable to pharmaceutical or biological products, will apply to companion diagnostics.

Outside the United States, ensuring adequate coverage and payment for any biological candidates we may develop will face challenges. Pricing of prescription pharmaceuticals is subject to governmental control in many countries. Pricing negotiations with governmental authorities can extend well beyond the receipt of regulatory marketing approval for a product and may require us to conduct a clinical trial that compares the cost effectiveness of any product candidates we may develop to other available therapies. The conduct of such a clinical trial could be expensive and result in delays in our commercialization efforts. In the European Union, pricing and reimbursement schemes vary widely from country to country. Some countries provide that products may be marketed only after a reimbursement price has been agreed. Some countries may require the completion of additional studies that compare the cost-effectiveness of a particular product candidate to currently available therapies (so called health technology assessments) in order to obtain reimbursement or pricing approval. For example, the European Union provides options for its member states to restrict the range of products for which their national health insurance systems provide reimbursement and to control the prices of medicinal products for human use. European Union member states may approve a specific price for a product or it may instead adopt a system of direct or indirect controls on the profitability of the company placing the product on the market. Other member states allow companies to fix their own prices for products but monitor and control prescription volumes and issue guidance to physicians to limit prescriptions. Recently, many countries in the European Union have increased the amount of discounts required on pharmaceuticals and these efforts could continue as countries attempt to manage healthcare expenditures, especially in light of the severe fiscal and debt crises experienced by many countries in the European Union. The downward pressure on healthcare costs in general, particularly prescription products, has become intense. As a result, increasingly high barriers are being erected to the entry of new products. Political, economic and regulatory developments may further complicate pricing negotiations, and

pricing negotiations may continue after reimbursement has been obtained. Reference pricing used by various European Union member states, and parallel trade (arbitrage between low-priced and high-priced member states), can further reduce prices. There can be no assurance that any country that has price controls or reimbursement limitations for pharmaceutical products will allow favorable reimbursement and pricing arrangements for any of our products, if approved in those countries.

## **Healthcare reform**

The United States and some foreign jurisdictions are considering or have enacted a number of reform proposals to change the healthcare system. There is significant interest in promoting changes in healthcare systems with the stated goals of containing healthcare costs, improving quality or expanding access. In the United States, the pharmaceutical industry has been a particular focus of these efforts and has been significantly affected by federal and state legislative initiatives, including those designed to limit the pricing, coverage and reimbursement of pharmaceutical and biopharmaceutical products, especially under government-funded healthcare programs, and increased governmental control of drug pricing.

The ACA, which was enacted in 2010, substantially changed the way healthcare is financed by both governmental and private insurers in the United States, and significantly affected the pharmaceutical industry. In addition, other legislative and regulatory changes have been proposed and adopted in the United States since the ACA was enacted:

- The Budget Control Act of 2011 and subsequent legislation, among other things, created measures for spending reductions by Congress that include aggregate reductions of Medicare payments to providers of 2% per fiscal year, which remain in effect through 2032. Due to the Statutory Pay-As-You-Go Act of 2010, estimated budget deficit increases resulting from the American Rescue Plan Act of 2021, and subsequent legislation, Medicare payments to providers will be further reduced starting in 2025 absent further legislation. The U.S. American Taxpayer Relief Act of 2012 further reduced Medicare payments to several types of providers and increased the statute of limitations period for the government to recover overpayments to providers from three to five years.
- On April 13, 2017, CMS published a final rule that gives states greater flexibility in setting benchmarks for insurers in the individual and small group marketplaces, which may have the effect of relaxing the essential health benefits required under the ACA for plans sold through such marketplaces.
- On May 23, 2019, CMS published a final rule to allow Medicare Advantage Plans the option of using step therapy for Part B drugs beginning January 1, 2020.

Additionally, there has been heightened governmental scrutiny over the manner in which manufacturers set prices for their marketed products, which has resulted in several U.S. presidential executive orders, Congressional inquiries and proposed and enacted federal legislation designed to, among other things, bring more transparency to drug pricing, reduce the cost of prescription drugs under Medicare, and review the relationship between pricing and manufacturer patient programs. The Inflation Reduction Act of 2022, or IRA, includes several provisions that may impact our business to varying degrees, including provisions that reduce the out-of-pocket spending cap for Medicare Part D beneficiaries from \$7,050 to \$2,000 starting in 2025, thereby effectively eliminating the coverage gap; impose new manufacturer financial liability on certain drugs under Medicare Part D, allow the U.S. government to negotiate Medicare Part B and Part D price caps for certain high-cost drugs and biologics without generic or biosimilar competition; require companies to pay rebates to Medicare for certain drug prices that increase faster than inflation; and delay until January 1, 2032 the implementation of the HHS rebate rule that would have limited the fees that pharmacy benefit managers can charge. The IRA permits HHS to implement many of these provisions through guidance, as opposed to regulation, for the initial

years. These provisions will take effect progressively starting in fiscal year 2023, although they may be subject to legal challenges. HHS has and will continue to issue and update guidance as these programs are implemented. Further, under the IRA, orphan drugs are exempted from the Medicare drug price negotiation program, but only if they have one rare disease designation and for which the only approved indication is for that disease or condition. If a product receives multiple rare disease designations or has multiple approved indications, it may not qualify for the orphan drug exemption. It is currently unclear how the IRA will be implemented but is likely to have a significant impact on the pharmaceutical industry.

In addition, in response to the Biden administration's October 2022 executive order, on February 14, 2023, HHS released a report outlining three new models for testing by the Center for Medicare and Medicaid Innovation which will be evaluated on their ability to lower the cost of drugs, promote accessibility, and improve quality of care. It is unclear whether the models will be utilized in any health reform measures in the future. Although a number of these and other proposed measures may require authorization through additional legislation to become effective, and the Biden administration may reverse or otherwise change these measures, both the Biden administration and Congress have indicated that they will continue to seek new legislative measures to control drug costs.

At the state level, legislatures have increasingly passed legislation and implemented regulations designed to control pharmaceutical product pricing, including price or patient reimbursement constraints, discounts, restrictions on certain product access and marketing cost disclosure and transparency measures, and, in some cases, designed to encourage importation from other countries and bulk purchasing.

## **Facilities**

In January 2022, we entered into a lease agreement for our New York headquarters located at 345 Park Avenue South, 6th Floor, New York, NY 10010, pursuant to which we lease a total of approximately 15,839 square feet of office and laboratory space used for our research and development, general and administrative, and other activities. This lease agreement began in April 2022 and expires in July 2029 with an additional five-year option to extend at the then prevailing effective market rental rate. We believe that our new facilities are sufficient to meet our current needs and that suitable additional space will be available as and when needed.

## **Employees and human capital resources**

As of April 30, 2023, we had 39 full-time employees and one part-time employee. Of our 40 full-time and part-time employees, 31 are engaged in research and development activities. None of our employees are represented by labor unions or covered by collective bargaining agreements. We consider our relationship with our employees to be good.

Our human capital resources objectives include, as applicable, identifying, recruiting, retaining, incentivizing and integrating our employees. We believe our success depends on our ability to attract, retain, develop and motivate diverse highly skilled personnel. In particular, we depend upon the personal efforts and abilities of the principal members of our senior management to partner effectively as a team, and to provide strategic direction, develop our business, manage our operations and maintain a cohesive and stable work environment. We also rely on qualified managers and skilled employees, such as scientists, medical professionals, engineers and laboratory technicians, with technical expertise in operations, scientific knowledge, engineering skills and quality management experience in order to operate our business successfully.

Our compensation program is designed to retain, motivate and, as needed, attract highly qualified employees. Accordingly, we use a mix of competitive base salary, performance-based equity compensation awards and other employee benefits.



## **Legal proceedings**

From time to time, we may become involved in legal proceedings arising in the ordinary course of our business. We are not currently subject to any material legal proceedings.

# Management

## Executive officers and directors

The following table provides information regarding our current executive officers and directors, including their ages as of March 31, 2023:

Name	Age	Position(s)
<b>Executive Officers</b>		
R. Nolan Townsend	43	Chief Executive Officer and Director
Eric Adler, M.D.	49	Chief Scientific Officer
Paul McCormac, Ph.D.	53	Chief Technical Officer
Jenny R. Robertson	49	Senior Vice President, General Counsel and Head of People
Micah Zajic	42	Chief Financial Officer
<b>Non-Employee Directors</b>		
Steven Altschuler, M.D.	69	Chairman of the Board of Directors
Mette Kirstine Agger	58	Director
Paula HJ Cholmondeley	76	Director
Bernard Davitian	62	Director
Reinaldo Diaz	69	Director

## Executive officers

**R. Nolan Townsend** has served as our Chief Executive Officer and as a member of our board of directors since January 2020. Before joining Lexeo, Mr. Townsend was at Pfizer Inc., a global pharmaceutical company, from 2008 to December 2019, where he held several roles of increasing responsibility. Most recently, he served as President, Pfizer Rare Disease for the North America region. In that role, Mr. Townsend was responsible for overseeing the division's overall strategy, cross functional organization and operating budget. Mr. Townsend received a B.A. in Economics from the University of Pennsylvania and an M.B.A. from the Harvard Business School. Our board of directors believes Mr. Townsend is qualified to serve as a director because of his role as our Chief Executive Officer and his experience as an executive in the biopharmaceutical industry.

**Eric Adler, M.D.**, has served as our Chief Scientific Officer since August 2022. Before joining Lexeo and since July 2011, Dr. Adler has worked at the University of California, San Diego, or UCSD, where he now serves as Professor and Section Head of Heart Failure. He is also the Director of the Strauss Center for Cardiomyopathy and Czarina M. and Humberto S. Lopez Endowed Chair in Cardiology at UCSD. His work has led to the development of a novel investigational gene therapy for Danon disease, which is currently in Phase 1 clinical development, and he serves on multiple advisory boards, including for Rocket Pharmaceuticals, Ionis Pharmaceuticals, and Sana Biotechnology. Dr. Adler's research is focused on the study and treatment of cardiomyopathy and he has published over 100 papers in peer-reviewed journals on the topic. He is currently an associate editor of *Circulation Heart Failure*, and he has served on leadership, grant review, and guideline committees for the American Heart Association, the Heart Failure Society of America, the International Society of Heart and Lung Transplant, and the National Institute of Health. Dr. Adler also performs clinical research on heart failure and has been a principal investigator in numerous cardiovascular clinical trials. Dr. Adler received a B.A. in English from Northwestern University and an M.D. from the Boston University School of Medicine.

**Paul McCormac, Ph.D.**, has served as our Chief Technical Officer since December 2021, prior to this he served as our Senior Vice President of Technical Operations since March 2021. Before joining Lexeo, Dr. McCormac

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was an Executive Director, Medicinal Sciences Category Lead for Pfizer Rare Disease, a unit of Pfizer, Inc., a global pharmaceutical company, from October 2016 until March 2021. In that role, he managed chemistry, manufacturing, and controls, or CMC, for product development with a focus on Pfizer's gene therapy CMC strategy. Prior to this, he served as Director/Team Leader for Pfizer's Bio-Manufacturing Sciences group from November 2008 until October 2016. Dr. McCormac received a B.S. in Analytical Science and a Ph.D. in Organic Chemistry from Dublin City University and completed a post-doctoral fellowship at Queen's University Belfast.

**Jenny R. Robertson, J.D.**, has served as our Senior Vice President, General Counsel and Head of People since January 2023. She previously served as General Counsel and Company Secretary from March 2022 through December 2022. Before joining Lexeo, from April 2010 to March 2022, Ms. Robertson served in various senior legal roles at Pfizer Inc., a global pharmaceutical company, most recently as Vice President and Chief Counsel to Pfizer's Oncology Business Unit. Prior to that role, Ms. Robertson served as Chief Counsel for Pfizer's Rare Disease Business Unit from April 2016 to September 2020. From 2000 – 2010, Ms. Robertson was in private practice with Hogan & Hartson LLP. Ms. Robertson received a B.A. in Political Science from Southern Illinois University and a J.D. from the Georgetown University Law Center.

**Micah Zajic** has served as our Chief Financial Officer since January 2023. Before joining Lexeo, Mr. Zajic served at SQZ Biotechnologies, Inc., where he led the business development, finance, investor relations and operations functions, as Chief Financial Officer from July 2022 to January 2023 and as Chief Business Officer from October 2020 to July 2022. Prior to that, Mr. Zajic served as Vice President of Corporate Development of MeiraGTx, a clinical-stage gene therapy company from April 2017 to September 2020, and served as Senior Director, Corporate Strategy of Alexion Pharmaceuticals, Inc., a global biopharmaceutical company, from September 2014 to April 2017. Mr. Zajic received a B.A. in Economics from Georgetown University.

### **Non-employee directors**

**Steven Altschuler, M.D.**, has served as the Chairman of our board of directors since January 2021. Dr. Altschuler has served as a Managing Director, Healthcare Ventures, of Ziff Capital Partners, a private investment firm, since May 2018. He previously served as a consultant to the University of Miami Health Care System from September 2017 through December 2017, the Chief Executive Officer of the University of Miami Health Care System and Executive Vice President for Healthcare at the University of Miami from January 2016 to September 2017, and the Chief Executive Officer of The Children's Hospital of Philadelphia, or CHOP, from April 2000 until June 2015. Prior to assuming the role of Chief Executive Officer, Dr. Altschuler held several positions at CHOP and the Perelman School of Medicine at the University of Pennsylvania, including Physician-in-Chief/Chair of Pediatrics and chief of the Division of Gastroenterology, Hepatology and Nutrition. Dr. Altschuler has served as a director of WW International, Inc. since September 2012, as a director of Orchard Therapeutics plc since January 2020 and as a director of 89bio, Inc. since March 2020. He previously served as Chair of the board of directors of Spark Therapeutics, Inc. from March 2013 to December 2019 and as a director of Adtalem Global Education Inc. from May 2018 to May 2020. Dr. Altschuler received a B.A. in Mathematics and an M.D. from Case Western Reserve University. Our board of directors believes Dr. Altschuler is qualified to serve on our board of directors based on his experience holding senior leadership positions within biotechnology companies and his role on public and private boards of directors, as well as his experience investing in healthcare companies.

**Mette Kirstine Agger** has served as a member of our board of directors since November 2020. Ms. Agger has served as Chief Executive Officer and Strategic Advisor of Ersum Biotech since March 2022. From 2009 to 2022, Ms. Agger served as a Senior Advisor and Managing Partner of Lundbeckfonden Biocapital (formerly Lundbeckfonden Ventures), a life sciences venture capital fund and, with its affiliates, a holder of more than 5% of our voting securities. Prior to joining Lundbeckfonden Biocapital, Ms. Agger co-founded 7TM A/S, a biotech

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company engaged in therapeutic drug discovery and development, in 2000, and served as its Chief Executive Officer from founding to 2009. Ms. Agger served on the board of Trevi Therapeutics, Inc., a public life sciences company, from July 2017 to June 2019 and Veloxis Pharmaceuticals A/S, a public pharmaceutical company from April 2010 to December 2019, and has served on the board of directors of Imara Inc., a public biopharmaceutical company since January 2016 and scPharmaceuticals Inc., a public pharmaceutical company, since March 2014. Ms. Agger received her M.Sc. in Biology from the University of Copenhagen and received her M.B.A. from Henley Business School at the University of Reading. Our board of directors believe Ms. Agger is qualified to serve on our board of directors based on her experience holding senior leadership positions within biotechnology companies and her role on public and private boards of directors, as well as her experience investing in healthcare companies.

**Paula HJ Cholmondeley** has served as a member of our board of directors since November 2021. Ms. Cholmondeley has been the Chief Executive Officer of the Sorrel Group, a management consulting firm, since January 2004 and has been a part time faculty member of the National Association of Corporate Directors, or NACD, a corporate governance association, since June 2008. Ms. Cholmondeley is a NACD Certified Director and has been elected to the NACD Directorship 100. She is a former Chief Financial Officer and Certified Public Accountant. She has also held profit and loss roles as a Divisional President. Ms. Cholmondeley has served on the boards of directors of the Bank OZK, a regional bank headquartered in Little Rock, Arkansas, since May 2016, of Terex Corporation, a manufacturing company since 2004, and of Nationwide Mutual Funds, an investment firm, from 2002 to 2022. Ms. Cholmondeley holds an M.S. and a B.S. in Accounting from the Wharton School of the University of Pennsylvania. Our board of directors believes Ms. Cholmondeley is qualified to serve on our board of directors because of her financial leadership experience and her leadership on public boards of directors.

**Bernard Davitian** has served as a member of our board of directors since November 2020. Mr. Davitian has been a Partner at Omega Funds, a life sciences-focused investment fund, since January 2020. Prior to that, Mr. Davitian served as Senior Vice President and Managing Director at Sanofi Ventures, the venture capital arm of Sanofi S.A., or Sanofi, for seven years from June 2012 to October 2019. Previously, he served as the Deputy of the Global Head of Business Development at Sanofi for two years. Prior to Sanofi, Mr. Davitian was Chief Financial Officer at Fovea Pharmaceuticals SA, a French biopharmaceutical R&D company specializing in ophthalmology (which was acquired by Sanofi in 2009), Chief Executive Officer at Neurotech Pharmaceuticals, Inc., a biotechnology company developing sight saving therapies for retinal diseases, and Chief Financial Officer at Transgene SA, a biopharmaceutical company. Previously, he served in various capacities at Institut Mérieux, including that of Corporate Chief Financial Officer. Prior to that, he was a senior auditor at Arthur Andersen LLP. Mr. Davitian has extensive experience in the life sciences and biotech industry, marked by a number of successful transactions involving financings, public offerings and acquisitions. A Certified Public Accountant in France, Mr. Davitian holds an M.Sc. in Management (M.B.A. equivalent) from the EM Lyon Business School (France) and an A.M.P. from the Wharton School of Business at the University of Pennsylvania. Our board of directors believes Mr. Davitian is qualified to serve as a director because of his experience and leadership in healthcare venture capital investing.

**Reinaldo Diaz** has served as a member of our board of directors since February 2022. Mr. Diaz has served as a Venture Partner at Longitude Capital, a healthcare venture capital firm since 2015, and he currently serves as the Chief Business Officer of Opna-Immuno Oncology, SA, a Longitude Capital portfolio company. Mr. Diaz has also served as a Managing Director of DA Advisors, LLC since 2005, providing strategic and financial advice primarily to life science companies. From 2008 to 2018, Mr. Diaz served as a managing director at Auen Therapeutics, a private equity firm focusing on life science companies. From 1996 to 2005, Mr. Diaz served as a managing member and co-founder of Diaz & Altschul Capital Management, LLC, an asset management firm focusing on healthcare companies. Prior to that, Mr. Diaz served as a managing director and head of the

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healthcare group at Schroder Wertheim & Co., Inc., and in various roles at PaineWebber Development Corporation, including as president. Mr. Diaz currently serves on the board of directors of the public pharmaceutical company, Inozyme Pharma, Inc., where he chairs the compensation committee. Mr. Diaz received a B.A. in Economics from Harvard University and an M.B.A. from Harvard Business School. Our board of directors believes Mr. Diaz is qualified to serve as a director because of his experience in the life sciences industry.

### **Board composition**

Our business and affairs are managed under the direction of our board of directors, which currently consists of six members. Our directors were elected to, and currently serve on, the board pursuant to a voting agreement among us and all of our stockholders and voting rights granted by our current amended and restated certificate of incorporation. The voting agreement will terminate upon the closing of this offering, after which there will be no further contractual obligations regarding the election of our directors.

In accordance with our amended and restated certificate of incorporation that will be in effect upon the closing of this offering, our board of directors will be divided into three classes, each of which will consist, as nearly as possible, of one-third of the total number of directors constituting our entire board and which will serve staggered three-year terms. At each annual meeting of stockholders, the successors to directors whose terms then expire will be elected to serve from the time of election and qualification until the third annual meeting following election. Our directors will be divided among the three classes as follows:

- Class I, which will consist of \_\_\_\_\_ and \_\_\_\_\_, and their terms will expire at our first annual meeting of stockholders to be held after the closing of this offering;
- Class II, which will consist of \_\_\_\_\_ and \_\_\_\_\_, and their terms will expire at our second annual meeting of stockholders to be held after the closing of this offering; and
- Class III, which will consist of \_\_\_\_\_ and \_\_\_\_\_, and their terms will expire at our third annual meeting of stockholders to be held after the closing of this offering.

Our amended and restated bylaws, which will become effective upon the closing of this offering, will provide that the authorized number of directors may be changed only by resolution approved by a majority of our board of directors. Any additional directorships resulting from an increase in the number of directors will be distributed among the three classes so that, as nearly as possible, each class will consist of one-third of the directors. The division of our board of directors into three classes with staggered three-year terms may delay or prevent a change of our management or a change of control.

### **Director independence**

Applicable Nasdaq rules, or the Nasdaq Listing Rules, require a majority of a listed company's board of directors to be comprised of independent directors within one year of listing. In addition, the Nasdaq Listing Rules require that, subject to specified exceptions, each member of a listed company's audit, compensation and nominating and corporate governance committees be independent and that audit committee members also satisfy independence criteria set forth in Rule 10A-3 under the Exchange Act of 1934, as amended, or the Exchange Act. The Nasdaq independence definition includes a series of objective tests, such as that the director is not, and has not been for at least three years, one of our employees, that neither the director nor any of his family members has engaged in various types of business dealings with us and that the director is not associated with the holders of more than 5% of our common stock. In addition, under applicable Nasdaq rules, a director will only qualify as an "independent director" if, in the opinion of the listed company's board of directors, that person does not have a relationship that would interfere with the exercise of independent judgment in carrying out the responsibilities of a director.

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Our board of directors has determined that all of our directors other than \_\_\_\_\_, representing \_\_\_\_\_ of our \_\_\_\_\_ directors, are “independent directors” as defined under applicable Nasdaq rules. In making such determination, our board of directors considered the current and prior relationships that each such director has with our company and all other facts and circumstances that our board of directors deemed relevant in determining his or her independence, including the beneficial ownership of our capital stock by each director and the transactions described in the section titled “Certain Relationships and Related Party Transactions.”

There are no family relationships among any of our directors or executive officers.

### **Role of the board in risk oversight**

One of the key functions of our board of directors is informed oversight of our risk management process. Our board of directors does not have a standing risk management committee, but rather administers this oversight function directly through the board of directors as a whole, as well as through various standing committees of our board of directors that address risks inherent in their respective areas of oversight. In particular, our board of directors is responsible for monitoring and assessing strategic risk exposure. Following the completion of this offering, we intend for our audit committee to have the responsibility to consider and discuss our major financial risk exposures and the steps our management has taken to monitor and control these exposures, including guidelines and policies to govern the process by which risk assessment and management is undertaken. The audit committee will also monitor compliance with legal and regulatory requirements.

### **Board committees**

Our board of directors has established an audit committee, compensation committee and a nominating and corporate governance committee, each of which operate pursuant to a committee charter. Our board of directors may establish other committees to facilitate the management of our business. The composition and functions of each committee are described below.

#### ***Audit committee***

Upon the completion of this offering, our audit committee will consist of \_\_\_\_\_, \_\_\_\_\_ and \_\_\_\_\_, with \_\_\_\_\_ serving as chair of the audit committee. Our board of directors has determined that each of these individuals meets the independence requirements of Rule 10A-3 under the Securities Exchange Act of 1934, or the Exchange Act, and the applicable listing standards of Nasdaq. Each member of our audit committee can read and understand fundamental financial statements in accordance with Nasdaq audit committee requirements. Our board of directors has also determined that \_\_\_\_\_ qualifies as an audit committee financial expert within the meaning of SEC regulations and meets the financial sophistication requirements of the Nasdaq Listing Rules. In arriving at these determinations, the board has examined each audit committee member’s scope of experience and the nature of their prior and/or current employment.

The functions of this committee include, among other things:

- helping our board of directors oversee our corporate accounting and financial reporting processes;
- managing the selection, engagement, qualifications, independence and performance of a qualified firm to serve as the independent registered public accounting firm to audit our financial statements;
- discussing the scope and results of the audit with the independent registered public accounting firm, and reviewing, with management and the independent accountants, our interim and year-end operating results;

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- developing procedures for employees to submit concerns anonymously about questionable accounting or audit matters;
- reviewing related person transactions;
- obtaining and reviewing a report by the independent registered public accounting firm at least annually that describes our internal quality control procedures, any material issues with such procedures and any steps taken to deal with such issues when required by applicable law;
- approving or, as permitted, pre-approving, audit and permissible non-audit services to be performed by the independent registered public accounting firm;
- reviewing and evaluating on an annual basis the performance of the audit committee and the audit committee charter; and
- any other responsibility delegated to it by the board of directors from time to time.

We believe that the composition and functioning of our audit committee will comply with all applicable SEC and Nasdaq rules and regulations. We intend to comply with future requirements to the extent they become applicable to us.

### ***Compensation committee***

Upon the completion of this offering, our compensation committee will consist of \_\_\_\_\_, \_\_\_\_\_ and \_\_\_\_\_, with \_\_\_\_\_ serving as chair of the compensation committee. Each of these individuals is a non-employee director, as defined in Rule 16b-3 promulgated under the Exchange Act and is an “outside director,” as defined pursuant to Section 162(m) of the Internal Revenue Code of 1986, as amended, or the Code. Our board of directors has determined that each of these individuals is “independent” as defined under the applicable listing standards of Nasdaq, including the standards specific to members of a compensation committee. The functions of this committee include, among other things:

- reviewing, modifying and approving (or if it deems appropriate, making recommendations to the full board of directors regarding) our overall compensation strategy and policies;
- making recommendations to the full board of directors regarding the compensation and other terms of employment of our executive officers;
- reviewing and making recommendations to the full board of directors regarding performance goals and objectives relevant to the compensation of our executive officers and assessing their performance against these goals and objectives;
- reviewing and approving (or if it deems it appropriate, making recommendations to the full board of directors regarding) the equity incentive plans, compensation plans and similar programs advisable for us, as well as modifying, amending or terminating existing plans and programs;
- evaluating risks associated with our compensation policies and practices and assessing whether risks arising from our compensation policies and practices for our employees are reasonably likely to have a material adverse effect on us;
- reviewing and making recommendations to the full board of directors regarding the type and amount of compensation to be paid or awarded to our non-employee board members;
- establishing policies with respect to votes by our stockholders to approve executive compensation to the extent required by Section 14A of the Exchange Act and, if applicable, determining our recommendations regarding the frequency of advisory votes on executive compensation;

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- reviewing and assessing the independence of compensation consultants, legal counsel and other advisors as required by Section 10C of the Exchange Act;
- administering our equity incentive plans;
- establishing policies with respect to equity compensation arrangements;
- reviewing the competitiveness of our executive compensation programs and evaluating the effectiveness of our compensation policy and strategy in achieving expected benefits to us;
- reviewing and making recommendations to the full board of directors regarding the terms of any employment agreements, severance arrangements, change in control protections and any other compensatory arrangements for our executive officers;
- reviewing with management and approving our disclosures under the caption “Compensation Discussion and Analysis” in our periodic reports or proxy statements to be filed with the SEC, to the extent such caption is included in any such report or proxy statement;
- preparing the report that the SEC requires in our annual proxy statement;
- reviewing and evaluating on an annual basis the performance of the compensation committee and the compensation committee charter; and
- any other responsibility delegated to it by the board of directors from time to time.

We believe that the composition and functioning of our compensation committee will comply with all applicable SEC and Nasdaq rules and regulations. We intend to comply with future requirements to the extent they become applicable to us.

### ***Nominating and corporate governance committee***

Upon the completion of this offering, our nominating and corporate governance committee will consist of \_\_\_\_\_, \_\_\_\_\_ and \_\_\_\_\_, with \_\_\_\_\_ serving as chair of the nominating and corporate governance committee. Our board of directors has determined that each of these individuals is “independent” as defined under the applicable listing standards of Nasdaq and SEC rules and regulations. The functions of this committee include, among other things:

- identifying, reviewing and evaluating candidates to serve on our board of directors;
- determining the minimum qualifications for service on our board of directors;
- evaluating director performance on the board and applicable committees of the board and determining whether continued service on our board is appropriate;
- evaluating, nominating and recommending individuals for membership on our board of directors;
- evaluating nominations by stockholders of candidates for election to our board of directors;
- considering and assessing the independence of members of our board of directors;
- developing a set of corporate governance policies and principles and recommending to our board of directors any changes to such policies and principles;
- reviewing and making recommendations to the board of directors with respect to management succession planning;
- considering questions of possible conflicts of interest of directors as such questions arise;



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- reviewing and evaluating on an annual basis the performance of the nominating and corporate governance committee and the nominating and corporate governance committee charter; and
- any other responsibility delegated to it by the board of directors from time to time.

We believe that the composition and functioning of our nominating and corporate governance committee will comply with all applicable SEC and Nasdaq rules and regulations. We intend to comply with future requirements to the extent they become applicable to us.

### Compensation committee interlocks and insider participation

None of our directors who serve as a member of our compensation committee is, or has at any time during the past year been, one of our officers or employees. None of our executive officers currently serves, or in the past year has served, as a member of the board of directors or compensation committee of any other entity that has one or more executive officers serving on our board of directors or compensation committee.

### Code of business conduct and ethics

Effective upon the closing of this offering, we will adopt a Code of Business Conduct and Ethics, or the Code of Conduct, applicable to all of our employees, executive officers and directors. This includes our principal executive officer, principal financial officer and principal accounting officer or controller, or persons performing similar functions. Following the closing of this offering, the full text of the Code of Conduct will be available on our website at [www.lexeotx.com](http://www.lexeotx.com). We intend to post on our website all disclosures that are required by law or the listing standards of the Nasdaq Stock Market concerning any amendments to, or waivers from, any provision of the Code of Conduct. Information contained on, or that can be accessed through, our website is not incorporated by reference into this prospectus, and you should not consider information on our website to be part of this prospectus.

### Non-employee director compensation

The following table shows information concerning the compensation that our non-employee directors earned during the last completed fiscal year ended December 31, 2022. A director who is also our employee receives no additional compensation for his or her services as a director.

Name	Fees Earned or Paid in Cash (\$) <sup>(1)</sup>	Option Awards (\$) <sup>(2)(3)</sup>	Total (\$)
Sandip Agarwala <sup>(4)</sup>	\$ 20,250	—	—
Mette Kirstine Agger	—	—	—
Steven Altschuler, M.D.	\$ 150,000	\$ 162,524	\$ 312,524
Paula HJ Cholmondeley	\$ 37,000	\$ 49,260	\$ 86,260
Bernard Davitian	—	—	—
Reinaldo Diaz <sup>(5)</sup>	—	—	—

(1) Represents actual fees paid in fiscal year 2022. For Dr. Altschuler and Ms. Cholmondeley, this includes fees paid for board services. Mr. Agarwala, Ms. Agger, Mr. Davitian and Mr. Diaz did not receive cash compensation for their service to us as non-employee directors in fiscal year 2022.

(2) Option awards in this column are reported at the aggregate grant date fair value in accordance with FASB ASC Topic 718. Option awards under this column are discussed in greater detail below.

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(3) As of December 31, 2022, our then non-employee directors held the following option awards:

Name	Grant Date	Option Awards		Exercise Price (\$)	Expiration Date
		Unexercisable	Exercisable		
Sandip Agarwala	—	—	—	—	—
Mette Kirstine Agger	—	—	—	—	—
Steven Altschuler, M.D.	2/16/2021	—	1,189,097	0.22	02/15/2031
Paula HJ Cholmondeley	—	—	—	—	—
Bernard Davitian	—	—	—	—	—
Reinaldo Diaz	—	—	—	—	—

(4) Sandip Agarwala resigned from our board of directors effective January 28, 2022.

(5) Reinaldo Diaz was appointed to our board of directors effective February 2, 2022.

### **Stock option awards**

On February 16, 2021, Steven Altschuler, M.D. received a grant of 1,189,097 stock options, with a vesting start date of January 5, 2021, and on November 15, 2021, Paula Cholmondeley received a grant of 165,565 stock options. The stock options granted to Dr. Altschuler and Ms. Cholmondeley were exercisable on the date of grant and vest with respect to 25% of the underlying shares on the first anniversary of the vesting start date and thereafter in equal monthly installments over three years.

## Executive compensation

Our named executive officers for the period ended December 31, 2022, consisting of our principal executive officer and the next two most highly compensated executive officers, were:

- R. Nolan Townsend, who currently serves as our Chief Executive Officer and as a member of our board of directors;
- Jenny R. Robertson, J.D., who currently serves as our Senior Vice President, General Counsel and Head of People; and
- Jay A. Barth, M.D., our former Executive Vice President and Chief Medical Officer, who resigned on January 13, 2023.

### Summary compensation table

The following table sets forth information regarding compensation awarded to, earned by and paid to our named executive officers with respect to the period ended December 31, 2022.

<b>Name and Principal Position</b>	<b>Year</b>	<b>Salary (\$)</b>	<b>Bonus (\$)<sup>(1)</sup></b>	<b>Option Awards (\$)<sup>(2)</sup></b>	<b>Non-Equity Incentive Plan Compensation (\$)<sup>(3)</sup></b>	<b>All Other Compensation (\$)</b>	<b>Total (\$)</b>
R. Nolan Townsend	2022	\$465,000	—	—	\$ 267,375	\$ 36,545	\$ 768,920
<i>Chief Executive Officer and Director</i>	2021	\$410,000	—	\$1,382,160	\$ 267,375	\$ 1,500	\$2,061,869
Jenny R. Robertson, J.D. <sup>(4)</sup>	2022	\$257,813	\$99,000 <sup>(3)</sup>	\$ 690,997	\$ 14,850	\$ 2,588	\$1,065,247
<i>Senior Vice President, General Counsel and Head of People</i>	2021	—	—	—	—	—	—
Jay A. Barth, M.D. <sup>(5)</sup>	2022	\$420,000	—	—	\$ 151,200	\$ 21,370	\$ 592,570
<i>Executive Vice President and Chief Medical Officer</i>	2021	\$399,167	—	\$ 816,045	\$ 151,200	\$ —	\$1,366,411

(1) This column reflects an amount awarded as a signing bonus to Ms. Robertson in connection with her joining our company on March 21, 2022.

(2) Option awards in this column are reported at the aggregate grant date fair value in accordance with FASB ASC Topic 718 for awards granted to the named executive officer under the Existing Plan. Assumptions used in the calculation of these amounts are included in Note to our audited financial statements included elsewhere in this prospectus.

(3) Non-equity incentive plan compensation amounts represent actual amounts earned during 2022, but paid in 2023. The amount paid to each named executive officer is equal to a percentage of the executive's target annual bonus pursuant to the applicable employment agreement.

(4) Ms. Robertson joined our company on March 21, 2022, and thus she was not a named executive officer for fiscal year 2021.

(5) Dr. Barth resigned as our Executive Vice President and Chief Medical Officer on January 13, 2023.

## Outstanding equity awards at December 31, 2022

The following table sets forth certain information about outstanding equity awards granted to our named executive officers that were outstanding as of December 31, 2022.

Name	Grant Date	Vesting Commencement Date	Number of Securities Underlying Unexercised Options Exercisable	Option Exercise Price (\$) <sup>(1)</sup>	Option Expiration Date
R. Nolan Townsend	2/16/21	11/21/2020	1,627,848 <sup>(2)</sup>	\$ 0.22	2/15/31
<i>Chief Executive Officer</i>	11/15/21	8/11/2021	5,500,000 <sup>(3)</sup>	\$ 0.46	11/14/2031
Jenny R. Robertson, J.D.	5/6/22	3/21/2022	729,680 <sup>(4)</sup>	\$ 1.43	3/21/31
<i>Senior Vice President, General Counsel and Head of People</i>	—	—	—	—	—
Jay A. Barth, M.D.	2/16/21	11/21/2020	1,783,645 <sup>(5)(7)</sup>	\$ 0.22	4/13/2023
<i>Executive Vice President and Chief Medical Officer</i>	11/15/21	8/11/2021	2,920,855 <sup>(6)(7)</sup>	\$ 0.46	4/13/2023

- (1) All of the option awards were granted under the Existing Plan. All of the option awards were granted with a per share exercise price equal to the fair market value of one share of our common stock on the date of grant.
- (2) All of the 2,170,463 shares originally subject to this option were exercisable as of the date of grant, and 542,615 shares were exercised on May 14, 2021. Of the total shares subject to this option, 25% will vest one year after the vesting commencement date, and the remaining shares vest in 36 equal monthly installments thereafter, subject to continuous service through each applicable vesting date. All unvested shares will vest immediately prior to the consummation of a change in control of the Company. As of December 31, 2022, 1,113,449 shares subject to this option have vested, of which 542,615 shares have been exercised and 587,834 vested shares remain unexercised.
- (3) All of the shares subject to these options were exercisable as of the date of grant. 25% of the total shares subject to these options will vest one year after the vesting commencement date, and the remaining shares vest in 36 equal monthly installments thereafter, subject to continuous service through each applicable vesting date. All unvested shares will vest immediately prior to the consummation of a change in control of the Company. As of December 31, 2022, 1,833,333 shares subject to these options have vested, and all vested shares remain unexercised.
- (4) All of the shares subject to this option were exercisable as of the date of grant. Of the total shares subject to this option, 25% will vest one year after the vesting commencement date, and the remaining shares vest in 36 equal monthly installments thereafter, subject to continuous service through each applicable vesting date. All unvested shares will vest immediately prior to the consummation of a change in control of the Company. As of December 31, 2022, zero shares subject to these options have vested.
- (5) All of the shares subject to these options were exercisable as of the date of grant. 25% of the total shares subject to these options will vest one year after the vesting commencement date, and the remaining shares vest in 36 equal monthly installments thereafter, subject to continuous service through each applicable vesting date. All unvested shares will vest immediately prior to the consummation of a change in control of the Company. As of December 31, 2022, 213,215 shares subject to these options have vested, and all vested shares remain unexercised.
- (6) All of the shares subject to these options were exercisable as of the date of grant. 25% of the total shares subject to these options will vest one year after the vesting commencement date, and the remaining shares vest in 36 equal monthly installments thereafter, subject to continuous service through each applicable vesting date. All unvested shares will vest immediately prior to the consummation of a change in control of the Company. As of December 31, 2022, 295,614 shares subject to these options have vested, and all vested shares remain unexercised.
- (7) Mr. Barth resigned as our Executive Vice President and Chief Medical Officer on January 13, 2023. The post termination exercise period for all exercisable options expired on April 13, 2023. As such, all shares underlying these options have returned to our equity incentive plan share reserve.

## Agreements with our named executive officers

In connection with this offering, we intend to enter into amended and restated employment agreements with each of our named executive officers.

## Potential payments upon termination or change in control

As noted above, in connection with this offering, we intend to enter into amended and restated employment agreements with each of our named executive officers. It is expected that those employment agreements will contain provisions with respect to severance upon certain terminations of employment.

