

**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION  
Washington, D.C. 20549  
FORM 10-Q**

(Mark One)

**QUARTERLY REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934**

For the quarterly period ended June 30, 2025

OR

**TRANSITION REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934**

For the transition period from to

Commission file number 001-40445

**CENTESSA PHARMACEUTICALS PLC**

(Exact name of registrant as specified in its charter)

**England and Wales**

(State or other jurisdiction of  
incorporation or organization)

**98-1612294**

(I.R.S. Employer Identification No.)

**3rd Floor  
1 Ashley Road  
Altrincham  
Cheshire  
United Kingdom**

(Address of principal executive offices)

**WA14 2DT**

(Zip code)

**+1 (617) 468-5770**

(Registrant's telephone number, including area code)

Securities registered pursuant to Section 12(b) of the Act:

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Ordinary shares, nominal value £0.002 per share	CNTA	Nasdaq Stock Market, LLC*
American Depositary Shares, each representing one ordinary share, nominal value £0.002 per share	CNTA	Nasdaq Stock Market, LLC

\*Not for trading, but only in connection with the listing of the American Depositary Shares on The Nasdaq Stock Market, LLC.

Indicate by check mark whether the registrant (1) has filed all reports required to be filed by Section 13 or 15(d) of the Securities Exchange Act of 1934 during the preceding 12 months (or for such shorter period that the registrant was required to file such reports), and (2) has been subject to such filing requirements for the past 90 days. Yes  No

Indicate by check mark whether the registrant has submitted electronically every Interactive Data File required to be submitted pursuant to Rule 405 of Regulation S-T (§232.405 of this chapter) during the preceding 12 months (or for such shorter period that the registrant was required to submit such files). Yes  No

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, a 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	<input type="checkbox"/>	Accelerated filer	<input type="checkbox"/>
Non-accelerated filer	<input checked="" type="checkbox"/>	Smaller reporting company	<input checked="" type="checkbox"/>
		Emerging growth company	<input checked="" type="checkbox"/>

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 13(a) of the Exchange Act.

Indicate by check mark whether the registrant is a shell company (as defined in Rule 12b-2 of the Exchange Act). Yes  No

### **Summary of the Material Risks Associated with Our Business**

Our business is subject to numerous risks and uncertainties that you should be aware of in evaluating our business. These risks are described more fully in Item 1A - Risk Factors, and include, but are not limited to, the following:

- We may not be successful in our efforts to use our differentiated asset-centric approach to drug discovery and development to build a pipeline of product candidates with commercial value.
  - A single or limited number of programs, developmental assets or product candidates may comprise a large proportion of our value.
  - We face challenges, risks and expenses related to our operations as well as the management of the expected growth in the scale and complexity of our operations.
  - We, and our Centessa Subsidiaries, have incurred net losses since inception, and we expect to continue to incur losses for the foreseeable future and may never achieve or maintain profitability.
  - We will need substantial additional funds to advance development of our product candidates, and we cannot guarantee that we will have sufficient funds available in the future to develop and commercialize our current or future product candidates.
  - Our loan facility and payment obligations under the Loan and Security Agreement with Oxford Finance contain operating and financial covenants that restrict our business and financing activities, are subject to acceleration in specified circumstances and may adversely affect our financial position or results of operations and our ability to raise additional capital which in turn may increase our vulnerability to adverse regulatory developments or economic or business downturns or which may result in Oxford Finance taking possession of our assets and disposing of any collateral.
  - Our product candidates are in various stages of development, including several in discovery and preclinical stages, and may fail in development or suffer delays that materially adversely affect their commercial viability and we may fail to differentiate our molecules, including ORX750, ORX142, ORX489 and other orexin agonist molecules, LockBody molecules and/or the LockBody technology platform from other available treatment options including other molecules in development.
  - We may not be successful in our efforts to identify, discover, in-license or otherwise acquire additional product candidates and may fail to capitalize on programs or product candidates that may be a greater commercial opportunity or for which there is a greater likelihood of success.
  - Success in preclinical studies or early clinical trials may not be indicative of results obtained in later trials.
  - We may encounter substantial delays or challenges in the initiation, conduct or completion of our clinical trials, and the results of clinical development are uncertain.
  - Even if we complete the necessary preclinical studies and clinical trials, the marketing approval process is expensive, time-consuming and uncertain and may prevent us from obtaining approvals for the commercialization of our product candidates.
  - We may be unable to obtain U.S. or foreign regulatory approval and, as a result, unable to develop and/or commercialize our product candidates.
  - We rely, and expect to continue to rely, on third parties to conduct our preclinical studies and clinical trials and if these third parties perform in an unsatisfactory manner, our business could be substantially harmed.
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**Summary of the Material Risks Associated with Our Business (continued)**

- Preclinical and clinical development is a long, expensive and uncertain process, and we may terminate one or more of our current preclinical and/or clinical development programs.
- We could experience manufacturing problems that result in delays or other material disruptions in our development or commercialization of our programs or otherwise harm our business.
- Business interruptions resulting from the Russia-Ukraine war, the Middle East conflict(s), tensions in U.S.-China relations, changes in trade policy, including the imposition of tariffs and trade wars, or similar geo-political conflicts or tensions could cause a disruption to our business activities including the development of our product candidates and the conduct of clinical trials thereby adversely impacting our business and results of operations.
- If we are unable to obtain and maintain sufficient patent and other intellectual property protection for our product candidates and technology or other product candidates that may be identified, or if the scope of the intellectual property protection obtained is not sufficiently broad, our competitors could develop and commercialize product candidates similar or identical to the product candidates, and our ability to successfully commercialize the product candidates and other product candidates that we may pursue may be impaired.
- The patent protection we obtain for our product candidates and technology may be challenged or not sufficient enough to provide us with any competitive advantage.
- A number of our programs and associated product candidates are heavily dependent on licensed intellectual property. If we were to lose our rights to licensed intellectual property, we may not be able to continue developing or commercializing our product candidates, if approved. If we breach any of the agreements under which we license the use, development and commercialization rights to our product candidates or technology from third parties or, in certain cases, we fail to meet certain development deadlines, we could lose license rights that are important to our business.
- We have never commercialized a product candidate and we may lack the necessary expertise, personnel and resources to successfully commercialize any of our products that receive regulatory approval on our own or together with collaborators.
- Our international operations may expose us to business, regulatory, legal, political, operational, financial, pricing and reimbursement risks associated with doing business across multiple jurisdictions outside of the United States.
- We are an emerging growth company and a smaller reporting company, and we cannot be certain if the reduced reporting requirements applicable to emerging growth companies and smaller reporting companies will make our ADSs less attractive to investors.
- We previously had material weaknesses in our internal control systems over financial reporting, which have been remediated. We may identify new material weaknesses in the future that may cause us to fail to meet our reporting obligations or result in material misstatements in our financial statements. If we fail to remediate any new material weaknesses, we may not be able to report our financial results accurately or to prevent fraud.
- If we fail to develop or maintain an effective system of disclosure controls and internal control over financial reporting, our ability to produce timely and accurate financial statements or comply with applicable regulations could be impaired.
- Holders of ADSs may be subject to limitations on the transfer of their ADSs and the withdrawal of the underlying ordinary shares.
- While we do not believe we were a “passive foreign investment company” (“PFIC”) in 2024, there is uncertainty as to whether we are or will be a PFIC in the past or in the future. If we are a PFIC, there could be material adverse U.S. federal income tax consequences to U.S. holders.