## Other compensation and benefits

We maintain broad-based employee benefit plans and programs for the benefit of our employees, in which our named executive officers are entitled to participate. All of our current named executive officers are eligible to participate in our employee benefit plans, including our medical, dental, vision, and accidental death and dismemberment insurance plans, in each case on the same basis as all of our other employees.

We currently maintain a 401(k) retirement savings plan for our employees, including our named executive officers, who satisfy certain eligibility requirements. The 401(k) plan is intended to qualify as a tax-qualified plan under the Internal Revenue Code. Our named executive officers are eligible to participate in the 401(k) plan on the same basis as our other employees. The Internal Revenue Code allows eligible employees to defer a portion of their compensation, within prescribed limits, on a pre-tax basis through contributions to the 401(k) plan. Currently, we provide matching contributions of 50% on the first 6% of each participating employee's deferral up to the contribution limit permitted under the Internal Revenue Code.

## Equity incentive plans

### *2023 equity incentive plan*

Our board of directors adopted the 2023 Equity Incentive Plan, or 2023 Plan, in 2023, and our stockholders approved the 2023 Plan in 2023. The 2023 Plan will become effective upon the execution of the underwriting agreement for this offering. Once the 2023 Plan becomes effective, no further grants will be made under the 2021 Equity Incentive Plan, or the Existing Plan.

*Types of Awards.* Our 2023 Plan provides for the grant of incentive stock options, or ISOs, nonstatutory stock options, or NSOs, stock appreciation rights, restricted stock awards, restricted stock unit awards, performance-based awards, and other awards, or collectively, awards. ISOs may be granted only to our employees, including our officers, and the employees of our affiliates. All other awards may be granted to our employees, including our officers, our non-employee directors, consultants, and the employees and consultants of our affiliates.

*Authorized Shares.* The maximum number of shares of common stock that may be issued under our 2023 Plan is \_\_\_\_\_ shares, which is the sum of (i) \_\_\_\_\_ new shares, plus (ii) the Existing Plan's available reserve, plus (iii) the number of returning shares, if any, as such shares become available from time to time. In addition, the number of shares of common stock reserved for issuance under our 2023 Plan will automatically increase on January 1 of each year, beginning on January 1, 2024, and continuing through and including January 1, 2033, by \_\_\_\_\_ % of the total number of shares of common stock outstanding on December 31 of the immediately preceding calendar year, or a lesser number of shares determined by our board of directors prior to the applicable January 1. The maximum number of shares that may be issued upon the exercise of ISOs under our 2023 Plan is \_\_\_\_\_ shares.

Shares issued under our 2023 Plan will be authorized but unissued or reacquired shares of common stock. Shares subject to awards granted under our 2023 Plan that expire or terminate without being exercised in full, or that are paid out in cash rather than in shares, will not reduce the number of shares available for issuance under our 2023 Plan. Additionally, shares issued pursuant to awards under our 2023 Plan that we repurchase or that are forfeited, as well as shares used to pay the exercise price of an award or to satisfy the tax withholding obligations to an award, will become available for future grant under our 2023 Plan.

The maximum number of shares of common stock subject to stock awards granted under the 2023 Plan or otherwise during any calendar year beginning in 2024 to any non-employee director, taken together with any cash fees paid by us to such non-employee director during such calendar year for service on the board of

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directors, will not exceed \$ \_\_\_\_\_ in total value (calculating the value of any such stock awards based on the grant date fair value of such stock awards for financial reporting purposes), or, with respect to the calendar year in which a non-employee director is first appointed or elected to our board of directors, \$ \_\_\_\_\_.

*Plan administration.* Our board of directors, or a duly authorized committee of our board, may administer our 2023 Plan. We expect that our board of directors will delegate concurrent authority to administer our 2023 Plan to the compensation committee under the terms of the compensation committee's charter. We sometimes refer to the board of directors, or the applicable committee with the power to administer our equity incentive plans, as the administrator. The administrator may also delegate to one or more of our officers the authority to (1) designate employees (other than officers) to receive specified awards, and (2) determine the number of shares subject to such awards.

The administrator has the authority to determine the terms of awards, including recipients, the exercise, purchase or strike price of awards, if any, the number of shares subject to each award, the fair market value of a share of common stock, the vesting schedule applicable to the awards, together with any vesting acceleration, and the form of consideration, if any, payable upon exercise or settlement of the award and the terms of the award agreements for use under our 2023 Plan.

In addition, subject to the terms of the 2023 Plan, the administrator also has the power to modify outstanding awards under our 2023 Plan, including the authority to reprice any outstanding option or stock appreciation right, cancel and re-grant any outstanding option or stock appreciation right in exchange for new stock awards, cash or other consideration, or take any other action that is treated as a repricing under generally accepted accounting principles, with the consent of any materially adversely affected participant.

*Stock options.* ISOs and NSOs are granted pursuant to stock option agreements adopted by the administrator. The administrator determines the exercise price for a stock option, within the terms and conditions of the 2023 Plan, provided that the exercise price of a stock option generally cannot be less than 100% of the fair market value of our common stock on the date of grant. Options granted under the 2023 Plan vest at the rate specified in the stock option agreement as specified in the stock option agreement by the administrator.

The administrator determines the term of stock options granted under the 2023 Plan, up to a maximum of ten years. Unless the terms of an optionholder's stock option agreement provide otherwise, if an optionholder's service relationship with us, or any of our affiliates, ceases for any reason other than disability, death, or cause, the optionholder may generally exercise any vested options for a period of three months following the cessation of service. The option term may be extended in the event that either an exercise of the option or an immediate sale of shares acquired upon exercise of the option following such a termination of service is prohibited by applicable securities laws or our insider trading policy. If an optionholder's service relationship with us or any of our affiliates ceases due to disability or death, or an optionholder dies within a certain period following cessation of service, the optionholder or a beneficiary may generally exercise any vested options for a period of 12 months in the event of disability and 18 months in the event of death. In the event of a termination for cause, options generally terminate immediately upon the termination of the individual for cause. In no event may an option be exercised beyond the expiration of its term.

Acceptable consideration for the purchase of common stock issued upon the exercise of a stock option will be determined by the administrator and may include (1) cash, check, bank draft, or money order, (2) a broker-assisted cashless exercise, (3) the tender of shares of common stock previously owned by the optionholder, (4) a net exercise of the option if it is an NSO, and (5) other legal consideration approved by the administrator.

Options may not be transferred to third-party financial institutions for value. Unless the administrator provides otherwise, options generally are not transferable except by will, the laws of descent and distribution or pursuant

to a domestic relations order. An optionholder may designate a beneficiary, however, who may exercise the option following the optionholder's death.

*Tax limitations on ISOs.* The aggregate fair market value, determined at the time of grant, of common stock with respect to ISOs that are exercisable for the first time by an option holder during any calendar year under all of our stock plans may not exceed \$100,000. Options or portions thereof that exceed such limit will be treated as NSOs. No ISOs may be granted to any person who, at the time of the grant, owns or is deemed to own stock possessing more than 10% of our total combined voting power or that of any of our parent or subsidiary corporations, unless (1) the option exercise price is at least 110% of the fair market value of the stock subject to the option on the date of grant and (2) the term of the ISO does not exceed five years from the date of grant.

*Restricted stock awards.* Restricted stock awards are granted pursuant to restricted stock award agreements adopted by the administrator. Restricted stock awards may be granted in consideration for cash, check, bank draft or money order, services rendered to us, or our affiliates or any other form of legal consideration. Common stock acquired under a restricted stock award may, but need not, be subject to a share repurchase option in our favor in accordance with a vesting schedule to be determined by the administrator. A restricted stock award may be transferred only upon such terms and conditions as set by the administrator. Except as otherwise provided in the applicable award agreement, restricted stock awards that have not vested may be forfeited or repurchased by us upon the participant's cessation of continuous service for any reason.

*Restricted stock unit awards.* Restricted stock unit awards are granted pursuant to restricted stock unit award agreements adopted by the administrator. Restricted stock unit awards may be granted in consideration for any form of legal consideration. A restricted stock unit award may be settled by cash, delivery of stock, a combination of cash and stock as deemed appropriate by the administrator, or in any other form of consideration set forth in the restricted stock unit award agreement. Additionally, dividend equivalents may be credited in respect of shares covered by a restricted stock unit award. Except as otherwise provided in the applicable award agreement, restricted stock units that have not vested will be forfeited upon the participant's cessation of continuous service for any reason.

*Stock appreciation rights.* Stock appreciation rights are granted pursuant to stock appreciation right grant agreements adopted by the administrator. The administrator determines the strike price for a stock appreciation right, which generally cannot be less than 100% of the fair market value of common stock on the date of grant. Upon the exercise of a stock appreciation right, we will pay the participant an amount equal to the product of (1) the excess of the per share fair market value of common stock on the date of exercise over the strike price, multiplied by (2) the number of shares of common stock with respect to which the stock appreciation right is exercised. A stock appreciation right granted under the 2023 Plan vests at the rate specified in the stock appreciation right agreement as determined by the administrator.

The administrator determines the term of stock appreciation rights granted under the 2023 Plan, up to a maximum of ten years. Unless the terms of a participant's stock appreciation right agreement provide otherwise, if a participant's service relationship with us or any of our affiliates ceases for any reason other than cause, disability, or death, the participant may generally exercise any vested stock appreciation right for a period of three months following the cessation of service. The stock appreciation right term may be further extended in the event that exercise of the stock appreciation right following such a termination of service is prohibited by applicable securities laws. If a participant's service relationship with us, or any of our affiliates, ceases due to disability or death, or a participant dies within a certain period following cessation of service, the participant or a beneficiary may generally exercise any vested stock appreciation right for a period of 12 months in the event of disability and 18 months in the event of death. In the event of a termination for cause, stock appreciation rights generally terminate immediately upon the occurrence of the event giving rise to the termination of the individual for cause. In no event may a stock appreciation right be exercised beyond the expiration of its term.

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*Performance awards.* Our 2023 Plan permits the grant of performance-based stock and cash awards. The administrator can structure such awards so that the stock or cash will be issued or paid pursuant to such award only following the achievement of certain pre-established performance goals during a designated performance period. Performance awards that are settled in cash or other property are not required to be valued in whole or in part by reference to, or otherwise based on, the common stock.

The performance goals may be based on any measure of performance selected by the board of directors. The administrator may establish performance goals on a company-wide basis, with respect to one or more business units, divisions, affiliates, or business segments, and in either absolute terms or relative to the performance of one or more comparable companies or the performance of one or more relevant indices. Unless specified otherwise (i) in the award agreement at the time the award is granted or (ii) in such other document setting forth the performance goals at the time the goals are established, the administrator will appropriately make adjustments in the method of calculating the attainment of the performance goals as follows: (1) to exclude restructuring and/or other nonrecurring charges; (2) to exclude exchange rate effects; (3) to exclude the effects of changes to generally accepted accounting principles; (4) to exclude the effects of any statutory adjustments to corporate tax rates; (5) to exclude the effects of items that are “unusual” in nature or occur “infrequently” as determined under generally accepted accounting principles; (6) to exclude the dilutive effects of acquisitions or joint ventures; (7) to assume that any business divested by us achieved performance objectives at targeted levels during the balance of a performance period following such divestiture; (8) to exclude the effect of any change in the outstanding shares of common stock by reason of any stock dividend or split, stock repurchase, reorganization, recapitalization, merger, consolidation, spin-off, combination or exchange of shares or other similar corporate change, or any distributions to common stockholders other than regular cash dividends; (9) to exclude the effects of stock-based compensation and the award of bonuses under our bonus plans; (10) to exclude costs incurred in connection with potential acquisitions or divestitures that are required to be expensed under generally accepted accounting principles; and (11) to exclude the goodwill and intangible asset impairment charges that are required to be recorded under generally accepted accounting principles.

*Other awards.* The administrator may grant other awards based in whole or in part by reference to common stock. The administrator will set the number of shares under the award and all other terms and conditions of such awards.

*Changes to capital structure.* In the event there is a specified type of change in our capital structure, such as a stock split, reverse stock split, or recapitalization, appropriate adjustments will be made to (1) the class and maximum number of shares reserved for issuance under the 2023 Plan; (2) the class and maximum number of shares by which the share reserve may increase automatically each year; (3) the class and maximum number of shares that may be issued upon the exercise of ISOs; and (4) the class and number of shares and exercise price, strike price, or purchase price, if applicable, of all outstanding awards.

*Corporate transactions.* The following applies to stock awards under the 2023 Plan in the event of a corporate transaction, unless otherwise provided in a participant’s stock award agreement or other written agreement with us or one of our affiliates or unless otherwise expressly provided by the administrator at the time of grant. Under the 2023 Plan, a corporate transaction is generally the consummation of (1) a sale or other disposition of all or substantially all of our assets, (2) a sale or other disposition of at least 50% of our outstanding securities, (3) a merger, consolidation or similar transaction following which we are not the surviving corporation, or (4) a merger, consolidation, or similar transaction following which we are the surviving corporation but the shares of common stock outstanding immediately prior to such transaction are converted or exchanged into other property by virtue of the transaction.

In the event of a corporate transaction, any stock awards outstanding under the 2023 Plan may be assumed, continued or substituted for by any surviving or acquiring corporation (or its parent company), and any



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reacquisition or repurchase rights held by us with respect to the stock award may be assigned to the successor (or its parent company). If the surviving or acquiring corporation (or its parent company) does not assume, continue, or substitute for such stock awards, then (i) with respect to any such stock awards that are held by participants whose continuous service has not terminated prior to the effective time of the corporate transaction, or current participants, the vesting (and exercisability, if applicable) of such stock awards will be accelerated in full to a date prior to the effective time of the corporate transaction (contingent upon the effectiveness of the corporate transaction), and such stock awards will terminate if not exercised (if applicable) at or prior to the effective time of the corporate transaction, and any reacquisition or repurchase rights held by us with respect to such stock awards will lapse (contingent upon the effectiveness of the corporate transaction), and (ii) any such stock awards that are held by persons other than current participants will terminate if not exercised (if applicable) prior to the effective time of the corporate transaction, except that any reacquisition or repurchase rights held by us with respect to such stock awards will not terminate and may continue to be exercised notwithstanding the corporate transaction. In addition, the plan administrator may also provide, in its sole discretion, that the holder of a stock award that will terminate upon the occurrence of a corporate transaction if not previously exercised will receive a payment, if any, equal to the excess of the value of the property the participant would have received upon exercise of the stock award over the exercise price otherwise payable in connection with the stock award.

A stock award may be subject to additional acceleration of vesting and exercisability upon or after a change in control as may be provided in an applicable award agreement or other written agreement, but in the absence of such provision, no such acceleration will occur.

*Transferability.* A participant may not transfer awards under our 2023 Plan other than by will, the laws of descent and distribution, or as otherwise provided under our 2023 Plan.

*Plan amendment or termination.* Our board has the authority to amend, suspend, or terminate our 2023 Plan, provided that such action does not materially impair the existing rights of any participant without such participant's written consent. Certain material amendments also require the approval of our stockholders. No ISOs may be granted after the tenth anniversary of the date our board adopted our 2023 Plan. No awards may be granted under our 2023 Plan while it is suspended or after it is terminated.

### **2023 employee stock purchase plan**

Our board of directors adopted the 2023 ESPP on \_\_\_\_\_, 2023, and our stockholders adopted the 2023 ESPP on \_\_\_\_\_, 2023. The 2023 ESPP will become effective upon the execution of the underwriting agreement for this offering. The purpose of the 2023 ESPP is to secure the services of new employees, to retain the services of existing employees and to provide incentives for such individuals to exert maximum efforts toward our success and that of our affiliates. The 2023 ESPP includes two components. One component is designed to allow our eligible U.S. employees to purchase common stock in a manner that may qualify for favorable tax treatment under Section 423 of the Code. In addition, purchase rights may be granted under a component that does not qualify for such favorable tax treatment when necessary or appropriate to permit participation by our eligible employees who are foreign nationals or employed outside of the United States while complying with applicable foreign laws.

*Authorized shares.* The maximum aggregate number of shares of common stock that may be issued under our 2023 ESPP is \_\_\_\_\_ shares. The number of shares of common stock reserved for issuance under our 2023 ESPP will automatically increase on January 1 of each calendar year, beginning on January 1, 2024 and continuing through and including January 1, 2033, by the lesser of (1) \_\_\_\_\_ % of the total number of shares of capital stock outstanding on December 31 of the preceding calendar year, (2) \_\_\_\_\_ shares and (3) a number of shares determined by our board. Shares subject to purchase rights granted under our 2023 ESPP that terminate without having been exercised in full will not reduce the number of shares available for issuance under our 2023 ESPP.

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*Plan administration.* Our board, or a duly authorized committee thereof, will administer our 2023 ESPP. We anticipate that our board will delegate concurrent authority to administer our 2023 ESPP to the compensation committee under the terms of the compensation committee's charter. The 2023 ESPP is implemented through a series of offerings with specific terms approved by the administrator and under which eligible employees are granted purchase rights to purchase shares of common stock on specified dates during such offerings. Under the 2023 ESPP, we may specify offerings with durations of not more than 27 months and may specify shorter purchase periods within each offering. Each offering will have one or more purchase dates on which shares of common stock will be purchased for our eligible employees participating in the offering. An offering under the 2023 ESPP may be terminated under certain circumstances.

*Payroll deductions.* Generally, all regular employees, including executive officers, employed by us or by any of our designated affiliates, may participate in the 2023 ESPP and may contribute, normally through payroll deductions, with a maximum dollar amount as designated by the board. Unless otherwise determined by the administrator, common stock will be purchased for the accounts of employees participating in the 2023 ESPP at a price per share equal to the lower of (a) 85% of the fair market value of a share of common stock on the first date of an offering or (b) 85% of the fair market value of a share of common stock on the date of purchase. For the initial offering, which we expect will commence upon the execution and delivery of the underwriting agreement relating to this offering, the fair market value on the first day of the initial offering will be the price at which shares are first sold to the public.

*Limitations.* Our employees, including executive officers, or any of our designated affiliates may have to satisfy one or more of the following service requirements before participating in our 2023 ESPP, as determined by the administrator: (1) customary employment with us or one of our affiliates for more than 20 hours per week and more than five months per calendar year, or (2) continuous employment with us or one of our affiliates for a minimum period of time, not to exceed two years, prior to the first date of an offering. An employee may not be granted rights to purchase stock under our 2023 ESPP if such employee (1) immediately after the grant would own stock possessing 5% or more of the total combined voting power or value of common stock, or (2) holds rights to purchase stock under our 2023 ESPP that would accrue at a rate that exceeds \$25,000 worth of our stock for each calendar year that the rights remain outstanding.

*Changes to capital structure.* In the event that there occurs a change in our capital structure through such actions as a stock split, merger, consolidation, reorganization, recapitalization, reincorporation, stock dividend, dividend in property other than cash, large nonrecurring cash dividend, liquidating dividend, combination of shares, exchange of shares, change in corporate structure or similar transaction, the board of directors will make appropriate adjustments to (1) the number of shares reserved under the 2023 ESPP, (2) the maximum number of shares by which the share reserve may increase automatically each year, (3) the number of shares and purchase price of all outstanding purchase rights, and (4) the number of shares that are subject to purchase limits under ongoing offerings.

*Corporate transactions.* In the event of certain corporate transactions, including: (1) a sale of all or substantially all of our assets, (2) the sale or disposition of 50% of our outstanding securities, (3) the consummation of a merger or consolidation where we do not survive the transaction, and (4) the consummation of a merger or consolidation where we do survive the transaction but the shares of our common stock outstanding immediately before such transaction are converted or exchanged into other property by virtue of the transaction, any then-outstanding rights to purchase our stock under the 2023 ESPP may be assumed, continued or substituted for by any surviving or acquiring entity (or its parent company). If the surviving or acquiring entity (or its parent company) elects not to assume, continue, or substitute for such purchase rights, then the participants' accumulated payroll contributions will be used to purchase shares of common stock within 10 business days (or such other period specified by the board) prior to such corporate transaction and such purchase rights will terminate immediately.

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Under the 2023 ESPP, a corporate transaction is generally the consummation of: (1) a sale of all or substantially all of our assets, (2) the sale or disposition of more than 50% of our outstanding securities, (3) a merger or consolidation where we do not survive the transaction, and (4) a merger or consolidation where we do survive the transaction but the shares of our common stock outstanding immediately before such transaction are converted or exchanged into other property by virtue of the transaction.

*2023 ESPP amendment or termination.* The administrator has the authority to amend or terminate our 2023 ESPP, provided that except in certain circumstances such amendment or termination may not materially impair any outstanding purchase rights without the holder's consent. We will obtain stockholder approval of any amendment to our 2023 ESPP as required by applicable law or listing requirements.

### **2021 equity incentive plan**

Our board of directors adopted the Existing Plan in February 2021, and our stockholders approved the Existing Plan in September 2021.

*Stock awards.* The Existing Plan provides for the grant of ISOs, NSOs, and restricted stock awards, or collectively, stock awards. ISOs may be granted only to our employees and the employees of any parent corporation or subsidiary corporation. All other awards may be granted to our service providers. We have granted ISOs and NSOs under the Existing Plan. As of March 31, 2023, options to purchase \_\_\_\_\_ shares of our common stock were outstanding with a weighted-average exercise price of \$ \_\_\_\_\_ per share and restricted stock awards to purchase \_\_\_\_\_ shares of our common were outstanding, and \_\_\_\_\_ shares of our common stock remained available for future awards under the Existing Plan.

*Share reserve.* Subject to certain capitalization adjustments, the aggregate number of shares of our common stock that has been reserved for issuance pursuant to stock awards under the Existing Plan is \_\_\_\_\_ shares.

Shares subject to stock awards granted under our Existing Plan that expire or are forfeited or become unexercisable without having been exercised in full, or are surrendered pursuant to an exchange program shall continue to be available under the Existing Plan for issuance pursuant to future awards. If any shares of common stock issued pursuant to a stock award are forfeited back to or repurchased by us for at the original purchase price, the shares that are forfeited or repurchased or reacquired will revert to and again become available for issuance under the Existing Plan. Any shares retained in satisfaction of tax withholding obligations or as consideration for the exercise or purchase price of a stock award will again become available for issuance under the Existing Plan.

*Plan administration.* Our board of directors administers and interprets the provisions of the Existing Plan. The board of directors may delegate its authority to a committee, or committees, of the board, referred to as the "administrator." The administrator may additionally delegate limited authority to specified directors or executive officers. Under our Existing Plan, the administrator has the authority to, among other things, approve award recipients, determine the numbers and types of stock awards to be granted, determine the applicable fair market value and the provisions of each stock award, including the period of their exercisability and the vesting schedule applicable to a stock award, construe and interpret the Existing Plan and awards granted thereunder, and prescribe, amend, modify, and rescind or terminate rules and regulations for the administration of the Existing Plan.

*Stock options.* ISOs and NSOs are granted under stock option agreements in such form and containing such provisions as approved by the administrator. The administrator determines the exercise price for stock options, within the terms and conditions of the Existing Plan, provided that the exercise price of a stock option generally will not be less than 100% of the fair market value of our common stock on the date of grant (or 110% of the fair market value for 10% stockholders as required by the Code). Stock options granted under the Existing Plan vest at the rate specified in the stock option agreements and option rules as determined by the administrator.

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The administrator determines the term of stock options granted under the Existing Plan, up to a maximum of 10 years (or five years for 10% stockholders as required by the Code). If an optionholder's service relationship with us or any of our affiliates ceases for any reason other than disability, death, or cause the optionholder may generally exercise any vested options for a period of up to three months following the cessation of service, or such other period of time set forth in the option agreement. If an optionholder's service relationship with us or any of our affiliates ceases due to death or disability (or the participant dies within three months after a termination other than for cause), then options vested as of the termination date may generally be exercised within 12 months following the date of termination, or such other period of time set forth in the option agreement. In no event may an option be exercised beyond the expiration of its term. If an optionholder's service relationship with us or any of our affiliates ceases due to termination for cause, the optionholder's vested options shall expire on the optionholder's termination date, or such later time as determined by the administrator.

The exercise price for shares issued under the Existing Plan are generally payable in cash, check, promissory note, surrender of previously owned shares, consideration received through a broker-assisted (or other) cashless exercise program (whether through a broker or otherwise) implemented by us, net exercise, or other forms of consideration determined by the administrator (or any combination thereof).

Unless the administrator provides otherwise, options generally are not transferable except by will, the laws of descent and distribution, or, with respect to NSOs for participants in the United States, by gift to a family member.

*Restricted stock.* The administrator determines to whom an offer of restricted stock will be made, the number of shares the person may purchase, the purchase price, the restrictions to which the shares will be subject, and other terms and conditions. If a participant's service relationship with us ends for any reason, we may receive any or all of the shares of common stock held by the participant that have not vested as of the date the participant terminates service with us through, but not limited to, a repurchase right.

*Changes to capital structure.* In the event of any stock split, reverse stock split, stock dividend, combination, consolidation, recapitalization (including a recapitalization through a large nonrecurring cash dividend) or reclassification of the shares, subdivision of the shares, a rights offering, a reorganization, merger, spin-off, split-up, repurchase, or exchange of our common stock or other securities of ours or other significant corporate transaction, or other change affecting our common stock, then in order to prevent diminution or enlargement of the benefits or potential benefits intended to be made available under the Existing Plan, the administrator will adjust the number, kind and class of securities that may be delivered under the Existing Plan and/or the number, class, kind and price of securities covered by each outstanding award.

*Change in control.* In the event of Change in Control under the Existing Plan shall be subject to the agreement evidencing the acquisition or other combination, which need not treat all outstanding awards in an identical manner. Such agreement, without the participant's consent, shall provide for one or more of the following with respect to all outstanding awards:

- the continuation of such outstanding awards;
- the assumption of such outstanding awards by the surviving corporation or its parent;
- the substitution by the surviving corporation or its parent of new options or other equity awards for such awards;
- the cancellation of such awards in exchange for a payment to the participants equal to the excess, if any, of (1) the fair market value of the shares subject to such awards as of the closing date of such Change in Control over (2) the exercise or purchase price paid or to be paid for the shares subject to the awards; provided that at the discretion of the administrator, such payment may be subject to the same conditions that apply to the consideration that will be paid to holders of shares in connection with the transaction; or

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- the opportunity for participants to exercise the options prior to the occurrence of the Change in Control and the termination (for no consideration) upon the consummation of such Change in Control of any options not exercised prior thereto.

Under the Existing Plan, a Change in Control is generally defined as (i) the consummation of a merger or consolidation of the company with or into another entity or any other corporate reorganization, if the company's stockholders immediately prior to such merger, consolidation or reorganization cease to directly or indirectly own immediately after such merger, consolidation or reorganization at least a majority of the combined voting power of the continuing or surviving entity's securities outstanding immediately after such merger, consolidation or reorganization; (ii) the consummation of the sale, transfer or other disposition of all or substantially all of the company's assets; (iii) a change in the effective control of the company; or (iv) the consummation of any transaction as a result of which any Person (as defined below) becomes the "beneficial owner" (as defined in Rule 13d-3 under the Exchange Act), directly or indirectly, of securities of the company representing at least fifty percent (50%) of the total voting power represented by the company's then outstanding voting securities. For purposes of this Section 2(h), the term "Person" shall have the same meaning as when used in Sections 13(d) and 14(d) of the Exchange Act but shall exclude:

- a trustee or other fiduciary holding securities under an employee benefit plan of the company or an affiliate of the company;
- a corporation or other entity owned directly or indirectly by the stockholders of the company in substantially the same proportions as their ownership of our common stock;
- the company; and
- a corporation or other entity of which at least a majority of its combined voting power is owned directly or indirectly by the company.

A transaction shall not constitute a Change in Control if its sole purpose is to change the state of the company's incorporation or to create a holding company that will be owned in substantially the same proportions by the persons who held the company's securities immediately before such transactions. In addition, if any Person (as defined above) is considered to be in effective control of the company, the acquisition of additional control of the company by the same Person will not be considered to cause a Change in Control. If required for compliance with Section 409A of the Code, in no event will a Change in Control be deemed to have occurred if such transaction is not also a "change in the ownership or effective control of" the company or "a change in the ownership of a substantial portion of the assets of" the company as determined under Treasury Regulation Section 1.409A-3(i)(5) (without regard to any alternative definition thereunder).

*Plan Amendment or Termination.* The board may at any time terminate, amend or suspend the Existing Plan and all outstanding options or restricted stock awards upon a dissolution or liquidation of us.

### **Limitations on liability and indemnification matters**

Upon the closing of this offering, our amended and restated certificate of incorporation will contain provisions that limit the liability of our current and former directors for monetary damages to the fullest extent permitted by Delaware law. Delaware law provides that directors of a corporation will not be personally liable for monetary damages for any breach of fiduciary duties as directors, except liability for:

- any breach of the director's duty of loyalty to the corporation or its stockholders;
- any act or omission not in good faith or that involves intentional misconduct or a knowing violation of law;

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- unlawful payments of dividends or unlawful stock repurchases or redemptions as provided in Section 174 of the Delaware General Corporation Law; or
- any transaction from which the director derived an improper personal benefit.

These limitations of liability do not apply to liabilities arising under federal securities laws and does not affect the availability of equitable remedies such as injunctive relief or rescission.

Our amended and restated certificate of incorporation and our amended and restated bylaws that will be in effect upon the closing of this offering will provide that we are required to indemnify our directors to the fullest extent permitted by Delaware law. Our amended and restated bylaws will also provide that, upon satisfaction of certain conditions, we are required to advance expenses incurred by a director in advance of the final disposition of any action or proceeding, and permit us to secure insurance on behalf of any officer, director, employee or other agent for any liability arising out of his or her actions in that capacity regardless of whether we would otherwise be permitted to indemnify him or her under the provisions of Delaware law. Our amended and restated bylaws will also provide our board of directors with discretion to indemnify our officers and employees when determined appropriate by the board.

We have entered into indemnification agreements with each of our directors, and we expect to enter into amended and restated indemnification agreements with each of our directors and enter into indemnification agreements with each of our executive officers prior to the closing of this offering. With certain exceptions, these agreements provide for indemnification for related expenses including, among other things, attorneys' fees, judgments, fines and settlement amounts incurred by any of these individuals in any action or proceeding. We believe that these bylaw provisions and indemnification agreements are necessary to attract and retain qualified persons as directors and executive officers. We also maintain customary directors' and officers' liability insurance.

The limitation of liability and indemnification provisions in our amended and restated certificate of incorporation and amended and restated bylaws may discourage stockholders from bringing a lawsuit against our directors for breach of their fiduciary duty. They may also reduce the likelihood of derivative litigation against our directors and officers, even though an action, if successful, might benefit us and other stockholders. Further, a stockholder's investment may be adversely affected to the extent that we pay the costs of settlement and damage awards against directors and officers as required by these indemnification provisions. At present, there is no pending litigation or proceeding involving any of our directors, officers or employees for which indemnification is sought and we are not aware of any threatened litigation that may result in claims for indemnification.

Insofar as indemnification for liabilities arising under the Securities Act may be permitted for our directors, executive officers or persons controlling us, we have been informed that, in the opinion of the SEC, such indemnification is against public policy as expressed in the Securities Act and is therefore unenforceable.

### **Rule 10b5-1 sales plans**

Our directors and executive officers may adopt written plans, known as Rule 10b5-1 plans, in which they will contract with a broker to buy or sell shares of our common stock on a periodic basis. Under a Rule 10b5-1 plan, a broker executes trades pursuant to parameters established by the director or officer when entering into the plan, without further direction from them. The director or officer may amend a Rule 10b5-1 plan in some circumstances and may terminate a plan at any time. Our directors and executive officers also may buy or sell additional shares outside of a Rule 10b5-1 plan when they are not in possession of material nonpublic information subject to compliance with the terms of our insider trading policy. Prior to 180 days after the date of this offering, subject to early termination, the sale of any shares under such plans would be prohibited by the lock-up agreement that the director or officer has entered into with the underwriters for this offering.

## **Other policies**

After we become a public company, if we are required to restate our financial results due to our material noncompliance with any financial reporting requirements under the federal securities laws as a result of misconduct, our Chief Executive Officer and Chief Financial Officer may be legally required to reimburse our Company for any bonus or other incentive-based or equity-based compensation they receive in accordance with the provisions of Section 304 of the Sarbanes-Oxley Act of 2002.

Additionally, we plan to adopt a formal clawback policy that complies with regulations mandated under the Dodd-Frank Wall Street Reform and Consumer Protection Act of 2010 when the applicable rules adopted by the SEC in October 2022 become effective.

## **Certain relationships and related party transactions**

The following is a description of transactions since January 1, 2020 to which we have been a participant in which the amount involved exceeded or will exceed \$120,000, and in which any of our directors, executive officers or holders of more than 5% of our voting securities, or any members of their immediate family, had or will have a direct or indirect material interest, other than compensation arrangements which are described under “Executive Compensation.”

### **Agreements with Ronald G. Crystal, M.D.**

#### ***Consulting agreement***

In October 2020, we entered into a consulting agreement, or the Crystal Consulting Agreement, with Ronald G. Crystal, M.D., our founder and Chief Scientific Advisor and a holder of more than 5% of our voting securities. Under the Crystal Consulting Agreement, in consideration for an annual consulting fee of \$250,000, Dr. Crystal will provide us with consulting services in relation to his extensive expertise in gene therapy technologies, including (a) consulting and advisory services with respect to matters related to gene therapies for diseases; (b) participation in corporate and research and development strategy sessions; and (c) advising on recruiting and interviewing, or the Services. The Crystal Consulting Agreement shall continue for five years and, unless sooner terminated in accordance with its terms, is renewable annually for one-year periods after the initial term of five years. The Crystal Consulting Agreement is terminable by either party with proper notice or by us for cause.

#### ***Stock option award agreement***

In March 2021, we granted Dr. Crystal an option to purchase 1,791,822 shares of our common stock at an exercise price per share of \$0.22 and entered into our standard Stock Option Award Agreement with Dr. Crystal, or the Crystal Option Agreement. Of the shares subject to the option, 25% vested on the one-year anniversary of June 30, 2021 and the remainder of the shares subject to the option vest monthly on the thirtieth day of the month, or if there is no such date, the last day of the month, subject in each case to Dr. Crystal continuing to provide services to us. All of the shares subject to the option were exercisable immediately upon grant, and any then-unvested shares subject to the option will become fully vested as of immediately prior to a change of control of Lexeo.

### **Agreements with Eric Adler, M.D.**

#### ***Consulting agreement***

In July 2021, in connection with the execution of the Stock Purchase Agreement with Stelios Therapeutics, Inc., or Stelios, we entered into a consulting agreement with Eric Adler, M.D., effective July 16, 2021, or the Adler Consulting Agreement, pursuant to which Dr. Adler agreed, among other things, to serve as a member of our advisory board and to advise us on TNNI3-associated hypertrophic cardiomyopathy, ARVC and other, to-be-determined cardiomyopathies. Dr. Adler agreed to provide consulting services of no more than six hours per week. In connection with the Adler Consulting Agreement, Dr. Adler received a cash payment of \$50,000, and we agreed to grant, subject to the approval of our board of directors, an option to purchase an aggregate of 505,366 shares to Dr. Adler. Dr. Adler provided consulting services until his appointment as the Company’s Chief Scientific Officer in August 2022.



### ***Consultant stock option award agreements***

In November 2021, we granted Dr. Adler options to purchase 416,184 shares of common stock at an exercise price of \$0.46 per share stock, and entered into the Stock Option Award Agreement, dated November 4, 2021, or the Adler Consulting Stock Option Agreement. Commencing July 16, 2021, the shares subject to the Adler Consulting Stock Option Agreement vest in 48 equal monthly installments on the first day of each month, subject to Dr. Adler's continuing to provide services to us.

Additionally, in November 2021, we granted Dr. Adler options to purchase 282,949 shares of common stock at an exercise price of \$0.46 per share, and entered into the Stock Option Award Agreement, dated November 9, 2021, or the Adler Performance Stock Option Agreement. All of the shares subject to the Adler Performance Stock Option Agreement vested upon our completion of the first milestone of an HCM- or ARVC-sponsored research program in June 2022.

All of the shares subject to the above options were exercisable immediately upon grant, and any then-unvested shares subject to the above options will become fully vested immediately prior to a change of control of the Company. In December 2022, our board of directors approved an amendment to Dr. Adler's Stock Option Award Agreements to extend the length of the post-termination exercise period from three months to 10 years.

### ***Employment stock option award agreement***

In March 2023, in connection with our employment agreement with Dr. Adler, dated December 21, 2022, we granted Dr. Adler an option to purchase 1,920,210 shares of our common stock at an exercise price per share of \$1.66 and entered into the Stock Option Award Agreement, dated March 14, 2023, or the Adler Employment Stock Option Award Agreement. Of the shares subject to the option, 25% will vest on the one-year anniversary of March 14, 2023 and the remainder of the shares subject to the option vest monthly on the 14th day of each month, subject in each case to Dr. Adler's continuing to provide services to us. All of the shares subject to the option were exercisable immediately upon grant, and any then-unvested shares subject to the option will become fully vested as of immediately prior to a change of control of the Company.

### **Stelios Therapeutics Inc. acquisition**

On July 16, 2021, we purchased all of the issued and outstanding capital stock of Stelios for initial cash consideration of \$7.0 million, with payments of up to an additional \$20.5 million due to the selling shareholders upon the achievement of the following milestones when and if we achieve them: clinical candidate selection for either the ARVC or the TNNI3-associated programs; first patient dosed in a Phase 1 clinical trial for any product in our CX43 program; first patient dosed in a Phase 1 clinical trial for any product in our PKP2 program; or first patient dosed in a Phase 1 clinical trial for any candidate in our TNNI3 program. In the third quarter of 2022, a development milestone associated with a clinical candidate selection was achieved resulting in an aggregate payment of \$2.0 million to the selling shareholders of Stelios. Dr. Eric Adler, our Chief Scientific Officer, was a co-founder of Stelios and a selling shareholder. Of the initial \$7.0 million cash consideration, Dr. Adler received approximately \$1.2 million, and of the \$2.0 million milestone payment, Dr. Adler received approximately \$0.4 million. On December 15, 2022, Stelios was merged into Lexeo Therapeutics, Inc. and ceased to exist. See the section titled "Management's Discussion and Analysis of Financial Condition and Results of Operations" for additional information related to our acquisition of Stelios.

### **Private placements of our securities**

#### ***Series A convertible preferred stock financing***

In November 2020, we entered into a preferred stock purchase agreement with certain investors, including beneficial owners of greater than 5% of our capital stock and affiliates of members of our board of directors. In

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connection with the initial closing of our Series A convertible preferred stock, we sold an aggregate of 34,495,725 shares of our Series A convertible preferred stock at a purchase price of \$1.00 (including approximately \$4 million in principal and accrued interest on outstanding convertible promissory notes issued to certain investors, the conversion of which resulted in issuance of 4,495,729 shares of our Series A convertible preferred stock) for aggregate gross proceeds of approximately \$34 million. In July 2021, we sold an aggregate of 50,999,997 shares of our Series A convertible preferred stock at a purchase price of \$1.00 for aggregate gross proceeds of approximately \$51 million.