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## SPECIAL NOTE REGARDING FORWARD-LOOKING STATEMENTS

This Quarterly Report on Form 10-Q (“10-Q”), contains express or implied forward-looking statements within the meaning of Section 27A of the Securities Act of 1933, as amended, and Section 21E of the Securities Exchange Act of 1934, as amended, that involve substantial risks and uncertainties. In some cases, forward-looking statements may be identified by the words “may,” “might,” “will,” “could,” “would,” “should,” “expect,” “intend,” “plan,” “objective,” “anticipate,” “believe,” “estimate,” “predict,” “potential,” “continue,” “ongoing,” “aim,” “seek,” “strive,” or the negative of these terms, or other comparable terminology intended to identify statements about the future. These statements involve known and unknown risks, uncertainties and other important factors that may cause our actual results, levels of activity, performance or achievements to be materially different from the information expressed or implied by these forward-looking statements. The forward-looking statements and opinions contained in this 10-Q are based upon information available to our management as of the date of this 10-Q, and while we believe such information forms a reasonable basis for such statements, such information may be limited or incomplete, and our statements should not be read to indicate that we have conducted an exhaustive inquiry into, or review of, all potentially available relevant information. Forward-looking statements contained in this 10-Q include, but are not limited to, statements about:

- the initiation, timing, progress and results (preliminary, interim or final) of our preclinical studies and clinical trials, and our research and development programs including ORX750, ORX142, ORX489 and other orexin agonist molecules, any future LockBody candidates, and the LockBody technology platform;
- our ability to execute our research and clinical development plans and our expectations on and the timing thereof;
- our expectations and ability to advance our pipeline and product candidates into, and successfully complete, clinical trials;
- our ability to identify screen, recruit and maintain a sufficient number of, or any, subjects in our existing and anticipated studies or clinical trials including the ongoing Phase 1 first-in-human, clinical trial and Phase 2 clinical trial of ORX750 and other orexin agonist molecules, and any other LockBody candidates;
- our ability to differentiate ORX750, ORX142, ORX489, other orexin agonist molecules and any future LockBody candidates from other treatment options;
- the development and therapeutic potential of ORX750, ORX142, ORX489 and other orexin agonist molecules, LockBody candidates, and the LockBody technology platform;
- our expectations relating to the ongoing Phase 1 first-in-human clinical trial and the ongoing Phase 2 clinical trial of ORX750, including the predicted timing of enrollment, the predicted efficacious doses of ORX750 and our ability to successfully conduct our clinical development of ORX750;
- the safety and tolerability profile of our product candidates;
- our reliance on the success of our product candidates and our pipeline programs;
- our ability to discover and develop transformational medicines for patients including identifying and advancing additional product candidates into clinical development;
- our ability to become the partner of choice to attract founder-subject matter experts with high conviction programs;
- the timing or likelihood of regulatory filings and approvals;
- the impact of inflation on increasing costs of labor, research, manufacturing and clinical trial expenses;
- the impact of the Russia-Ukraine war, the Middle East conflict(s) and tensions in U.S.-China relations on our business and operations;
- the commercialization of our product candidates, if approved;
- our ability to develop sales and marketing capabilities;
- the pricing, coverage and reimbursement of our product candidates, if approved;
- the implementation of our business model, strategic plans for our business, product candidates and technology;
- the scope of protection we are able to establish and maintain for intellectual property rights covering our product candidates and technology;
- our ability to operate our business without infringing the intellectual property rights and proprietary technology of third parties;
- cost associated with prosecuting and maintaining our intellectual property and with defending intellectual property infringement, product liability and other claims;
- legal and regulatory developments in the United States, the European Union, the United Kingdom and other jurisdictions;
- estimates of our expenses, future revenues, capital requirements and our needs for additional financing;
- the potential benefits of strategic collaboration agreements and our ability to negotiate and enter into strategic arrangements;
- our ability to identify collaboration opportunities and to establish and maintain collaborations;
- our ability to judiciously manage and allocate our cash;

- our expectations on our anticipated cash runway;
- our ability to obtain additional funding;
- our ability to fulfill our obligations under the Loan and Security Agreement with Oxford Finance LLC (as Collateral Agent and Lender), Oxford Finance Credit Fund II LP (as a Lender) and Oxford Finance Credit Fund III LP (as a Lender) (collectively “Oxford Finance”);
- the rate and degree of market acceptance of any approved products;
- developments relating to our competitors and our industry, including competing therapies and our ability to respond to such developments;
- our ability to effectively manage our anticipated growth;
- our ability to attract and retain qualified employees and key personnel;
- our expectations regarding the period during which we qualify as a smaller reporting company and as an emerging growth company under the JOBS Act;
- statements regarding future revenue, hiring plans, expenses, capital expenditures, capital requirements and share performance;
- our expected use of proceeds of our capital raises;
- the future trading price of the ADSs and impact of securities analysts’ reports on these prices; and
- other risks and uncertainties, including those listed under the caption “Risk Factors.”

You should refer to the section titled “[Item 1A. Risk Factors](#)” in this 10-Q 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. As a result of these factors, we cannot be assured that the forward-looking statements in this 10-Q will prove to be accurate. Furthermore, if our forward-looking statements prove to be inaccurate, the inaccuracy may be material. In light of the significant uncertainties in these forward-looking statements, these statements should not be regarded as a representation or warranty by us or any other person that we will achieve our objectives and plans in any specified time frame, or at all. We undertake no obligation to publicly update any forward-looking statements, whether as a result of new information, future events or otherwise, except as required by law.

You should read this 10-Q and the documents that we reference in this 10-Q and have filed as exhibits to this 10-Q 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.

**PART I. FINANCIAL INFORMATION**

**Item 1. Financial Statements**

**Centessa Pharmaceuticals plc**  
**Consolidated Balance Sheets**  
(unaudited)  
(amounts in thousands except share and per share data)

	June 30, 2025	December 31, 2024
<b>Assets</b>		
Current assets:		
Cash and cash equivalents	\$ 44,242	\$ 383,221
Short-term investments	206,482	98,956
Tax incentive receivable	40,704	43,768
Prepaid expenses and other current assets	8,935	10,464
Total current assets	300,363	536,409
Long-term investments	153,406	—
Property and equipment, net	733	744
Operating lease right-of-use assets	10,539	11,013
Deferred tax asset	25,366	26,586
Other non-current assets	1,720	2,046
Total assets	\$ 492,127	\$ 576,798
<b>Liabilities and shareholders' equity</b>		
Current liabilities:		
Accounts payable	\$ 5,692	\$ 7,143
Accrued expenses and other current liabilities	24,002	50,855
Total current liabilities	29,694	57,998
Long term debt	109,545	108,940
Operating lease liabilities	7,940	8,286
Other non-current liabilities	29	29
Total liabilities	147,208	175,253
Commitments and contingencies (Note 7)		
Shareholders' equity:		
Ordinary shares: £0.002 nominal value: 222,480,776 shares authorized, 133,913,556 issued and outstanding at June 30, 2025; 184,469,623 shares authorized, 132,631,587 issued and outstanding at December 31, 2024	363	359
Additional paid-in capital	1,408,145	1,385,675
Accumulated other comprehensive income	1,591	4,213
Accumulated deficit	(1,065,180)	(988,702)
Total shareholders' equity	344,919	401,545
Total liabilities and shareholders' equity	\$ 492,127	\$ 576,798

*The accompanying notes are an integral part of these unaudited interim consolidated financial statements.*

**Centessa Pharmaceuticals plc**  
**Consolidated Statements of Operations and Comprehensive Loss**  
(unaudited)  
(amounts in thousands except share and per share data)

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
License and other revenue	\$ —	\$ —	\$ 15,000	\$ —
Operating expenses:				
Research and development	42,741	32,815	76,184	55,467
General and administrative	11,912	11,165	24,246	24,603
Loss from operations	(54,653)	(43,980)	(85,430)	(80,070)
Interest and investment income	4,380	3,240	12,270	5,831
Interest expense	(2,884)	(2,525)	(5,761)	(5,054)
Other non-operating income (expense), net	3,592	154	4,618	(1,383)
Loss before income taxes	(49,565)	(43,111)	(74,303)	(80,676)
Income tax expense	778	705	2,175	1,186
Net loss	(50,343)	(43,816)	(76,478)	(81,862)
Other comprehensive (loss) income:				
Foreign currency translation adjustment	(479)	(61)	164	(86)
Unrealized (loss) gain on available for sale marketable securities, net of reclassification adjustment and tax	(5)	33	(2,786)	188
Other comprehensive (loss) income	(484)	(28)	(2,622)	102
Total comprehensive loss	\$ (50,827)	\$ (43,844)	\$ (79,100)	\$ (81,760)
Net loss per ordinary share - basic and diluted	\$ (0.38)	\$ (0.40)	\$ (0.57)	\$ (0.78)
Weighted average ordinary shares outstanding - basic and diluted	133,677,405	109,489,184	133,354,373	104,688,452

*The accompanying notes are an integral part of these unaudited interim consolidated financial statements.*

**Centessa Pharmaceuticals plc**  
**Consolidated Statements of Shareholders' Equity**  
(unaudited)  
(amounts in thousands except share data)

	Ordinary Shares		Additional paid-in capital	Accumulated Other Comprehensive Income (Loss)	Accumulated Deficit	Total
	Shares	Amount				
<b>Balance at April 1, 2025</b>	<b>133,554,598</b>	<b>\$ 362</b>	<b>\$ 1,398,647</b>	<b>\$ 2,075</b>	<b>\$ (1,014,837)</b>	<b>\$ 386,247</b>
Stock option exercises	311,610	1	1,646	—	—	1,647
Share-based compensation expense	—	—	7,949	—	—	7,949
Vesting of ordinary shares	54,409	—	—	—	—	—
Shares withheld to pay employee withholding tax on share-based compensation	(7,061)	—	(97)	—	—	(97)
Foreign currency translation adjustments	—	—	—	(479)	—	(479)
Unrealized loss on available for sale securities, net of tax	—	—	—	(5)	—	(5)
Net loss	—	—	—	—	(50,343)	(50,343)
<b>Balance at June 30, 2025</b>	<b>133,913,556</b>	<b>\$ 363</b>	<b>\$ 1,408,145</b>	<b>\$ 1,591</b>	<b>\$ (1,065,180)</b>	<b>\$ 344,919</b>
<b>Balance at April 1, 2024</b>	<b>100,481,993</b>	<b>\$ 277</b>	<b>\$ 1,004,254</b>	<b>\$ 1,623</b>	<b>\$ (790,991)</b>	<b>\$ 215,163</b>
Issuance of ordinary shares under share offering, net of issuance costs	12,390,254	31	107,241	—	—	107,272
Stock option exercises	125,976	—	692	—	—	692
Share-based compensation expense	—	—	8,611	—	—	8,611
Vesting of ordinary shares	282,225	1	(1)	—	—	—
Shares withheld to pay employee withholding tax on share-based compensation	(102,684)	—	(929)	—	—	(929)
Foreign currency translation adjustments	—	—	—	(61)	—	(61)
Unrealized gain on available for sale securities, net of tax	—	—	—	33	—	33
Net loss	—	—	—	—	(43,816)	(43,816)
<b>Balance at June 30, 2024</b>	<b>113,177,764</b>	<b>\$ 309</b>	<b>\$ 1,119,868</b>	<b>\$ 1,595</b>	<b>\$ (834,807)</b>	<b>\$ 286,965</b>

*The accompanying notes are an integral part of these unaudited interim consolidated financial statements.*

**Centessa Pharmaceuticals plc**  
**Consolidated Statements of Shareholders' Equity**  
(unaudited)  
(amounts in thousands except share data)