The table below sets forth the aggregate number of shares of Series A convertible preferred stock issued to our related parties in this financing:

<b>Name</b>	<b>Series A Convertible Preferred Stock (#)</b>	<b>Aggregate Purchase Price (\$)</b>
Entities affiliated with Janus Henderson Investors <sup>(1)</sup>	10,000,000	10,000,000
Longitude Venture Partners IV, L.P. <sup>(2)</sup>	18,999,999	18,999,999
Lundbeckfond Invest A/S <sup>(3)</sup>	13,999,999	13,999,999
Omega Fund VI, L.P. <sup>(4)</sup>	14,999,999	14,999,999
PBM LEX Holdings, LLC <sup>(5)</sup>	11,999,999	11,999,999

(1) Entities affiliated with Janus Henderson Investors hold more than 5% of our voting securities. Consists of (i) 3,643,715 shares of Series A preferred stock purchased by Janus Henderson Global Life Sciences Fund, (ii) 3,466,664 shares of Series A preferred stock purchased by Janus Henderson Biotech Innovation Master Fund Limited, (iii) 2,660,459 shares of Series A preferred stock purchased by Janus Henderson Capital Funds PLC on behalf of its Series Janus Henderson Global Life Sciences Fund and (iv) 229,162 shares of Series A preferred stock purchased by Janus Henderson Horizon Fund—Biotechnology Fund.

(2) Longitude Venture Partners IV, L.P., or LVP IV, holds more than 5% of our capital stock prior to this offering. Longitude Capital Partners IV, LLC is the general partner of LVP IV. Reinaldo Diaz was designated to serve as a member of our board of directors by LVP IV.

(3) Lundbeckfond Invest A/S, which is wholly owned by the Lundbeck Foundation, holds more than 5% of our capital stock prior to this offering. Mette Kirstine Agger is the former Managing Partner of Lundbeckfonden BioCapital, a department within Lundbeckfond Invest A/S, and was designated to serve as a member of our board of directors by Lundbeckfond Invest A/S.

(4) Omega Fund VI, L.P., or Omega Fund, holds more than 5% of our capital stock prior to this offering. Omega Fund VI GP, L.P., or Omega GP LP, is the sole general partner of the Omega Fund, and Omega Fund VI GP Manager, Ltd. is the sole general partner of Omega GP LP. Bernard Davitian was designated to serve as a member of our board of directors by Omega Fund.

(5) PBM LEX Holdings, LLC, or PBM LEX, holds more than 5% of our capital stock prior to this offering.

### ***Series B convertible preferred stock financing***

In August 2021, we entered into a preferred stock purchase agreement with certain investors, including beneficial owners of greater than 5% of our capital stock, affiliates of members of our board of directors and certain of our executive officers, pursuant to which we issued and sold to such investors an aggregate of 58,157,823 shares of our Series B convertible preferred stock at a purchase price of \$1.72049 per share for aggregate gross proceeds of approximately \$100 million.

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The table below sets forth the aggregate number of shares of Series B convertible preferred stock issued to our related parties in this financing:

<b>Name</b>	<b>Series B Convertible Preferred Stock (#)</b>	<b>Aggregate Purchase Price (\$)</b>
D1 Master Holdco I LLC <sup>(1)</sup>	20,343,041	34,999,999
Entities affiliated with Janus Henderson Investors <sup>(2)</sup>	2,324,918	3,999,998
Longitude Venture Partners IV, L.P. <sup>(3)</sup>	2,906,148	4,999,999
Lundbeckfond Invest A/S <sup>(4)</sup>	2,615,533	4,499,998
Mutual Fund Series Trust, On Behalf Of Eventide Healthcare & Life Sciences Fund <sup>(5)</sup>	8,718,446	14,999,999
Omega Fund VI, L.P. <sup>(6)</sup>	2,615,533	4,499,998
Paul McCormac, Ph.D. <sup>(7)</sup>	29,061	49,999

- (1) D1 Master Holdco I LLC, or Holdco I, holds more than 5% of our capital stock prior to this offering. Holdco I is a wholly owned subsidiary of D1 Capital Partners Master LP. D1 Capital Partners L.P. is a registered investment adviser and serves as the investment manager of private investment vehicles and accounts, including D1 Capital Partners Master LP, and may be deemed to beneficially own the shares held by Holdco I. Daniel Sundheim indirectly controls D1 Capital Partners L.P., and may be deemed to beneficially own the shares held by Holdco I. Paula HJ Cholmondeley was designated to serve as a member of our board of directors by Holdco I.
- (2) Consists of (i) 883,469 shares of Series B preferred stock purchased by Janus Henderson Global Life Sciences Fund and (ii) 1,441,449 shares of Series B preferred stock purchased by Janus Henderson Capital Funds PLC on behalf of its Series Janus Henderson Global Life Sciences Fund. Entities affiliated with Janus Henderson Investors hold more than 5% of our voting securities.
- (3) Longitude Venture Partners IV, L.P., or LVP IV, holds more than 5% of our capital stock prior to this offering. Longitude Capital Partners IV, LLC is the general partner of LVP IV. Reinaldo Diaz was designated to serve as a member of our board of directors by LVP IV.
- (4) Lundbeckfond Invest A/S holds more than 5% of our capital stock prior to this offering. Mette Kirstine Agger is the former Managing Partner of Lundbeckfonden BioCapital, a department within Lundbeckfond Invest A/S, and was designated to serve as a member of our board of directors by Lundbeckfond Invest A/S.
- (5) Mutual Fund Series Trust, on behalf of Eventide Healthcare & Life Sciences Fund, holds more than 5% of our capital stock prior to this offering.
- (6) Omega Fund holds more than 5% of our capital stock prior to this offering. Omega GP LP is the sole general partner of the Omega Fund, and Omega Fund VI GP Manager, Ltd. is the sole general partner of Omega GP LP. Bernard Davitian was designated to serve as a member of our board of directors by Omega Fund.
- (7) All shares are held by Paul McCormac, Ph.D., our Chief Technical Officer.

## **Investors' rights, voting and right of first refusal agreements**

In connection with the sales of convertible preferred stock described above, we entered into an amended and restated investors' rights agreement, an amended and restated voting agreement and an amended and restated right of first refusal and co-sale agreement containing registration rights, information rights, voting rights and rights of first refusal, among other things, with the holders of our convertible preferred stock, including funds affiliated with D1 Capital Partners, Eventide Healthcare & Life Sciences Fund, Janus Henderson Investors, Longitude Capital, Lundbeckfond Invest A/S, Omega Funds and PBM Capital. These agreements will terminate upon the closing of this offering, except for the registration rights granted under our amended and restated investors' rights agreement, as more fully described in the section of this prospectus titled "Description of Capital Stock—Registration Rights."

## **Employment arrangements**

We have entered into employment agreements or offer letter agreements with certain of our executive officers. For more information regarding such employment agreements, see "Executive Compensation—Employment Agreements with our Named Executive Officers."

## **Indemnification agreements**

Our amended and restated certificate of incorporation that will be in effect upon the closing of this offering will contain provisions limiting the liability of directors, and our amended and restated bylaws will provide that we will indemnify each of our directors to the fullest extent permitted under Delaware law. Our amended and restated certificate of incorporation and amended and restated bylaws will also provide our board of directors with discretion to indemnify our officers and employees when determined appropriate by the board.

In addition, we have entered into indemnification agreements with each of our directors, and we expect to enter into amended and restated indemnification agreements with each of our directors and enter into indemnification agreements with each of our executive officers prior to the closing of this offering. For more information regarding these agreements, see “Executive Compensation—Limitations on Liability and Indemnification Matters.”

## **Related person transaction policy**

Prior to this offering, we have not had a formal policy regarding approval of transactions with related parties. In connection with this offering, we have adopted a related person transaction policy that sets forth our procedures for the identification, review, consideration and approval or ratification of related person transactions, which policy will become effective immediately upon the execution of the underwriting agreement for this offering. For purposes of our policy only, a related person transaction will be a transaction, arrangement or relationship, or any series of similar transactions, arrangements or relationships, in which we and any related person are, were, or will be participants and in which the amount involved exceeds \$120,000. Transactions involving compensation for services provided to us as an employee or director will not be covered by this policy. A related person will be any executive officer, director or beneficial owner of more than 5% of any class of our voting securities, including any of their immediate family members and any entity owned or controlled by such persons.

Under the policy, if a transaction has been identified as a related person transaction, including any transaction that was not a related person transaction when originally consummated or any transaction that was not initially identified as a related person transaction prior to consummation, our management must present information regarding the related person transaction to our audit committee, or, if audit committee approval would be inappropriate, to another independent body of our board of directors, for review, consideration and approval or ratification. The presentation must include a description of, among other things, the material facts, the interests, direct and indirect, of the related persons, the benefits to us of the transaction and whether the transaction is on terms that are comparable to the terms available to or from, as the case may be, an unrelated third party or to or from employees generally. Under the policy, we will collect information that we deem reasonably necessary from each director, executive officer and, to the extent feasible, significant stockholder to enable us to identify any existing or potential related-person transactions and to effectuate the terms of the policy. In addition, under our Code of Conduct that we expect to adopt prior to the closing of this offering, our employees and directors will have an affirmative responsibility to disclose any transaction or relationship that reasonably could be expected to give rise to a conflict of interest. In considering related person transactions, our audit committee, or other independent body of our board of directors, will take into account the relevant available facts and circumstances including:

- the risks, costs and benefits to us;
- the impact on a director’s independence in the event that the related person is a director, immediate family member of a director or an entity with which a director is affiliated;

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- the availability of other sources for comparable services or products; and
- the terms available to or from, as the case may be, unrelated third parties or to or from employees generally.

The policy will require that, in determining whether to approve, ratify or reject a related person transaction, our audit committee, or other independent body of our board of directors, must consider, in light of known circumstances, whether the transaction is in, or is not inconsistent with, our best interests and those of our stockholders, as our audit committee, or other independent body of our board of directors, determines in the good faith exercise of its discretion.

All of the transactions described in this section were entered into prior to the adoption of this policy. Although we have not had a written policy for the review and approval of transactions with related persons, our board of directors has historically reviewed and approved any transaction where a director or officer had a financial interest, including the transactions described above. Prior to approving such a transaction, the material facts as to a director's or officer's relationship or interest in the agreement or transaction were disclosed to our board of directors. Our board of directors took this information into account when evaluating the transaction and in determining whether such transaction was fair to us and in the best interest of all our stockholders.

## Principal stockholders

The following table sets forth information regarding beneficial ownership of our capital stock by:

- each person, or group of affiliated persons, known by us to beneficially own more than 5% of our common stock;
- each of our directors;
- our named executive officers; and
- all of our current executive officers and directors as a group.

We have determined beneficial ownership in accordance with the rules of the SEC. Under these rules, beneficial ownership includes any shares of common stock as to which the individual or entity has sole or shared voting power or investment power. Applicable percentage ownership is based on \_\_\_\_\_ shares of common stock outstanding as of \_\_\_\_\_, 2023, after giving effect to the conversion of all of our preferred stock. In computing the number of shares beneficially owned by an individual or entity and the percentage ownership of that person, shares of common stock subject to options held by such person that are currently exercisable or will become exercisable within 60 days of \_\_\_\_\_, 2023 are considered outstanding, although these shares are not considered outstanding for purposes of computing the percentage ownership of any other person.

Unless noted otherwise, the address of all listed stockholders is c/o Lexeo Therapeutics, Inc., 345 Park Avenue South, Floor 6, New York, New York 10010.

Except as indicated by the footnotes below, we believe, based on information furnished to us, that each of the stockholders listed has sole voting and investment power with respect to the shares beneficially owned by the stockholder unless noted otherwise, subject to community property laws where applicable.

Name of Beneficial Owner	Number of Shares Beneficially Owned	Percentage of Shares Beneficially Owned	
		Before Offering	After Offering
<b>Greater than 5% stockholders</b>			
D1 Master Holdco I LLC <sup>(1)</sup>			%
PBM LEX Holdings, LLC <sup>(2)</sup>			
Entities affiliated with Janus Henderson Investors <sup>(3)</sup>			
Longitude Venture Partners IV, L.P. <sup>(4)</sup>			
Lundbeckfond Invest A/S <sup>(5)</sup>			
Mutual Fund Series Trust, on Behalf of Eventide Healthcare & Life Sciences Fund <sup>(6)</sup>			
Omega Fund VI, L.P. <sup>(7)</sup>			
Ronald G. Crystal, M.D. and affiliates <sup>(8)</sup>			
<b>Named Executive Officers and Directors</b>			
R. Nolan Townsend <sup>(9)</sup>			
Jenny R. Robertson			
Jay A. Barth			
Mette Kirstine Agger			
Steven Altschuler, M.D. <sup>(11)</sup>			
Paula HJ Cholmondeley <sup>(12)</sup>			
Bernard Davitian			
Reinaldo Diaz			
All current executive officers and directors as a group (ten persons) <sup>(13)</sup>			

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\* Represents beneficial ownership of less than one percent.

- (1) All of the shares are held by D1 Master Holdco I LLC. Consists of \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock. D1 Capital Partners L.P., or D1 Management Company, is a registered investment adviser and serves as the investment manager of private investment vehicles and accounts, including D1 Master Holdco I LLC, and may be deemed to beneficially own the shares of common stock held by D1 Master Holdco I LLC. Daniel Sundheim indirectly controls the D1 Management Company and may be deemed to beneficially own the shares of common stock held by D1 Master Holdco I LLC. The business address of each of D1 Master Holdco I LLC, D1 Capital Partners L.P. and Daniel Sundheim is 9 West 57th Street, 36th Floor, New York, New York 10019.
- (2) Consists of \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock held by PBM LEX Holdings, LLC. PBM LEX Holdings, LLC is majority owned by PBM Capital Investments II, LLC, which is managed by PBM Capital Group, LLC. Paul B. Manning is the Chief Executive Officer of PBM Capital Group, LLC, and has sole voting and investment power with respect to the shares held by PBM LEX Holdings, LLC. The business address for each person and entity named in this footnote is 200 Garrett Street, Suite S, Charlottesville, Virginia 22902.
- (3) Consists of (i) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock held by Janus Henderson Biotech Innovation Master Fund Limited, (ii) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock held by Janus Henderson Horizon Fund – Biotechnology Fund, (iii) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock held by Janus Henderson Capital Funds PLC On Behalf Of Its Series Janus Henderson Global Life Sciences Fund, (iv) \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock held by Janus Henderson Capital Funds PLC On Behalf Of Its Series Janus Henderson Global Life Sciences Fund, (v) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock held by Janus Henderson Global Life Sciences Fund and (vi) \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock held by Janus Henderson Global Life Sciences Fund. Such shares owned by the funds may be deemed to be beneficially owned by, Janus Henderson Investors US LLC, or Janus, an investment adviser registered under the Investment Advisers Act of 1940, who acts as investment adviser for the funds set forth above and has the ability to make decisions with respect to the voting and disposition of the shares subject to the oversight of the board of trustees (or similar entity) of each fund. Under the terms of its management contract with each fund, Janus has overall responsibility for directing the investments of the fund in accordance with the fund's investment objective, policies and limitations. Each fund has one or more portfolio managers appointed by and serving at the pleasure of Janus who makes decisions with respect to the disposition of the Shares. The address for Janus and each of the foregoing entities is 151 Detroit Street, Denver, Colorado 80206.
- (4) All of the shares are held by Longitude Venture Partners IV, L.P. (LVP IV). Consists of (i) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock and (ii) \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock. Longitude Capital Partners IV, LLC (LCP IV), is the general partner of LVP IV and may be deemed to have voting and investment power over the shares held by LVP IV. Patrick Enright and Juliet Tammenoms Bakker are managing members of LCP IV and may be deemed to share voting and investment power over the shares held by LVP IV. Each of LCP IV, Ms. Tammenoms Bakker and Mr. Enright disclaims beneficial ownership of such shares except to the extent of their respective pecuniary interests therein. The address for this entity is 2740 Sand Hill Road, 2nd Floor, Menlo Park, California 94025.
- (5) All of the shares are held by Lundbeckfond Invest A/S. Consists of (i) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock and (ii) \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock. Lundbeckfond Invest A/S is wholly-owned by the Lundbeck Foundation, and the board of directors of the Lundbeckfond Foundation consists of Steffen Kragh, Lars Holmqvist, Susanne Krüger Kjær, Michael Kjær, Peter Schütze, Gunhild Waldemar, Svend Andersen, Ludovic Tranholm Otterbein, Vagn Flink Møller Pedersen and Kristian Funding Andersen. The board of directors of the Lundbeck Foundation also serve as the board members of Lundbeckfond Invest A/S. No individual member of the Lundbeckfond Invest A/S board of directors is deemed to hold any beneficial ownership or reportable pecuniary interest in the shares held by Lundbeckfond Invest A/S. The board of directors of Lundbeckfond Invest A/S makes decisions with respect to investments made by Lundbeckfond Invest A/S, and the board of directors of Lundbeckfond Invest A/S and Lene Skole, the chief executive officer of Lundbeckfond Invest A/S, may be deemed to share voting and investment authority over the shares held by Lundbeckfond Invest A/S. The address of Lundbeckfond Invest A/S is Scherfigsvej 7 DK-2100, Copenhagen Ø, Denmark.
- (6) All of the shares are held by Mutual Fund Series Trust, On Behalf of Eventide Healthcare & Life Sciences Fund. Consists of \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock. Eventide Healthcare & Life Sciences Fund is a registered investment company for which Eventide Asset Management, LLC acts as investment advisor. Eventide Asset Management, LLC has voting and investment power with respect to the shares. The address for this entity is c/o Eventide Asset Management, LLC, One International Place, Suite 4210, Boston, Massachusetts 02110.
- (7) All of the shares are held by Omega Fund. Consists of (i) \_\_\_\_\_ shares of common stock issuable upon conversion of Series A convertible preferred stock and (ii) \_\_\_\_\_ shares of common stock issuable upon conversion of Series B convertible preferred stock. Omega Fund GP Manager, Ltd., or Omega Ltd., is the sole general partner of Omega Fund VI GP, L.P., or Omega GP, which is the sole general partner of Omega Fund; and each of Omega Ltd. and Omega GP may be deemed to own beneficially the shares held by Omega Fund. Claudio Nessi, Otello Stampacchia and Anne-Mari Paster are the managing directors of Omega Ltd. and, as a result, may be deemed to share voting and investment power over the shares held directly by Omega Fund. Each of Dr. Stampacchia, Dr. Nessi, and Ms. Paster, and Omega Ltd. and Omega GP disclaims beneficial ownership of the shares held by Omega Fund except to the extent of their pecuniary interest therein. The address for this entity is 888 Boylston Street, Suite 1111, Boston, Massachusetts 02199.
- (8) Consists of (i) \_\_\_\_\_ shares held by Ronald G. Crystal, M.D., (ii) \_\_\_\_\_ shares held by the Ronald G. Crystal 2021 Grantor Retained Annuity Trust over which Dr. Crystal retains sole voting and dispositive power and (iii) \_\_\_\_\_ shares subject to options held by Dr. Crystal that are exercisable within 60 days of \_\_\_\_\_, 2023.
- (9) Consists of (i) \_\_\_\_\_ shares of common stock and (ii) \_\_\_\_\_ shares subject to options that are exercisable within 60 days of \_\_\_\_\_, 2023.
- (10) Includes shares subject to options that are exercisable within 60 days of \_\_\_\_\_, 2023.
- (11) Includes shares subject to options that are exercisable within 60 days of \_\_\_\_\_, 2023.
- (12) Includes shares subject to options that are exercisable within 60 days of \_\_\_\_\_, 2023.
- (13) Consists of (i) \_\_\_\_\_ shares of common stock and (ii) \_\_\_\_\_ shares subject to options that are exercisable within 60 days of \_\_\_\_\_, 2023.

## Description of capital stock

*The following description of our capital stock, certain provisions of our amended and restated certificate of incorporation and amended and restated bylaws, as each will be in effect following the completion of this offering, and certain provisions of Delaware law are summaries. You should also refer to the amended and restated certificate of incorporation and the amended and restated bylaws, which are filed as exhibits to the registration statement of which this prospectus is part.*

### General

Upon the completion of this offering, our amended and restated certificate of incorporation will authorize us to issue up to \_\_\_\_\_ shares of common stock, \$0.0001 par value per share, and \_\_\_\_\_ shares of preferred stock, \$0.0001 par value per share, all of which shares of preferred stock will be undesignated. Our board of directors may establish the rights and preferences of the preferred stock from time to time. As of March 31, 2023, there were \_\_\_\_\_ shares of Series A convertible preferred stock outstanding and \_\_\_\_\_ shares of Series B convertible preferred stock outstanding, held by stockholders of record. All currently outstanding shares of convertible preferred stock will be converted into an aggregate of \_\_\_\_\_ shares of common stock upon the closing of this offering.

### Common stock

#### *Voting rights*

Each holder of our common stock is entitled to one vote for each share on all matters submitted to a vote of the stockholders, including the election of directors. The affirmative vote of holders of at least 66-2/3% of the voting power of all of the then-outstanding shares of capital stock, voting as a single class, will be required to amend certain provisions of our amended and restated certificate of incorporation, including provisions relating to amending our amended and restated bylaws, the classified board, the size of our board, removal of directors, director liability, vacancies on our board, special meetings, stockholder notices, actions by written consent and exclusive forum.

#### *Dividends*

Subject to preferences that may be applicable to any then-outstanding preferred stock, holders of common stock are entitled to receive ratably those dividends, if any, as may be declared from time to time by the board of directors out of legally available funds.

#### *Liquidation*

In the event of our liquidation, dissolution or winding up, holders of common stock will be entitled to share ratably in the net assets legally available for distribution to stockholders after the payment of all of our debts and other liabilities and the satisfaction of any liquidation preference granted to the holders of any then-outstanding shares of preferred stock.

#### *Rights and preferences*

Holders of common stock have no preemptive, conversion or subscription rights and there are no redemption or sinking fund provisions applicable to the common stock. The rights, preferences and privileges of the holders of common stock are subject to, and may be adversely affected by, the right of the holders of shares of any series of preferred stock that we may designate in the future.



## **Preferred stock**

Following the closing of this offering, our board of directors will have the authority under our amended and restated certificate of incorporation, without further action by our stockholders, to issue up to \_\_\_\_\_ shares of preferred stock in one or more series, to establish from time to time the number of shares to be included in each such series, to fix the rights, preferences and privileges of the shares of each wholly unissued series and any qualifications, limitations or restrictions thereon, and to increase or decrease the number of shares of any such series, but not below the number of shares of such series then outstanding.

Our board of directors may authorize the issuance of preferred stock with voting or conversion rights that could adversely affect the voting power or other rights of the holders of our common stock. The purpose of authorizing our board of directors to issue preferred stock and determine its rights and preferences is to eliminate delays associated with a stockholder vote on specific issuances. The issuance of preferred stock, while providing flexibility in connection with possible acquisitions and other corporate purposes, could, among other things, have the effect of delaying, deferring or preventing a change in control of us and may adversely affect the market price of our common stock and the voting and other rights of the holders of our common stock. It is not possible to state the actual effect of the issuance of any shares of preferred stock on the rights of holders of common stock until the board of directors determines the specific rights attached to that preferred stock.

We have no present plans to issue any shares of preferred stock following completion of this offering.

## **Options and restricted stock awards**

As of March 31, 2023, there were options to purchase \_\_\_\_\_ shares of common stock outstanding and no shares of common stock had been issued pursuant to restricted stock awards. For additional information regarding the terms of the Existing Plan, see “Executive Compensation—Equity Incentive Plans.”

## **Registration rights**

We, the holders of our existing convertible preferred stock and certain holders of our existing common stock have entered into an amended and restated investors’ rights agreement. The registration rights provisions of this agreement provide those holders with demand, piggyback and Form S-3 registration rights with respect to the shares of common stock currently held by them and issuable to them upon conversion of our convertible preferred stock in connection with our initial public offering. These shares are collectively referred to herein as registrable securities.

### ***Demand registration rights***

At any time beginning 180 days following the effective date of the registration statement of which this prospectus is a part, the holders of a majority of registrable securities then outstanding have the right to demand that we file a Form S-1 registration statement covering registrable securities then outstanding having an aggregate offering price in excess of \$10 million. These registration rights are subject to specified conditions and limitations, including the right of the underwriters, if any, to limit the number of shares included in any such registration under specified circumstances. Upon such a request, we are required to effect a Form S-1 registration statement as soon as practicable, but in any event no later than 60 days after the date such request is given by the initiating holders. An aggregate of \_\_\_\_\_ shares of common stock will be entitled to these demand registration rights.

### ***Piggyback registration rights***

If we propose to register any of our securities under the Securities Act either for our own account or for the account of other stockholders, the holders of registrable securities will each be entitled to notice of the

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registration and will be entitled to include their shares of common stock in the registration statement. These piggyback registration rights are subject to specified conditions and limitations, including the right of the underwriters to limit the number of shares included in any such registration under specified circumstances. An aggregate of \_\_\_\_\_ shares of common stock will be entitled to these piggyback registration rights.

### ***Registration on Form S-3***

At any time after we become eligible to file a registration statement on Form S-3, the holders of at least 10% of registrable securities then outstanding will be entitled to request to have such shares registered by us on a Form S-3 registration statement. These Form S-3 registration rights are subject to other specified conditions and limitations, including the condition that the anticipated aggregate offering price, net of certain selling expenses, is at least \$1 million. Upon receipt of this request, the holders of registrable securities will each be entitled to participate in this registration. An aggregate of \_\_\_\_\_ shares of common stock will be entitled to these Form S-3 registration rights.

### ***Expenses of registration***

We are required to pay all expenses, including reasonable fees and expenses of one counsel to represent the selling stockholders, relating to any demand, piggyback or Form S-3 registration, other than underwriting discounts and commissions, stock transfer taxes and any additional fees of counsel for the selling stockholders, subject to specified conditions and limitations. We are not required to pay registration expenses if a demand registration request is withdrawn at the request of a majority of holders of registrable securities to be registered, unless holders of a majority of the registrable securities agree to forfeit their right to one demand registration.

The amended and restated investors' rights agreement contains customary cross-indemnification provisions, pursuant to which we are obligated to indemnify the selling stockholders in the event of material misstatements or omissions in the applicable registration statement attributable to us, and the selling stockholders are obligated to indemnify us for material misstatements or omissions in the registration statement attributable to them, subject to certain limitations.

### ***Termination of registration rights***

The registration rights granted under the investors' rights agreement will terminate with respect to any particular stockholder upon the earlier of (a) the closing of a deemed liquidation event, as defined in our certificate of incorporation, (b) with respect to each stockholder, at such time such stockholder is able to sell all of its shares pursuant to Rule 144 or another similar exemption under the Securities Act during a three-month period without registration and (c) the fifth anniversary of the closing of this offering or such later date that is 180 days following the expiration of all deferrals of our obligations with respect to any registration rights exercised by holders of registrable securities as of such fifth anniversary.

## **Anti-takeover provisions**

### ***Section 203 of the Delaware General Corporation Law***

We are subject to Section 203 of the Delaware General Corporation Law, which prohibits a Delaware corporation from engaging in any business combination with any interested stockholder for a period of three years after the date that such stockholder became an interested stockholder, with the following exceptions:

- before such date, the board of directors of the corporation approved either the business combination or the transaction that resulted in the stockholder becoming an interested stockholder;

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- upon completion of the transaction that resulted in the stockholder becoming an interested stockholder, the interested stockholder owned at least 85% of the voting stock of the corporation outstanding at the time the transaction began, excluding for purposes of determining the voting stock outstanding, but not the outstanding voting stock owned by the interested stockholder, those shares owned (i) by persons who are directors and also officers and (ii) employee stock plans in which employee participants do not have the right to determine confidentially whether shares held subject to the plan will be tendered in a tender or exchange offer; or
- on or after such date, the business combination is approved by the board of directors and authorized at an annual or special meeting of the stockholders, and not by written consent, by the affirmative vote of at least 66-2/3% of the outstanding voting stock that is not owned by the interested stockholder.

In general, Section 203 defines a “business combination” to include the following:

- any merger or consolidation involving the corporation or any direct or indirect majority-owned subsidiary of the corporation and the interested stockholder;
- any sale, transfer, pledge or other disposition of 10% or more of the assets of the corporation involving the interested stockholder (in one transaction or a series of transactions);
- subject to certain exceptions, any transaction that results in the issuance or transfer by the corporation or by any direct or indirect majority-owned subsidiary of the corporation of any stock of the corporation or of such subsidiary to the interested stockholder;
- any transaction involving the corporation or any direct or indirect majority-owned subsidiary of the corporation that has the effect of increasing the proportionate share of the stock or any class or series of the corporation beneficially owned by the interested stockholder; or
- the receipt by the interested stockholder of the benefit of any loans, advances, guarantees, pledges or other financial benefits by or through the corporation.

In general, Section 203 defines an “interested stockholder” as an entity or person who, together with the person’s affiliates and associates, beneficially owns, or within three years prior to the time of determination of interested stockholder status did own, 15% or more of the outstanding voting stock of the corporation.

### ***Amended and restated certificate of incorporation and amended and restated bylaws***

Our amended and restated certificate of incorporation to be in effect upon the completion of this offering, or our restated certificate, will provide for our board of directors to be divided into three classes with staggered three-year terms. Only one class of directors will be elected at each annual meeting of our stockholders, with the other classes continuing for the remainder of their respective three-year terms. Because our stockholders do not have cumulative voting rights, stockholders holding a majority of the shares of common stock outstanding will be able to elect all of our directors. Our restated certificate and our amended and restated bylaws to be effective upon the completion of this offering, or our restated bylaws, will also provide that directors may be removed by the stockholders only for cause upon the vote of 66-2/3% or more of our outstanding common stock. Furthermore, the authorized number of directors may be changed only by resolution of the board of directors, and vacancies and newly created directorships on the board of directors may, except as otherwise required by law or determined by the board, only be filled by a majority vote of the directors then serving on the board, even though less than a quorum.

Under our restated certificate of incorporation and amended and restated bylaws our stockholders will not have cumulative voting rights. Because of this, the holders of a majority of the shares of common stock entitled to vote in any election of directors can elect all of the directors standing for election, if they should so choose.

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Our restated certificate and restated bylaws will also provide that all stockholder actions must be effected at a duly called meeting of stockholders and will eliminate the right of stockholders to act by written consent without a meeting. Our restated bylaws will also provide that only our Chairman of the board, Chief Executive Officer or the board of directors pursuant to a resolution adopted by a majority of the total number of authorized directors may call a special meeting of stockholders.

Our restated bylaws will also provide that stockholders seeking to present proposals before a meeting of stockholders to nominate candidates for election as directors at a meeting of stockholders must provide timely advance notice in writing, and will specify requirements as to the form and content of a stockholder's notice.

Our restated certificate and restated bylaws will provide that the stockholders cannot amend many of the provisions described above except by a vote of 66-2/3% or more of our outstanding common stock.

As described in "—Preferred Stock" above, our restated certificate will give our board of directors the authority, without further action by our stockholders, to issue up to \_\_\_\_\_ shares of preferred stock in one or more series, with any rights, preferences and privileges as they may designate, including the right to approve an acquisition or other change in control.

The combination of these provisions will make it more difficult for our existing stockholders to replace our board of directors as well as for another party to obtain control of us by replacing our board of directors. Since our board of directors has the power to retain and discharge our officers, these provisions could also make it more difficult for existing stockholders or another party to effect a change in management. In addition, the authorization of undesignated preferred stock makes it possible for our board of directors to issue preferred stock with voting or other rights or preferences that could impede the success of any attempt to change our control.

These provisions are intended to enhance the likelihood of continued stability in the composition of our board of directors and its policies and to discourage coercive takeover practices and inadequate takeover bids. These provisions are also designed to reduce our vulnerability to hostile takeovers and to discourage certain tactics that may be used in proxy fights. However, such provisions could have the effect of discouraging others from making tender offers for our shares and may have the effect of delaying changes in our control or management. As a consequence, these provisions may also inhibit fluctuations in the market price of our stock that could result from actual or rumored takeover attempts. We believe that the benefits of these provisions, including increased protection of our potential ability to negotiate with the proponent of an unfriendly or unsolicited proposal to acquire or restructure our company, outweigh the disadvantages of discouraging takeover proposals, because negotiation of takeover proposals could result in an improvement of their terms.

### **Choice of forum**

Our amended and restated certificate of incorporation will provide that the Court of Chancery of the state of Delaware will be the exclusive forum for the following types of actions or proceedings under Delaware statutory or common law:

- any derivative action or proceeding brought on our behalf;
- any action asserting a breach of fiduciary duty;
- any action asserting a claim against us arising pursuant to the Delaware General Corporation Law, our restated certificate, or our amended and restated bylaws; or
- any action asserting a claim against us that is governed by the internal affairs doctrine.

The provision would not apply to suits brought to enforce a duty or liability created by the Exchange Act. Furthermore, Section 22 of the Securities Act creates concurrent jurisdiction for federal and state courts over all

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such Securities Act actions. Accordingly, both state and federal courts have jurisdiction to entertain such claims. To prevent having to litigate claims in multiple jurisdictions and the threat of inconsistent or contrary rulings by different courts, among other considerations, our amended and restated certificate of incorporation will also provide that unless we consent in writing to the selection of an alternative forum, the federal district courts of the United States shall be the exclusive forum for the resolution of any complaint asserting a cause of action arising under the Securities Act.

While the Delaware courts have determined that such choice of forum provisions are facially valid, a stockholder may nevertheless seek to bring a claim in a venue other than those designated in the exclusive forum provisions. In such instance, we would expect to vigorously assert the validity and enforceability of the exclusive forum provisions of our restated certificate of incorporation. This may require significant additional costs associated with resolving such action in other jurisdictions and there can be no assurance that the provisions will be enforced by a court in those other jurisdictions.

These exclusive forum provisions may limit a stockholder's ability to bring a claim in a judicial forum that it finds favorable for disputes with us or our directors, officers, or other employees, which may discourage lawsuits against us and our directors, officers and other employees. If a court were to find either exclusive-forum provision in our amended and restated certificate of incorporation to be inapplicable or unenforceable in an action, we may incur further significant additional costs associated with resolving the dispute in other jurisdictions, all of which could seriously harm our business.

Our amended and restated certificate of incorporation will further provide that the federal district courts of the United States of America will be the exclusive forum for resolving any complaint asserting a cause of action arising under the Securities Act, subject to and contingent upon a final adjudication in the State of Delaware of the enforceability of such exclusive forum provision.

### **Transfer agent and registrar**

The transfer agent and registrar for our common stock is . The transfer agent's address is .

### **Listing**

We intend to apply for listing of our common stock on the Nasdaq Global Market under the trading symbol "LXEO."

## Shares eligible for future sale

Prior to this offering, no public market existed for our common stock. Future sales of our common stock in the public market, or the availability of such shares for sale in the public market, could adversely affect market prices prevailing from time to time. As described below, only a limited number of shares will be available for sale shortly after this offering due to contractual and legal restrictions on resale. Nevertheless, sales of our common stock in the public market after such restrictions lapse, or the perception that those sales may occur, could adversely affect the prevailing market price at such time and our ability to raise equity capital in the future.

Based on the number of shares outstanding as of March 31, 2023, upon the closing of this offering and assuming no exercise of the underwriters' option to purchase additional shares, \_\_\_\_\_ shares of common stock will be outstanding, assuming no outstanding options are exercised. All of the shares of common stock sold in this offering will be freely tradable without restrictions or further registration under the Securities Act, except for any shares sold to our "affiliates," as that term is defined under Rule 144 under the Securities Act. The remaining \_\_\_\_\_ shares of common stock held by existing stockholders are "restricted securities," as that term is defined in Rule 144 under the Securities Act. Restricted securities may be sold in the public market only if registered or if their resale qualifies for exemption from registration described below under Rule 144 promulgated under the Securities Act or another available exemption.

As a result of the lock-up agreements described below and the provisions of Rules 144 and 701 under the Securities Act, the shares of common stock that will be deemed restricted securities after this offering will be available for sale in the public market as follows:

- none of the existing shares will be eligible for immediate sale upon the completion of this offering; and
- \_\_\_\_\_ shares will be eligible for sale in the public market upon expiration of lock-up agreements 180 days after the date of this prospectus, subject in certain circumstances to the volume, manner of sale and other limitations under Rule 144 and Rule 701 under the Securities Act, which are summarized below.

### Rule 144

In general, non-affiliate persons who have beneficially owned restricted shares of our common stock for at least six months, and any affiliate of the company who owns either restricted or unrestricted shares of our common stock, are entitled to sell their securities without registration with the SEC under an exemption from registration provided by Rule 144 under the Securities Act.

#### *Non-affiliates*

Any person who is not deemed to have been one of our affiliates at the time of, or at any time during the three months preceding, a sale may sell an unlimited number of restricted securities under Rule 144 if:

- the restricted securities have been held for at least six months, including the holding period of any prior owner other than one of our affiliates (subject to certain exceptions);
- we have been subject to the Exchange Act periodic reporting requirements for at least 90 days before the sale; and
- we are current in our Exchange Act reporting at the time of sale.

Any person who is not deemed to have been an affiliate of ours at the time of, or at any time during the three months preceding, a sale and has held the restricted securities for at least one year, including the holding period

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of any prior owner other than one of our affiliates, will be entitled to sell an unlimited number of restricted securities without regard to the length of time we have been subject to Exchange Act periodic reporting or whether we are current in our Exchange Act reporting. Non-affiliate resales are not subject to the manner of sale, volume limitation or notice filing provisions of Rule 144.

### ***Affiliates***

Persons seeking to sell restricted securities who are our affiliates at the time of, or any time during the three months preceding, a sale, would be subject to the restrictions described above. They are also subject to additional restrictions, by which such person would be required to comply with the manner of sale and notice provisions of Rule 144 and would be entitled to sell within any three-month period only that number of securities that does not exceed the greater of either of the following:

- 1% of the number of shares of our common stock then outstanding, which will equal approximately \_\_\_\_\_ shares immediately after the completion of this offering based on the number of shares outstanding as of March 31, 2023; or
- the average weekly trading volume of our common stock on the stock exchange on which our shares are listed during the four calendar weeks preceding the filing of a notice on Form 144 with respect to the sale.

Additionally, persons who are our affiliates at the time of, or any time during the three months preceding, a sale may sell unrestricted securities under the requirements of Rule 144 described above, without regard to the six-month holding period of Rule 144, which does not apply to sales of unrestricted securities.

### **Rule 701**

Rule 701 under the Securities Act, as in effect on the date of this prospectus, permits resales of shares in reliance upon Rule 144 but without compliance with certain restrictions of Rule 144, including the holding period requirement. Most of our employees, executive officers or directors who purchased shares under a written compensatory plan or contract may be entitled to rely on the resale provisions of Rule 701, but all holders of Rule 701 shares are required to wait until 90 days after the date of this prospectus before selling their shares. However, substantially all Rule 701 shares are subject to lock-up agreements as described below and in the section titled “Underwriting” and will become eligible for sale upon the expiration of the restrictions set forth in those agreements.