	Ordinary Shares		Additional paid-in capital	Accumulated Other Comprehensive Income (Loss)	Accumulated Deficit	Total
	Shares	Amount				
<b>Balance at January 1, 2025</b>	<b>132,631,587</b>	<b>\$ 359</b>	<b>\$ 1,385,675</b>	<b>\$ 4,213</b>	<b>\$ (988,702)</b>	<b>\$ 401,545</b>
Issuance of ordinary shares under ATM program, net of issuance costs	372,538	1	6,108	—	—	6,109
Stock option exercises	592,326	2	3,525	—	—	3,527
Share-based compensation expense	—	—	15,832	—	—	15,832
Vesting of ordinary shares	495,318	1	(1)	—	—	—
Shares withheld to pay employee withholding tax on share-based compensation	(178,213)	—	(2,994)	—	—	(2,994)
Foreign currency translation adjustments	—	—	—	164	—	164
Unrealized gain on available for sale securities, net of tax	—	—	—	1	—	1
Reclassification adjustment for realized net gain on available for sale marketable securities included in net loss, net of tax of \$0.8 million	—	—	—	(2,787)	—	(2,787)
Net loss	—	—	—	—	(76,478)	(76,478)
<b>Balance at June 30, 2025</b>	<b>133,913,556</b>	<b>\$ 363</b>	<b>\$ 1,408,145</b>	<b>\$ 1,591</b>	<b>\$ (1,065,180)</b>	<b>\$ 344,919</b>
<b>Balance at January 1, 2024</b>	<b>98,774,827</b>	<b>\$ 273</b>	<b>\$ 987,423</b>	<b>\$ 1,493</b>	<b>\$ (752,945)</b>	<b>\$ 236,244</b>
Issuance of ordinary shares under share offering, net of issuance costs	12,390,254	31	107,241	—	—	107,272
Issuance of ordinary shares under ATM program, net of issuance costs	1,250,000	3	9,652	—	—	9,655
Stock option exercises	302,436	—	1,647	—	—	1,647
Share-based compensation expense	—	—	16,765	—	—	16,765
Vesting of ordinary shares	763,151	2	(2)	—	—	—
Shares withheld to pay employee withholding tax on share-based compensation	(302,904)	—	(2,858)	—	—	(2,858)
Foreign currency translation adjustments	—	—	—	(86)	—	(86)
Unrealized gain on available for sale securities, net of tax	—	—	—	1,406	—	1,406
Reclassification adjustment for realized net gain on available for sale marketable securities included in net loss, net of tax of \$0.4 million	—	—	—	(1,218)	—	(1,218)
Net loss	—	—	—	—	(81,862)	(81,862)
<b>Balance at June 30, 2024</b>	<b>113,177,764</b>	<b>\$ 309</b>	<b>\$ 1,119,868</b>	<b>\$ 1,595</b>	<b>\$ (834,807)</b>	<b>\$ 286,965</b>

*The accompanying notes are an integral part of these unaudited interim consolidated financial statements.*

**Centessa Pharmaceuticals plc**  
**Consolidated Statements of Cash Flows**  
(unaudited)  
(amounts in thousands)

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
<b>Cash flows from operating activities:</b>		
Net loss	\$ (76,478)	\$ (81,862)
Adjustments to reconcile net loss to net cash used in operating activities:		
Share-based compensation expense	15,832	16,765
Depreciation and amortization	443	476
Amortization of discount of marketable securities	(2,135)	—
Amortization of debt issuance costs and exit fee of long-term debt	605	—
Change in fair value of financial instruments	—	800
Change in deferred taxes	2,021	1,001
Changes in operating assets and liabilities:		
Tax incentive receivable	6,448	(4,349)
Prepaid expenses and other assets	2,025	1,718
Operating leases, net	128	151
Accounts payable	(1,918)	(6,434)
Accrued expenses and other liabilities	(26,669)	(6,132)
Net cash used in operating activities	(79,698)	(77,866)
<b>Cash flows from investing activities:</b>		
Purchases of investments in marketable securities	(380,537)	(140,533)
Proceeds from redemption of investments in marketable securities	118,321	101,836
Purchases of property and equipment	(126)	—
Net cash used in investing activities	(262,342)	(38,697)
<b>Cash flows from financing activities:</b>		
Proceeds from issuance of shares under offerings, net of issuance costs	—	107,272
Proceeds from issuance of shares under ATM program, net of issuance costs	6,109	9,655
Proceeds from option exercises	3,468	1,640
Payment of employee withholding taxes related to withheld shares for stock based compensation plans	(4,490)	(2,761)
Other, net	(396)	—
Net cash provided by financing activities	4,691	115,806
Effect of exchange rate on cash and cash equivalents	(1,630)	99
Net decrease in cash and cash equivalents	(338,979)	(658)
Cash and cash equivalents at beginning of period	383,221	128,030
Cash and cash equivalents at end of period	\$ 44,242	\$ 127,372
<b>Supplemental disclosure:</b>		
Interest paid	\$ 4,302	\$ 5,053
Income taxes paid	\$ 990	\$ 1
Operating lease payments reducing operating lease liabilities	\$ 817	\$ 801

*The accompanying notes are an integral part of these unaudited interim consolidated financial statements.*

**Centessa Pharmaceuticals plc**  
**Notes to the Unaudited Interim Consolidated Financial Statements**

## **1. Organization and Description of Business**

Centessa Pharmaceuticals plc (“Centessa” or “the Company”) is a clinical-stage pharmaceutical company that aims to discover, develop and ultimately deliver medicines that are transformational for patients. Centessa was incorporated on October 26, 2020 as a limited liability company under the laws of England and Wales. In connection with the Company’s initial public offering, or IPO, the Company re-registered Centessa Pharmaceuticals Limited as an English public limited company and renamed it as Centessa Pharmaceuticals plc.

### *Risks and Liquidity*

The Company is subject to risks common to other life science companies in various stages of development including, but not limited to, uncertainty of product development and commercialization, lack of marketing and sales history, development by its competitors of new technological innovations, dependence on key personnel, market acceptance of products, product liability, protection of proprietary technology, ability to raise additional financing and compliance with government regulations in the markets in which the Company is seeking approvals, including U.S. Food and Drug Administration (“FDA”) regulations. If the Company does not successfully advance its programs, into and through human clinical trials and/or enter into collaborations for its programs and commercialize any of its product candidates, it may be unable to produce product revenue or achieve profitability.