### **Form S-8 registration statements**

We intend to file one or more registration statements on Form S-8 under the Securities Act to register all shares of common stock subject to outstanding stock options and common stock issued or issuable under our equity plans. We expect to file the registration statement covering shares offered pursuant to our stock plans as soon as practicable after the closing of this offering, permitting the resale of such shares by non-affiliates in the public market without restriction under the Securities Act and the sale by affiliates in the public market, subject to compliance with the resale provisions of Rule 144 and expiration or release from the terms of the lock-up agreements described above.

### **Lock-up agreements**

We, our executive officers and directors and substantially all of the holders of our common stock outstanding on the date of this prospectus have entered into lock-up agreements with the underwriters or otherwise agreed, subject to limited exceptions, that we and they will not, offer, pledge, announce the intention to sell, sell, contract

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to sell, sell any option or contract to purchase, purchase any option or contract to sell, grant any option, right or warrant to purchase or otherwise dispose of, directly or indirectly, or enter into any swap or other agreement that transfers, in whole or in part, any of the economic consequences of ownership of our common stock, or any securities convertible into, or exchangeable for or that represent the right to receive shares of our common stock, without the prior written consent of J.P. Morgan Securities LLC and SVB Securities LLC for a period of 180 days from the date of this prospectus. See “Underwriting” for additional information on these lock-up agreements.

In addition to the restrictions contained in the lock-up agreements described above, we have entered into an agreement with the holders of our preferred stock that contains market stand-off provisions imposing restrictions on the ability of such security holders to sell or otherwise transfer or dispose of any registrable securities for a period of 180 days following the date of this prospectus.

### **Registration rights**

Upon the closing of this offering, the holders of \_\_\_\_\_ shares of our common stock, including common stock issuable upon the conversion of our preferred stock, or their transferees, will be entitled to specified rights with respect to the registration of their registrable shares under the Securities Act, subject to certain limitations and the expiration, waiver or termination of the lock-up agreements. Registration of these shares under the Securities Act would result in the shares becoming freely tradable without restriction under the Securities Act immediately upon effectiveness of the registration. See “Description of Capital Stock—Registration Rights” for additional information.



## Material U.S. federal income tax consequences to non-U.S. holders

The following is a summary of material U.S. federal income tax consequences to non-U.S. holders (as defined below) of the purchase, ownership and disposition of our common stock offered pursuant to this prospectus. This discussion is not a complete analysis of all potential U.S. federal income tax consequences relating thereto, does not address the potential application of the Medicare contribution tax on net investment income, the alternative minimum tax, or the special tax accounting rules under Section 451(b) of the Code, and does not address any U.S. federal non-income tax consequences such as estate or gift tax consequences or any tax consequences arising under any state, local, or non-U.S. tax laws. This discussion is based on the Code and applicable Treasury Regulations promulgated thereunder, judicial decisions and published rulings, and administrative pronouncements of the Internal Revenue Service, or the IRS, all as in effect as of the date hereof. These authorities are subject to differing interpretations and may change, possibly retroactively, resulting in U.S. federal income tax consequences different from those discussed below. We have not requested, and do not intend to request, a ruling from the IRS with respect to the U.S. federal income tax consequences discussed below, and there can be no assurance that the IRS or a court will agree with such tax consequences.

This discussion is further limited to non-U.S. holders that hold our common stock as a “capital asset” within the meaning of Section 1221 of the Code (generally, property held for investment). This discussion also does not address all U.S. federal income tax consequences that may be relevant to a particular non-U.S. holder in light of such non-U.S. holder’s particular circumstances. Finally, this discussion does not address any specific facts or circumstances that may be relevant to non-U.S. holders subject to special rules under U.S. federal income tax laws, including:

- certain former citizens or long-term residents of the United States;
- partnerships, S corporations, or other entities or arrangements treated as partnerships, pass-through entities, or disregarded entities (including hybrid entities) for U.S. federal income tax purposes (and investors therein);
- “controlled foreign corporations” within the meaning of Section 957(a) of the Code;
- “passive foreign investment companies” within the meaning of Section 1297(a) of the Code;
- corporations that accumulate earnings to avoid U.S. federal income tax;
- banks or other financial institutions;
- real estate investment trusts or regulated investment companies;
- insurance companies;
- brokers or dealers in securities;
- persons that have elected to mark securities to market;
- tax-exempt organizations (including private foundations), governmental organizations, or international organizations;
- tax-qualified retirement plans;
- persons that acquired our common stock through the exercise of employee stock options or otherwise as compensation;

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- persons that acquired our common stock through the exercise of warrants or conversion rights under convertible instruments;
- persons that hold our common stock as “qualified small business stock” under Section 1202 of the Code or “Section 1244 stock” under Section 1244 of the Code;
- persons that acquired our common stock in a transaction subject to the gain rollover provisions of the Code (including Section 1045 of the Code);
- persons that own, or have owned, actually or constructively, more than 5% of our common stock;
- “qualified foreign pension funds” within the meaning of Section 897(l)(2) of the Code and entities all of the interests of which are held by qualified foreign pension funds; and
- persons that hold our common stock as part of a hedging or conversion transaction, straddle, a constructive sale, or any other risk reduction strategy or integrated investment.

If a partnership (or an entity or arrangement that is treated as a partnership for U.S. federal income tax purposes) holds our common stock, the U.S. federal income tax treatment of a partner in such partnership will generally depend on the status of the partner and the activities of the partnership. Partnerships that hold our common stock and the partners in such partnerships are urged to consult their tax advisors about the particular U.S. federal income tax consequences to them of holding and disposing of our common stock.

**THIS DISCUSSION IS FOR INFORMATIONAL PURPOSES ONLY AND IS NOT TAX ADVICE. PROSPECTIVE INVESTORS SHOULD CONSULT THEIR TAX ADVISORS REGARDING THE PARTICULAR U.S. FEDERAL INCOME TAX CONSEQUENCES TO THEM OF ACQUIRING, OWNING, AND DISPOSING OF OUR COMMON STOCK, AS WELL AS ANY TAX CONSEQUENCES ARISING UNDER ANY STATE, LOCAL, OR NON-U.S. TAX LAWS AND ANY U.S. FEDERAL NON-INCOME TAX LAWS, OR UNDER ANY APPLICABLE INCOME TAX TREATY.**

### **Definition of non-U.S. holder**

For purposes of this discussion, a non-U.S. holder is any beneficial owner of our common stock that is not a “United States person” or a partnership (including any entity or arrangement treated as a partnership) for U.S. federal income tax purposes. A U.S. person is any person that, for U.S. federal income tax purposes, is or is treated as any of the following:

- an individual who is a citizen or resident of the United States;
- a corporation (or any entity treated as a corporation for U.S. federal income tax purposes) created or organized under the laws of the United States, any state thereof, or the District of Columbia;
- an estate, the income of which is subject to U.S. federal income tax regardless of its source; or
- a trust (1) whose administration is subject to the primary supervision of a U.S. court and which has one or more “United States persons” within the meaning of Section 7701(a)(30) of the Code who have the authority to control all substantial decisions of the trust or (2) that has a valid election in effect under applicable Treasury Regulations to be treated as a United States person.

### **Distributions on our common stock**

As described in the section titled “Dividend Policy” above, we have not paid and do not anticipate paying dividends in the foreseeable future. However, if we make cash or other property distributions on our common

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stock, such distributions will constitute dividends for U.S. federal income tax purposes to the extent paid from our current or accumulated earnings and profits, as determined under U.S. federal income tax principles. Any portion of a distribution that exceeds our current and accumulated earnings and profits will constitute a return of capital and first be applied against and reduce the non-U.S. holder's tax basis in our common stock, but not below zero. Any amount distributed in excess of tax basis will be treated as gain realized on the sale or other disposition of our common stock and, therefore, treated as described in the section titled "—Gain on Disposition of Our Common Stock" below.

Subject to the discussions below regarding effectively connected income, backup withholding, and Sections 1471 through 1474 of the Code, commonly referred to as the Foreign Account Tax Compliance Act (FATCA), dividends paid to a non-U.S. holder generally will be subject to U.S. federal withholding tax at a rate of 30% (or such lower rate specified by an applicable income tax treaty) of the gross amount of the dividends. To receive the benefit of a lower treaty rate, a non-U.S. holder must furnish the applicable withholding agent with a valid IRS Form W-8BEN or IRS Form W-8BEN-E (or applicable successor form) certifying such non-U.S. holder's qualification for such rate. This certification must be provided to the applicable withholding agent before the payment of dividends and updated periodically. If the non-U.S. holder holds our common stock through a financial institution or other agent acting on the non-U.S. holder's behalf, the non-U.S. holder will be required to provide appropriate documentation to the financial institution or agent, which then will be required to provide certification to the applicable withholding agent, either directly or through other intermediaries.

Non-U.S. holders that do not provide the required certification on a timely basis, but that qualify for a reduced treaty rate, may obtain a refund of any excess amounts withheld by timely filing an appropriate claim for refund with the IRS.

If a non-U.S. holder holds our common stock in connection with the conduct of a trade or business in the United States, and dividends paid on our common stock are effectively connected with such non-U.S. holder's U.S. trade or business (and are attributable to such non-U.S. holder's permanent establishment in the United States, if required by an applicable tax treaty), the non-U.S. holder generally will be exempt from U.S. federal withholding tax. To claim the exemption, the non-U.S. holder must generally furnish a valid IRS Form W-8ECI (or applicable successor form) to the applicable withholding agent.

However, any such effectively connected dividends paid on our common stock generally will be subject to U.S. federal income tax on a net income basis at regular U.S. federal income tax rates in the same manner as if the non-U.S. holder were a resident of the United States. A non-U.S. holder that is a foreign corporation also may be subject to an additional branch profits tax equal to 30% (or such lower rate specified by an applicable income tax treaty) of its effectively connected earnings and profits for the taxable year, as adjusted for certain items. Non-U.S. holders are urged to consult their tax advisors about any applicable income tax treaties that may provide for different rules.

### **Gain on disposition of our common stock**

Subject to the discussions below regarding backup withholding and FATCA, a non-U.S. holder generally will not be subject to U.S. federal income tax on any gain realized on the sale or other disposition of our common stock, unless:

- the gain is effectively connected with the non-U.S. holder's conduct of a trade or business in the United States and, if required by an applicable income tax treaty, is attributable to the non-U.S. holder's permanent establishment in the United States;
- the non-U.S. holder is a nonresident alien individual present in the United States for 183 days or more during the taxable year of the sale or other disposition, and certain other requirements are met; or

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- our common stock constitutes a “United States real property interest” by reason of our status as a United States real property holding corporation, or USRPHC, for U.S. federal income tax purposes at any time within the shorter of the five-year period preceding the sale or other disposition or the non-U.S. holder’s holding period for our common stock, and our common stock is not regularly traded on an established securities market as defined by applicable Treasury Regulations.

Determining whether we are a USRPHC depends on the fair market value of our U.S. real property interests relative to the fair market value of our other trade or business assets and our foreign real property interests. We do not believe that we are, or have been, and do not anticipate becoming, a USRPHC for U.S. federal income tax purposes, although there can be no assurance that we will not in the future become a USRPHC. Even if we are or were to become a USRPHC, gain arising from the sale or other disposition by a non-U.S. holder of our common stock may not be subject to U.S. federal income tax if our common stock is “regularly traded,” as defined by applicable Treasury Regulations, on an established securities market.

Gain described in the first bullet point above generally will be subject to U.S. federal income tax on a net income basis at regular U.S. federal income tax rates in the same manner as if the non-U.S. holder were a resident of the United States. A non-U.S. holder that is a foreign corporation also may be subject to an additional branch profits tax equal to 30% (or such lower rate specified by an applicable income tax treaty) of its effectively connected earnings and profits for the taxable year, as adjusted for certain items. A non-U.S. holder described in the second bullet point above will be subject to U.S. federal income tax at a flat 30% rate (or such lower rate specified by an applicable income tax treaty) on gain realized upon the sale or other taxable disposition of our common stock which may be offset by certain U.S.-source capital losses (even though the individual is not considered a resident of the United States), provided that the non-U.S. holder has timely filed U.S. federal income tax returns with respect to such losses. If we are or become a USRPHC during the period described in the third bullet point above and our common stock is not regularly traded for purposes of the relevant rules, gain arising from the sale or other taxable disposition of our common stock by a non-U.S. holder will generally be subject to U.S. federal income tax in the same manner as gain that is effectively connected with the conduct of a U.S. trade or business, except that the branch profits tax generally will not apply. Non-U.S. holders are urged to consult their tax advisors about any applicable income tax treaties that may provide for different rules.

### **Information reporting and backup withholding**

Annual reports are required to be filed with the IRS and provided to each non-U.S. holder indicating the amount of distributions on our common stock paid to such non-U.S. holder and the amount of any tax withheld with respect to those distributions. These information reporting requirements apply even if no withholding was required (because the distributions were effectively connected with the non-U.S. holder’s conduct of a U.S. trade or business, or withholding was reduced or eliminated by an applicable income tax treaty) and regardless of whether such distributions constitute dividends. This information also may be made available under a specific treaty or agreement with the tax authorities in the country in which the non-U.S. holder resides or is established. Backup withholding, currently at a 24% rate, generally will not apply to payments to a non-U.S. holder of dividends on or the gross proceeds of a sale or other disposition of our common stock provided that the non-U.S. holder furnishes the required certification for its non-U.S. status, such as by providing a valid IRS Form W-8BEN, IRS Form W-8BEN-E, or IRS Form W-8ECI, or certain other requirements are met. Backup withholding may apply if the payor has actual knowledge, or reason to know, that the non-U.S. holder is a United States person who is not an exempt recipient.

Backup withholding is not an additional tax. If any amount is withheld under the backup withholding rules, non-U.S. holders are urged to consult their tax advisors about the possibility of and procedure for obtaining a refund or a credit against the non-U.S. holder’s U.S. federal income tax liability, if any.

## **Foreign account tax compliance act**

FATCA imposes a U.S. federal withholding tax of 30% on certain payments made to a “foreign financial institution” (as specially defined under the FATCA rules) unless such institution enters into an agreement with the U.S. government to withhold on certain payments and to collect and provide to the U.S. tax authorities substantial information regarding certain U.S. account holders of such institution (which includes certain equity and debt holders of such institution, as well as certain account holders that are foreign entities with U.S. owners) or an exemption applies. FATCA also generally will impose a U.S. federal withholding tax of 30% on certain payments made to a non-financial foreign entity unless such entity provides the withholding agent a certification identifying certain direct and indirect U.S. owners of the entity or an exemption applies. An intergovernmental agreement between the United States and an applicable non-U.S. country may modify these requirements. Under certain circumstances, a non-U.S. holder might be eligible for refunds or credits of such taxes. FATCA applies to dividends paid on our common stock and, subject to the proposed Treasury Regulations described below, also applies to gross proceeds from sales or other dispositions of our common stock. The U.S. Treasury Department released proposed Treasury Regulations which, if finalized in their present form, would eliminate the U.S. federal withholding tax of 30% applicable to the gross proceeds of a sale or other disposition of our common stock. In its preamble to such proposed Treasury Regulations, the U.S. Treasury Department stated that taxpayers may generally rely on the proposed Treasury Regulations until final regulations are issued.

Prospective investors are urged to consult with their tax advisors about the possible implications of FATCA on their investment in our common stock.

## Underwriting

We are offering the shares of common stock described in this prospectus through a number of underwriters. J.P. Morgan Securities LLC, SVB Securities LLC, Stifel, Nicolaus & Company, Incorporated and RBC Capital Markets, LLC are acting as joint book-running managers of the offering and as representatives of the underwriters. We have entered into an underwriting agreement with the underwriters. Subject to the terms and conditions of the underwriting agreement, we have agreed to sell to the underwriters, and each underwriter has severally agreed to purchase, at the public offering price less the underwriting discounts and commissions set forth on the cover page of this prospectus, the number of shares of common stock listed next to its name in the following table:

Name	Number of Shares
J.P. Morgan Securities LLC	
SVB Securities LLC	
Stifel, Nicolaus & Company, Incorporated	
RBC Capital Markets, LLC	
Chardan Capital Markets, LLC	
Total	

The underwriters are committed to purchase all the common stock offered by us if they purchase any shares. The underwriting agreement also provides that if an underwriter defaults, the purchase commitments of non-defaulting underwriters may also be increased or the offering may be terminated.

The underwriters propose to offer the common stock directly to the public at the initial public offering price set forth on the cover page of this prospectus and to certain dealers at that price less a concession not in excess of \$ \_\_\_\_\_ per share. Any such dealers may resell shares to certain other brokers or dealers at a discount of up to \$ \_\_\_\_\_ per share from the initial public offering price. After the initial offering of the shares to the public, if all of the common stock is not sold at the initial public offering price, the underwriters may change the offering price and the other selling terms. Sales of any shares made outside of the United States may be made by affiliates of the underwriters.

The underwriters have an option to buy up to \_\_\_\_\_ additional shares of common stock from us to cover sales of shares by the underwriters which exceed the number of shares specified in the table above. The underwriters have 30 days from the date of this prospectus to exercise this option to purchase additional shares. If any shares are purchased with this option to purchase additional shares, the underwriters will purchase shares in approximately the same proportion as shown in the table above. If any additional shares of common stock are purchased, the underwriters will offer the additional shares on the same terms as those on which the shares are being offered.

The underwriting fee is equal to the public offering price per share of common stock less the amount paid by the underwriters to us per share of common stock. The underwriting fee is \$ \_\_\_\_\_ per share. The following table shows the per share and total underwriting discounts and commissions to be paid to the underwriters assuming both no exercise and full exercise of the underwriters' option to purchase additional shares.

	Without Option to Purchase Additional Shares Exercise	With Full Option to Purchase Additional Shares Exercise
Per Share	\$	\$
Total	\$	\$

We estimate that the total expenses of this offering, including registration, filing and listing fees, printing fees and legal and accounting expenses, but excluding the underwriting discounts and commissions, will be approximately \$ \_\_\_\_\_.

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A prospectus in electronic format may be made available on the web sites maintained by one or more underwriters, or selling group members, if any, participating in the offering. The underwriters may agree to allocate a number of shares to underwriters and selling group members for sale to their online brokerage account holders. Internet distributions will be allocated by the representatives to underwriters and selling group members that may make Internet distributions on the same basis as other allocations.

We have agreed that we will not (i) offer, pledge, sell, contract to sell, sell any option or contract to purchase, purchase any option or contract to sell, grant any option, right or warrant to purchase, lend or otherwise transfer or dispose of, directly or indirectly, or submit to, or file with, the Securities and Exchange Commission a registration statement under the Securities Act relating to, any shares of our common stock or securities convertible into or exercisable or exchangeable for any shares of our common stock, or publicly disclose the intention to make any offer, sale, pledge, loan, disposition or filing, or (ii) enter into any swap or other arrangement that transfers all or a portion of the economic consequences associated with the ownership of any shares of common stock or any such other securities (regardless of whether any of these transactions are to be settled by the delivery of shares of common stock or such other securities, in cash or otherwise), in each case without the prior written consent of J.P. Morgan Securities LLC and SVB Securities LLC for a period of 180 days after the date of this prospectus, other than the shares of our common stock to be sold in this offering.

The restrictions on our actions, as described above, do not apply to: (i) the issuance of shares of common stock or securities convertible into or exercisable for shares of our common stock pursuant to the conversion or exchange of convertible or exchangeable securities or the exercise of warrants or options (including net exercise) or the settlement of RSUs (including net settlement), in each case outstanding on the date of the underwriting agreement and described in this prospectus; (ii) grants of stock options, stock awards, restricted stock, RSUs, or other equity awards and the issuance of shares of our common stock or securities convertible into or exercisable or exchangeable for shares of our common stock (whether upon the exercise of stock options or otherwise) to our employees, officers, directors, advisors, or consultants pursuant to the terms of an equity compensation plan in effect as of the closing of this offering and described in this prospectus, provided that such recipients enter into a lock-up agreement with the underwriters; (iii) the issuance of up to % of the outstanding shares of our common stock, or securities convertible into, exercisable for, or which are otherwise exchangeable for, our common stock, immediately following the closing of this offering, in acquisitions or other similar strategic transactions, provided that such recipients enter into a lock-up agreement with the underwriters; or (iv) our filing of any registration statement on Form S-8 relating to securities granted or to be granted pursuant to any plan in effect on the date of the underwriting agreement and described in this prospectus or any assumed benefit plan pursuant to an acquisition or similar strategic transaction.

Our directors and executive officers, and substantially all of our shareholders (such persons, the “lock-up parties”) have entered into lock up agreements with the underwriters prior to the commencement of this offering pursuant to which each lock-up party, with limited exceptions, for a period of 180 days after the date of this prospectus (such period, the “restricted period”), may not (and may not cause any of their direct or indirect affiliates to), without the prior written consent of J.P. Morgan Securities LLC and SVB Securities LLC, (1) offer, pledge, sell, contract to sell, sell any option or contract to purchase, purchase any option or contract to sell, grant any option, right or warrant to purchase, lend or otherwise transfer or dispose of, directly or indirectly, any shares of our common stock or any securities convertible into or exercisable or exchangeable for our common stock (including, without limitation, common stock or such other securities which may be deemed to be beneficially owned by such lock-up parties in accordance with the rules and regulations of the SEC and securities which may be issued upon exercise of a stock option or warrant (collectively with the common stock, the “lock-up securities”)), (2) enter into any hedging, swap or other agreement or transaction that transfers, in whole or in part, any of the economic consequences of ownership of the lock-up securities, whether any such transaction described in clause (1) or (2) above is to be settled by delivery of lock-up securities, in cash or otherwise, (3) make any

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demand for, or exercise any right with respect to, the registration of any lock-up securities, or (4) publicly disclose the intention to do any of the foregoing. Such persons or entities have further acknowledged that these undertakings preclude them from engaging in any hedging or other transactions or arrangements (including, without limitation, any short sale or the purchase or sale of, or entry into, any put or call option, or combination thereof, forward, swap or any other derivative transaction or instrument, however described or defined) designed or intended, or which could reasonably be expected to lead to or result in, a sale or disposition or transfer (by any person or entity, whether or not a signatory to such agreement) of any economic consequences of ownership, in whole or in part, directly or indirectly, of any lock-up securities, whether any such transaction or arrangement (or instrument provided for thereunder) would be settled by delivery of lock-up securities, in cash or otherwise.

The restrictions described in the immediately preceding paragraph and contained in the lock-up agreements between the underwriters and the lock-up parties do not apply, subject in certain cases to various conditions, to certain transactions, including (a) transfers of lock-up securities: (i) as bona fide gifts, or for bona fide estate planning purposes; (ii) by will or intestacy; (iii) to any trust for the direct or indirect benefit of the lock-up party or any immediate family member, or if the lock-up party is a trust, to a trustor, trustee (or co-trustee) or beneficiary of the trust or to the estate of a beneficiary of the trust; (iv) to a partnership, limited liability company or other entity of which the lock-up party and its immediate family members are the legal and beneficial owner of all of the outstanding equity securities or similar interests; (v) to a nominee or custodian of a person or entity to whom a disposition or transfer would be permissible under clauses (i) through (iv); (vi) in the case of a corporation, partnership, limited liability company, trust or other business entity, (A) to another corporation, partnership, limited liability company, trust or other business entity that is an affiliate of the lock-up party, or to any investment fund or other entity controlling, controlled by, managing or managed by or under common control with the lock-up party or its affiliates or (B) as part of a distribution, transfer or disposition to stockholders, partners, members, beneficiaries or other equity holders of the lock-up party; (vii) by operation of law; (viii) to us from an employee upon death, disability or termination of employment of such employee; (ix) as part of a sale of lock-up securities acquired in this offering or in open market transactions after the completion of this offering; (x) to us in connection with the vesting, settlement or exercise of restricted stock units, options, warrants or other rights to purchase shares of our common stock (including “net” or “cashless” exercise), including for the payment of exercise price and tax and remittance payments; (xi) pursuant to a bona fide third-party tender offer, merger, consolidation or other similar transaction approved by our board of directors and made to all shareholders involving a change in control, provided that if such transaction is not completed, all such lock-up securities would remain subject to the restrictions in the immediately preceding paragraph; or (xii) to an immediate family member of the lock-up party; (b) exercise of the options, settlement of RSUs or other equity awards, or the exercise of warrants granted pursuant to plans described in this prospectus, provided that any lock-up securities received upon such exercise, vesting or settlement would be subject to restrictions similar to those in the immediately preceding paragraph; (c) the conversion of outstanding preferred stock, warrants to acquire preferred stock, or convertible securities into shares of our common stock or warrants to acquire shares of our common stock, provided that any common stock or warrant received upon such conversion would be subject to restrictions similar to those in the immediately preceding paragraph; and (d) the establishment by lock-up parties of trading plans under Rule 10b5-1 under the Exchange Act, provided that such plan does not provide for the transfer of lock-up securities during the restricted period.

J.P. Morgan Securities LLC and SVB Securities LLC, in their sole discretion, may release the securities subject to any of the lock-up agreements with the underwriters described above, in whole or in part at any time.

We have agreed to indemnify the underwriters against certain liabilities, including liabilities under the Securities Act.

We intend to apply to have our common stock approved for listing on the Nasdaq Global Market under the symbol “LXEO”.



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In connection with this offering, the underwriters may engage in stabilizing transactions, which involves making bids for, purchasing and selling shares of common stock in the open market for the purpose of preventing or retarding a decline in the market price of the common stock while this offering is in progress. These stabilizing transactions may include making short sales of common stock, which involves the sale by the underwriters of a greater number of shares of common stock than they are required to purchase in this offering, and purchasing shares of common stock on the open market to cover positions created by short sales. Short sales may be “covered” shorts, which are short positions in an amount not greater than the underwriters’ option to purchase additional shares referred to above, or may be “naked” shorts, which are short positions in excess of that amount. The underwriters may close out any covered short position either by exercising their option to purchase additional shares, in whole or in part, or by purchasing shares in the open market. In making this determination, the underwriters will consider, among other things, the price of shares available for purchase in the open market compared to the price at which the underwriters may purchase shares through the option to purchase additional shares. A naked short position is more likely to be created if the underwriters are concerned that there may be downward pressure on the price of the common stock in the open market that could adversely affect investors who purchase in this offering. To the extent that the underwriters create a naked short position, they will purchase shares in the open market to cover the position.

The underwriters have advised us that, pursuant to Regulation M of the Securities Act, they may also engage in other activities that stabilize, maintain or otherwise affect the price of the common stock, including the imposition of penalty bids. This means that if the representatives of the underwriters purchase common stock in the open market in stabilizing transactions or to cover short sales, the representatives can require the underwriters that sold those shares as part of this offering to repay the underwriting discount received by them.

These activities may have the effect of raising or maintaining the market price of the common stock or preventing or retarding a decline in the market price of the common stock, and, as a result, the price of the common stock may be higher than the price that otherwise might exist in the open market. If the underwriters commence these activities, they may discontinue them at any time. The underwriters may carry out these transactions on the Nasdaq Stock Market, in the over-the-counter market or otherwise.

In addition, in connection with this offering certain of the underwriters (and selling group members) may engage in passive market making transactions in our common stock on the Nasdaq Stock Market prior to the pricing and completion of this offering. Passive market making consists of displaying bids on the Nasdaq Stock Market no higher than the bid prices of independent market makers and making purchases at prices no higher than these independent bids and effected in response to order flow. Net purchases by a passive market maker on each day are generally limited to a specified percentage of the passive market maker’s average daily trading volume in the common stock during a specified period and must be discontinued when such limit is reached. Passive market making may cause the price of our common stock to be higher than the price that otherwise would exist in the open market in the absence of these transactions. If passive market making is commenced, it may be discontinued at any time.

Prior to this offering, there has been no public market for our common stock. The initial public offering price will be determined by negotiations between us and the representatives of the underwriters. In determining the initial public offering price, we and the representatives of the underwriters expect to consider a number of factors including:

- the information set forth in this prospectus and otherwise available to the representatives;
- our prospects and the history and prospects for the industry in which we compete;
- an assessment of our management;

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- our prospects for future earnings;
- the general condition of the securities markets at the time of this offering;
- the recent market prices of, and demand for, publicly traded common stock of generally comparable companies; and
- other factors deemed relevant by the underwriters and us.

Neither we nor the underwriters can assure investors that an active trading market will develop for our common stock, or that the shares will trade in the public market at or above the initial public offering price.

Other than in the United States, no action has been taken by us or the underwriters that would permit a public offering of the securities offered by this prospectus in any jurisdiction where action for that purpose is required. The securities offered by this prospectus may not be offered or sold, directly or indirectly, nor may this prospectus or any other offering material or advertisements in connection with the offer and sale of any such securities be distributed or published in any jurisdiction, except under circumstances that will result in compliance with the applicable rules and regulations of that jurisdiction. Persons into whose possession this prospectus comes are advised to inform themselves about and to observe any restrictions relating to the offering and the distribution of this prospectus. This prospectus does not constitute an offer to sell or a solicitation of an offer to buy any securities offered by this prospectus in any jurisdiction in which such an offer or a solicitation is unlawful.

### **Notice to prospective investors in the European Economic Area and the United Kingdom**

In relation to each Member State of the European Economic Area and the United Kingdom, each a Relevant State, no shares of our common stock have been offered or will be offered pursuant to the offering to the public in that Relevant State prior to the publication of a prospectus in relation to the shares of our common stock which has been approved by the competent authority in that Relevant State or, where appropriate, approved in another Relevant State and notified to the competent authority in that Relevant State, all in accordance with the Prospectus Regulation, except that offers of Shares may be made to the public in that Relevant State other than at any time under the following exemptions under the Prospectus Regulation:

- a) to any legal entity which is a qualified investor as defined under the Prospectus Regulation;
- b) to fewer than 150 natural or legal persons (other than qualified investors as defined under the Prospectus Regulation), subject to obtaining the prior consent of the underwriters for any such offer; or
- c) in any other circumstances falling within Article 1(4) of the Prospectus Regulation,

provided that no such offer of shares of common stock shall require us or any underwriter to publish a prospectus pursuant to Article 3 the Prospectus Regulation or supplement a prospectus pursuant to Article 23 of the Prospectus Regulation and each person who initially acquires any shares of common stock or to whom any offer is made will be deemed to have represented, acknowledged and agreed to and with each of the underwriters and us that it is a “qualified investor” implementing Article 2(e) of the Prospectus Regulation.

In the case of any shares of common stock being offered to a financial intermediary as that term is used in the Prospectus Regulation, each such financial intermediary will be deemed to have represented, acknowledged and agreed that the shares of common stock acquired by it in the offer have not been acquired on a non-discretionary basis on behalf of, nor have they been acquired with a view to their offer or resale to, persons in circumstances which may give rise to an offer of any shares of common stock to the public other than their offer or resale in a Relevant State to qualified investors as so defined or in circumstances in which the prior consent of the underwriters has been obtained to each such proposed offer or resale.

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For the purposes of this provision, the expression an “offer to the public” in relation to any shares of common stock in any Relevant State means the communication in any form and by any means of sufficient information on the terms of the offer and any Shares to be offered so as to enable an investor to decide to purchase or subscribe for any Shares, and the expression “Prospectus Regulation” means Regulation (EU) 2017/1129. References to the Prospectus Regulation includes, in relation to the United Kingdom, the Prospectus Regulation as it forms part of United Kingdom domestic law by virtue of the European Union (Withdrawal) Act 2018.

### **Notice to prospective investors in the United Kingdom**

In addition, in the United Kingdom, this document is being distributed only to, and is directed only at, and any offer subsequently made may only be directed at persons who are “qualified investors” (as defined in the Prospectus Regulation) (i) who have professional experience in matters relating to investments falling within Article 19(5) of the Financial Services and Markets Act 2000 (Financial Promotion) Order 2005, as amended, or the Order, and/or (ii) who are high net worth companies (or persons to whom it may otherwise be lawfully communicated) falling within Article 49(2)(a) to (d) of the Order (all such persons together being referred to as “relevant persons”) or otherwise in circumstances which have not resulted and will not result in an offer to the public of the Shares in the United Kingdom within the meaning of the Financial Services and Markets Act 2000.

Any person in the United Kingdom that is not a relevant person should not act or rely on the information included in this document or use it as basis for taking any action. In the United Kingdom, any investment or investment activity that this document relates to may be made or taken exclusively by relevant persons.

### **Notice to prospective investors in Canada**

The shares of common stock may be sold only to purchasers purchasing, or deemed to be purchasing, as principal that are accredited investors, as defined in National Instrument 45-106 Prospectus Exemptions or subsection 73.3(1) of the Securities Act (Ontario), and are permitted clients, as defined in National Instrument 31-103 Registration Requirements, Exemptions and Ongoing Registrant Obligations. Any resale of the shares of common stock must be made in accordance with an exemption from, or in a transaction not subject to, the prospectus requirements of applicable securities laws.

Securities legislation in certain provinces or territories of Canada may provide a purchaser with remedies for rescission or damages if this prospectus (including any amendment thereto) contains a misrepresentation, provided that the remedies for rescission or damages are exercised by the purchaser within the time limit prescribed by the securities legislation of the purchaser’s province or territory. The purchaser should refer to any applicable provisions of the securities legislation of the purchaser’s province or territory for particulars of these rights or consult with a legal advisor.

Pursuant to section 3A.3 of National Instrument 33-105 Underwriting Conflicts, or NI 33-105, the underwriters are not required to comply with the disclosure requirements of NI 33-105 regarding underwriter conflicts of interest in connection with this offering.

### **Notice to prospective investors in Switzerland**

The shares of common stock may not be publicly offered in Switzerland and will not be listed on the SIX Swiss Exchange, or the SIX, or on any other stock exchange or regulated trading facility in Switzerland. This document does not constitute a prospectus within the meaning of, and has been prepared without regard to the disclosure standards for issuance prospectuses under art. 652a or art. 1156 of the Swiss Code of Obligations or the disclosure standards for listing prospectuses under art. 27 ff. of the SIX Listing Rules or the listing rules of any other stock exchange or regulated trading facility in Switzerland. Neither this document nor any other offering or

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marketing material relating to the shares of common stock or the offering may be publicly distributed or otherwise made publicly available in Switzerland.

Neither this document nor any other offering or marketing material relating to the offering, us, or the shares of common stock have been or will be filed with or approved by any Swiss regulatory authority. In particular, this document will not be filed with, and the offer of shares of common stock will not be supervised by, the Swiss Financial Market Supervisory Authority, or FINMA, and the offer of shares of common stock has not been and will not be authorized under the Swiss Federal Act on Collective Investment Schemes, or CISA. The investor protection afforded to acquirers of interests in collective investment schemes under the CISA does not extend to acquirers of shares of common stock.

### **Notice to prospective investors in Hong Kong**

The shares of common stock have not been offered or sold, and will not be offered or sold, in Hong Kong, by means of any document, other than (a) to “professional investors” as defined in the Securities and Futures Ordinance (Cap. 571 of the laws of Hong Kong), or the SFO, and any rules made thereunder; or (b) in other circumstances which do not result in the document being a “prospectus” as defined in the Companies (Winding Up and Miscellaneous Provisions) Ordinance (Cap. 32) of Hong Kong, or the CO, or which do not constitute an offer to the public within the meaning of the CO. No advertisement, invitation or document relating to the shares of common stock has been or may be issued or has been or may be in the possession of any person for the purposes of issue, whether in Hong Kong or elsewhere, which is directed at, or the contents of which are likely to be accessed or read by, the public of Hong Kong (except if permitted to do so under the securities laws of Hong Kong) other than with respect to shares of common stock which are or are intended to be disposed of only to persons outside Hong Kong or only to “professional investors” as defined in the SFO and any rules made thereunder.

### **Notice to prospective investors in Singapore**

Each representative has acknowledged that this prospectus has not been registered as a prospectus with the Monetary Authority of Singapore. Accordingly, each representative has represented and agreed that it has not offered or sold any shares of common stock or caused the shares of common stock to be made the subject of an invitation for subscription or purchase and will not offer or sell any shares of common stock or cause the shares of common stock to be made the subject of an invitation for subscription or purchase, and has not circulated or distributed, nor will it circulate or distribute, this prospectus or any other document or material in connection with the offer or sale, or invitation for subscription or purchase, of the shares of common stock, whether directly or indirectly, to any person in Singapore other than:

- (a) to an institutional investor (as defined in Section 4A of the Securities and Futures Act (Chapter 289) of Singapore, as modified or amended from time to time, or the SFA) pursuant to Section 274 of the SFA;
- (b) to a relevant person (as defined in Section 275(2) of the SFA) pursuant to Section 275(1) of the SFA, or any person pursuant to Section 275(1A) of the SFA, and in accordance with the conditions specified in Section 275 of the SFA; or
- (c) otherwise pursuant to, and in accordance with the conditions of, any other applicable provision of the SFA.

Where the shares of common stock are subscribed or purchased under Section 275 of the SFA by a relevant person which is:

- (a) a corporation (which is not an accredited investor (as defined in Section 4A of the SFA)) the sole business of which is to hold investments and the entire share capital of which is owned by one or more individuals, each of whom is an accredited investor; or

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- (b) a trust (where the trustee is not an accredited investor) whose sole purpose is to hold investments and each beneficiary of the trust is an individual who is an accredited investor, securities or securities-based derivatives contracts (each term as defined in Section 2(1) of the SFA) of that corporation or the beneficiaries' rights and interest (howsoever described) in that trust shall not be transferred within six months after that corporation or that trust has acquired the shares of common stock pursuant to an offer made under Section 275 of the SFA except:
- (i) to an institutional investor or to a relevant person, or to any person arising from an offer referred to in Section 275(1A) or Section 276(4)(i) (B) of the SFA;
  - (ii) where no consideration is or will be given for the transfer;
  - (iii) where the transfer is by operation of law;
  - (iv) as specified in Section 276(7) of the SFA; or
  - (v) as specified in Regulation 37A of the Securities and Futures (Offers of Investments) (Securities and Securities-based Derivatives Contracts) Regulations 2018.

Singapore SFA Product Classification—In connection with Section 309B of the SFA and the CMP Regulations 2018, unless otherwise specified before an offer of the shares of common stock, we have determined, and hereby notify all relevant persons (as defined in Section 309A(1) of the SFA), that the shares of common stock are “prescribed capital markets products” (as defined in the CMP Regulations 2018) and Excluded Investment Products (as defined in MAS Notice SFA 04-N12: Notice on the Sale of Investment Products and MAS Notice FAA-N16: Notice on Recommendations on Investment Products).