The Company has incurred losses and negative cash flows from operations since inception and the Company had an accumulated deficit of \$1.07 billion as of June 30, 2025. The Company anticipates incurring additional losses until such time, if ever, that it can generate significant sales of the product candidates currently in development. Substantial additional capital will be needed by the Company to fund its operations and to develop its product candidates.

The Company entered into a Sales Agreement, dated January 27, 2023, by and between Centessa Pharmaceuticals plc and Leerink Partners LLC (formerly SVB Securities LLC). As sales agent, Leerink Partners LLC will provide for the issuance and sale by the Company of up to \$125.0 million of its ordinary shares represented by American Depository Shares (“ADSs”) from time to time in “at-the-market” offerings (“ATM Program”). In the six months ended June 30, 2025, the Company sold 372,538 ordinary shares under the ATM Program, resulting in net proceeds of \$6.1 million, all of which were collected during the first quarter of 2025. Since the inception of the ATM program in 2023, as of June 30, 2025, the Company has sold 4,663,354 ordinary shares under the ATM Program, resulting in net proceeds to us of approximately \$36.6 million.

The Company expects its existing cash, cash equivalents and investments as of June 30, 2025 of \$404.1 million will be sufficient to fund its expected operating expenses and capital expenditure requirements for at least the next twelve months from the date of issuance of these unaudited interim consolidated financial statements.

### *Emerging Growth Company and Smaller Reporting Company status*

The Company is an emerging growth company, as defined in the JOBS Act. Under the JOBS Act, emerging growth companies can delay the adoption of new or revised accounting standards issued subsequent to the enactment of the JOBS Act until such time as those standards apply to private companies. Other exemptions and reduced reporting requirements under the JOBS Act for emerging growth companies include presentation of only two years of audited financial statements in a registration statement for an initial public offering, an exemption from the requirement to provide an auditor’s report on internal controls over financial reporting pursuant to Section 404 of the Sarbanes-Oxley Act of 2002, as amended, an exemption from any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation and less extensive disclosure about our executive compensation arrangements. The Company has elected to use the extended transition period for complying with new or revised accounting standards that have different effective dates for public and private companies until the earlier of the date that (i) it is no longer an emerging growth company or (ii) the Company affirmatively and irrevocably opts out of the extended transition period provided in the JOBS Act.

The Company will remain an emerging growth company until the earliest of (i) the last day of its first fiscal year in which it has total annual gross revenues of \$1.235 billion or more, (ii) the last day of its first fiscal year following the fifth anniversary of the closing of its initial public offering, (iii) the date on which it is deemed to be a “large accelerated filer,” under the rules of the SEC, which means the market value of equity securities that is held by non-affiliates exceeds \$700.0 million as

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of the prior June 30th and (iv) the date on which we have issued more than \$1.0 billion in non-convertible debt securities during the prior three-year period.

The Company is also a “smaller reporting company” as defined in the Securities Exchange Act of 1934 (the “Exchange Act”). If it is a smaller reporting company at the time we cease to be an emerging growth company, the Company may continue to rely on exemptions from certain disclosure requirements that are available to smaller reporting companies. Specifically, as a smaller reporting company it may choose to present only the two most recent fiscal years of audited financial statements in its Annual Report on Form 10-K and, similar to emerging growth companies, smaller reporting companies have reduced disclosure obligations regarding executive compensation.

On the last business day of its second quarter in 2025, the aggregate market value of the Company’s shares of common stock held by non-affiliates exceeded \$700 million. As a result, as of December 31, 2025, the Company will be considered a “large accelerated filer” as defined in Rule 12b-2 under the Exchange Act, and will cease to be an “emerging growth company” as defined in the JOBS Act and a “smaller reporting company” as defined in the Exchange Act. The Company will no longer be exempt from the auditor attestation requirements of Section 404(b) of the Sarbanes-Oxley Act, and its independent registered public accounting firm will evaluate and report on the effectiveness of internal control over financial reporting. The Company will also no longer be permitted to take advantage of reduced reporting requirements for smaller reporting companies.

## **2. Summary of Significant Accounting Policies**

The accompanying unaudited interim consolidated financial statements should be read in conjunction with the annual audited consolidated financial statements of Centessa Pharmaceuticals plc and related notes which can be found in our Annual Report on Form 10-K for the year ended December 31, 2024 (the “2024 Annual Report”). The Summary of Significant Accounting Policies included in the Company’s annual financial statements have not materially changed, except as set forth below.

### *Basis of Presentation and Consolidation*

The accompanying unaudited interim consolidated financial statements have been prepared in conformity with accounting principles generally accepted in the United States of America (“GAAP”). Any reference in these notes to applicable guidance is meant to refer to GAAP as found in the Accounting Standards Codification (“ASC”) and Accounting Standards Updates (“ASUs”) promulgated by the Financial Accounting Standards Board (“FASB”).

In the opinion of management, the accompanying unaudited interim consolidated financial statements include all normal and recurring adjustments (which consist primarily of accruals, estimates and assumptions that impact the financial statements) considered necessary to present fairly:

- the Company’s financial position as of June 30, 2025 and as of December 31, 2024;
- the Company’s results of operations for the three and six months ended June 30, 2025 and June 30, 2024; and
- the Company’s cash flows for the six months ended June 30, 2025 and June 30, 2024.

Operating results for the Company for the three and six months ended June 30, 2025 are not necessarily indicative of the results that may be expected for the full year. The unaudited interim consolidated financial statements presented herein do not contain all of the required disclosures under U.S. GAAP for annual financial statements. Notes to the financial statements which would substantially duplicate the disclosure contained in the audited financial statements for fiscal 2024 as reported in the 2024 Annual Report have been omitted. Therefore, these unaudited interim consolidated financial statements should be read in conjunction with the annual audited consolidated financial statements and related notes for Centessa Pharmaceuticals plc found in the Form 10-K for the year ended December 31, 2024 filed with the SEC.

The Company’s unaudited interim consolidated financial statements include the accounts of Centessa Pharmaceuticals plc, and its wholly owned subsidiaries. All intercompany accounts and transactions have been eliminated in consolidation.

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*Investments in Marketable Securities*

The Company invests its excess cash in money market funds, U.S. Treasury securities, U.S. government agency securities, corporate notes and commercial paper. The Company's investment policy limits investments to money market funds, certain types of debt securities issued by the U.S. Government and its agencies, corporate notes and commercial paper, and places restrictions on the credit ratings, maturities and concentration by type and issuer. Securities with original maturities of three months or less when purchased are included in cash and cash equivalents. We consider investments with original maturities greater than three months and remaining maturities less than one year to be short-term investments, while remaining maturities greater than one year are classified as long-term investments.

As of June 30, 2025, investments in U.S. government Treasury and agency securities, corporate notes and commercial paper were classified as available-for-sale securities, which are recorded at fair value. Unrealized holding gains and losses on available-for-sale securities are reported net of related income taxes in accumulated other comprehensive income until realized. Purchase premiums and discounts are amortized to interest income over the terms of the related securities.

*Reclassifications*

Certain amounts in the prior period financial statements have been reclassified to conform to the presentation of the current period financial statements. These reclassifications had no effect on previously reported net loss or comprehensive loss.

*Use of Estimates*

The preparation of unaudited interim consolidated financial statements in conformity 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 as of the date of the unaudited interim consolidated financial statements and the reported amounts of expenses during the reporting period. Actual results could differ from those estimates.

Estimates and assumptions are periodically reviewed and the effects of revisions are reflected in the unaudited interim consolidated financial statements in the period they are determined to be necessary. On an ongoing basis, an evaluation of estimates and judgments are required, including those related to accrued and prepaid research and development expenses, share-based compensation expense and tax related matters. Estimates are based on historical experience, known trends and events, and various other factors that are believed to be 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. Changes in estimates are recorded in the period in which they become known. Actual results may differ from these estimates under different assumptions or conditions.

*License and other revenue*

The Company recognizes revenues from collaboration, license or other research or sale arrangements when or as performance obligations are satisfied. For milestone payments, the Company assesses, at contract inception, whether the milestones are considered probable of being achieved. If it is probable that a significant revenue reversal will occur, the Company will not record revenue until the uncertainty has been resolved. Milestone payments that are contingent upon regulatory approval are not considered probable until the approvals are obtained as it is outside of the control of the Company. If it is probable that significant revenue reversal will not occur, the Company will estimate the milestone payments using the most likely amount method. The Company reassesses the milestones each reporting period to determine the probability of

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achievement. Any potential consideration received in the form of royalty or sales-based milestones will be recorded when the customer's subsequent sales or usages occur.

*Net Loss Per Ordinary Share*

The following potentially dilutive securities have been excluded from the computation of diluted weighted-average ordinary shares outstanding for the six months ended June 30, 2025 and June 30, 2024, as they would be anti-dilutive:

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
Unvested ordinary shares	—	196,985
Restricted stock units	1,103,383	2,164,448
Stock options	21,111,837	18,764,967
	<u>22,215,220</u>	<u>21,126,400</u>

### 3. Genmab License Agreement

*Background*

On February 14, 2025, the Company entered into a license agreement (the "License Agreement") with Genmab A/S ("Genmab") pursuant to which the Company granted to Genmab an exclusive worldwide license to leverage its proprietary LockBody technology platform to perform research on up to three undisclosed targets selected during a multi-year research period. Additionally, the License Agreement provides Genmab with the option to obtain an exclusive commercial license for worldwide development and commercialization of products against each of the selected three targets. Genmab will conduct all research and development activities under the License Agreement, which may combine the Company's LockBody technology with Genmab's proprietary antibody technologies. The LockBody technology platform is designed to improve the therapeutic index of therapies by allowing the conditional activation of potent cell killing mechanisms in diseased tissue, but not in non-diseased tissue.

Under the terms of the License Agreement, the Company received a non-refundable upfront payment of \$15.0 million related to the license of the LockBody technology platform. Additionally, the Company may receive, in the future, an aggregate of up to \$15.0 million when (and if) Genmab exercises the options to acquire the exclusive commercial licenses as discussed above. Further, the Company is eligible to receive potential payouts of up to \$234.0 million in development, regulatory and sales milestones per product, as well as tiered royalties ranging in the mid single-digits on annual global net licensed product sales.

The License Agreement includes various representations, warranties, covenants, indemnities, and other customary provisions. Unless earlier terminated in accordance with its terms, the License Agreement will expire upon expiration of the last royalty term for the last licensed product. Genmab may terminate the License Agreement or on a target-by-target basis for convenience upon specified time periods. On a target-by-target basis, if Genmab elects not to exercise its option for an exclusive commercial license for worldwide development and commercialization of products against the applicable target (a "Reserved Target"), then the License Agreement will automatically terminate with respect to such Reserved Target. Subject to the terms and specified exceptions set forth in the License Agreement, either party may terminate the License Agreement for the other party's uncured material breach or insolvency upon a specified notice period.

*Accounting Treatment*

The Company assessed the License Agreement in accordance with ASC 606, *Revenue with Contracts from Customers*, and concluded that the promises in the License Agreement represent a transaction with a customer. The following promises within the License Agreement were identified: (i) the exclusive license of the Lockbody technology platform and the ability to perform research on up to three targets, (ii) initial and continuing know-how transfer for the Lockbody technology platform, and (iii) participation in a joint steering committee ("JSC"). The promises related to the know-how transfer and JSC participation were determined to be immaterial in the context of the contract as the time commitment and related cost is

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expected to be inconsequential to the total contract consideration. Accordingly, they were not assessed as performance obligations.

Additionally, the Company also evaluated the options to acquire exclusive commercial licenses contained within the License Agreement to determine if they provide Genmab with any material rights. The Company concluded that the options were not issued at a significant and incremental discount and therefore do not provide Genmab with a material right. As such, these options were excluded as performance obligations and will be accounted for when (and if) they are exercised.

As a result, the Company determined that the promise to provide the exclusive license constitutes a single performance obligation within the License Agreement. Further, the exclusive license represents functional intellectual property given the Company will not be providing any additional services to Genmab outside of the right to use the licensed intellectual property. Accordingly, the \$15.0 million up front payment was recognized as revenue at a point in time during the first quarter of 2025, upon delivery of the exclusive license to Genmab at contract inception.

As of June 30, 2025, the Company has constrained all variable consideration related to potential milestone payments associated with the License Agreement given the level of uncertainty associated with their achievement. Accordingly, no revenue relating to these milestones has been recognized.

#### **4. Fair Value Measurement**

Fair value is the price that could be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants. Fair value determination in accordance with applicable accounting guidance requires that a number of significant judgments be made. Additionally, fair value is used on a nonrecurring basis to evaluate assets for impairment or as required for disclosure purposes by applicable accounting guidance on disclosures about fair value of financial instruments. Depending on the nature of the assets and liabilities, various valuation techniques and assumptions are used when estimating fair value. The carrying amounts of certain of the Company's financial instruments, including cash, prepaid expenses, tax incentive receivable and accounts payable, are shown at cost, which approximates fair value due to the short-term nature of these instruments.