### **Notice to prospective investors in Japan**

The shares of common stock have not been and will not be registered pursuant to Article 4, Paragraph 1 of the Financial Instruments and Exchange Act. Accordingly, none of the shares of common stock nor any interest therein may be offered or sold, directly or indirectly, in Japan or to, or for the benefit of, any “resident” of Japan (which term as used herein means any person resident in Japan, including any corporation or other entity organized under the laws of Japan), or to others for re-offering or resale, directly or indirectly, in Japan or to or for the benefit of a resident of Japan, except pursuant to an exemption from the registration requirements of, and otherwise in compliance with, the Financial Instruments and Exchange Act and any other applicable laws, regulations and ministerial guidelines of Japan in effect at the relevant time.

### **Notice to prospective investors in the United Arab Emirates**

The shares of common stock have not been, and are not being, publicly offered, sold, promoted or advertised in the United Arab Emirates (including the Dubai International Financial Centre, or the DIFC) other than in compliance with the laws of the United Arab Emirates (and the DIFC) governing the issue, offering and sale of securities. Further, this prospectus does not constitute a public offer of securities in the United Arab Emirates (including the DIFC) and is not intended to be a public offer. This prospectus has not been approved by or filed with the Central Bank of the United Arab Emirates, the Securities and Commodities Authority or the Dubai Financial Services Authority, or the DFSA.

### **Notice to prospective investors in Israel**

In the State of Israel this prospectus shall not be regarded as an offer to the public to purchase shares of common stock under the Israeli Securities Law, 5728—1968, which requires a prospectus to be published and authorized

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by the Israel Securities Authority, if it complies with certain provisions of Section 15 of the Israeli Securities Law, 5728–1968, including, inter alia, if: (i) the offer is made, distributed or directed to not more than 35 investors, subject to certain conditions, or the Addressed Investors; or (ii) the offer is made, distributed or directed to certain qualified investors defined in the First Addendum of the Israeli Securities Law, 5728–1968, subject to certain conditions, or the Qualified Investors. The Qualified Investors shall not be taken into account in the count of the Addressed Investors and may be offered to purchase securities in addition to the 35 Addressed Investors. We have not and will not take any action that would require it to publish a prospectus in accordance with and subject to the Israeli Securities Law, 5728–1968. We have not and will not distribute this prospectus or make, distribute or direct an offer to subscribe for our shares of common stock to any person within the State of Israel, other than to Qualified Investors and up to 35 Addressed Investors.

Qualified Investors may have to submit written evidence that they meet the definitions set out in of the First Addendum to the Israeli Securities Law, 5728–1968. In particular, we may request, as a condition to be offered shares of common stock, that Qualified Investors will each represent, warrant and certify to us and/or to anyone acting on our behalf: (i) that it is an investor falling within one of the categories listed in the First Addendum to the Israeli Securities Law, 5728–1968; (ii) which of the categories listed in the First Addendum to the Israeli Securities Law, 5728–1968 regarding Qualified Investors is applicable to it; (iii) that it will abide by all provisions set forth in the Israeli Securities Law, 5728–1968 and the regulations promulgated thereunder in connection with the offer to be issued shares of common stock; (iv) that the shares of common stock that it will be issued are, subject to exemptions available under the Israeli Securities Law, 5728–1968: (a) for its own account; (b) for investment purposes only; and (c) not issued with a view to resale within the State of Israel, other than in accordance with the provisions of the Israeli Securities Law, 5728–1968; and (v) that it is willing to provide further evidence of its Qualified Investor status. Addressed Investors may have to submit written evidence in respect of their identity and may have to sign and submit a declaration containing, inter alia, the Addressed Investor’s name, address and passport number or Israeli identification number.

### **Notice to prospective investors in Australia**

This prospectus:

- (a) does not constitute a disclosure document or a prospectus under Chapter 6D.2 of the Corporations Act 2001 (Cth), or the Corporations Act;
- (b) has not been, and will not be, lodged with the Australian Securities and Investments Commission, or ASIC, as a disclosure document for the purposes of the Corporations Act and does not purport to include the information required of a disclosure document for the purposes of the Corporations Act; and
- (c) may only be provided in Australia to select investors who are able to demonstrate that they fall within one or more of the categories of investors, or Exempt Investors, available under section 708 of the Corporations Act.

The shares of common stock may not be directly or indirectly offered for subscription or purchased or sold, and no invitations to subscribe for or buy the shares of common stock may be issued, and no draft or definitive offering memorandum, advertisement or other offering material relating to any shares of common stock may be distributed in Australia, except where disclosure to investors is not required under Chapter 6D of the Corporations Act or is otherwise in compliance with all applicable Australian laws and regulations. By submitting an application for the shares of common stock, you represent and warrant to us that you are an Exempt Investor.

As any offer of shares of common stock under this document will be made without disclosure in Australia under Chapter 6D.2 of the Corporations Act, the offer of those securities for resale in Australia within 12 months may,

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under section 707 of the Corporations Act, require disclosure to investors under Chapter 6D.2 if none of the exemptions in section 708 applies to that resale. By applying for the shares of common stock you undertake to us that you will not, for a period of 12 months from the date of issue of the shares of common stock, offer, transfer, assign or otherwise alienate those shares of common stock to investors in Australia except in circumstances where disclosure to investors is not required under Chapter 6D.2 of the Corporations Act or where a compliant disclosure document is prepared and lodged with ASIC.

### **Notice to prospective investors in China**

This prospectus will not be circulated or distributed in the PRC and the shares of common stock will not be offered or sold, and will not be offered or sold to any person for re-offering or resale directly or indirectly to any residents of the PRC except pursuant to any applicable laws and regulations of the PRC. Neither this prospectus nor any advertisement or other offering material may be distributed or published in the PRC, except under circumstances that will result in compliance with applicable laws and regulations.

### **Notice to prospective investors in Korea**

The shares of common stock have not been and will not be registered under the Financial Investments Services and Capital Markets Act of Korea and the decrees and regulations thereunder, or the FSCMA, and the shares of common stock have been and will be offered in Korea as a private placement under the FSCMA. None of the shares of common stock may be offered, sold or delivered directly or indirectly, or offered or sold to any person for re-offering or resale, directly or indirectly, in Korea or to any resident of Korea except pursuant to the applicable laws and regulations of Korea, including the FSCMA and the Foreign Exchange Transaction Law of Korea and the decrees and regulations thereunder, or FETL. The shares have not been listed on any of the securities exchanges in the world including, without limitation, the Korea Exchange in Korea. Furthermore, the purchaser of the shares of common stock shall comply with all applicable regulatory requirements (including but not limited to requirements under the FETL) in connection with the purchase of the shares of common stock. By the purchase of the shares of common stock, the relevant holder thereof will be deemed to represent and warrant that if it is in Korea or is a resident of Korea, it purchased the shares of common stock pursuant to the applicable laws and regulations of Korea.

### **Notice to prospective investors in Saudi Arabia**

This document may not be distributed in the Kingdom of Saudi Arabia except to such persons as are permitted under the Offers of Securities Regulations as issued by the board of the Saudi Arabian Capital Market Authority, or CMA, pursuant to resolution number 2-11-2004 dated 4 October 2004 as amended by resolution number 1-28-2008, as amended, or the CMA Regulations. The CMA does not make any representation as to the accuracy or completeness of this document and expressly disclaims any liability whatsoever for any loss arising from, or incurred in reliance upon, any part of this document. Prospective purchasers of the securities offered hereby should conduct their own due diligence on the accuracy of the information relating to the securities. If you do not understand the contents of this document, you should consult an authorized financial adviser.

### **Notice to prospective investors in the Dubai International Financial Centre**

This document relates to an Exempt Offer in accordance with the Markets Rules 2012 of the DFSA. This document is intended for distribution only to persons of a type specified in the Offered Securities Rules. It must not be delivered to, or relied on by, any other person. The DFSA has no responsibility for reviewing or verifying any documents in connection with Exempt Offers. The DFSA has not approved this prospectus supplement nor taken steps to verify the information set forth herein and has no responsibility for this document. The securities to

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which this document relates may be illiquid and/or subject to restrictions on their resale. Prospective purchasers of the securities offered should conduct their own due diligence on the securities. If you do not understand the contents of this document, you should consult an authorized financial advisor.

In relation to its use in the DIFC, this document is strictly private and confidential and is being distributed to a limited number of investors and must not be provided to any person other than the original recipient, and may not be reproduced or used for any other purpose. The interests in the securities may not be offered or sold directly or indirectly to the public in the DIFC.

### **Notice to prospective investors in Bermuda**

Shares of common stock may be offered or sold in Bermuda only in compliance with the provisions of the Investment Business Act of 2003 of Bermuda which regulates the sale of securities in Bermuda. Additionally, non-Bermudian persons (including companies) may not carry on or engage in any trade or business in Bermuda unless such persons are permitted to do so under applicable Bermuda legislation.

### **Notice to prospective investors in the British Virgin Islands**

The shares of common stock are not being, and may not be offered, to the public or to any person in the British Virgin Islands for purchase or subscription by or on behalf of the Company. The shares of common stock may be offered to companies incorporated under the BVI Business Companies Act, 2004 (British Virgin Islands), or the BVI Companies, but only where the offer will be made to, and received by, the relevant BVI Company entirely outside of the British Virgin Islands. This prospectus has not been, and will not be, registered with the Financial Services Commission of the British Virgin Islands. No registered prospectus has been or will be prepared in respect of the shares for the purposes of the Securities and Investment Business Act, 2010 or the Public Issuers Code of the British Virgin Islands.

### **Notice to prospective investors in Malaysia**

No prospectus or other offering material or document in connection with the offer and sale of the shares of common stock has been or will be registered with the Securities Commission of Malaysia, or the Commission, for the Commission's approval pursuant to the Capital Markets and Services Act 2007. Accordingly, this prospectus and any other document or material in connection with the offer or sale, or invitation for subscription or purchase, of the shares of common stock may not be circulated or distributed, nor may the shares of common stock be offered or sold, or be made the subject of an invitation for subscription or purchase, whether directly or indirectly, to persons in Malaysia other than (i) a closed end fund approved by the Commission, (ii) a holder of a Capital Markets Services Licence, (iii) a person who acquires the shares of common stock, as principal, if the offer is on terms that the shares of common stock may only be acquired at a consideration of not less than RM250,000 (or its equivalent in foreign currencies) for each transaction, (iv) an individual whose total net personal assets or total net joint assets with his or her spouse exceeds RM3 million (or its equivalent in foreign currencies), excluding the value of the primary residence of the individual, (v) an individual who has a gross annual income exceeding RM300,000 (or its equivalent in foreign currencies) per annum in the preceding twelve months, (vi) an individual who, jointly with his or her spouse, has a gross annual income of RM400,000 (or its equivalent in foreign currencies), per annum in the preceding twelve months, (vii) a corporation with total net assets exceeding RM10 million (or its equivalent in a foreign currencies) based on the last audited accounts, (viii) a partnership with total net assets exceeding RM10 million (or its equivalent in foreign currencies), (ix) a bank licensee or insurance licensee as defined in the Labuan Financial Services and Securities Act 2010, (x) an Islamic bank licensee or takaful licensee as defined in the Labuan Financial Services and Securities Act 2010, and (xi) any other person as may be specified by the Commission; provided that, in the each of the preceding



categories (i) to (xi), the distribution of the shares of common stock is made by a holder of a Capital Markets Services Licence who carries on the business of dealing in securities. The distribution in Malaysia of this prospectus is subject to Malaysian laws. This prospectus does not constitute and may not be used for the purpose of public offering or an issue, offer for subscription or purchase, invitation to subscribe for or purchase any securities requiring the registration of a prospectus with the Commission under the Capital Markets and Services Act 2007.

### **Notice to prospective investors in Taiwan**

The shares of common stock have not been and will not be registered with the Financial Supervisory Commission of Taiwan pursuant to relevant securities laws and regulations and may not be sold, issued or offered within Taiwan through a public offering or in circumstances which constitutes an offer within the meaning of the Securities and Exchange Act of Taiwan that requires a registration or approval of the Financial Supervisory Commission of Taiwan. No person or entity in Taiwan has been authorized to offer, sell, give advice regarding or otherwise intermediate the offering and sale of the shares of common stock in Taiwan.

### **Notice to prospective investors in South Africa**

Due to restrictions under the securities laws of South Africa, no “offer to the public” (as such term is defined in the South African Companies Act, No. 71 of 2008 (as amended or re-enacted), or the South African Companies Act) is being made in connection with the issue of the shares of common stock in South Africa. Accordingly, this document does not, nor is it intended to, constitute a “registered prospectus” (as that term is defined in the South African Companies Act) prepared and registered under the South African Companies Act and has not been approved by, and/or filed with, the South African Companies and Intellectual Property Commission or any other regulatory authority in South Africa. The shares of common stock are not offered, and the offer shall not be transferred, sold, renounced or delivered, in South Africa or to a person with an address in South Africa, unless one or other of the following exemptions stipulated in Section 96(1) applies:

Section 96(1)(a): the offer, transfer, sale, renunciation or delivery is to:

- (i) persons whose ordinary business, or part of whose ordinary business, is to deal in securities, as principal or agent;
- (ii) the South African Public Investment Corporation;
- (iii) persons or entities regulated by the Reserve Bank of South Africa;
- (iv) authorised financial service providers under South African law;
- (v) financial institutions recognised as such under South African law;
- (vi) a wholly-owned subsidiary of any person or entity contemplated in (iii), (iv) or (v), acting as agent in the capacity of an authorized portfolio manager for a pension fund, or as manager for a collective investment scheme (in each case duly registered as such under South African law); or
- (vii) any combination of the persons in (i) to (vi); or

Section 96(1)(b): the total contemplated acquisition cost of the securities, for a single addressee acting as principal is equal to or greater than ZAR1,000,000 or such higher amount as may be promulgated by notice in the Government Gazette of South Africa pursuant to section 96(2)(a) of the South African Companies Act.

Information made available in this prospectus should not be considered as “advice” as defined in the South African Financial Advisory and Intermediary Services Act, 2002.

## **Other Relationships**

Certain of the underwriters and their affiliates have provided in the past to us and our affiliates and may provide from time to time in the future certain commercial banking, financial advisory, investment banking and other services for us and such affiliates in the ordinary course of their business, for which they have received and may continue to receive customary fees and commissions. In addition, from time to time, certain of the underwriters and their affiliates may effect transactions for their own account or the account of customers, and hold on behalf of themselves or their customers, long or short positions in our debt or equity securities or loans, and may do so in the future.

## Legal matters

The validity of the shares of common stock offered hereby will be passed upon for us by Cooley LLP, New York, New York. Certain legal matters will be passed upon for the underwriters by Goodwin Procter LLP, New York, New York.

## Experts

The financial statements of Lexeo Therapeutics, Inc. as of December 31, 2022 and 2021, and for the years then ended have been included herein in reliance upon the report of KPMG LLP, independent registered public accounting firm, appearing elsewhere herein, and upon the authority of said firm as experts in accounting and auditing.

## Where you can find additional information

We have filed with the SEC a registration statement on Form S-1 under the Securities Act with respect to the shares of common stock being offered by this prospectus. This prospectus, which constitutes part of the registration statement, does not contain all of the information in the registration statement and its exhibits. For further information with respect to our company and the common stock offered by this prospectus, we refer you to the registration statement and its exhibits. Statements contained in this prospectus as to the contents of any contract or any other document referred to are not necessarily complete, and in each instance, we refer you to the copy of the contract or other document filed as an exhibit to the registration statement. Each of these statements is qualified in all respects by this reference.

You can read our SEC filings, including the registration statement, over the internet at the SEC's website at [www.sec.gov](http://www.sec.gov). Upon completion of this offering, we will be subject to the information reporting requirements of the Exchange Act, and we will file reports, proxy statements and other information with the SEC. These reports, proxy statements and other information will be available at [www.sec.gov](http://www.sec.gov).

We also maintain a website at [www.lexeotx.com](http://www.lexeotx.com), at which you may access these materials free of charge as soon as reasonably practicable after they are electronically filed with, or furnished to, the SEC. The information contained in, or that can be accessed through, our website is not part of, and is not incorporated into, this prospectus, and the inclusion of our website address in this prospectus is only as an inactive textual reference.

# Lexeo Therapeutics, Inc.

## Unaudited condensed consolidated financial statements

For the period ended March 31, 2023

### Unaudited Financial Statements

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# Lexeo Therapeutics, Inc.

## Condensed Consolidated Balance Sheets

(Unaudited)

	March 31, 2023	December 31, 2022
<b>Assets</b>		
Current assets:		
Cash and cash equivalents	\$ 58,853,899	\$ 77,334,973
Prepaid expenses	1,818,123	2,342,550
Other current assets	550,215	370,541
Total current assets	61,222,237	80,048,064
Restricted cash	3,261,611	3,253,246
Property and equipment, net	952,746	998,350
Lease right-of-use assets - finance, net	1,999,809	2,068,633
Lease right-of-use assets - operating	10,387,470	10,708,144
Total assets	\$ 77,823,873	\$ 97,076,437
<b>Liabilities, Convertible Preferred Stock and Stockholders' Deficit</b>		
Current liabilities:		
Accounts payable	\$ 3,912,209	\$ 2,677,800
Taxes payable	448,697	505,681
Accrued expenses and other current liabilities	5,887,665	6,420,538
Accrued bonuses	691,274	1,911,369
Current portion of lease liabilities - finance	523,157	547,294
Current portion of lease liabilities - operating	2,230,981	2,287,109
Total current liabilities	13,693,983	14,349,791
Non-current liabilities		
Non-current portion of lease liabilities - finance	1,554,503	1,647,898
Non-current portion of lease liabilities - operating	8,723,572	8,999,643
Total liabilities	23,972,058	24,997,332
<b>Commitments and contingencies (Note 11)</b>		
Series A convertible preferred stock, \$0.0001 par value; 85,495,722 shares authorized, issued, and outstanding as of March 31, 2023 and December 31, 2022; aggregate liquidation value of \$85,495,722 as of March 31, 2023 and December 31, 2022	85,268,513	85,268,513
Series B convertible preferred stock, \$0.0001 par value; 58,157,824 shares authorized as of March 31, 2023 and December 31, 2022; 58,157,823 shares issued and outstanding as of March 31, 2023 and December 31, 2022; aggregate liquidation value of \$100,059,953 as of March 31, 2023 and December 31, 2022	99,764,986	99,764,986
<b>Stockholders' and members' deficit:</b>		
Common stock, \$0.0001 par value, 192,020,973 shares authorized as of March 31, 2023 and December 31, 2022; 17,482,991 shares issued and 17,099,158 shares outstanding as of March 31, 2023; 17,467,148 shares issued and 17,026,915 shares outstanding as of December 31, 2022	1,710	1,703
Additional paid-in capital	2,920,353	2,490,168
Accumulated deficit	(134,103,747)	(115,446,265)
Total stockholders' deficit	(131,181,684)	(112,954,394)
Total liabilities, convertible preferred stock, stockholders' deficit	\$ 77,823,873	\$ 97,076,437

The accompanying notes are an integral part of these unaudited condensed consolidated financial statements.

# Lexeo Therapeutics, Inc.

## Condensed Consolidated Statements of Operations and Comprehensive Loss

(Unaudited)

	Three Months Ended	
	2023	March 31, 2022
Revenue		
Grant revenue	\$ —	\$ 85,550
Total revenue	—	85,550
Operating expenses		
Research and development	16,438,110	10,146,371
General and administrative	2,852,379	2,458,438
Total operating expenses	19,290,489	12,604,809
Operating loss	(19,290,489)	(12,519,259)
Other income (expense)		
Other income (expense)	(4,521)	146
Interest expense	(49,967)	—
Interest income	687,495	17,283
Total other income (expense)	633,007	17,429
Loss from operations before income taxes	(18,657,482)	(12,501,830)
Income taxes	—	—
Net loss and comprehensive loss	\$ (18,657,482)	\$ (12,501,830)
Net loss per common share, basic and diluted	\$ (1.09)	\$ (0.74)
Weighted average number of shares outstanding used in computation of net loss per common share, basic and diluted	17,069,994	16,863,270

The accompanying notes are an integral part of these unaudited condensed consolidated financial statements.

# Lexeo Therapeutics, Inc.

## Condensed Consolidated Statements of Convertible Preferred Stock and Stockholders' Deficit

(Unaudited)

	Convertible Preferred Stock		Common Stock		Additional Paid-in Capital	Accumulated Deficit	Total Stockholders' Deficit
	Shares	Amount	Shares	Amount			
Balances at January 1, 2023	143,653,545	\$ 185,033,499	17,026,915	\$ 1,703	\$ 2,490,168	\$ (115,446,265)	\$ (112,954,394)
Exercise of stock options	—	—	15,843	1	12,137	—	12,138
Amounts reclassified from deposit liabilities upon the vesting of early-exercised stock options previously subject to repurchase, net of proceeds received from early exercise of unvested stock options subject to repurchase and recorded as deposit liabilities	—	—	56,400	6	20,793	—	20,799
Stock-based compensation expense	—	—	—	—	397,255	—	397,255
Net loss	—	—	—	—	—	(18,657,482)	(18,657,482)
Balances at March 31, 2023	143,653,545	\$ 185,033,499	17,099,158	\$ 1,710	\$ 2,920,353	\$ (134,103,747)	\$ (131,181,684)

	Convertible Preferred Stock		Common Stock		Additional Paid-in Capital	Accumulated Deficit	Total Stockholders' Deficit
	Shares	Amount	Shares	Amount			
Balances at January 1, 2022	143,653,545	\$ 185,012,970	16,745,827	\$ 1,675	\$ 566,290	\$ (56,169,538)	\$ (55,601,573)
Proceeds from issuance of series A convertible preferred stock	—	20,529	—	—	—	—	—
Exercise of stock options	—	—	511,866	51	235,407	—	235,457
Proceeds received from early exercise of unvested stock options subject to repurchase and recorded as deposit liabilities	—	—	(639,242)	(64)	(235,394)	—	(235,458)
Stock-based compensation expense	—	—	—	—	274,723	—	274,724
Net loss	—	—	—	—	—	(12,501,830)	(12,501,830)
Balances at March 31, 2022	143,653,545	\$ 185,033,499	16,618,451	\$ 1,662	\$ 841,026	\$ (68,671,368)	\$ (67,828,680)

The accompanying notes are an integral part of these unaudited condensed consolidated financial statements.

# Lexeo Therapeutics Inc.

## Condensed Consolidated Statements of Cash Flows

(Unaudited)

	Three Months Ended	
	2023	March 31, 2022
<b>Cash flows from operating activities:</b>		
Net loss	\$ (18,657,482)	\$ (12,501,830)
Adjustments to reconcile net loss to net cash (used in) provided by operating activities:		
Amortization of right-of-use asset, operating	320,674	—
Amortization of right-of-use assets, finance	68,824	—
Stock based compensation expense	397,255	274,723
Depreciation and amortization expense	57,438	6,702
Changes in operating assets and liabilities:		
Prepaid expenses and other current assets	524,444	(1,632,655)
Accounts payable	1,042,884	(90,898)
Change in deferred income	—	(46,460)
Accrued expenses and other current liabilities	(1,732,169)	(1,104,377)
Tax payable	(56,984)	—
Lease liability, operating	(332,199)	—
Lease liabilities, finance	(7,954)	—
Net cash used in operating activities	<u>(18,375,269)</u>	<u>(15,094,795)</u>
<b>Cash flows from investing activities:</b>		
Purchase of property and equipment	—	(173,625)
Net cash used in investing activities	<u>—</u>	<u>(173,625)</u>
<b>Cash flows from financing activities:</b>		
Proceeds from issuance of series A convertible preferred stock	—	20,529
Proceeds from exercise of stock options	12,138	235,458
Payments on finance leases	(109,578)	—
Net cash (used in) provided by financing activities	<u>(97,440)</u>	<u>255,987</u>
<b>Net change in cash, cash equivalents and restricted cash</b>	<b>(18,472,709)</b>	<b>(15,012,433)</b>
Total cash, cash equivalents and restricted cash at beginning of period	80,588,219	135,860,060
Total cash, cash equivalents and restricted cash at end of period	<u>\$ 62,115,510</u>	<u>\$ 120,847,627</u>
Cash and cash equivalents at beginning of period	77,334,973	135,860,060
Restricted cash included in long-term assets at beginning of period	3,253,246	—
Total cash, cash equivalents and restricted cash at beginning of period	<u>\$ 80,588,219</u>	<u>\$ 135,860,060</u>
Cash and cash equivalents at end of period	58,853,899	117,864,303
Restricted cash included in long-term assets at end of period	3,261,611	2,983,324
Total cash, cash equivalents and restricted cash at end of period	<u>\$ 62,115,510</u>	<u>\$ 120,847,627</u>
<b>Supplemental disclosure of non-cash investing and financing activities</b>		
Deferred offering costs included in accounts payable and accrued expenses	\$ 179,691	\$ —
Purchase of property and equipment included in accounts payable and accrued expenses	\$ 11,834	\$ —
Amounts reclassified from deposit liabilities upon the vesting of early-exercised stock options previously subject to repurchase, net of (proceeds received from early exercise of unvested stock options subject to repurchase and recorded as deposit liabilities)	\$ 20,799	\$ (235,458)

The accompanying notes are an integral part of these unaudited condensed consolidated financial statements.



**LEXEO THERAPEUTICS, INC.**  
**NOTES TO CONDENSED CONSOLIDATED FINANCIAL STATEMENTS (UNAUDITED)**

**1. DESCRIPTION OF BUSINESS AND BASIS OF PRESENTATION**

**Description of Business**—Lexeo Therapeutics Inc. (“Lexeo” or the “Company”), together with its consolidated subsidiary, Stelios Therapeutics Inc., until such subsidiary was merged into the Company on December 15, 2022 and ceased to exist, is a clinical-stage gene therapy company focused on addressing genetically defined cardiovascular diseases and a genetically defined sub-population of Alzheimer’s disease affecting both larger-rare and prevalent patient populations. The Company is advancing a pipeline of adeno-associated virus (“AAV”)–based gene therapy candidates utilizing its modular approach that integrates a clinically validated vector, a disease area strategy targeting defined patient sub-groups, and scalable manufacturing, which is designed to overcome many of the challenges facing the field of gene therapy.

The Company is located in New York, NY and was first formed on February 17, 2017, as an LLC under the laws of the State of Delaware under the legal name Lexeo Therapeutics, LLC. The Company filed and executed a certificate of conversion to corporation on November 20, 2020, to convert the LLC to Lexeo Therapeutics, Inc, a Delaware corporation. All of the Company’s tangible assets are held in the United States (“U.S.”).

**Basis of Presentation and Principles of Consolidation**—The Company’s fiscal year ends on December 31, and its fiscal quarters end on March 31, June 30, and September 30.

These unaudited condensed consolidated financial statements and accompanying notes reflect the operations of the Company and its wholly owned subsidiary that have been prepared in conformity with generally accepted accounting principles in the United States of America (“GAAP”) for interim financial information. Accordingly, they do not include all of the information and disclosures required by GAAP for complete financial statements. These unaudited condensed consolidated financial statements and accompanying notes should be read in conjunction with the Company’s audited consolidated financial statements and notes thereto for the year ended December 31, 2022 (the “Annual Financial Statements”). The unaudited condensed consolidated balance sheet at December 31, 2022 has been derived from the audited consolidated financial statements at that date. In the opinion of management, all adjustments (consisting of normal recurring adjustments) considered necessary to present fairly the financial position of the Company and its results of operations and cash flows for the periods presented have been included. Operating results for the three months ended March 31, 2023 are not necessarily indicative of the results that may be expected for the year ending December 31, 2023, for any other interim period, or for any other future year.

Any reference in these notes to applicable guidance is meant to refer to the authoritative GAAP as found in the Accounting Standards Codification (“ASC”) and Accounting Standards Updates (“ASU”) of the Financial Accounting Standards Board (“FASB”). The accounts of the wholly owned subsidiary are included in these unaudited condensed consolidated financial statements and accompanying notes. The Company has no unconsolidated subsidiaries. All intercompany balances and transactions have been eliminated in consolidation.

**Need for Additional Capital**—Since inception, the Company has incurred net losses and negative cash flows from operations, including net losses of \$18.7 million and \$12.5 million during the three months ended March 31, 2023 and March 31, 2022, respectively. As of March 31, 2023, the Company had an accumulated deficit of \$134.1 million and expects to incur substantial operating losses and negative cash flows from operations for the foreseeable future. The Company had \$58.9 million and \$77.3 million of cash and cash equivalents at March 31, 2023 and December 31, 2022, respectively. During the year ended December 31, 2021 the Company raised \$51.0 million and \$100.1 million of gross proceeds in connection with the issuance of its

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Series A convertible preferred stock and Series B convertible preferred stock (see Note 7), respectively, and which management believes is sufficient to fund its operating expenses and capital expenditure requirements for at least 12 months from the issuance date of these unaudited condensed consolidated financial statements. The future viability of the Company beyond that point is dependent on its ability to raise additional capital to finance operations. The Company is seeking to complete an initial public offering (“IPO”) of its common stock. In the event the Company does not complete an IPO, the Company expects to seek additional funding through private equity financings, debt financings or other capital sources, including collaborations with other companies or other strategic transactions. The Company may not be able to obtain financing on acceptable terms, or at all. The terms of any financing may adversely affect the holdings or the rights of the Company’s stockholders.

If the Company is unable to obtain funding, the Company will be forced to delay, reduce or eliminate some or all of its research and development programs which could adversely affect its business prospects, or the Company may be unable to continue operations. Although management continues to pursue these plans, there is no assurance that the Company will be successful in obtaining sufficient funding on terms acceptable to the Company to fund continuing operations, if at all.

**Risks and Uncertainties**—The Company is subject to risks and uncertainties common to early-stage companies in the biopharmaceutical industry, including, but not limited to, successful discovery and development of its product candidates, development by competitors of new technological innovations, dependence on key personnel, the ability to attract and retain qualified employees, protection of proprietary technology, compliance with governmental regulations, the ability to secure additional capital to fund operations, and commercial success of its product candidates. Any of the Company’s current product candidates and future product candidates that it may develop will require extensive nonclinical and clinical testing and regulatory approval prior to commercialization. These efforts require significant amounts of additional capital, adequate personnel, infrastructure and extensive compliance-reporting capabilities. Even if the Company’s product development efforts are successful, it is uncertain when, if ever, the Company will realize significant revenue from product sales.

## 2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES

There have been no significant changes in the Company’s accounting policies from those disclosed in its Annual Financial Statements.

**Use of Estimates**—The preparation of the financial statements in accordance with U.S. GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities and disclosure of contingent assets and liabilities at the date of the financial statements, and the reported amounts of income and expense during the reporting period. The most significant estimates relate to the accruals of research and development costs, accruals of research contract costs, and assumptions used to estimate the fair value of the Company’s stock option awards and the determination of fair value of its common stock. Management evaluates its estimates and assumptions on an ongoing basis using historical experience and other factors, including the current economic environment, and makes adjustments when facts and circumstances dictate. These estimates are based on information available as of the date of the financial statements; therefore, actual results could differ from those estimates.

**Net Loss per Share**—The Company follows the two-class method when computing net income (loss) per common share as the Company has issued shares that meet the definition of participating securities. The two-class method determines net income (loss) per common share for each class of common and participating securities according to dividends declared or accumulated and participation rights in undistributed earnings. The two-class method requires income (loss) available to common stockholders for the period to be allocated between common and participating securities based upon their respective rights to receive dividends as if all income for

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the period had been distributed. The Company considers its convertible preferred stock to be participating securities as, in the event a dividend is paid on common stock, the holders of these securities would be entitled to receive dividends on a basis consistent with the common stockholders. The Company also considers the shares issued upon the early exercise of stock options that are subject to repurchase to be participating securities because holders of such shares have non-forfeitable dividend rights in the event a dividend is paid on common stock. There is no allocation required under the two-class method during periods of loss since the participating securities do not have a contractual obligation to share in the losses of the Company.

Basic net income (loss) per common share is computed by dividing the net income (loss) per common share by the weighted-average number of common shares outstanding for the period. Diluted net income (loss) per common share is computed by adjusting net income (loss) to reallocate undistributed earnings based on the potential impact of dilutive securities. Diluted net loss per common share is computed by dividing the diluted net loss by the weighted-average number of common shares outstanding for the period, including potential dilutive common shares. For the purposes of this calculation, shares of convertible preferred stock are considered potential dilutive common shares.

In periods in which the Company reported a net loss, diluted net loss per common share was the same as basic net loss per common share, since dilutive common shares were not assumed to have been issued if their effect was anti-dilutive. During the three months ended March 31, 2023 and March 31, 2022, the Company excluded 85,495,722 potential common shares related to the conversion of Series A convertible preferred stock and 58,157,823 potential common shares related to the conversion of Series B convertible preferred stock, and during the three months ended March 31, 2023 and March 31, 2022 excluded 22,293,421 and 19,874,512 potential common shares, respectively, related to the exercise of stock options outstanding, from the computation of diluted net loss per common share because including them would have had an anti-dilutive effect and the Company reported net losses for those periods.

**Concentrations of Credit Risk**—Financial instruments that potentially subject the Company to concentrations of credit risk consist primarily of cash and cash equivalents and restricted cash. The Company’s cash and restricted cash balances exceed Federal Deposit Insurance Corporation (“FDIC”) insurance limits, and the Company’s cash equivalents consist of U.S. government money market funds. The majority of the Company’s cash and cash equivalents and restricted cash is held with a large financial institution that management believes to be of high credit quality. To date, the Company has not recognized any losses caused by uninsured balances.

**Recent Accounting Pronouncements Not Yet Adopted**—In October 2021, the FASB issued ASU 2021-08, *Business Combinations (“Topic 805”): Accounting for Contract Assets and Contract Liabilities from Contracts with Customers (“ASU 2021-08”)*. ASU 2021-08 requires an acquirer in a business combination to recognize and measure deferred revenue from acquired contracts using the revenue recognition guidance in Accounting Standards Codification Topic 606, rather than the prior requirement to record deferred revenue at fair value. ASU 2021-08 allows for immediate adoption on a retrospective basis for all business combinations that have occurred since the beginning of the annual period that includes the interim period of adoption. ASU 2021-08 is effective for annual private company reporting periods, and interim periods within those years, beginning after December 15, 2023. The Company is currently in the process of evaluating the impact of the adoption of ASU 2021-08 but does not expect a material impact to the Company’s consolidated financial statements or disclosures as it does not currently have revenue or deferred revenue from acquired contracts.

### 3. FAIR VALUE MEASUREMENTS

Fair value is defined as the price that would be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants at the measurement date. The three levels of inputs that may be used to measure fair value are as follows:

*Level 1*—Inputs are quoted prices (unadjusted) in active markets for identical assets or liabilities that the Company has the ability to access at the measurement date.

*Level 2*—Inputs are observable, unadjusted quoted prices in active markets for similar assets or liabilities, unadjusted quoted prices for identical or similar assets or liabilities in markets that are not active, or other inputs that are observable or can be corroborated by observable market data for substantially the full term of the related assets or liabilities.

*Level 3*—Inputs are unobservable inputs for the asset or liability.

The Company's cash equivalents consist of U.S. government money market funds that are reported at carrying value, which approximates fair value and is based on quoted prices in active markets for identical securities. Cash is stated at carrying value, which approximates fair value due to its short-term nature. The carrying values of prepaid expenses, other current assets, accounts payable and accrued expenses approximate their fair values due to the short-term nature of these assets and liabilities.

The following table presents information about the Company's financial assets and liabilities measured at fair value on a recurring basis and indicates the level of the fair value hierarchy utilized to determine such fair values:

	As of March 31, 2023:			
	Level 1	Level 2	Level 3	Total
Assets:				
Cash equivalents (money market)	\$ 52,990,023	\$ —	\$ —	\$ 52,990,023
	\$ 52,990,023	\$ —	\$ —	\$ 52,990,023

	As of December 31, 2022:			
	Level 1	Level 2	Level 3	Total
Assets:				
Cash equivalents (money market)	\$74,802,103	\$ —	\$ —	\$74,802,103
	\$74,802,103	\$ —	\$ —	\$74,802,103

#### 4. PROPERTY AND EQUIPMENT

The following is the summary of the Company's property and equipment and related accumulated depreciation and amortization as of March 31, 2023 and December 31, 2022:

	Useful Life	March 31, 2023	December 31, 2022
Internal use software	3 years	245,195	245,195
Furniture and fixtures	5 years	375,311	375,311
Lab equipment	7 years	290,582	278,749
Leasehold improvements	7 years	231,939	231,939
Total property and equipment		1,143,027	1,131,194
Less: accumulated depreciation and amortization		(190,281)	(132,844)
Total property and equipment, net		\$ 952,746	\$ 998,350

#### 5. ACCRUED EXPENSES AND OTHER CURRENT LIABILITIES

Accrued expenses and other current liabilities consisted of the following:

	March 31, 2023	December 31, 2022
Accrued research and development expenses	\$ 4,265,560	\$ 4,755,095
Accrued payroll expenses	682,579	796,250
Accrued professional fees	694,359	629,350
Other current liabilities	245,167	239,843
Total accrued expenses and other current liabilities	\$ 5,887,665	\$ 6,420,538

#### 6. LEASES

##### *Operating Lease Right-of-Use Asset*

In February 2022, the Company entered into a lease agreement for an office facility and laboratory space in New York, New York that commenced in April 2022 and ends in July 2029 with an additional five-year option to extend the lease beyond July 2029 at the then-prevailing effective market rental rate. Upon commencement of this lease, the Company recorded operating lease right-of-use assets and operating lease liabilities of approximately \$11.6 million based on the present value of payments over the lease term using an estimated incremental borrowing rate of 8.53% in accordance with the provisions of ASC Topic 842, *Leases* ("ASC 842"). In connection with the Company's lease of office space and laboratory space, the Company provided a security deposit to the landlord in the form of a letter of credit totaling \$1.2 million. The cash collateralizing the letter of credit was included in long-term restricted cash in the Company's condensed consolidated balance sheets as of March 31, 2023 and December 31, 2022. This lease was classified as an operating lease in accordance with the provisions of Topic 842. The Company did not recognize any right-of-use assets and lease liabilities associated with the potential option to renew or extend. The Company's operating lease agreement does not contain any significant residual value guarantees or restrictive covenants.