The Company follows the provisions of FASB ASC Topic 820, *Fair Value Measurement*, for financial assets and liabilities measured on a recurring basis. The guidance requires fair value measurements be classified and disclosed in one of the following three categories:

- Level 1:* Unadjusted quoted prices in active markets that are accessible at the measurement date for identical, unrestricted assets or liabilities.
- Level 2:* Quoted prices in markets that are not active, or inputs which are observable, either directly or indirectly, for substantially the full term of the asset or liabilities.
- Level 3:* Prices or valuation techniques that require inputs that are both significant to the fair value measurement and unobservable (i.e., supported by little or no market activity).

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The following fair value hierarchy table presents information about the Company's financial assets measured and carried at fair value (amounts in thousands):

	Fair value measurement at reporting date using			Total
	Quoted prices in active markets for identical assets (Level 1)	Significant other observable inputs (Level 2)	Significant unobservable inputs (Level 3)	
<b>June 30, 2025</b>				
Assets				
Money Market fund	\$ 18,364	\$ —	\$ —	\$ 18,364
U.S. Treasury and government agency securities	97,999	112,537	—	210,536
Corporate debt securities	—	109,974	—	109,974
Commercial Paper	—	39,378	—	39,378
Total	<u>\$ 116,363</u>	<u>\$ 261,889</u>	<u>\$ —</u>	<u>\$ 378,252</u>
<b>December 31, 2024</b>				
Assets				
Money Market fund	\$ 217,515	\$ —	\$ —	\$ 217,515
U.S. Treasury securities	171,885	—	—	171,885
Total	<u>\$ 389,400</u>	<u>\$ —</u>	<u>\$ —</u>	<u>\$ 389,400</u>

We classify our investments in available-for-sale U.S. Treasury securities and the money market fund into Level 1 of the ASC Topic 820 hierarchy because fair values represent quoted market prices for identical or comparable instruments. The Company estimates the fair value of its U.S. government agency securities, corporate debt securities and commercial paper by taking into consideration valuations obtained from third-party pricing services. The pricing services utilize industry standard valuation models, including both income and market-based approaches, for which all significant inputs are observable, either directly or indirectly, to estimate fair value. These inputs include reported trades of and broker/dealer quotes on the same or similar securities, issuer credit spreads; benchmark securities; prepayment/default projections based on historical data; and other observable inputs.

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The following represents the amortized cost bases and fair values of the Company's fixed income investments as of June 30, 2025 (amounts in thousands):

	Amortized Cost	Gross Unrealized Gains	Gross Unrealized Losses	Fair Value
Money Market fund, included in Cash and cash equivalents	\$ 18,364	\$ —	\$ —	\$ 18,364
U.S. Treasury and government agency securities, included in:				
Short-term investments	144,962	—	(98)	144,864
Long-term investments	65,710	32	(70)	65,672
Total U.S. Treasury and government agency securities	\$ 210,672	\$ 32	\$ (168)	\$ 210,536
Corporate debt securities, included in:				
Short-term investments	\$ 22,228	\$ 17	\$ (5)	\$ 22,240
Long-term investments	87,597	163	(26)	87,734
Total Corporate debt securities	\$ 109,825	\$ 180	\$ (31)	\$ 109,974
Commercial paper, included in:				
Short-term investments	39,389	—	(11)	39,378
Total Commercial paper	\$ 39,389	\$ —	\$ (11)	\$ 39,378
In aggregate:				
Investments above with no unrealized loss	\$ 107,842	\$ 212	\$ —	\$ 108,054
Investments above with unrealized loss	270,408	—	(210)	270,198
Total investments	\$ 378,250	\$ 212	\$ (210)	\$ 378,252

At the end of June 30, 2025, the Company held approximately 53 debt securities, of which 33 were in a unrealized loss position as of the end of the period, with an aggregate fair value of \$270.2 million. The Company has not recorded any allowance for credit losses as of June 30, 2025 as it believes the decline in fair value below amortized cost is not related to credit losses. These investments in an unrealized loss position were in this position for less than 12 months and there has been no change in the credit risk of such securities during the period. The Company does not intend to sell its investments and it is not more likely than not that the Company will be required to sell the securities before recovery of the amortized cost basis of its debt securities. Securities are evaluated at the end of each reporting period for evidence of the credit-related impairment.

## 5. Balance Sheet Components

Prepaid expenses and other current assets consist of the following (amounts in thousands):

	June 30, 2025	December 31, 2024
Research and development expenses	\$ 3,691	\$ 6,314
Interest receivable	2,249	287
Value added tax receivable	1,143	1,528
Insurance related expenses	233	840
Other	1,619	1,495
	\$ 8,935	\$ 10,464

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Accrued expenses and other current liabilities consist of the following (amounts in thousands):

	June 30, 2025	December 31, 2024
Research and development expenses*	\$ 16,085	\$ 38,063
Personnel related expenses	4,045	9,465
Professional fees	1,952	1,782
Interest payable	855	—
Operating lease	656	602
Income tax liability	257	835
Other	152	108
	<u>\$ 24,002</u>	<u>\$ 50,855</u>

\*Includes accrued contract termination costs of \$1.7 million as of June 30, 2025 and \$30.3 million as of December 31, 2024. See Note 8.