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Components of this operating lease are as follows:

	<b>2023</b>
<b>Period ended March 31</b>	
Operating lease expense	\$ 536,854
Variable lease expense	69,296
Total operating lease expense	\$ 606,150
<b>Other information</b>	
Remaining lease term and payment term (in years)	6.3
Incremental borrowing rate	8.53%
Cash paid for amounts included in the measurement of lease liabilities, included in operating cash flows	\$ 548,379

The following table provides a reconciliation of the Company's remaining undiscounted contractual rent obligations due within each year ended December 31 to the operating lease liabilities recognized as of March 31, 2023:

	<b>Operating Leases</b>
<b>Years ended December 31</b>	
2023	\$ 1,759,037
2024	2,099,723
2025	2,152,216
2026	2,206,022
2027	2,261,172
Thereafter	3,689,208
Total lease payments	14,167,378
Less: present value adjustment	(3,212,825)
Total operating lease liabilities	\$ 10,954,553
<b>Included in the consolidated balance sheet:</b>	
Current portion of lease liabilities - operating	2,230,981
Non-current portion of lease liabilities - operating	8,723,572
Total operating lease liabilities	\$ 10,954,553

### *Equipment Finance Leases*

Commencing in April 2022, the Company leases certain laboratory equipment under financing arrangements accounted for as finance leases in accordance with the provision of ASC Topic 842 that are classified in the Company's condensed consolidated balance sheet as finance lease liabilities with related right-of-use assets recorded and depreciated on a straight-line basis over the estimated useful life of 7 years. The total gross, accumulated amortization, and net book values of equipment finance lease right-of-use assets capitalized under such finance lease arrangements at March 31, 2023 were approximately \$2.2 million, \$0.2 million and \$2.0 million, respectively. Under the terms of the equipment finance lease agreements executed through the issuance date of these unaudited condensed consolidated financial statements, the principal balances plus interest for the equipment are to be repaid in full after 60 monthly installments following lease commencement, with lease commencement dates ranging from April 1, 2022 to April 1, 2023, annual imputed interest rates ranging from 7.90% to 9.30%, and monthly installment payment amounts ranging from approximately \$4,000 to \$18,000. As of March 31, 2023, the total aggregate monthly installment payment amount was approximately \$48,000 for

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equipment finance lease agreements executed through the issuance date of these unaudited condensed consolidated financial statements. Total equipment finance right of use assets and lease liabilities of approximately \$187,000 were recorded in the Company's condensed consolidated balance sheet at March 31, 2023 for which lease periods had not yet commenced.

Components of the equipment finance leases were as follows:

	2023
Period ended March 31	
Finance lease amortization of right-of-use assets	\$ 68,824
Interest on finance lease liabilities	34,428
Total finance lease expense	\$ 103,252
Other information	
Weighted Average Remaining Lease Payment Term (in years)	4.4
Weighted Average Remaining Amortization Term (in years)	6.4
Weighted Average Effective Interest Rate	8.66%
Cash paid for amounts included in the measurement of lease liabilities, included in financing cash flows	\$ 159,610

The following table provides a reconciliation of the Company's remaining equipment finance lease obligations due within each year ending December 31 to the equipment finance lease liabilities recognized at March 31, 2023:

	Equipment Finance Leases
Years ended December 31	
2023	\$ 390,353
2024	579,734
2025	579,734
2026	579,734
2027	357,653
Thereafter	11,591
Total lease payments	2,498,799
Less: imputed interest	(421,139)
Total finance lease liabilities	\$ 2,077,660
Included in the consolidated balance sheet:	
Current portion of lease liabilities—finance	523,157
Non-current portion of lease liabilities—finance	1,554,503
Total finance lease liabilities	\$ 2,077,660

## 7. PREFERRED STOCK

As of March 31, 2023, the Company's certificate of incorporation authorized the Company to issue 143,653,546 million shares of series A and series B convertible preferred stock at par value of \$0.0001.

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As of March 31, 2023 and December 31, 2022, preferred stock consisted of the following:

	<b>Convertible Preferred Stock Authorized</b>	<b>Convertible Preferred Stock Issued and Outstanding</b>	<b>Proceeds Received</b>	<b>Liquidation Preference</b>	<b>Common Stock Issuable Upon Conversion</b>
Series A convertible preferred stock	85,495,722	85,495,722	\$ 85,495,722	\$ 85,495,722	85,495,722
Series B convertible preferred stock	58,157,824	58,157,823	100,059,953	100,059,953	58,157,825
	143,653,546	143,653,545	\$ 185,555,675	\$ 185,555,675	143,653,547

As of March 31, 2023, the rights and privileges of the holders of the convertible preferred stock were as follows:

*Voting* – Holders of preferred stock shall vote together with the holders of common stock as a single class and on an as converted to common stock basis.

*Dividends* – The holders of shares of series A convertible preferred stock are entitled to receive, as declared by the Company’s board of directors, dividends at the rate of 8% per share of preferred stock, prior and in preference to any declaration or payment of any other dividend (other than dividends on shares of common stock payable in shares of common stock).

*Deemed Liquidation Event* – Each of the following events shall be considered a “Deemed Liquidation Event” unless the holders of a majority of the outstanding shares of preferred stock elect otherwise by written notice sent to the Company at least five business days prior to the effective date of any such event; (a) A merger or consolidation in which the Company is a constituent party or a subsidiary of the Company is a constituent party and the Company issues shares of its capital stock pursuant to such merger or consolidation. Except any such merger or consolidation involving the Company or a subsidiary in which the shares of capital stock of the Company outstanding immediately prior to such merger or consolidation continue to represent, or are converted into or exchanged for shares of capital stock that represent, immediately following such merger or consolidation, at least a majority of the capital stock of (1) the surviving or resulting company; or (2) if the surviving or resulting company is a wholly owned subsidiary of another company immediately following such merger or consolidation, the parent company of such surviving or resulting corporation; or (b) (1) the sale, lease, transfer, exclusive license or other disposition, in a single transaction or series of related transactions, by the Company or any subsidiary of the Company of all or substantially all the assets of the Company and its subsidiaries taken as a whole, or (2) the sale or disposition (whether by merger, consolidation or otherwise, and whether in a single transaction or a series of related transactions) of one or more subsidiaries of the Company if substantially all of the assets of the Company and its subsidiaries taken as a whole are held by such subsidiary or subsidiaries, except where such sale, lease, transfer, exclusive license or other disposition is to a wholly owned subsidiary of the Company.

*Liquidation Preference* – In the event of any voluntary or involuntary liquidation, dissolution, or winding up of the Company, the holders of shares of preferred stock shall be entitled to be paid out of the assets of the Company or, in the case of a Deemed Liquidation Event, out of the consideration payable to stockholders or the available proceeds, before any payment shall be made to the holders of common stock, an amount per share equal to the series B convertible preferred stock issue price of \$1.72 per share and the series A convertible preferred stock issue price of \$1.00 per share, plus any dividends declared but unpaid. If upon any such liquidation, dissolution or winding up of the Company or Deemed Liquidation Event, the assets of the Company available for distribution to its stockholders shall be insufficient to pay the holders of shares of preferred stock the full amount to which they shall share ratably in any distribution of the assets available for distribution in proportion to the



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respective amounts which would otherwise be payable in respect of the shares held by them upon such distribution if all amounts payable on or with respect to such shares were paid in full (collectively, the “Liquidation Preference Per Convertible Preferred Stock”).

In the event that there are additional assets to be distributed, the holders of the series A convertible preferred stock and series B convertible preferred stock will share in the distribution along with common stockholders as if the shares of the series A convertible preferred stock and series B convertible preferred stock had converted to common stock immediately prior to the distribution up to an amount taken in consideration with the Liquidation Preference Per Convertible Preferred Stock not to exceed \$3.00 per share allocated to series A convertible preferred stockholders and \$5.16147 per share allocated to series B convertible preferred stockholders.

*Redemption* – The preferred stock does not contain a mandatory redemption provision. Upon a Deemed Liquidation Event, the holders will be paid their preference amounts according to their priority. That is, the preferred stockholders will be entitled to an amount up to \$1.00 per share, plus any accrued dividends. Accretion to redemption value will only be required if a Deemed Liquidation Event is probable.

*Conversion* – In the event that any Noteholder, or an affiliate of any Noteholder, the investor of Notes did not participate in any Milestones then each share of preferred stock held by the Noteholders would have automatically been converted into the number of shares of common stock equal to the \$1.00 per share divided by the conversion price in effect immediately prior to the consummation of the Milestones closings multiplied by ten.

Upon either (a) the closing of the sale of shares of common stock to the public at a price of at least \$3.00 per share (subject to appropriate adjustment in the event of any stock dividend, stock split, combination or other similar recapitalization with respect to the common stock), in a firm-commitment underwritten public offering pursuant to an effective registration statement under the Securities Act of 1933, as amended, resulting in at least \$50.0 million of gross proceeds to the Company and in connection with the offering of the common stock listed for trading on the Nasdaq Stock Market’s National Market, the New York Stock Exchange or another exchange or marketplace approved by the Company’s board of directors, including the approval of at least two (2) preferred directors (a “Qualified IPO”) or (b) the date and time, or the occurrence of an event specified by vote or written consent of the Noteholders, then (i) all outstanding shares of preferred stock shall automatically be converted into shares of common stock, at the then effective conversion rate.

## **8. COMMON STOCK**

As of March 31, 2023, the Company’s certificate of incorporation authorized the Company to issue 192,020,973 shares of common stock with a par value of \$0.0001 per share. The voting, dividend and liquidation rights of the holders of the Company’s common shares are subject to and qualified by the rights, powers and preferences of the holders of the convertible preferred stock set forth in Note 7. Each common share entitles the holder to one vote on all matters submitted to a vote of the Company’s stockholders.

Common stockholders are entitled to receive dividends, if any, as may be declared by the Company’s board of directors, subject to the preferential dividend rights of the convertible preferred stock. Through March 31, 2023, no cash dividends have been declared or paid by the Company.

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The Company has reserved the following number of shares of common stock for the exercise of outstanding stock options, future issuance of stock-based awards, and conversion of outstanding shares of convertible preferred stock:

	March 31, 2023	December 31, 2022
Shares of series A convertible preferred stock	85,495,722	85,495,722
Shares of series B convertible preferred stock	58,157,823	58,157,823
Options to purchase shares of common stock	22,293,421	20,300,861
Shares available for issuance under the 2021 Plan	8,591,016	10,599,419
Total shares of common stock reserved for future issuance	174,537,982	174,553,825

## 9. STOCK-BASED COMPENSATION

In February 2021, the Company adopted the 2021 Equity Incentive Plan (the “Plan”) for the issuance of stock options to the Company’s key directors, officers, employees and consultants, as a means to secure the benefits arising from capital stock ownership. The purposes of the Plan are to promote the alignment of the interests of key directors, officers, employees, and consultants with the success of the Company and to provide compensation opportunities to attract, retain and motivate directors, officers, employees, and consultants of the Company.

Under the Plan, the Company initially authorized 8,918,225 shares of its common stock for issuance upon exercise of options granted under the Plan. In August 2021, the Company’s Board of Directors amended the Plan to increase the aggregate number of shares of its common stock reserved for issuance pursuant to the Plan by 20,019,725 shares, from 8,918,225 shares to a new total of 28,937,950 shares. In November 2021, the Company’s Board of Directors further amended the Plan to further increase the aggregate number of shares of its common stock reserved for issuance pursuant to the Plan by 3,388,594 shares, from 28,937,950 shares to a new total of 32,326,544 shares. As of March 31, 2023, 8,591,016 shares were available for future grants.

### *Stock option activity*

Stock options granted generally (i) are subject to requisite service requirements, (ii) vest over a four-year period with 25% of the options granted vesting after one year and the remainder vesting in equal monthly installments over the following 36 months, and (iii) allow for early exercise subject to repurchase. Stock options granted to certain of the Company’s non-employees vest in equal monthly installments over a four-year period or vested upon the achievement of a certain milestone event. The Company did not repurchase any stock options during the three months ended March 31, 2023 and March 31, 2022.

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The following table summarizes the stock option activity under the Plan for the three months ended March 31, 2023:

	Number of Shares	Weighted-Average Exercise Price	Weighted-Average Grant Date Fair Value	Weighted-Average Remaining Contractual Term (in years)	Aggregate Intrinsic Value (in thousands)
Outstanding as of December 31, 2022	20,300,861	\$ 0.44	\$ 0.22	8.65	\$ 20,130
Granted	4,922,272	1.66	1.03		
Exercised	(15,843)	0.77	0.50		11
Forfeited	(2,913,869)	0.45	0.29		
Outstanding as of March 31, 2023	22,293,421	\$ 0.71	\$ 0.45	8.01	\$ 17,256
Vested options outstanding and exercisable as of March 31, 2023	8,186,308	\$ 0.39	\$ 0.25	6.35	\$ 8,554
Unvested options outstanding and exercisable as of March 31, 2023	14,107,113	\$ 0.89	\$ 0.56	8.97	\$ 8,701

The aggregate intrinsic value of stock options is calculated as the difference between the exercise price of the stock options and the fair value of the Company's common stock for those stock options that had exercise prices lower than the fair value of the Company's common stock.

The total grant date fair values of options vested during the three months ended March 31, 2023 and March 31, 2022 were \$0.5 million and \$0.1 million, respectively.

The Company estimated the fair value of options granted using a Black-Scholes option pricing model with the following assumptions presented on a weighted average basis during the following periods presented:

	Three Months Ended March 31,	
	2023	2022
Weighted average risk-free interest rate	3.76%	1.74%
Expected term (in years)	5.90	6.06
Expected volatility	86.80%	72.50%
Expected dividend yield	0.00%	0.00%

The expected dividend yields are 0.00% as the Company has never paid cash dividends and does not expect to pay any cash dividends in the foreseeable future.

### *Stock-based compensation expense*

The Company recorded total stock-based compensation of \$0.4 million and \$0.3 million for the three months ended March 31, 2023, and March 31, 2022, respectively. Stock-based compensation expense was classified as follows in the Company's unaudited condensed consolidated statements of operations and comprehensive loss:

	Three Months Ended March 31,	
	2023	2022
Research and development expense	\$ 158,214	\$ 131,976
General and administrative expense	239,041	142,747
Total stock-based compensation expense	\$ 397,255	\$ 274,723

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As of March 31, 2023 there was \$7.8 million of unrecognized stock-based compensation expense related to unvested stock options estimated to be recognized over a weighted-average period of 2.75 years.

### 10. NET LOSS PER SHARE

Basic and diluted net loss per common share attributable to common stockholders was calculated as follows:

	Three Months Ended	
	March 31,	
	2023	2022
Numerator:		
Net loss attributable to common stockholders	\$ (18,657,482)	\$ (12,501,830)
Denominator:		
Weighted-average common shares outstanding, basic and diluted	17,069,994	16,863,270
Net loss per share attributable to common stockholders, basic and diluted	\$ (1.09)	\$ (0.74)

### 11. COMMITMENTS AND CONTINGENCIES

**Leases**—As of March 31, 2023, the Company had entered into commitments under lease agreements to rent office space and finance equipment (see Note 6).

**Commitments**—As of December 31, 2022, the Company had entered into commitments under license, acquisition, research collaboration and sponsored research agreements with third parties (see Note 12). In addition, the Company has entered into services agreements with third parties for pharmaceutical manufacturing and research activities in the normal course of business, which can generally be terminated by the Company with 30- to 60-days' written notice, unless otherwise indicated. Further, certain of the Company's manufacturing agreements could require early termination and wind-down payments due from the Company upon either the termination of its clinical trials or if the Company terminates such agreements for convenience.

**Contingencies**—From time to time, the Company may have certain contingent liabilities that arise in the ordinary course of business. The Company accrues a liability for such matters when it is probable that future expenditures will be made, and such expenditures can be reasonably estimated. For all periods presented, the Company was not a party to any pending material litigation or other material legal proceedings.

**Indemnification Agreements**—In the ordinary course of business, the Company may provide indemnification of varying scope and terms to vendors, lessors, business partners and other parties with respect to certain matters including, but not limited to, losses arising out of breach of such agreements or from intellectual property infringement claims made by third parties. In addition, the Company has entered into indemnification agreements with members of its board of directors and executive officers that will require the Company, among other things, to indemnify them against certain liabilities that may arise by reason of their status or service as directors or officers. The maximum potential amount of future payments the Company could be required to make under these indemnification agreements is, in many cases, unlimited. For all periods presented, the Company has not incurred any material costs as a result of such indemnifications.

### 12. LICENSE, ACQUISITION, RESEARCH COLLABORATION AND SPONSORED RESEARCH AGREEMENTS

**Adverum Biotechnologies**—On January 25, 2021, the Company entered into an exclusive license agreement with Adverum Biotechnologies Inc. ("Adverum") to in-license materials and technology related to the treatment of Cardiomyopathy due to Friedrich's Ataxia (the "Adverum Agreement"). In connection with the Adverum

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Agreement, the Company gained access to a portfolio of inventions, patent rights, technology, and licensed methods that the Company continues to develop, and the Company will assume all development and commercialization activities worldwide. Pursuant to the Adverum Agreement, the Company paid a one-time up-front non-refundable fee of \$7.5 million, and is obligated to pay aggregate development and regulatory milestones of up to \$17.5 million including a \$3.5 million development milestone that was achieved and paid in the first quarter of 2023, aggregate sales event and commercialization milestones of up to \$49.0 million, and royalty payments ranging from 8% to 10% based on annual aggregate worldwide net sales of Products, subject to reductions upon the expiration of valid claims in licensed patents and third-party licenses. As of March 31, 2023, there were no research and development expenses recorded by the Company or payments made to Adverum under the terms of the Adverum Agreement other than the one-time up-front non-refundable fee of \$7.5 million and the \$3.5 million development milestone that was achieved and paid in the first quarter of 2023.

The Adverum Agreement remains in effect until termination at the date of the last royalty term to expire. The Company can terminate the Adverum Agreement with 120 days' written notice. The Adverum Agreement can also be terminated as a result of a patent challenge, material breach of contractual terms, or insolvency by either party.

**Cornell University**—On May 27, 2020, the Company entered into two exclusive license agreements with Cornell University (“Cornell”) (the “Cornell First License Agreement” and the “Cornell Second License Agreement,” collectively “the Cornell License Agreements”). The Cornell First License Agreement is for the in-license of technology related to portfolios for APOE, Alzheimer’s disease, and Anti-Tau, although the Company’s license is not restricted by such indications and it includes assignment to the Company of Cornell’s IND for the use of AAVrh10.hAPOE2 vector to treat APOE4 homozygous patients who are at risk for or have Alzheimer’s disease to support development of the Company’s LX1001 program. The Cornell Second License Agreement is for the in-license of technology related to portfolios for infantile neuronal ceroid lipofuscinosis (also called CLN2 Batten disease) and Friedreich’s ataxia although the Company’s license is not restricted by such indications, and it includes assignment to the Company of Cornell’s IND for the use of AAVrh.10cUhCLN2 to treat children with CLN2 Batten disease to support development of the Company’s LX1004 program. Through the Cornell License Agreements, the Company gains access to a portfolio of inventions, patent rights, technology, and licensed methods that the Company continues to develop. Under the terms of the Cornell License Agreements, the Company has assumed all development and commercialization activities worldwide with respect to the licensed technology.

As initial consideration for the Cornell License Agreements, the Company paid Cornell an upfront payment in cash of \$0.3 million and issued \$1.3 million of notes (“Notes”). In November 2020, Notes with outstanding principal of \$1.3 million were cancelled in exchange for 1,337,610 shares of Series A convertible preferred stock. As additional consideration, the Company is required to pay Cornell up to \$8.4 million for each portfolio upon the achievement of specific clinical and regulatory milestones under the Cornell First License Agreement and up to \$4.3 million in two portfolios and up to \$0.6 million for a third portfolio upon the achievement of specific clinical and regulatory milestones under the Cornell Second License Agreement. In the second quarter of 2022, a clinical and regulatory milestone of \$0.1 million was recognized and paid to Cornell in connection with the Cornell Second License Agreement. The Company is also required to pay Cornell a flat royalty in the mid-single-digits based on net sales of the products covered by the licenses, subject to certain adjustments.

Upon expiration of the royalty term of a given licensed product in a country, the respective license becomes non-exclusive and royalty-free. In addition, each of the Cornell License Agreements may be terminated by the Company for any reason upon ninety (90) days’ advance notice to Cornell and by Cornell upon the Company’s material uncured breach, and all licenses and rights granted by either party under such agreement will concurrently terminate.

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During the three months ended March 31, 2023 and March 31, 2022, the Company did not incur any research and development expenses or make any payments to Cornell in connection with the Cornell License Agreements.

**Stelios Therapeutics, Inc.**—Stelios Therapeutics, Inc. (“Stelios”) was an early-stage company developing novel adeno-associated AAV-based gene therapies for rare cardiac conditions including arrhythmogenic right ventricular cardiomyopathy and TNNI3-associated hypertrophic cardiomyopathy. On July 16, 2021, the Company acquired 100% of the outstanding stock of Stelios that was accounted for as an asset acquisition pursuant FASB ASC 805, *Business Combinations*. The Company is required to pay up to an aggregate of \$20.5 million to the selling shareholders of Stelios upon the achievement of certain development milestones, including a \$2.0 million development milestone that was achieved and paid in the third quarter of 2022.

**Regents of the University of California, San Diego**—Stelios entered into exclusive worldwide license agreements on April 23, 2020 and August 6, 2020 (the “First UCSD Agreement” and the “Second UCSD Agreement”, respectively) with the Regents of UCSD to in-license materials and intellectual property related to gene therapies for arrhythmogenic right ventricular cardiomyopathy and hypertrophic cardiomyopathy, respectively. The First UCSD Agreement and the Second UCSD Agreement relate to the Company’s development efforts for its LX2021 and LX2022 programs, respectively. In connection with the First UCSD Agreement and the Second UCSD Agreement, the Company gained access to inventions, patent rights, technology, and licensed methods that it continues to develop, and it has assumed all worldwide development and commercialization activities with respect to the licensed technologies. The First UCSD Agreement and Second UCSD Agreement required Stelios to pay one-time up-front non-refundable cash fees of \$20,000 for each agreement and requires the Company to pay aggregate development and commercialization milestones of up to \$4.75 million and \$2.4 million, respectively, and low- to mid-single digit royalties and low-single digit royalties, respectively, based on aggregate net sales. The only research and development expenses incurred by Stelios or the Company and payments made to the Regents of UCSD to-date under the terms of the First UCSD Agreement and the Second UCSD Agreement were the one-time up-front non-refundable cash fees of \$20,000 for each agreement. The Company has the right to terminate the First UCSD Agreement and the Second UCSD Agreement at any time upon sixty (60)-days’ written notice to the Regents of UCSD.

On October 4, 2021, the Company entered into an exclusive worldwide license agreement (the “Third UCSD Agreement” and collectively with the First UCSD Agreement and the Second UCSD Agreement, the “UCSD Agreements”) with the Regents of UCSD to in-license materials and intellectual property related to a gene therapy for arrhythmogenic right ventricular cardiomyopathy. The Third UCSD Agreement relates to the Company’s development efforts for its LX2020 program. In connection with the Third UCSD Agreement, the Company gained access to inventions, patent rights, technology, and licensed methods that it continues to develop, and it has assumed all worldwide development and commercialization activities with respect to the licensed technology. The Third UCSD Agreement required the Company to pay a one-time up-front non-refundable cash fee of \$20,000 and requires the Company to pay aggregate development and commercialization milestones of up to \$4.0 million, and low- to mid-single digit royalties based on aggregate net sales. The only research and development expenses incurred by the Company and payments made to the Regents of UCSD to-date under the terms of the Third UCSD Agreement were the one-time up-front non-refundable cash fee of \$20,000. The Company has the right to terminate the Third UCSD Agreement at any time upon sixty (60)-days’ written notice to the Regents of UCSD.

On December 3, 2021, the Company entered into two sponsored research agreements with the Regents of UCSD (collectively, the “UCSD SRAs”) for the Company’s LX2020, LX2021 and LX2022 programs in connection with the UCSD Agreements. Under the terms of the UCSD SRAs, the Company has the first rights to obtain non-exclusive or exclusive, sublicensable, royalty-bearing, perpetual and transferable worldwide licenses in any resulting inventions owned by the Regents of UCSD or resulting jointly owned inventions between the Company and the Regents of UCSD, and the Company retains the rights to any resulting inventions owned by the Company. The UCSD SRAs have a two-year term and may be terminated early by the Company at any time

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upon the giving of thirty (30) days' written notice to the Regents of UCSD. The total costs to be invoiced to the Company over the terms of the UCSD SRAs are approximately \$5.0 million, of which \$0.2 million and \$0.7 million were recorded in research and development expenses during the three months ended March 31, 2023 and March 31, 2022, respectively, and of which a cumulative total of \$2.6 million has been recorded in research and development expenses to-date as of March 31, 2023. The Company paid totals of \$0.2 million and \$1.4 million to the Regents of UCSD during the three months ended March 31, 2023 and March 31, 2022, respectively, and has paid a cumulative total of \$3.0 million to the Regents of UCSD to-date as of March 31, 2023, in connection with the UCSD SRAs.

**Weill Cornell Medical College**—On February 2, 2021, the Company entered into a Research Collaboration Agreement with Weill Cornell Medical College (“WCM” and the “WCM Agreement”) in conjunction with the Cornell License Agreements entered on May 27, 2020. The Company committed to fund scientific research at WCM to investigate further and potentially enhance the technology licensed to the Company pursuant to the License Agreements.

Under the terms of the WCM Agreement, the Company is required pay WCM at least \$3.0 million over the course of 3 years, with equal \$0.25 million payments due each quarter starting 10 days after the execution of the WCM Agreement. With respect to each WCM invention, joint invention, and related joint results for which an Improvement, as defined in the WCM Agreement, applies and the Company has made an election to amend the Cornell License Agreements, the Company has the first option to negotiate in good faith with WCM for royalty-bearing, worldwide license, under Cornell patent rights, Cornell rights, and Cornell's interest in joint patent rights, to develop, make, have made, use, offer for sale, sell, have sold, and import derived products in the field. Research and development costs incurred by the Company and paid to WCM related to the WCM Agreement were \$0.7 million and \$1.6 million, respectively, during the three months ended March 31, 2023. Research and development costs incurred by the Company and paid to WCM related to the WCM Agreement were \$1.1 million and \$0.8 million, respectively, during the three months ended March 31, 2022. Cumulatively, the Company has incurred total research and development costs of \$9.6 million and paid a total of \$9.4 million to WCM to-date in connection with the WCM Agreement as of March 31, 2023.

The WCM Agreement terminates upon the expiration of the agreed upon 3-year term. In addition, the WCM Agreement may be terminated by the Company in the event that the WCM principal investigator ceases to supervise the research and WCM is unable to or declines to find a substitute.

### **13. SUBSEQUENT EVENTS**

Subsequent events have been evaluated through May 11, 2023, which is the date that these unaudited condensed consolidated financial statements were issued and were available to be issued.

# Lexeo Therapeutics, Inc.

## Audited consolidated financial statements

For the years ended December 31, 2022 and 2021

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KPMG LLP  
345 Park Avenue  
New York, NY 10154-0102

## Report of Independent Registered Public Accounting Firm

To the Stockholders and Board of Directors  
Lexeo Therapeutics, Inc.:

### *Opinion on the Consolidated Financial Statements*

We have audited the accompanying consolidated balance sheets of Lexeo Therapeutics, Inc. and subsidiary (the Company) as of December 31, 2022 and 2021, the related consolidated statements of operations and comprehensive loss, convertible preferred stock and stockholders' deficit, and cash flows for the years then ended, and the related notes (collectively, the consolidated financial statements). In our opinion, the consolidated financial statements present fairly, in all material respects, the financial position of the Company as of December 31, 2022 and 2021, and the results of its operations and its cash flows for the years then ended, in conformity with U.S. generally accepted accounting principles.

### *Change in Accounting Principle*

As discussed in Note 2 to the consolidated financial statements, the Company has changed its method of accounting for leases as of January 1, 2022, due to the adoption of Accounting Standards Codification Topic 842, Leases.

### *Basis for Opinion*

These consolidated financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on these consolidated financial statements based on our audits. We are a public accounting firm registered with the Public Company Accounting Oversight Board (United States) (PCAOB) and are required to be independent with respect to the Company in accordance with the U.S. federal securities laws and the applicable rules and regulations of the Securities and Exchange Commission and the PCAOB.

We conducted our audits in accordance with the standards of the PCAOB and in accordance with auditing standards generally accepted in the United States of America. Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the consolidated financial statements are free of material misstatement, whether due to error or fraud. Our audits included performing procedures to assess the risks of material misstatement of the consolidated financial statements, whether due to error or fraud, and performing procedures that respond to those risks. Such procedures included examining, on a test basis, evidence regarding the amounts and disclosures in the consolidated financial statements. Our audits also included evaluating the accounting principles used and significant estimates made by management, as well as evaluating the overall presentation of the consolidated financial statements. We believe that our audits provide a reasonable basis for our opinion.

**KPMG LLP**

We have served as the Company's auditor since 2021.

New York, New York  
March 31, 2023

KPMG LLP, a Delaware limited liability partnership and a member firm of the KPMG global organization of independent member firms affiliated with KPMG International Limited, a private English company limited by guarantee.

## Lexeo Therapeutics, Inc. Consolidated Balance Sheets

	December 31,	
	2022	2021
<b>Assets</b>		
Current assets:		
Cash and cash equivalents	\$ 77,334,973	\$ 135,860,060
Prepaid expenses	2,342,550	128,089
Other current assets	370,541	695,899
Total current assets	80,048,064	136,684,048
Security deposits	—	2,500
Restricted cash	3,253,246	—
Property and equipment, net	998,356	174,266
Lease right-of-use assets—finance, net	2,068,633	—
Lease right-of-use assets—operating	10,708,144	—
Total assets	\$ 97,076,437	\$ 136,860,814
<b>Liabilities, Convertible Preferred Stock and Stockholders' Deficit</b>		
Current liabilities:		
Accounts payable	\$ 2,677,800	\$ 3,060,755
Taxes payable	505,681	527,823
Deferred income	—	46,460
Accrued expenses and other current liabilities	6,420,538	2,883,413
Accrued bonuses	1,911,369	930,966
Current portion of lease liabilities—finance	547,294	—
Current portion of lease liabilities—operating	2,287,109	—
Total current liabilities	14,349,791	7,449,417
Non-current liabilities		
Non-current portion of lease liabilities—finance	1,647,898	—
Non-current portion of lease liabilities—operating	8,999,643	—
Total liabilities	24,997,332	7,449,417
Commitments and contingencies (Note 12)		
Series A convertible preferred stock, \$0.0001 par value; 85,495,722 shares authorized, issued, and outstanding as of December 31, 2022 and December 31, 2021; aggregate liquidation value of \$85,495,722 as of December 31, 2022 and December 31, 2021	85,268,513	85,247,984
Series B convertible preferred stock, \$0.0001 par value; 58,157,824 shares authorized as of December 31, 2022 and December 31, 2022; 58,157,823 shares issued and outstanding as of December 31, 2022 and December 31, 2021; aggregate liquidation value of \$100,059,953 as of December 31, 2022 and December 31, 2021	99,764,986	99,764,986
Stockholders' and members' deficit:		
Common stock, \$0.0001 par value, 192,020,973 shares authorized as of December 31, 2022 and December 31, 2021; 17,467,148 shares issued and 17,026,915 shares outstanding as of December 31, 2022; 16,745,827 shares issued and outstanding as of December 31, 2021	1,703	1,675
Additional paid-in capital	2,490,168	566,290
Accumulated deficit	(115,446,265)	(56,169,538)
Total stockholders' deficit	(112,954,394)	(55,601,573)
Total liabilities, convertible preferred stock, stockholders' deficit	\$ 97,076,437	\$ 136,860,814

The accompanying notes are an integral part of these consolidated financial statements.

## Lexeo Therapeutics, Inc.

### Consolidated Statements of Operations and Comprehensive Loss

	Years ended December 31,	
	2022	2021
Revenue		
Grant revenue	\$ 653,956	\$ 1,657,122
Total revenue	653,956	1,657,122
Operating expenses		
Research and development	49,161,540	45,120,874
General and administrative	12,001,097	7,173,026
Total operating expenses	61,162,637	52,293,900
Operating loss	(60,508,681)	(50,636,778)
Other income and expense		
Other expense, net	1,852	155
Interest expense	90,874	—
Interest income	1,324,680	15,220
Total other income and expense	1,231,954	15,065
Loss from operations before income taxes	(59,276,727)	(50,621,713)
Income taxes	—	—
Net loss and comprehensive loss	\$ (59,276,727)	\$ (50,621,713)
Net loss per common share, basic and diluted	(3.43)	(3.08)
Weighted average number of shares outstanding used in computation of net loss per common share, basic and diluted	17,272,352	16,449,814

The accompanying notes are an integral part of these consolidated financial statements.

## Lexeo Therapeutics, Inc.

### Consolidated Statements of Convertible Preferred Stock and Stockholders' Deficit

	Convertible Preferred Stock		Common Stock		Additional Paid-in Capital	Accumulated Deficit	Total Stockholders' Deficit
	Shares	Amount	Shares	Amount			
Balances at January 1, 2021	34,495,725	\$ 31,237,986	16,040,884	\$ 1,604	\$ 9,794	\$ (5,547,825)	\$ (5,536,427)
Stock-based compensation expense	—	—	—	—	—	—	—
Tranche Obligation Settlement	—	3,010,000	—	—	—	—	—
Issuance of Series A convertible preferred stock, 1st milestone and 2nd milestone	50,999,997	50,999,998	—	—	—	—	—
Issuance of Series B convertible preferred stock, net of issuance costs of \$294,978	58,157,823	99,764,986	—	—	—	—	—
Exercise of stock options	—	—	704,943	71	169,861	—	169,932
Stock-based compensation expense	—	—	—	—	386,635	—	386,635
Net loss	—	—	—	—	—	(50,621,713)	(50,621,713)
<b>Balances at December 31, 2021</b>	<b>143,653,545</b>	<b>\$ 185,012,970</b>	<b>16,745,827</b>	<b>\$ 1,675</b>	<b>\$ 566,290</b>	<b>\$ (56,169,538)</b>	<b>\$ (55,601,573)</b>
Proceeds from issuance of series A convertible preferred stock	—	20,529	—	—	—	—	—
Exercise of stock options	—	—	721,321	72	331,736	—	331,808
Proceeds received from early exercise of unvested stock options subject to repurchase and recorded as deposit liabilities	—	—	(440,233)	(44)	(188,423)	—	(188,467)
Stock-based compensation expense	—	—	—	—	1,780,566	—	1,780,566
Net loss	—	—	—	—	—	(59,276,727)	(59,276,727)
<b>Balances at December 31, 2022</b>	<b>143,653,545</b>	<b>\$ 185,033,499</b>	<b>17,026,915</b>	<b>\$ 1,703</b>	<b>\$ 2,490,168</b>	<b>\$ (115,446,265)</b>	<b>\$ (112,954,394)</b>

The accompanying notes are an integral part of these financial statements.

# Lexeo Therapeutics Inc.

## Consolidated Statements of Cash Flows

	Years ended December 31,	
	2022	2021
<b>Cash flows from operating activities:</b>		
Net loss	\$ (59,276,727)	\$ (50,621,713)
Adjustments to reconcile net loss to net cash (used in) provided by operating activities:		
Amortization of right-of-use asset, operating	877,269	—
Amortization of right-of-use assets, finance	147,807	—
Stock based compensation expense	1,780,566	386,635
Depreciation and amortization expense	119,441	13,405
Changes in operating assets and liabilities:		
Prepaid expenses and other current assets	(1,889,103)	(543,733)
Security deposits	2,500	—
Accounts payable	(425,943)	3,015,273
Issuance costs payable	—	(223,587)
Change in deferred income	(46,460)	(93,924)
Accrued expenses and other current liabilities	4,476,496	3,395,162
Tax payable	(22,143)	489,519
Lease liability, operating	(298,661)	—
Lease liabilities, finance	(5,635)	—
Net cash used in operating activities	<u>(54,560,593)</u>	<u>(44,182,963)</u>
<b>Cash flows from investing activities:</b>		
Purchase of internal use software	(228,438)	—
Purchase of property and equipment	(672,098)	(187,671)
Net cash used in investing activities	<u>(900,536)</u>	<u>(187,671)</u>
<b>Cash flows from financing activities:</b>		
Proceeds from issuance of series A/B convertible preferred stock	20,529	151,059,962
Series B convertible preferred stock issuance costs	—	(294,978)
Payment of costs related to initial public offering	—	(77,431)
Proceeds from exercise of stock options	331,808	169,932
Payments on finance leases	(163,049)	—
Net cash provided by financing activities	<u>189,288</u>	<u>150,857,485</u>
<b>Net change in cash, cash equivalents and restricted cash</b>	<u>(55,271,841)</u>	<u>106,486,851</u>
Total cash, cash equivalents and restricted cash at beginning of period	<u>135,860,060</u>	<u>29,373,209</u>
Total cash, cash equivalents and restricted cash at end of period	<u>\$ 80,588,219</u>	<u>\$ 135,860,060</u>
Cash and cash equivalents at beginning of period	<u>135,860,060</u>	<u>29,373,209</u>
Restricted cash included in long-term assets at beginning of period	<u>—</u>	<u>—</u>
Total cash, cash equivalents and restricted cash at beginning of period	<u>135,860,060</u>	<u>29,373,209</u>
Cash and cash equivalents at end of period	<u>77,334,973</u>	<u>135,860,060</u>
Restricted cash included in long-term assets at end of period	<u>3,253,246</u>	<u>—</u>
Total cash, cash equivalents and restricted cash at end of period	<u>\$ 80,588,219</u>	<u>\$ 135,860,060</u>
<b>Supplemental disclosures of cash flow information</b>		
Interest paid	\$ 77,211	\$ —
Income taxes paid	\$ —	\$ —
<b>Supplemental disclosure of non-cash investing and financing activities</b>		
Deferred offering costs included in accounts payable and accrued expenses	\$ —	\$ 169,253
Purchase of property and equipment included in accounts payable and accrued expenses	\$ 42,988	\$ —
Operating lease right-of-use assets and operating lease liabilities recognized	\$ 11,585,413	\$ —
Finance lease right-of use assets and finance lease liabilities recognized	\$ 2,363,876	\$ —
Proceeds received from early exercise of unvested stock options subject to repurchase and recorded as deposit liabilities	\$ 188,467	\$ —

The accompanying notes are an integral part of these consolidated financial statements.

# Lexeo Therapeutics, Inc.

## Notes to consolidated financial statements

### 1. DESCRIPTION OF BUSINESS AND BASIS OF PRESENTATION

**Description of Business**—Lexeo Therapeutics Inc. (“Lexeo” or the “Company”), together with its consolidated subsidiary, Stelios Therapeutics Inc., until such subsidiary was merged into the Company on December 15, 2022 and ceased to exist, is a clinical stage gene therapy biotechnology company with a focus on hereditary and acquired diseases of high unmet need. The Company’s investigational therapies have the potential to offer gene therapy-based treatments to address many diseases that have eluded today’s existing drug delivery platforms. Lexeo utilizes adeno-associated viruses (“AAV”) that have been engineered to transfer genes to patients. The Company’s therapeutic investigational treatments include gene therapies primarily in the early clinical and late pre-clinical stages of research and development.

The Company is located in New York, NY and was first formed on February 17, 2017, as an LLC under the laws of the State of Delaware under the legal name Lexeo Therapeutics, LLC. The Company filed and executed a certificate of conversion to corporation on November 20, 2020, to convert the LLC to Lexeo Therapeutics, Inc, a Delaware corporation. All of the Company’s tangible assets are held in the United States (“U.S.”).

**Basis of Presentation and Principles of Consolidation**—The accompanying consolidated financial statements reflect the operations of the Company and its wholly-owned subsidiary that have been prepared in conformity with generally accepted accounting principles in the United States of America (“GAAP”). Any reference in these notes to applicable guidance is meant to refer to the authoritative GAAP as found in the Accounting Standards Codification (“ASC”) and Accounting Standards Updates (“ASU”) of the Financial Accounting Standards Board (“FASB”). The accounts of the wholly owned subsidiary are included in these consolidated financial statements. All intercompany balances and transactions have been eliminated in consolidation.

**Need for Additional Capital**—Since inception, the Company has incurred net losses and negative cash flows from operations, including net losses of \$59.3 million and \$50.6 million during the years ended December 31, 2022 and 2021, respectively. As of December 31, 2022 and December 31, 2021, the Company had an accumulated deficit of \$115.4 million and \$56.2 million, respectively, and expects to incur substantial operating losses and negative cash flows from operations for the foreseeable future. The Company had \$77.3 million and \$135.9 million of cash and cash equivalents at December 31, 2022 and December 31, 2021, respectively. During the year ended December 31, 2021 the Company raised \$51.0 million and \$100.1 million of gross proceeds in connection with the issuance of its Series A convertible preferred stock and Series B convertible preferred stock (see Note 6), respectively, and which management believes is sufficient to fund its operating expenses and capital expenditure requirements for at least 12 months from the issuance date of these financial statements. The future viability of the Company beyond that point is dependent on its ability to raise additional capital to finance operations. The Company is seeking to complete an initial public offering (“IPO”) of its common stock. In the event the Company does not complete an IPO, the Company expects to seek additional funding through private equity financings, debt financings or other capital sources, including collaborations with other companies or other strategic transactions. The Company may not be able to obtain financing on acceptable terms, or at all. The terms of any financing may adversely affect the holdings or the rights of the Company’s stockholders.

If the Company is unable to obtain funding, the Company will be forced to delay, reduce or eliminate some or all of its research and development programs which could adversely affect its business prospects, or the Company may be unable to continue operations. Although management continues to pursue these plans, there is no assurance that the Company will be successful in obtaining sufficient funding on terms acceptable to the Company to fund continuing operations, if at all.

**Risks and Uncertainties**—The Company is subject to risks and uncertainties common to early-stage companies in the biopharmaceutical industry, including, but not limited to, successful discovery and development of its product candidates, development by competitors of new technological innovations, dependence on key personnel, the ability to attract and retain qualified employees, protection of proprietary technology, compliance with governmental regulations, the ability to secure additional capital to fund operations, and commercial success of its product candidates. Any of the Company’s current product candidates and future product candidates that it may develop will require extensive nonclinical and clinical testing and regulatory approval prior to commercialization. These efforts require significant amounts of additional capital, adequate personnel, infrastructure and extensive compliance-reporting capabilities. Even if the Company’s product development efforts are successful, it is uncertain when, if ever, the Company will realize significant revenue from product sales.

## 2. SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES

**Use of Estimates**—The preparation of the financial statements in accordance with U.S. GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities and disclosure of contingent assets and liabilities at the date of the financial statements, and the reported amounts of income and expense during the reporting period. The most significant estimates relate to the accruals of research and development costs, accruals of research contract costs, assumptions used to estimate the fair value of its stock option awards, and determination of fair value of the Company’s common stock. Management evaluates its estimates and assumptions on an ongoing basis using historical experience and other factors, including the current economic environment, and makes adjustments when facts and circumstances dictate. These estimates are based on information available as of the date of the financial statements; therefore, actual results could differ from those estimates.

**Cash and Cash Equivalents and Restricted Cash**—The Company considers all highly liquid investments with a remaining maturity when purchased of three months or less to be cash equivalents. Cash equivalents are reported at carrying values, which approximated their fair values and are based on quoted prices in active markets for identical securities. At December 31, 2022 and December 31, 2021, the Company’s cash equivalents are in money market funds. As of December 31, 2022, and 2021, cash consists of cash on deposit with U.S. banks denominated in U.S. dollars, and restricted cash consists of cash on deposit with U.S. banks denominated in U.S. dollars for which its use is restricted and is classified as non-current in the Company’s consolidated balance sheet according to the timing of maturity, and is associated with collateral for letters of credit issued in connection with its operating lease right-of-use and finance lease right-of-use assets and corresponding lease liabilities. Cash is stated at its historical carrying amount, which approximates fair value due to its short-term nature. Restricted cash is stated at its historical carrying amount, which approximates fair value. The Company regularly maintains cash and cash equivalents and restricted cash balances with financial institutions that exceed Federal Deposit Insurance Corporation (“FDIC”) insurance limits.

**Net Loss per Share**—The Company follows the two-class method when computing net income (loss) per common share as the Company has issued shares that meet the definition of participating securities. The two-class method determines net income (loss) per common share for each class of common and participating securities according to dividends declared or accumulated and participation rights in undistributed earnings. The two-class method requires income (loss) available to common stockholders for the period to be allocated between common and participating securities based upon their respective rights to receive dividends as if all income for the period had been distributed. The Company considers its convertible preferred stock to be participating securities as, in the event a dividend is paid on common stock, the holders of these securities would be entitled to receive dividends on a basis consistent with the common stockholders. The Company also considers the shares issued upon the early exercise of stock options that are subject to repurchase to be participating securities.

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because holders of such shares have non-forfeitable dividend rights in the event a dividend is paid on common stock. There is no allocation required under the two-class method during periods of loss since the participating securities do not have a contractual obligation to share in the losses of the Company.

Basic net income (loss) per common share is computed by dividing the net income (loss) per common share by the weighted-average number of common shares outstanding for the period. Diluted net income (loss) per common share is computed by adjusting net income (loss) to reallocate undistributed earnings based on the potential impact of dilutive securities. Diluted net loss per common share is computed by dividing the diluted net loss by the weighted-average number of common shares outstanding for the period, including potential dilutive common shares. For purpose of this calculation, convertible preferred stock are considered potential dilutive common shares.

In periods in which the Company reported a net loss, diluted net loss per common share was the same as basic net loss per common share, since dilutive common shares were not assumed to have been issued if their effect was anti-dilutive. During the years ended December 31, 2022 and 2021, the Company excluded 85,495,722 potential common shares related to the conversion of Series A convertible preferred stock and 58,157,823 potential common shares related to the conversion of Series B convertible preferred stock, and during the years ended December 31, 2022 and 2021 excluded 20,300,861, and 19,922,318 potential common shares, respectively, related to the exercise of stock options outstanding, from the computation of diluted net loss per common share because including them would have had an anti-dilutive effect and the Company reported net losses for those years.

**Deferred Offering Costs**—The Company capitalizes certain legal, accounting and other third-party fees that are directly associated with in-process equity financings as deferred offering costs until such financings are consummated. After consummation of the equity financing, these costs are recorded as a reduction of the proceeds from the offering, either as a reduction of the carrying value of preferred stock or in stockholders' deficit as a reduction of additional paid-in capital generated as a result of the offering. Should the in-process equity financing be abandoned, the deferred offering costs would be expensed immediately as a charge to operating expenses in the consolidated statement of operations and comprehensive loss. The Company had no deferred offering costs recorded as of December 31, 2022. As of December 31, 2021, the Company had deferred offering costs totaling \$0.2 million in other current assets in its consolidated balance sheet, which were charged to general and administrative expenses during the year ended December 31, 2022.

**Accounts Receivable and Allowance for Doubtful Accounts**—Accounts receivable are recorded net of allowances for doubtful accounts. The Company does not have any accounts receivable but in certain cases had a grant receivable balance. For the Company to be in a receivable position related to grants, the cumulative total spend on grants would have to exceed the cumulative cash receipts in the event there remains additional award money to be collected.

**Classification of Convertible Preferred Stock**—The holders of the series A and series B convertible preferred stock have certain liquidation rights in the event of a deemed liquidation that, in certain situations, are not solely within the control of the Company and would call for the redemption of the then outstanding series A convertible preferred stock. Therefore, the series A convertible preferred stock are classified outside of the stockholders' deficit on the Company's consolidated balance sheets. The carrying value of the convertible preferred stock is not subsequently remeasured to the redemption value until the contingent redemption events are considered to be probable of occurring.

**Revenue Recognition**—On January 1, 2020, the Company adopted ASU 2014-09, *Revenue from Contracts with Customers* ("Topic 606"), on a modified retrospective basis. Topic 606 establishes a principle for recognizing revenue upon the transfer of promised goods or services to customers, in an amount that reflects the expected consideration received in exchange for those goods or services. The standard also provides guidance on the recognition of costs related to obtaining and fulfilling customer contracts. Additionally, the standard requires



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disclosures regarding the nature, amount, timing and uncertainty of revenue and cash flows arising from contracts with customers. To determine revenue recognition for arrangements that the Company determines are within the scope of Topic 606, the Company performs the following five steps: (i) identify the contract(s) with a customer; (ii) identify the performance obligations in the contract; (iii) determine the transaction price; (iv) allocate the transaction price to the performance obligations in the contract; and (v) recognize revenue when (or as) the Company satisfies a performance obligation. The Company only applies the five-step model to contracts when it is probable that it will collect the consideration it is entitled to in exchange for the goods and services it transfers to the customer. At contract inception, the Company assesses the goods or services promised within each contract that falls under the scope of Topic 606, determines those that are performance obligations and assesses whether each promised good or service is distinct. The Company then recognizes as revenue the amount of the transaction price that is allocated to the respective performance obligation when (or as) the performance obligation is satisfied. As of December 31, 2022, and December 31, 2021, there is no revenue from contracts with customers.

**Grant Revenue**—Grants received are recognized as grant revenue in the statements of operations as and when they are earned for the specific research and development projects for which these grants are designated. Grant payments received in excess of grant revenue earned are recognized as deferred grant revenue on the balance sheet and grant revenue earned in excess of grant payments received is recognized as grant receivable on the balance sheets.

**Research and Development**—Research and development costs are expensed as incurred. Research and development expenses consist of costs incurred to discover, research, and develop drug candidates, including personnel expenses, allocated facility-related and depreciation expenses and third-party license fees in connection with nonclinical and preclinical studies. Costs incurred in obtaining technology licenses are charged immediately to research and development expense if the technology licensed has not reached technological feasibility and has no alternative future use.

**General and Administrative**—General and administrative expenses consist primarily of the cost of employees to engage in corporate functions, such as finance and accounting, information technology, human resources, legal and executive management. General and administrative expense also includes rent occupancy costs, office expenses, recruiting expenses, entertainment allocations, depreciation and amortization, other general overhead costs, insurance premiums, professional service fees, and cost related to regulatory and litigation matters.

**Acquisitions**—The Company evaluates acquisitions under the accounting framework in Accounting Standards Codification (“ASC”) Topic 805, Business Combinations, to determine whether the transaction is a business combination or an asset acquisition. In determining whether an acquisition should be accounted for as a business combination or an asset acquisition, the Company first performs a screen test to determine whether substantially all of the fair value of the gross assets acquired is concentrated in a single identifiable asset or a group of similar identifiable assets. If this is the case, the acquired set is not deemed to be a business and is instead accounted for as an asset acquisition. If this is not the case, the Company then further evaluates whether the acquired set includes, at a minimum, an input and a substantive process that together significantly contribute to the ability to create outputs. If so, the Company concludes that the acquired set is a business.

**Stock Based Compensation Expense**—The Company accounts for stock-based payment awards granted to employees and non-employees as stock-based compensation expense at fair value. The Company’s stock-based payments include stock options and grants of common stock, including common stock subject to vesting. The measurement date for employee awards is the date of grant, and stock-based compensation costs are recognized as expense over the employees’ requisite service period, which is the vesting period, on a straight-line basis. The measurement date for non-employee awards is the date of grant without changes in the fair value of the award. Stock-based compensation costs for non-employees are recognized as expense over the vesting period on a

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straight-line basis. Stock-based compensation expense is classified in the accompanying statement of operations based on the function to which the related services are provided. As the Company is permitted to repurchase shares legally issued for unvested stock options exercised at their exercise price, (i) cash received for unvested stock options exercised is recorded as a deposit liability as a reclassification from additional paid-in capital in the Company's consolidated balance sheet, which is relieved to additional paid-in capital as such awards vest, (ii) stock based compensation expense is recognized over the requisite service period for each such award, and (iii) the amount and number of shares of common stock outstanding in the Company's consolidated balance sheet and consolidated statement of convertible preferred stock and stockholders' deficit are reduced until such awards vest. The Company recognizes stock-based compensation expense for the portion of awards that have vested. Forfeitures are recorded as they occur.

The fair value of each stock option grant is estimated on the date of grant using the Black-Scholes option-pricing model. The Company has historically been a private company and lacks company-specific historical and implied volatility information. Therefore, it estimates its expected stock volatility based on the historical volatility of a publicly traded set of peer companies and expects to continue to do so until such time as it has adequate historical data regarding the volatility of its own traded stock price. The expected term of the Company's stock options has been determined utilizing the "simplified" method for awards that qualify as "plain-vanilla" options. The risk-free interest rate is determined by reference to the U.S. Treasury yield curve in effect at the time of grant of the award for time periods approximately equal to the expected term of the award. The expected dividend yield is zero as the Company has never paid cash dividends on its common stock and does not expect to pay any cash dividends in the foreseeable future.

The Company recognizes stock-based compensation expense on a straight-line basis over the requisite service or vesting period. The Company recognizes the cumulative effect of changes in the probability outcomes in the period in which the changes occur.

**Income Taxes**—Prior to November 20, 2020, the Company was an LLC entity and elected to be treated under the Partnership provisions of the Internal Revenue Code. Accordingly, the LLC entity was not viewed as a tax-paying entity in any jurisdiction and all income and deductions of the LLC entity flowed through to the individual members and therefore no income taxes were recorded by the Company. On November 20, 2020, the Company elected to convert to a C Corporation.

The Company accounts for income taxes under the asset and liability method pursuant to ASC 740, Income Taxes. Under this method, the Company recognizes deferred tax assets and liabilities for the future tax consequences attributable to differences between the financial statement carrying amounts of existing assets and liabilities and their respective tax bases and operating loss and tax credit carryforwards. Deferred tax assets and liabilities are measured using enacted tax rates expected to apply to taxable income in the years in which those temporary differences are expected to be recovered or settled. The effect on deferred tax assets and liabilities of a change in tax rates is recognized in income in the period that includes the enactment date. A valuation allowance is recorded for deferred tax assets if it is more likely than not that some portion or all of the deferred tax assets will not be realized based on all available positive and negative evidence. Under the Tax Cuts and Jobs Act of 2017, the Company is required to capitalize, and subsequently amortize, research and development expenses over five years for research activities conducted in the U.S. and over fifteen years for research activities conducted outside of the U.S commencing in 2022. The capitalization of research and development expenses during the year ended December 31, 2022 resulted in a decrease to our current year operating loss generation (see Note 11). As of December 31, 2022, the Company continues to maintain a full valuation allowance against its deferred tax assets.

The Company recognizes a tax benefit only if it is more likely than not the tax position will be sustained on examination by the local taxing authorities, based on the technical merits of the position. The tax benefits

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recognized in the financial statements from such positions are then measured based on the largest benefit greater than 50% likelihood of being realized upon settlement with the related tax authority. The changes in recognition or measurement are reflected in the period in which the change in judgment occurs. As of December 31, 2022, the Company has not identified any uncertain tax positions.

The Company records interest and penalties related to unrecognized tax benefits in the provision for income taxes.

**Fair Value Measurements**—Fair value is defined as the price that would be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants at the measurement date. The three levels of inputs that may be used to measure fair value are described below:

*Level 1*—Inputs are quoted prices (unadjusted) in active markets for identical assets or liabilities that the Company has the ability to access at the measurement date.

*Level 2*—Inputs are observable, unadjusted quoted prices in active markets for similar assets or liabilities, unadjusted quoted prices for identical or similar assets or liabilities in markets that are not active, or other inputs that are observable or can be corroborated by observable market data for substantially the full term of the related assets or liabilities.

*Level 3*—Inputs are unobservable inputs for the asset or liability.

**Concentrations of Credit Risk**—Financial instruments that potentially subject the Company to concentrations of credit risk consist primarily of cash and cash equivalents and restricted cash. The Company's cash balances exceed those that are federally insured. The majority of the Company's cash and cash equivalents and restricted cash is held with a large financial institution that management believes to be of high credit quality. To date, the Company has not recognized any losses caused by uninsured balances.

**Segment Information**—The Company manages its operations as a single segment for the purposes of assessing performance and allocating resources. The Company is focused on preclinical and clinical stage gene therapies, and specifically on hereditary and acquired diseases of high unmet need. The Company's chief operating decision maker reviews the Company's financial information on an aggregated basis for purposes of assessing performance and allocating resources. All assets are in the United States. The Company has not earned any product revenue through December 31, 2022.

**Property and Equipment**—Property and equipment are stated at cost less accumulated depreciation and amortization. Depreciation and amortization is recognized using the straight-line method over the estimated useful life of each asset. Such costs are periodically reviewed for recoverability when impairment indicators are present. Such indicators include, among other factors, unused capacity, market value declines, and technology obsolescence. Recorded values of asset groups of equipment that are not expected to be recovered through undiscounted future net cash flows are written down to current fair value, which generally is determined from estimated discounted future net cash flows (assets held for use) or net realizable value (assets held for sale).

Costs for capital assets not yet placed into service are capitalized as construction-in-progress and depreciated once placed into service. Upon retirement or sale, the cost of assets disposed of and the related accumulated depreciation and amortization are removed from the accounts and any resulting gain or loss is included in loss from operations. Expenditures for repairs and maintenance that do not improve or extend the life of the respective assets are charged to expense in the period incurred. See Note 5 for information regarding the Company's capitalized operating and finance lease right-of-use assets.

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The following is the summary of property and equipment and related accumulated depreciation as of December 31, 2022, and December 31, 2021:

	Useful life	December 31, 2022	December 31, 2021
Internal use software	3 years	245,195	—
Furniture and fixtures	5 years	375,311	—
Lab equipment	7 years	278,749	187,671
Leasehold improvements	7 years	231,939	—
Total property and equipment		1,131,194	187,671
Less: accumulated depreciation and amortization		(132,844)	(13,405)
Total property and equipment, net		\$ 998,350	\$ 174,266

**Leases**—In accordance with ASC 842, Leases, which the Company adopted in 2022, the Company determines if an arrangement is or contains a lease at inception. A contract is or contains a lease if the contract conveys the right to control the use of an identified asset for a period of time in exchange for consideration. The Company classifies leases at the lease commencement date as operating or finance leases and records a right-of-use asset and a lease liability on the consolidated balance sheet for all leases with an initial lease term of greater than 12 months. Leases with an initial term of 12 months or less are not recorded in the balance sheet, but payments are recognized as expense on a straight-line basis over the lease term. The Company has elected not to recognize leases with terms of 12 months or less.

A lease qualifies as a finance lease if any of the following criteria are met at the inception of the lease: (i) there is a transfer of ownership of the leased asset to the Company by the end of the lease term, (ii) the Company holds an option to purchase the leased asset that it is reasonably certain to exercise, (iii) the lease term is for a major part of the remaining economic life of the leased asset, (iv) the present value of the sum of lease payments equals or exceeds substantially all of the fair value of the leased asset, or (v) the nature of the leased asset is specialized to the point that it is expected to provide the lessor no alternative use at the end of the lease term. All other leases are recorded as operating leases.

The Company enters into contracts that contain both lease and non-lease components. Non-lease components may include maintenance, utilities, and other operating costs. The Company combines the lease and non-lease components of fixed costs in its lease arrangements as a single lease component. Variable costs, such as utilities or maintenance costs, are not included in the measurement of right-of-use assets and lease liabilities, but rather are expensed when the event determining the amount of variable consideration to be paid occurs.

Operating lease right-of-use assets and lease liabilities are recognized at the lease commencement date based on the present value of the lease payments over the lease term using the discount rate implicit in the lease. If the rate implicit is not readily determinable, the Company utilizes an estimate of its incremental borrowing rate based upon the available information at the operating lease commencement date. Operating lease right-of-use assets are further adjusted for prepaid or accrued lease payments. Operating lease payments are expensed using the straight-line method as an operating expense over the lease term. The Company's lease terms may include options to extend or terminate the lease when it is reasonably certain that the Company will exercise that option. Finance lease right-of-use assets and lease liabilities are recognized upon receipt of the assets based on their fair values, and are amortized to depreciation expense using the straight-line method over the estimated useful lives of the related assets. Finance lease liability payments are bifurcated into (i) a portion that is recorded as interest expense and (ii) a portion that reduces the finance liability associated with the lease, according to the interest rates implicit in the leases.

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Certain of the Company's leases include options to extend or terminate the lease. The amounts determined for the Company's right-of-use assets and lease liabilities generally do not assume that renewal options or early-termination provisions, if any, are exercised, unless it is reasonably certain that the Company will exercise such options.

**Recent Accounting Pronouncements Not Yet Adopted**—In June 2016, the FASB issued ASU 2016-13, *Financial Instruments-Credit Losses* (“Topic 326”): *Measurement of Credit Losses on Financial Instruments* (“ASU 2016-13”), which requires the measurement and recognition of expected credit losses for financial assets held at amortized cost. ASU 2016-13 replaces the existing incurred loss impairment model with an expected loss methodology, which will result in more timely recognition of credit losses. ASU 2016-13 and ASU 2022-02 are effective for annual private company reporting periods, and interim periods within those years, beginning after December 15, 2022. The Company is currently in the process of evaluating the impact of the adoptions of ASU 2016-13 and ASU 2022-02 but does not expect a material impact to its consolidated financial statements or disclosures as it does not currently maintain financial assets held at amortized cost.

In October 2021, the FASB issued ASU 2021-08, *Business Combinations* (“Topic 805”): *Accounting for Contract Assets and Contract Liabilities from Contracts with Customers* (“ASU 2021-08”). ASU 2021-08 requires an acquirer in a business combination to recognize and measure deferred revenue from acquired contracts using the revenue recognition guidance in Accounting Standards Codification Topic 606, rather than the prior requirement to record deferred revenue at fair value. ASU 2021-08 allows for immediate adoption on a retrospective basis for all business combinations that have occurred since the beginning of the annual period that includes the interim period of adoption. ASU 2021-08 is effective for annual private company reporting periods, and interim periods within those years, beginning after December 15, 2023. The Company is currently in the process of evaluating the impact of the adoption of ASU 2021-08 but does not expect a material impact to its consolidated financial statements or disclosures as it does not currently have revenue or deferred revenue from acquired contracts.

### 3. ACQUISITIONS

#### *Stelios Therapeutics, Inc.*

Stelios Therapeutics, Inc. (“Stelios”) is an early-stage company developing novel adeno-associated virus (“AAV”) based gene therapies for rare cardiac conditions. On July 16, 2021, the Company acquired 100% of the outstanding stock of Stelios, for initial consideration of \$7.0 million, with up to an aggregate of \$20.5 million payable upon the achievement of certain development milestones and concurrently entered into consulting agreements with three consultants who assist with research on Stelios' therapies. The Company accounted for the acquisition of Stelios as an asset acquisition pursuant FASB ASC 805, *Business Combinations*. The Company acquired in-process research and development (“IPR&D”) assets from Stelios related to arrhythmogenic right ventricular cardiomyopathy and TNNI3-associated hypertrophic cardiomyopathy programs. The fair value of the IPR&D acquired was \$7.0 million and was charged to research and development expense as it had no alternative future use at the time of the acquisition. In the third quarter of 2022, a development milestone was achieved resulting in a \$2.0 million payment to the selling shareholders of Stelios.

#### 4. FAIR VALUE MEASUREMENTS

The following table presents information about the Company’s financial assets and liabilities measured at fair value on a recurring basis and indicates the level of the fair value hierarchy utilized to determine such fair values:

	As of December 31, 2022:			
	Level 1	Level 2	Level 3	Total
<b>Assets:</b>				
Cash equivalents (money market)	\$ 74,802,103	\$ —	\$ —	\$ 74,802,103
	\$ 74,802,103	\$ —	\$ —	\$ 74,802,103

	As of December 31, 2021:			
	Level 1	Level 2	Level 3	Total
<b>Assets:</b>				
Cash equivalents (money market)	\$128,216,205	\$ —	\$ —	\$128,216,205
	\$128,216,205	\$ —	\$ —	\$128,216,205

There have been no transfers between fair value levels during the years ended December 31, 2022 and 2021. The carrying values of prepaid expenses, other current assets, accounts payable and accrued expenses approximate their fair values due to the short-term nature of these assets and liabilities.

##### *Preferred stock tranche obligation*

In 2020, the Company obtained series A funding from investors with two subsequent milestone closing dates. Each of the two milestones represented a separate tranche obligation as they could have been exercised separately and an investor could have transferred part of any rights under the agreement, including the right/obligation to participate in an individual tranche. The tranche obligation was recorded at fair value at the issuance date and a portion of the proceeds from the first closing equal to that amount was allocated to the tranche obligation. The Company determined that its obligation to issue additional shares at a fixed price (i.e. the issuance price) in subsequent tranches (collectively, the “Milestones”; see Note 6) following the initial closing of the series A convertible preferred stock financings represented two freestanding financial instruments (together, the “Tranche Obligation”). The Tranche Obligation was previously carried at fair value determined according to Level 3 inputs in the fair value hierarchy. On July 30, 2021, the Company waived all Milestone events causing the Milestone closings to occur. This resulted in the derecognition of the Tranche Obligation liabilities and the recognition of the cash proceeds received. The Company determined that the value of the Tranche Obligations at issuance would be equivalent at settlement and, as such, no mark-to-market adjustment was recorded upon settlement.

The following Level 3 table provides a roll forward of the aggregate fair value of the Company’s Tranche Obligation:

	December 31, 2022	December 31, 2021
Balance as of January 1	\$ —	\$ 3,010,000
Extinguishment of Tranche Obligation	—	(3,010,000)
Balance as of December 31	\$ —	\$ —

## 5. LEASES

Effective January 1, 2022, the Company adopted the US GAAP accounting rules in ASC Topic 842, *Leases* (“ASC 842”). The Company elected to follow the package of practical expedients provided under the transition guidance within ASC 842, and accordingly, elected to combine lease and non-lease components for its real estate and financed equipment leases. Upon commencement of its real estate lease, the Company recorded an operating lease right-of-use asset and related current and non-current lease liabilities in its consolidated balance sheet, which were recognized based on the present value of remaining lease payments over the lease term. Upon receipt of the financed equipment, the Company recorded finance lease right-of-use assets and lease liabilities in its consolidated balance sheet, which were based on the fair values of the equipment financed with implicit interest rates provided in the terms of the lease agreements. As the Company’s operating lease right-of-use asset does not provide an implicit rate, an estimated incremental borrowing rate was used based on the information available at the adoption date in determining the present value of lease payments. Operating lease right-of-use asset expense is recognized on a straight-line basis over the lease term. Variable lease costs such as common area costs and other operating costs are expensed as incurred. Leases with an initial term of 12 months or less are not recorded on the Company’s balance sheet.

### *Operating lease right-of-use asset*

In January 2022, the Company entered into a lease agreement for an office facility and laboratory space in New York, New York that commenced in April 2022 and ends in July 2029 and there is an additional five-year option to extend the lease beyond July 2029 at the then prevailing effective market rental rate. Upon commencement of this lease, the Company recorded operating lease right-of-use assets and operating lease liabilities of approximately \$11.6 million based on the present value of payments over the lease term using an estimated incremental borrowing rate of 8.53%. In connection with the Company’s lease of office space and laboratory space, the Company provided a security deposit to the landlord in the form of a letter of credit totaling \$1.2 million. The cash collateralizing the letter of credit is included in long-term restricted cash in the Company’s consolidated balance sheet as of December 31, 2022. This lease was classified as an operating lease in accordance with the provisions of Topic 842. The Company did not recognize any right-of-use assets and lease liabilities associated with the potential option to renew or extend. The Company’s operating lease agreement does not contain any significant residual value guarantees or restrictive covenants.

Components of the operating lease are as follows:

<b>Year ended December 31</b>	<b>2022</b>
Operating lease expense	\$ 1,610,561
Variable lease expense	184,788
Total operating lease expense	\$ 1,795,350
Other information	
Remaining lease term and payment term (in years)	6.6
Incremental borrowing rate	8.53%
Cash paid for amounts included in the measurement of lease liabilities, included in operating cash flows	\$ 1,031,953

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The following table provides a reconciliation of the Company's remaining undiscounted contractual rent obligations due within each year ended December 31 to the operating lease liabilities recognized as of December 31, 2022:

	<b>Operating leases</b>
Years ended December 31	
2023	\$ 2,307,415
2024	2,099,723
2025	2,152,216
2026	2,206,022
2027	2,261,172
Thereafter	3,689,208
Total lease payments	14,715,758
Less: present value adjustment	(3,429,005)
Total operating lease liabilities	<u>\$ 11,286,752</u>
Included in the consolidated balance sheet:	
Current portion of lease liabilities—operating	2,287,109
Non-current portion of lease liabilities—operating	8,999,643
Total operating lease liabilities	<u>\$ 11,286,752</u>

### *Equipment finance leases*

Commencing in April 2022, the Company leases certain laboratory equipment under financing arrangements accounted for as finance leases classified in the Company's balance sheet as finance lease liabilities with related right-of-use assets recorded and depreciated on a straight-line basis over the estimated useful life of 7 years. The total amount of equipment finance lease right-of-use assets capitalized was approximately \$2.2 million during the year ended December 31, 2022. The total gross, accumulated amortization, and net book values of equipment finance lease right-of-use assets capitalized under such finance lease arrangements at December 31, 2022 were approximately \$2.2 million, \$0.1 million and \$2.1 million, respectively. The equipment finance leases entered into also included approximately \$0.1 million of research and development laboratory supplies received that were not capital in nature and were recorded to research and development expense in the Company's consolidated statement of operations and comprehensive loss during the year ended December 31, 2022. Under the terms of the equipment finance lease agreements executed through the issuance date of these financial statements, the principal balances plus interest for the equipment are to be repaid in full after 60 monthly installments following lease commencement, with lease commencement dates ranging from April 1, 2022 to January 1, 2023, annual imputed interest rates ranging from 7.90% to 9.30%, and monthly installment payment amounts ranging from approximately \$7,000 to \$18,000. As of December 31, 2022, the total aggregate monthly installment payment amount was approximately \$45,000 for equipment finance lease agreements executed through the issuance date of these financial statements. Total equipment finance right of use assets and lease liabilities of approximately \$534,000 were recorded in the Company's consolidated balance sheet at December 31, 2022 for which lease periods had not yet commenced. In connection with the Company's leases of laboratory equipment, the Company provided a security deposit to the lessor in the form of a letter of credit totaling \$1.9 million and assigned all rights and interests in the equipment to the lessor. The cash collateralizing the letter of credit is included in long-term restricted cash in the Company's consolidated balance sheet as of December 31, 2022.



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Components of the equipment finance leases are as follows:

<b>Year ended December 31</b>	<b>2022</b>
Finance lease amortization of right-of-use assets	\$ 147,807
Interest on finance lease liabilities	46,300
<b>Total finance lease expense</b>	<b>\$ 194,107</b>
Other information	
Weighted Average Remaining Lease Payment Term (in years)	4.6
Weighted Average Remaining Amortization Term (in years)	6.6
Weighted Average Effective Interest Rate	8.66%
Cash paid for amounts included in the measurement of lease liabilities, included in financing cash flows	\$ 240,261

The following table provides a reconciliation of the Company's remaining equipment finance lease obligations due within each year ending December 31 to the equipment finance lease liabilities recognized at December 31, 2022, including estimated amounts due for equipment received by December 31, 2022 under equipment finance leases anticipated to be executed following the issuance date of these financial statements:

	<b>Equipment finance leases</b>
<b>Years ended December 31</b>	
2023	\$ 549,963
2024	579,734
2025	579,734
2026	579,734
2027	357,653
Thereafter	11,591
Total lease payments	2,658,408
Less: imputed interest	(463,216)
Total finance lease liabilities	\$ 2,195,192
Included in the consolidated balance sheet:	
Current portion of lease liabilities—finance	547,294
Non-current portion of lease liabilities—finance	1,647,898
Total finance lease liabilities	\$ 2,195,192

## **6. PREFERRED STOCK**

As of December 31, 2022, the Company's certificate of incorporation authorized the Company to issue 143,653,546 million shares of series A and series B convertible preferred stock at par value of \$0.0001.

On November 20, 2020, the Company entered into a series A convertible preferred stock purchase agreement (the "Series A Purchase Agreement") whereby the Company issued 29,999,996 shares of series A convertible preferred stock at \$1 per share for an aggregate purchase price of \$30.0 million and incurred \$0.2 million of issuance costs. Additionally, on November 20, 2020, the Company issued additional series A convertible preferred stock to the holders of the Company's Notes (the "Noteholders"). In connection with the conversion, the Company issued 4,495,719 shares of series A convertible preferred stock.

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Pursuant to the Series A Purchase Agreement, the shareholders were eligible to receive additional shares at the date of certain contractual Milestone events. The first Milestone event was the successful Federal Drug Administration (“FDA”) clearance to proceed into human clinical studies of one of the Company’s therapeutics; the second Milestone event was the successful dosing of the therapeutic in five human patients with all relevant safety data supportive of future development.

On July 30, 2021, the Company’s board of directors voted to waive the Milestones related to both the second and third tranches of the series A convertible preferred stock and issued 50,999,997 shares for an aggregate consideration of \$51.0 million.

On August 10, 2021, the Company entered into a series B convertible preferred stock purchase agreement whereby the Company issued 58,157,823 shares of series B convertible preferred stock at \$1.72 per share for an aggregate purchase price of approximately \$100.1 million.

As of December 31, 2022 and 2021, preferred stock consisted of the following:

	<b>Convertible Preferred Stock Authorized</b>	<b>Convertible Preferred Stock Issued and Outstanding</b>	<b>Proceeds Received</b>	<b>Liquidation Preference</b>	<b>Common Stock Issuable Upon Conversion</b>
Series A convertible preferred stock	85,495,722	85,495,722	\$ 85,495,722	\$ 85,495,722	85,495,722
Series B convertible preferred stock	58,157,824	58,157,823	100,059,953	100,059,953	58,157,825
	143,653,546	143,653,545	\$ 185,555,675	\$ 185,555,675	143,653,547

As of December 31, 2022, the rights and privileges of the holders of the convertible preferred stock were as follows:

*Voting*—Holders of preferred stock shall vote together with the holders of common stock as a single class and on an as converted to common stock basis.

*Dividends*—The holders of shares of series A convertible preferred stock are entitled to receive, as declared by the Company’s board of directors, dividends at the rate of 8% per share of preferred stock, prior and in preference to any declaration or payment of any other dividend (other than dividends on shares of common stock payable in shares of common stock).

*Deemed liquidation event*—Each of the following events shall be considered a “Deemed Liquidation Event” unless the holders of a majority of the outstanding shares of preferred stock elect otherwise by written notice sent to the Company at least five business days prior to the effective date of any such event; (a) A merger or consolidation in which the Company is a constituent party or a subsidiary of the Company is a constituent party and the Company issues shares of its capital stock pursuant to such merger or consolidation. Except any such merger or consolidation involving the Company or a subsidiary in which the shares of capital stock of the Company outstanding immediately prior to such merger or consolidation continue to represent, or are converted into or exchanged for shares of capital stock that represent, immediately following such merger or consolidation, at least a majority of the capital stock of (1) the surviving or resulting company; or (2) if the surviving or resulting company is a wholly owned subsidiary of another company immediately following such merger or consolidation, the parent company of such surviving or resulting corporation; or (b) (1) the sale, lease, transfer, exclusive license or other disposition, in a single transaction or series of related transactions, by the Company or any subsidiary of the Company of all or substantially all the assets of the Company and its subsidiaries taken as a

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whole, or (2) the sale or disposition (whether by merger, consolidation or otherwise, and whether in a single transaction or a series of related transactions) of one or more subsidiaries of the Company if substantially all of the assets of the Company and its subsidiaries taken as a whole are held by such subsidiary or subsidiaries, except where such sale, lease, transfer, exclusive license or other disposition is to a wholly owned subsidiary of the Company.

*Liquidation preference*—In the event of any voluntary or involuntary liquidation, dissolution, or winding up of the Company, the holders of shares of preferred stock shall be entitled to be paid out of the assets of the Company or, in the case of a Deemed Liquidation Event, out of the consideration payable to stockholders or the available proceeds, before any payment shall be made to the holders of common stock, an amount per share equal to the series B convertible preferred stock issue price of \$1.72 per share and the series A convertible preferred stock issue price of \$1.00 per share, plus any dividends declared but unpaid. If upon any such liquidation, dissolution or winding up of the Company or Deemed Liquidation Event, the assets of the Company available for distribution to its stockholders shall be insufficient to pay the holders of shares of preferred stock the full amount to which they shall share ratably in any distribution of the assets available for distribution in proportion to the respective amounts which would otherwise be payable in respect of the shares held by them upon such distribution if all amounts payable on or with respect to such shares were paid in full (collectively, the “Liquidation Preference Per Convertible Preferred Stock”).

In the event that there are additional assets to be distributed, the holders of the series A convertible preferred stock and series B convertible preferred stock will share in the distribution along with common stockholders as if the shares of the series A convertible preferred stock and series B convertible preferred stock had converted to common stock immediately prior to the distribution up to an amount taken in consideration with the Liquidation Preference Per Convertible Preferred Stock not to exceed \$3.00 per share allocated to series A convertible preferred stockholders and \$5.16147 per share allocated to series B convertible preferred stockholders.

*Redemption*—The preferred stock does not contain a mandatory redemption provision. Upon a Deemed Liquidation Event, the holders will be paid their preference amounts according to their priority. That is, the preferred stockholders will be entitled to an amount up to \$1.00 per share, plus any accrued dividends. Accretion to redemption value will only be required if a Deemed Liquidation Event is probable.

*Conversion*—In the event that any Noteholder, or an affiliate of any Noteholder, the investor of Notes did not participate in any Milestones then each share of preferred stock held by the Noteholders would have automatically been converted into the number of shares of common stock equal to the \$1.00 per share divided by the conversion price in effect immediately prior to the consummation of the Milestones closings multiplied by ten.

Upon either (a) the closing of the sale of shares of common stock to the public at a price of at least \$3.00 per share (subject to appropriate adjustment in the event of any stock dividend, stock split, combination or other similar recapitalization with respect to the common stock), in a firm-commitment underwritten public offering pursuant to an effective registration statement under the Securities Act of 1933, as amended, resulting in at least \$50.0 million of gross proceeds to the Company and in connection with the offering of the common stock listed for trading on the Nasdaq Stock Market’s National Market, the New York Stock Exchange or another exchange or marketplace approved by the Company’s board of directors, including the approval of at least two (2) preferred directors (a “Qualified IPO”) or (b) the date and time, or the occurrence of an event specified by vote or written consent of the Noteholders, then (i) all outstanding shares of preferred stock shall automatically be converted into shares of common stock, at the then effective conversion rate.

## 7. COMMON STOCK

As of December 31, 2022, the Company's certificate of incorporation authorized the Company to issue 192,020,973 shares of common stock with a par value of \$0.0001 per share. The voting, dividend and liquidation rights of the holders of the Company's common shares are subject to and qualified by the rights, powers and preferences of the holders of the convertible preferred stock set forth above.

Each common share entitles the holder to one vote on all matters submitted to a vote of the Company's stockholders. Common stockholders are entitled to receive dividends, if any, as may be declared by the Company's board of directors, subject to the preferential dividend rights of the convertible preferred stock. Through December 31, 2022, no cash dividends have been declared or paid by the Company.

The Company has reserved the following number of shares of common stock for the exercise of outstanding stock options, future issuance of stock-based awards, and conversion of outstanding shares of convertible preferred stock:

	Year ended December 31,	
	2022	2021
Shares of series A convertible preferred stock	85,495,722	85,495,722
Shares of series B convertible preferred stock	58,157,823	58,157,823
Options to purchase shares of common stock	20,300,861	19,922,318
Shares available for issuance under the 2021 Plan	10,599,419	11,699,283
Total shares of common stock reserved for future issuance	174,553,825	175,275,146

## 8. NET LOSS PER SHARE

Basic and diluted net loss per common share attributable to common stockholders was calculated as follows (in thousands, except share and per share amounts):

	Year ended December 31,	
	2022	2021
Numerator:		
Net loss attributable to common stockholders	(59,276,727)	(50,621,713)
Denominator:		
Weighted-average common shares outstanding, basic and diluted	17,272,352	16,449,814
Net loss per share attributable to common stockholders, basic and diluted	\$ (3.43)	\$ (3.08)

## 9. ACCRUED EXPENSES AND OTHER CURRENT LIABILITIES

Accrued expenses and other current liabilities consisted of the following:

	December 31, 2022	December 31, 2021
Accrued research and development expenses	\$ 4,755,095	\$ 2,588,441
Accrued payroll expenses	796,250	97,055
Accrued professional fees	629,350	168,090
Other current liabilities	239,843	29,827
Total accrued expenses and other current liabilities	\$ 6,420,538	\$ 2,883,413

## 10. STOCK-BASED COMPENSATION

In February 2021, the Company adopted the 2021 Equity Incentive Plan (the “Plan”) for the issuance of stock options to the Company’s key directors, officers, employees and consultants, as a means to secure the benefits arising from capital stock ownership. The purposes of the Plan are to promote the alignment of the interests of key directors, officers, employees, and consultants with the success of the Company and to provide compensation opportunities to attract, retain and motivate directors, officers, employees, and consultants of the Company.

Under the Plan, the Company initially authorized 8,918,225 shares of its common stock for issuance upon exercise of options granted under the Plan. In August 2021, the Company’s Board of Directors amended the Plan to increase the aggregate number of shares of its common stock reserved for issuance pursuant to the Plan by 20,019,725 shares, from 8,918,225 shares to a new total of 28,937,950 shares. In November 2021, the Company’s Board of Directors further amended the Plan to further increase the aggregate number of shares of its common stock reserved for issuance pursuant to the Plan by 3,388,594 shares, from 28,937,950 shares to a new total of 32,326,544 shares. As of December 31, 2022, 10,599,419 shares were available for future grants.

### *Stock option activity*

Stock options granted during the years ended December 31, 2022 and 2021 generally (i) are subject to requisite service requirements, (ii) vest over a four-year period with 25% of the options granted vesting after one year and the remainder vesting in equal monthly installments over the following 36 months, and (iii) allow for early exercise subject to repurchase. Stock options granted to certain of the Company’s non-employees vest in equal monthly installments over a four-year period or vested upon the achievement of a certain milestone event. The Company did not repurchase any stock options during the years ended December 31, 2022 and 2021.

The following table summarizes the stock option activity under the Plan for years ended December 31, 2022 and 2021:

	Number of shares	Weighted-average exercise price	Weighted-average grant date fair value	Weighted-average remaining contractual term (in years)	Aggregate intrinsic value (in thousands)
Outstanding as of December 31, 2021	19,922,318	\$ 0.37	\$ 0.24	9.61	\$ 19,787
Granted	1,728,638	1.42	0.94		
Exercised	(721,321)	0.46	0.30		700
Forfeited	(628,774)	0.97	0.63		
Outstanding as of December 31, 2022	20,300,861	\$ 0.44	\$ 0.22	8.65	\$ 20,130
Vested options outstanding and exercisable as of December 31, 2021	591,478	\$ 0.25	\$ 0.15	9.20	\$ 671
Vested options outstanding and exercisable as of December 31, 2022	7,005,610	\$ 0.35	\$ 0.23	8.55	\$ 7,540
Unvested options outstanding and exercisable as of December 31, 2021	18,988,439	\$ 0.37	\$ 0.24	9.61	\$ 19,117
Unvested options outstanding and exercisable as of December 31, 2022	13,295,251	\$ 0.48	\$ 0.31	8.70	\$ 12,590

The aggregate intrinsic value of stock options is calculated as the difference between the exercise price of the stock options and the fair value of the Company’s common stock for those stock options that had exercise prices lower than the fair value of the Company’s common stock.

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The total grant date fair values of options vested during the years ended December 31, 2022 and 2021 were \$1.6 million and \$0.2 million, respectively, including \$0.1 million for a total of 342,404 stock options granted to certain nonemployees that vested upon the achievement of a certain milestone event during the year ended December 31, 2022.

The Company estimated the fair value of options granted using a Black-Scholes option pricing model with the following assumptions presented on a weighted average basis:

	Year ended December 31, 2022	Year ended December 31, 2021
Weighted average risk-free interest rate	2.76%	1.04%
Expected term (in years)	6.03	5.37
Expected volatility	73.36%	61.36%
Expected dividend yield	0.00%	0.00%

Expected dividend yield is based on the fact that the Company has never paid cash dividends and does not expect to pay any cash dividends in the foreseeable future.

### *Stock-based compensation expense*

The Company recorded total stock-based compensation of \$1.8 million and \$0.4 million for the years ended December 31, 2022, and 2021, respectively. Stock-based compensation expense was classified as follows in the consolidated statements of operations and comprehensive loss:

	Year ended December 31,	
	2022	2021
Research and development expense	\$ 965,407	\$ 252,422
General and administrative expense	815,159	134,213
Total stock-based compensation expense	\$ 1,780,566	\$ 386,635

As of December 31, 2022 there was \$4.0 million of unrecognized stock-based compensation expense related to unvested stock options estimated to be recognized over a weighted-average period of 2.54 years.

## **11. INCOME TAXES**

A reconciliation of the Company's statutory income tax rate to the Company's effective income tax rate is as follows for the years ended December 31, 2022, and 2021:

	December 31,	
	2022	2021
Federal statutory rate	21.00%	21.00%
State taxes, net of federal benefit	11.77	9.06
Change in valuation allowance	(36.43)	(30.38)
Stelios IPR&D	—	(2.90)
Permanent differences	0.08	(0.37)
Tax credits	3.66	3.37
Other	(0.08)	0.21
Effective income tax rate	0.00%	0.00%

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The principal components of the Company's deferred tax assets and liabilities at December 31, 2022 and 2021 are as follows:

	Year ended December 31,	
	2022	2021
<b>Deferred tax assets:</b>		
Net operating loss carryforwards	\$ 15,379,000	\$ 11,496,000
Tax credits	3,883,000	1,710,000
Intangibles	3,407,000	2,838,000
Capitalized R&D expenses	13,823,000	—
Lease liabilities	4,423,000	—
Other deferred tax assets	1,309,000	355,000
Total deferred tax assets	42,224,000	16,399,000
Valuation allowance	(37,992,000)	(16,399,000)
Net deferred tax assets	\$ 4,232,000	\$ —

	Year ended December 31,	
	2022	2021
<b>Deferred tax liabilities:</b>		
Right of use asset	\$(4,214,000)	\$ —
Other deferred tax liabilities	(18,000)	—
Total deferred tax liabilities	(4,232,000)	—
Net deferred tax assets (liabilities)	\$ —	\$ —

As of December 31, 2022 and 2021, the Company had federal net operating loss carryforwards of approximately \$48.6 million and \$36.7 million, respectively, which can be carried forward indefinitely. As of December 31, 2022 and 2021, the Company had state and local net operating loss carryforwards of approximately \$109.3 million and \$73.7 million, respectively, which begin to expire in 2040.

As of December 31, 2022, the Company had federal tax credits of \$3.9 million, which begin to expire in 2041.

Future realization of the tax benefits of existing temporary differences and net operating loss carryforwards ultimately depends on the existence of sufficient taxable income within the carryforward period. As of December 31, 2022 and 2021, the Company performed an evaluation to determine whether a valuation allowance was needed. The Company considered all available evidence, both positive and negative, which included the results of operations for the current and preceding years. The Company determined that it was not possible to reasonably quantify future taxable income and determined that it is more likely than not that all of the deferred tax assets will not be realized. Accordingly, the Company maintained a full valuation allowance as of December 31, 2022 and 2021.

Under Internal Revenue Code Section 382, if a corporation undergoes an "ownership change," the corporation's ability to use its pre-change NOL carryforwards and other pre-change tax attributes to offset its post-change income may be limited. We have not completed a study to assess whether an "ownership change" has occurred or whether there have been multiple ownership changes since we became a "loss corporation" as defined in Section 382. Future changes in our stock ownership, which may be outside of our control, may trigger an "ownership change." In addition, future equity offerings or acquisitions that have equity as a component of the purchase price could result in an "ownership change." If an "ownership change" has occurred or does occur in

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the future, utilization of the NOL carryforwards or other tax attributes may be limited, which could potentially result in increased future tax liability to us.

The calculation of our tax liabilities involves dealing with uncertainties in the application of complex tax laws and regulations for both federal taxes and the many states in which we operate or do business in. ASC 740 states that a tax benefit from an uncertain tax position may be recognized when it is more likely than not that the position will be sustained upon examination, including resolutions of any related appeals or litigation processes, on the basis of the technical merits.

We record uncertain tax positions as liabilities in accordance with ASC 740 and adjust these liabilities when our judgment changes as a result of the evaluation of new information not previously available. Because of the complexity of some of these uncertainties, the ultimate resolution may result in a payment that is materially different from our current estimate of the unrecognized tax benefit liabilities. These differences will be reflected as increases or decreases to income tax expense in the period in which new information is available. As of December 31, 2021 and 2022 we have not recorded any uncertain tax positions in our consolidated financial statements.

We recognize interest and penalties related to unrecognized tax benefits on the income tax expense line in the accompanying consolidated statement of operations. As of December 31, 2022 and 2021, no accrued interest or penalties are included on the related tax liability line in the consolidated balance sheet.

The Company files tax returns as prescribed by the tax laws of the jurisdictions in which it operates. In the normal course of business, the Company is subject to examination by federal and state jurisdictions, where applicable. The resolution of tax matters is not expected to have a material effect on the Company's consolidated financial statements.

## 12. COMMITMENTS AND CONTINGENCIES

**Leases**—As of December 31, 2022, the Company had entered into commitments under lease agreements to rent office space and finance equipment (see Note 5).

**Commitments**—As of December 31, 2022, the Company had entered into commitments under license, acquisition, research collaboration and sponsored research agreements with third parties (see Notes 3 and 13). In addition, the Company has entered into services agreements with third parties for pharmaceutical manufacturing and research activities in the normal course of business, which can generally be terminated by the Company with 30- or 60-days' written notice, unless otherwise indicated. Further, certain of the Company's manufacturing agreements could require early termination and wind-down payments due from the Company upon either the termination of its clinical trials or if the Company terminates such agreements for convenience.

**Contingencies**—From time to time, the Company may have certain contingent liabilities that arise in the ordinary course of business. The Company accrues a liability for such matters when it is probable that future expenditures will be made, and such expenditures can be reasonably estimated. For all periods presented, the Company was not a party to any pending material litigation or other material legal proceedings.

**Indemnification agreements**— In the ordinary course of business, the Company may provide indemnification of varying scope and terms to vendors, lessors, business partners and other parties with respect to certain matters including, but not limited to, losses arising out of breach of such agreements or from intellectual property infringement claims made by third parties. In addition, the Company has entered into indemnification agreements with members of its board of directors and executive officers that will require the Company, among other things, to indemnify them against certain liabilities that may arise by reason of their status or service as directors or



officers. The maximum potential amount of future payments the Company could be required to make under these indemnification agreements is, in many cases, unlimited. As of December 31, 2022, and 2021, the Company had not incurred any material costs as a result of such indemnifications.

### **13. LICENSE, RESEARCH COLLABORATION AND SPONSORED RESEARCH AGREEMENTS**

**Adverum Biotechnologies**—On January 25, 2021, the Company entered into an exclusive license agreement with Adverum Biotechnologies Inc. (“Adverum”) (the “Adverum Agreement”) to in-license materials and technology related to the treatment of Cardiomyopathy due to Friedrich’s Ataxia. In connection with the Adverum Agreement, the Company gained access to a portfolio of inventions, patent rights, technology, and licensed methods that the Company continues to develop, and the Company will assume all development and commercialization activities worldwide. Pursuant to the Adverum Agreement the Company paid a one-time up-front non-refundable fee of \$7.5 million, and is obligated to pay aggregate development and regulatory milestones of up to \$17.5 million including a \$3.5 million development milestone that was achieved in the first quarter of 2023, aggregate sales event and commercialization milestones of up to \$49.0 million, and royalty payments ranging from 8% of aggregate net sales less than or equal to \$500 million in a calendar year to 10% of aggregate net sales greater than \$500 million in a calendar year subject to reductions upon the expiration of valid claims in licensed patents and third-party licenses. As of December 31, 2022, there were no research and development expenses recorded by the Company or payments made to Adverum under the terms of the Adverum Agreement other than the one-time up-front non-refundable fee of \$7.5 million.

The Adverum Agreement remains in effect until termination at the date of the last royalty term to expire. The Company can terminate the Adverum Agreement with 120 days written notice. The Adverum Agreement can also be terminated as a result of a patent challenge, material breach of contractual terms, or insolvency by either party.

**Cornell University**—On May 27, 2020, the Company entered into two exclusive license agreements with Cornell University (“Cornell”) (the “Cornell First License Agreement” and the “Cornell Second License Agreement,” collectively “the Cornell License Agreements”). The Cornell First License Agreement is for the in-license of technology related to portfolios for APOE, Alzheimer’s disease, and Anti-Tau, although our license is not restricted by such indications and it includes assignment to us of Cornell’s IND for the use of AAVrh10.hAPOE2 vector to treat APOE4 homozygous patients who are at risk for or have Alzheimer’s disease to support development of the Company’s LX1001 program. The Cornell Second License Agreement is for the in-license of technology related to portfolios for infantile neuronal ceroid lipofuscinosis (also called CLN2 Batten disease) and Friedreich’s ataxia although our license is not restricted by such indications, and it includes assignment to us of Cornell’s IND for the use of AAVrh.10cUhCLN2 to treat children with CLN2 Batten disease to support development of the Company’s LX1004 program. Through the Cornell License Agreements, the Company gains access to a portfolio of inventions, patent rights, technology, and licensed methods that the Company continues to develop. Under the terms of the Cornell License Agreements, the Company has assumed all development and commercialization activities worldwide with respect to the licensed technology.

As initial consideration for the Cornell License Agreements, the Company paid Cornell an upfront payment in cash of \$0.3 million and issued \$1.3 million of Notes. In November 2020, Notes with outstanding principal of \$1.3 million were cancelled in exchange for 1,337,610 shares of Series A convertible preferred stock. As additional consideration, the Company is required to pay Cornell up to \$8.4 million upon the achievement of specific clinical and regulatory milestones under the Cornell First License Agreement and up to \$4.3 million upon the achievement of specific clinical and regulatory milestones under the Cornell Second License Agreement. The Company is also required to pay Cornell a flat royalty in the mid-single-digits based on net sales of the products covered by the licenses, subject to certain adjustments.

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Upon expiration of the royalty term of a given licensed product in a country, the respective license becomes non-exclusive and royalty-free. In addition, each of the Cornell License Agreements may be terminated by the Company for any reason upon 90 days' advance notice to Cornell and by Cornell upon the Company's material uncured breach, and all licenses and rights granted by either party under such agreement will concurrently terminate.

In the second quarter of 2022, the Company paid Cornell \$0.1 million for a research and development milestone achieved in the first quarter of 2022 under the Cornell Second License Agreement. In connection with the Cornell License Agreements, the Company recorded \$0.1 million of research and development expenses within its consolidated statement of operations and comprehensive loss during the year ended December 31, 2022, which were paid during the year ended December 31, 2022. The Company did not record any research and development expenses in its consolidated statement of operations and comprehensive loss or make any payments to Cornell under the terms of the Cornell License Agreements during the year ended December 31, 2021.

**Regents of the University of California, San Diego**—Stelios, which was acquired by the Company in August 2021, entered into exclusive worldwide license agreements on April 23, 2020 and August 6, 2020 (the "First UCSD Agreement" and the "Second UCSD Agreement", respectively) with the Regents of UCSD to in-license materials and intellectual property related to gene therapies for arrhythmogenic right ventricular cardiomyopathy and hypertrophic cardiomyopathy, respectively. The First UCSD Agreement and the Second UCSD Agreement relate to the Company's development efforts for its LX2021 and LX2022 programs, respectively. In connection with the First UCSD Agreement and the Second UCSD Agreement, the Company gained access to inventions, patent rights, technology, and licensed methods that it continues to develop, and it has assumed all worldwide development and commercialization activities with respect to the licensed technologies. The First UCSD Agreement and Second UCSD Agreement required Stelios to pay one-time up-front non-refundable cash fees of \$20,000 for each agreement, and required the Company to pay aggregate development and commercialization milestones of up to \$4.75 million and \$2.4 million, respectively, and low- to mid-single digit royalties and low-single digit royalties, respectively, based on aggregate net sales. The only research and development expenses incurred by Stelios or the Company and payments made to the Regents of UCSD to-date under the terms of the First UCSD Agreement and the Second UCSD Agreement were the one-time up-front non-refundable cash fees of \$20,000 for each agreement. The Company has the right to terminate the First UCSD Agreement and the Second UCSD Agreement at any time upon a sixty (60)-day written notice to the Regents of UCSD.

On October 4, 2021, the Company entered into an exclusive worldwide license agreement (the "Third UCSD Agreement" and collectively with the First UCSD Agreement and the Second UCSD Agreement, the "UCSD Agreements") with the Regents of UCSD to in-license materials and intellectual property related to a gene therapy for arrhythmogenic right ventricular cardiomyopathy. The Third UCSD Agreement relates to the Company's development efforts for its LX2020 program. In connection with the Third UCSD Agreement, the Company gained access to inventions, patent rights, technology, and licensed methods that it continues to develop, and it has assumed all worldwide development and commercialization activities with respect to the licensed technology. The Third UCSD Agreement required the Company to pay a one-time up-front non-refundable cash fee of \$20,000, aggregate development and commercialization milestones of up to \$4.0 million, and low- to mid-single digit royalties based on aggregate net sales. The only research and development expenses incurred by the Company and payments made to the Regents of UCSD to-date under the terms of the Third UCSD Agreement were the one-time up-front non-refundable cash fee of \$20,000. The Company has the right to terminate the Third UCSD Agreement at any time upon a sixty (60)-day written notice to the Regents of UCSD.

On December 3, 2021, the Company entered into two sponsored research agreements with the Regents of UCSD (collectively, the "UCSD SRAs") for the Company's LX2020, LX2021 and LX2022 programs in connection with the UCSD Agreements. Under the terms of the UCSD SRAs, the Company has the first rights to obtain

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non-exclusive or exclusive, sublicensable, royalty-bearing, perpetual and transferable worldwide licenses in any resulting inventions owned by the Regents of UCSD or resulting jointly owned inventions between the Company and the Regents of UCSD, and the Company retains the rights to any resulting inventions owned by the Company. The UCSD SRAs have a two-year term and may be terminated early by the Company at any time upon the giving of thirty (30) days written notice to the Regents of UCSD. The total costs to be invoiced to the Company under the terms of the UCSD SRAs are approximately \$5.5 million, of which \$2.3 million and \$0.2 million were recorded in research and development expenses in the Company's consolidated statements of operations and comprehensive loss during the years ended December 31, 2022 and 2021, respectively, and of which \$2.8 million was paid for the year ended December 31, 2022. The Company did not make any payments to UCSD during the year ended December 31, 2021.

**Weill Cornell Medical College**—On February 2, 2021, the Company entered into a Research Collaboration Agreement with Weill Cornell Medical College ("WCM") (the "WCM Agreement") in conjunction with the Cornell License Agreements entered on May 27, 2020. The Company committed to fund scientific research at WCM to investigate further and potentially enhance the technology licensed to the Company pursuant to the License Agreements.

Under the terms of the WCM Agreement, the Company will pay WCM at least \$3.0 million over the course of 3 years, with equal \$0.25 million payments due each quarter starting 10 days after the execution of the WCM Agreement. With respect to each WCM invention, joint invention, and related joint results for which an Improvement applies and the Company has made an election to amend the Agreements, the Company has the first option to negotiate in good faith with WCM for royalty-bearing, worldwide license, under Cornell patent rights, Cornell rights, and Cornell's interest in joint patent rights, to develop, make, have made, use, offer for sale, sell, have sold, and import derived products in the field. Research and development costs incurred and paid by the Company related to the WCM Agreement were \$6.1 million and \$5.5 million, respectively, during the year ended December 31, 2022. Research and development costs incurred and paid by the Company related to the WCM Agreement were \$2.8 million and \$2.3 million during the year ended December 31, 2021.

The WCM Agreement terminates upon the expiration of the agreed upon 3-year term. In addition, the WCM Agreement may be terminated by the Company in the event that the WCM principal investigator ceases to supervise the research and WCM is unable to or declines to find a substitute.

## **14. SUBSEQUENT EVENTS**

Subsequent events have been evaluated through March 31, 2023, which is the date that these consolidated financial statements were issued and were available to be issued.

*Shares*

*Common Stock*



**J.P. Morgan**

**SVB Securities**

**Stifel**

**RBC Capital Markets**

**Chardan**

Through and including \_\_\_\_\_, 2023 (the 25th day after the date of this prospectus), all dealers that buy, sell or trade shares of our common stock, whether or not participating in this offering, may be required to deliver a prospectus. This delivery requirement is in addition to the dealers' obligation to deliver a prospectus when acting as underwriters and with respect to their unsold allotments or subscriptions.

## Part II

### Information Not Required in Prospectus

#### Item 13. Other Expenses of Issuance and Distribution.

The following table indicates the expenses to be incurred in connection with the offering described in this registration statement, other than underwriting discounts and commissions, all of which will be paid by us. All amounts are estimated except the Securities and Exchange Commission, or SEC, registration fee, the Financial Industry Regulatory Authority, Inc., or FINRA, filing fee and the Nasdaq Global Market initial listing fee.

	<b>Amount</b>
SEC registration fee	\$ *
FINRA filing fee	*
Nasdaq Global Market initial listing fee	*
Accountants' fees and expenses	*
Legal fees and expenses	*
Transfer agent's fees and expenses	*
Printing and engraving expenses	*
Miscellaneous	*
<b>Total expenses</b>	<b>\$ *</b>

\* To be provided by amendment

#### Item 14. Indemnification of Directors and Officers.

We are incorporated under the laws of the State of Delaware. Section 102 of the Delaware General Corporation Law permits a corporation to eliminate the personal liability of directors of a corporation to the corporation or its stockholders for monetary damages for a breach of fiduciary duty as a director, except where the director breached his duty of loyalty, failed to act in good faith, engaged in intentional misconduct or knowingly violated a law, authorized the payment of a dividend or approved a stock repurchase in violation of Delaware corporate law or obtained an improper personal benefit.

Section 145 of the Delaware General Corporation Law provides that a corporation has the power to indemnify a director, officer, employee or agent of the corporation and certain other persons serving at the request of the corporation in related capacities against expenses (including attorneys' fees), judgments, fines and amounts paid in settlements actually and reasonably incurred by the person in connection with an action, suit or proceeding to which he is or is threatened to be made a party by reason of such position, if such person acted in good faith and in a manner he reasonably believed to be in or not opposed to the best interests of the corporation, and, in any criminal action or proceeding, had no reasonable cause to believe his conduct was unlawful, except that, in the case of actions brought by or in the right of the corporation, no indemnification shall be made with respect to any claim, issue or matter as to which such person shall have been adjudged to be liable to the corporation unless and only to the extent that the Court of Chancery or other adjudicating court determines that, despite the adjudication of liability but in view of all of the circumstances of the case, such person is fairly and reasonably entitled to indemnity for such expenses which the Court of Chancery or such other court shall deem proper.

As permitted by the Delaware General Corporation Law, our amended and restated certificate of incorporation and bylaws to be in effect upon the closing of this offering will provide that: (i) we are required to indemnify our

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directors to the fullest extent permitted by the Delaware General Corporation Law; (ii) we may, in our discretion, indemnify our officers, employees and agents as set forth in the Delaware General Corporation Law; (iii) we are required, upon satisfaction of certain conditions, to advance all expenses incurred by our directors in connection with certain legal proceedings; (iv) the rights conferred in the bylaws are not exclusive; and (v) we are authorized to enter into indemnification agreements with our directors, officers, employees and agents.

In connection with this offering, we expect to enter into indemnification agreements with each of our directors and executive officers that require us to indemnify them against expenses, judgments, fines, settlements and other amounts that any such person becomes legally obligated to pay (including with respect to a derivative action) in connection with any proceeding, whether actual or threatened, to which such person may be made a party by reason of the fact that such person is or was a director or officer of us or any of our affiliates, provided such person acted in good faith and in a manner such person reasonably believed to be in, or not opposed to, our best interests. The indemnification agreements will also set forth certain procedures that will apply in the event of a claim for indemnification thereunder. We intend to enter into similar indemnification agreements with our executive officers prior to the completion of this offering. At present, no litigation or proceeding is pending that involves any of our directors or officers regarding which indemnification is sought, nor are we aware of any threatened litigation that may result in claims for indemnification.

We maintain a directors' and officers' liability insurance policy. The policy insures directors and officers against unindemnified losses arising from certain wrongful acts in their capacities as directors and officers and reimburses us for those losses for which we have lawfully indemnified the directors and officers. The policy contains various exclusions.

In addition, the underwriting agreement filed as Exhibit 1.1 to this Registration Statement provides for indemnification by the underwriters of us and our officers and directors for certain liabilities arising under the Securities Act, or otherwise. Our amended and restated investor rights agreement with certain investors also provides for cross-indemnification in connection with the registration of our common stock on behalf of such investors.

### **Item 15. Recent Sales of Unregistered Securities.**

The following list sets forth information regarding all unregistered securities sold by us since our inception through the date of the prospectus that forms a part of this registration statement.

#### ***Issuances of Limited Liability Interests***

In connection with our formation in February 2017, we issued 95 limited liability common units to our founder and 5 limited liability common units to a service provider that is an accredited investor.

In January 2020, we issued 500 limited liability company incentive units to our chief executive officer.

In May 2020, we issued an aggregate of 206 limited liability company incentive units to two of our employees.

#### ***Issuances of Common Stock***

In November 2020, we issued an aggregate of 16,040,884 shares of our common stock to five individuals and accredited investors, including our founder and our chief executive officer, in connection with the conversion of all outstanding units to shares of common stock and the conversion of our company from a limited liability company to a corporation.

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### ***Issuances of Convertible Promissory Notes***

In May 2020, we issued a convertible promissory note to Cornell University in the principal amount of \$650,000.

In May 2020, we issued a convertible promissory note to Cornell University in the principal amount of \$650,000.

In June 2020, we issued a convertible promissory note to the Alzheimer's Drug Discovery Foundation in the principal amount of \$1,977,336.

In August 2020, we issued a convertible promissory note to the Alchemy LP in the principal amount of \$250,000.

In August 2020, we issued a convertible promissory note to Thomas McWilliams in the principal amount of \$250,000.

In September 2020, we issued a convertible promissory note to Nova Venture Holdings LLC in the principal amount of \$125,000.

### ***Issuances of Preferred Stock***

In November 2020, we issued an aggregate of 31,337,606 shares of our Series A convertible preferred stock to 13 investors at a purchase price of \$1.00 per share, for aggregate cash consideration of approximately \$30 million and pursuant to the conversion of certain convertible promissory notes and related cancellation of indebtedness of approximately \$1.3 million. We also issued an aggregate of 4,495,729 shares of our Series A convertible preferred stock to four investors at a price of \$0.85 per share, pursuant to the conversion of certain convertible promissory notes and related cancellation of indebtedness of approximately \$2.7 million.

In July 2021, we issued an aggregate of 50,999,997 shares of our Series A convertible preferred stock to 11 investors at a purchase price of \$1.00 per share, for aggregate consideration of approximately \$51 million.

In August 2021, we issued an aggregate of 58,157,823 shares of our Series B convertible preferred stock to 32 investors at a purchase price of \$1.72049 per share, for aggregate consideration of over \$100 million.

### ***Issuances Pursuant to our Equity Plans***

From February 2017 (the date of our inception) through the date of this registration statement, we granted options under the Existing Plan to purchase an aggregate of \_\_\_\_\_ shares of common stock, at an exercise price of \$ \_\_\_\_\_ per share, to our employees and consultants. Of these, \_\_\_\_\_ shares have been issued upon the exercise of options, and \_\_\_\_\_ options have been forfeited, expired or cancelled.

None of the foregoing transactions involved any underwriters, underwriting discounts or commissions, or any public offering. Unless otherwise specified above, we believe these transactions were exempt from registration under the Securities Act in reliance on Sections 3(a)(9) and 4(a)(2) of the Securities Act (and Regulation D or Regulation S promulgated thereunder) or Rule 701 promulgated under Section 3(b) of the Securities Act as transactions by an issuer not involving any public offering or under benefit plans and contracts relating to compensation as provided under Rule 701. The recipients of the securities in each of these transactions represented their intentions to acquire the securities for investment only and not with a view to or for sale in connection with any distribution thereof, and appropriate legends were placed on the share certificates issued in these transactions. All recipients had adequate access, through their relationships with us, to information about us. The sales of these securities were made without any general solicitation or advertising.

**Item 16. Exhibits and Financial Statement Schedules.**

(a) Exhibits.

The exhibits listed below are filed as part of this registration statement.

<b>Exhibit Number</b>	<b>Description of Exhibit</b>
1.1**	Form of Underwriting Agreement
3.1*	Amended and Restated Certificate of Incorporation of the Registrant (as amended and currently in effect)
3.2*	Bylaws of the Registrant (currently in effect)
3.3**	Form of Amended and Restated Certificate of Incorporation of the Registrant (to be effective upon the closing of this offering)
3.4**	Form of Amended and Restated Bylaws of the Registrant (to be effective upon the closing of this offering)
4.1**	Amended and Restated Investors' Rights Agreement, by and among the Registrant and certain of its stockholders, dated August 10, 2021
5.1**	Opinion of Cooley LLP
10.1+**	2021 Equity Incentive Plan and Forms of Option Grant Notice and Agreement, Exercise Notice, Early Exercise Notice and Restricted Stock Award Notice
10.2+**	2023 Equity Incentive Plan and Forms of Option Grant Notice and Agreement, Exercise Notice, Early Exercise Notice and Restricted Stock Award Notice
10.3+**	2023 Employee Stock Purchase Plan
10.4+**	Form of Indemnification Agreement with Executive Officers and Directors
10.5†**	Stock Purchase Agreement, dated July 16, 2021, by and among LEXEO Therapeutics, Inc., The Cystinosis Research Foundation, Eric Adler, M.D., Farah Sheikh, Ph.D., Jeffrey M. Ostrove, Ph.D., and Stephanie Cherqui, Ph.D., and Jeffery M. Ostrove, Ph.D., as Stockholders' Representative.
10.6†**	License Agreement, dated October 4, 2021, by and between LEXEO Therapeutics, Inc. and the Regents of the University of California
10.7†**	License Agreement, dated April 23, 2020, by and between Stelios Therapeutics, Inc. (as successor-in-interest to ARVC Therapeutics, Inc.) and the Regents of the University of California
10.8†**	License Agreement, dated August 6, 2020, by and between Stelios Therapeutics, Inc. and the Regents of the University of California
10.9†**	ARVC Research Agreement, dated December 3, 2021, between LEXEO Therapeutics, Inc. and the Regents of the University of California
10.10†**	TNNI3 Research Agreement, dated December 3, 2021, between LEXEO Therapeutics, Inc. and the Regents of the University of California
10.11†**	Amendment No. 1, dated April 19, 2023, to the TNNI3 Research Agreement, dated December 3, 2021, between LEXEO Therapeutics, Inc. and the Regents of the University of California
10.12†*	First License Agreement, dated May 28, 2020, between LEXEO Therapeutics, LLC and Cornell University



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<b>Exhibit Number</b>	<b>Description of Exhibit</b>
10.13†*	Second License Agreement, dated May 28, 2020, between LEXEO Therapeutics, LLC and Cornell University
10.14†**	Amendment No. 1, dated January 13, 2022, to the Second License Agreement, dated May 28, 2020, by and between LEXEO Therapeutics, Inc. and Cornell University
10.15†**	Amendment No. 1, dated July 1, 2022, to the First License Agreement, dated May 28, 2020, by and between LEXEO Therapeutics, Inc. and Cornell University and Amendment No. 2, dated July 1, 2022, to the Second License Agreement, dated May 28, 2020 (as amended by Amendment No. 1 to the Second License Agreement, dated January 13, 2022)
10.16†**	Amendment No. 2, dated September 28, 2022, to the First License Agreement, dated May 28, 2020, by and between LEXEO Therapeutics, Inc. and Cornell University (as amended by Amendment No. 1 to the First License Agreement, dated May 28, 2020)
10.17†**	Research Collaboration Agreement, dated February 2, 2021, by and between Cornell University and LEXEO Therapeutics, Inc.
10.18†**	License Agreement, dated January 19, 2021, between Adverum Biotechnologies, Inc. and LEXEO Therapeutics, LLC, as amended on February 28, 2022
10.19†**	Consulting Agreement, dated October 9, 2020, between LEXEO Therapeutics, LLC and Ronald G. Crystal, M.D.
10.20†**	Consulting Agreement, dated July 16, 2021, between LEXEO Therapeutics, LLC and Eric Adler, M.D.
10.21†**	Consulting Agreement, dated July 16, 2021, between LEXEO Therapeutics, Inc. and Farah Sheikh, Ph.D.
23.1**	Consent of KPMG LLP, independent registered public accounting firm
23.2**	Consent of Cooley LLP (included in Exhibit 5.1)
24.1**	Power of Attorney (included on signature page)
107**	Filing Fee Table

+ Indicates management contract or compensatory plan.

† Confidential treatment will be requested for portions of this agreement.

\* Previously filed.

\*\* To be filed by amendment.

### (b) Financial Statement Schedules.

Schedules not listed above have been omitted because the information required to be set forth therein is not applicable or is shown in the financial statements or notes thereto.

## **Item 17. Undertakings.**

The undersigned Registrant hereby undertakes to provide to the underwriters at the closing specified in the underwriting agreement, certificates in such denominations and registered in such names as required by the underwriters to permit prompt delivery to each purchaser.

Insofar as indemnification for liabilities arising under the Securities Act may be permitted to directors, officers and controlling persons of the Registrant pursuant to the foregoing provisions, or otherwise, the Registrant has been advised that in the opinion of the SEC such indemnification is against public policy as expressed in the

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Securities Act and is, therefore, unenforceable. In the event that a claim for indemnification against such liabilities (other than the payment by the Registrant of expenses incurred or paid by a director, officer or controlling person of the Registrant in the successful defense of any action, suit or proceeding) is asserted by such director, officer or controlling person in connection with the securities being registered, the Registrant will, unless in the opinion of its counsel the matter has been settled by controlling precedent, submit to a court of appropriate jurisdiction the question whether such indemnification by it is against public policy as expressed in the Securities Act and will be governed by the final adjudication of such issue.

The undersigned Registrant hereby undertakes that:

- (1) For purposes of determining any liability under the Securities Act, the information omitted from the form of prospectus filed as part of this Registration Statement in reliance upon Rule 430A and contained in a form of prospectus filed by the Registrant pursuant to Rule 424(b)(1) or (4) or 497(h) under the Securities Act shall be deemed to be part of this Registration Statement as of the time it was declared effective.
- (2) For the purpose of determining any liability under the Securities Act, each post-effective amendment that contains a form of prospectus shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial bona fide offering thereof.

## Signatures

Pursuant to the requirements of the Securities Act of 1933, the registrant has duly caused this registration statement to be signed on its behalf by the undersigned, thereunto duly authorized in the City of New York, State of New York, on this    day of                   , 2023.

**LEXEO THERAPEUTICS, INC.**

By: \_\_\_\_\_  
R. Nolan Townsend  
Chief Executive Officer

KNOW ALL BY THESE PRESENTS, that each person whose signature appears below hereby constitutes and appoints R. Nolan Townsend as his or her true and lawful agent, proxy and attorney-in-fact, with full power of substitution and resubstitution, for him or her and in his or her name, place and stead, in any and all capacities, to (i) act on, sign and file with the Securities and Exchange Commission any and all amendments (including post-effective amendments) to this registration statement together with all schedules and exhibits thereto and any subsequent registration statement filed pursuant to Rule 462(b) under the Securities Act of 1933, as amended, together with all schedules and exhibits thereto, (ii) act on, sign and file such certificates, instruments, agreements and other documents as may be necessary or appropriate in connection therewith, (iii) act on and file any supplement to any prospectus included in this registration statement or any such amendment or any subsequent registration statement filed pursuant to Rule 462(b) under the Securities Act of 1933, as amended, and (iv) take any and all actions which may be necessary or appropriate to be done, as fully for all intents and purposes as he might or could do in person, hereby approving, ratifying and confirming all that such agent, proxy and attorney-in-fact or any of his substitutes may lawfully do or cause to be done by virtue thereof.

Pursuant to the requirements of the Securities Act of 1933, this registration statement has been signed by the following persons in the capacities and on the dates indicated.

<u>Signature</u>	<u>Title</u>	<u>Date</u>
_____ R. Nolan Townsend	Chief Executive Officer and Director <i>(Principal Executive Officer)</i>	, 2023
_____ Micah Zajic	Chief Financial Officer <i>(Principal Financial and Accounting Officer)</i>	, 2023
_____ Mette Kirstine Agger	Director	, 2023
_____ Steven Altschuler, M.D.	Director	, 2023
_____ Paula HJ Cholmondeley	Director	, 2023
_____ Bernard Davitian	Director	, 2023
_____ Reinaldo Diaz	Director	, 2023