Property and equipment, net consisted of the following (amounts in thousands):

	June 30, 2025	December 31, 2024
Computer equipment	\$ 894	\$ 768
Office furniture	724	724
Other	43	43
Property and equipment, at cost	1,661	1,535
Less: Accumulated depreciation	(928)	(791)
Property and equipment, net	<u>\$ 733</u>	<u>\$ 744</u>

The following table provides a reconciliation of changes, net of applicable income taxes, in other accumulated comprehensive income (amounts in thousands):

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
<b>Foreign currency translation adjustment:</b>		
Beginning balance	\$ 1,426	\$ 203
Foreign currency translation adjustment during period	164	(86)
Ending balance	<u>\$ 1,590</u>	<u>\$ 117</u>
<b>Unrealized gain on available for sale securities:</b>		
Beginning balance	\$ 2,787	\$ 1,290
Unrealized holding gain on available for sale securities during period, net of tax of \$0.4 million in 2024	1	1,406
Reclassification adjustment for realized net gain on available for sale marketable securities included in net loss, net of tax of \$0.8 million in 2025 and \$0.4 million in 2024	(2,787)	(1,218)
Ending balance	<u>\$ 1</u>	<u>\$ 1,478</u>

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## 6. Debt

Debt as of June 30, 2025 and December 31, 2024 (in thousands):

	June 30, 2025	December 31, 2024
<b>Loan and Security Agreement (“LSA”):</b>		
Principal amount	\$ 110,000	\$ 110,000
End of term charge	5,500	5,500
Unamortized debt issuance costs and unamortized end of term charge	(5,955)	(6,560)
<b>Loan and Security Agreement (carried at amortized cost)</b>	<b>\$ 109,545</b>	<b>\$ 108,940</b>
The fair value of LSA, classified as Level 2 under ASC Topic 820, <i>Fair Value Measurement</i>	<u>\$ 107,965</u>	<u>\$ 109,450</u>

Future principal payments, including the end of term charge, are as follows (in thousands):

Year ending:	Payments
2025	\$ —
2026	—
2027	—
2028	—
2029	115,500
<b>Total principal payments, including End of Term Charge</b>	<b>\$ 115,500</b>

## 7. Commitments and Contingencies

### Commitments

As of June 30, 2025, the Company had non-cancellable commitments for purchase of clinical materials, contract manufacturing, maintenance, and committed funding of up to \$10.4 million, of which the Company expects to pay \$8.6 million within one year and \$1.9 million over one to five years. The amount and timing of these payments vary depending on the rate of progress of development. Future clinical trial expenses have not been included within the purchase commitments because they are contingent on enrollment in clinical trials and the activities required to be performed by the clinical sites.

### Leases

The following table provides balance sheet information related to leases as of June 30, 2025 (amounts in thousands):

	June 30, 2025
<b>Assets:</b>	
Operating lease, right-of-use asset	\$ 10,539
<b>Liabilities:</b>	
Current portion of operating lease liabilities	\$ 656
Operating lease liabilities, net of current portion	7,940
<b>Total operating lease liabilities</b>	<b>\$ 8,596</b>

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In calculating the present value of the lease payments, the Company elected to utilize its incremental borrowing rate based on the original term of the lease. The following table summarizes supplemental information related to leases as of June 30, 2025:

	June 30, 2025
Weighted-average remaining lease term	7.5 years
Weighted-average discount rate	11.97 %

The components of the Company's lease costs are classified on its consolidated statements of operations as follows (amounts in thousands):

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
Operating lease cost	\$ 999	\$ 999
Variable lease cost	55	2
Short term lease cost	12	4
Sublease income	(175)	(175)
<b>Total operating lease cost</b>	<b>\$ 891</b>	<b>\$ 830</b>

Future lease payments and sublease income under non-cancelable operating leases as of June 30, 2025 were as follows (amounts in thousands):

	Operating Leases	Sublease Income
Year ending:		
2025	\$ 817	\$ 178
2026	1,667	362
2027	1,700	370
2028	1,734	345
2029	1,769	—
Thereafter	5,521	—
Total	\$ 13,208	\$ 1,255
Less: Imputed interest	(4,612)	
Present value of lease liabilities	\$ 8,596	
Less: current portion	(656)	
Lease liabilities, net of current portion	\$ 7,940	

### Contingencies

From time to time, the Company may have certain contingent liabilities that arise in the ordinary course of its business activities. 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.

### 8. Program Termination

On November 12, 2024, the Company announced the discontinuation of the global clinical development program for SerpinPC, a novel inhibitor of activated protein C (APC) that was being progressed for the treatment of hemophilia B. The strategic decision was made to prioritize capital towards the development of our growing OX2R agonist franchise based on the

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strength of the interim Phase 1 data for ORX750, coupled with the outcome of a planned interim analysis of Part 1 of the PREsent-2 study of SerpinPC, which was evaluated in the context of the evolving treatment and market landscape for hemophilia B, including the recent FDA approval of a competing product.

In the fourth quarter of 2024, as a result of the discontinuation of the program, we incurred a one-time charge of \$31.5 million, consisting of contract termination costs of \$30.3 million and employee severance related costs of \$1.2 million, which was recorded within research and development expenses. The contract termination costs related to firm commitments for the manufacture of registrational materials and other costs. The contract termination and employee severance costs associated with this program discontinuation were predominately accrued as of December 31, 2024. The Company expects the remaining balance as of June 30, 2025 will be fully extinguished by the end of December 2025.

The following table provides a roll-forward of the program termination liability as of June 30, 2025 (in thousands):

	Contract Termination	Severance-Related	Total
Balance as of January 1, 2025	\$ 30,283	\$ 1,199	\$ 31,482
Cash payments	(26,406)	(1,014)	(27,420)
Reversal of charge	(2,555)	—	(2,555)
Foreign currency translation changes	329	—	329
Balance as of June 30, 2025	<u>\$ 1,651</u>	<u>\$ 185</u>	<u>\$ 1,836</u>

## 9. Share-based Compensation

### *Centessa Pharmaceuticals plc Stock Option and Incentive Plan*

Remaining shares available for future grants as of June 30, 2025 under the 2021 Stock Option and Incentive Plan (the “2021 Plan”) were 11,665,913.

### *Share-based Compensation Expense*

The Company recorded share-based compensation expense in the following expense categories in the unaudited interim consolidated statements of operations and comprehensive loss (amounts in thousands):

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
Research and development	\$ 4,115	\$ 4,006	\$ 8,272	\$ 7,731
General and administrative	3,834	4,605	7,560	9,034
	<u>\$ 7,949</u>	<u>\$ 8,611</u>	<u>\$ 15,832</u>	<u>\$ 16,765</u>

Share-based compensation expense by award type was as follows included within the unaudited interim consolidated statements of operations and comprehensive loss: (amounts in thousands):

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
Stock options	\$ 6,889	\$ 5,775	\$ 13,022	\$ 11,207
Restricted share awards and units	1,060	2,836	2,810	5,558
	<u>\$ 7,949</u>	<u>\$ 8,611</u>	<u>\$ 15,832</u>	<u>\$ 16,765</u>

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*Stock Options*

The following table summarizes stock option activity for the six months ended June 30, 2025:

	Number of Shares	Weighted-Average Exercise Price Per Share	Weighted-Average Remaining Contractual Term	Aggregate Intrinsic Value (in millions)
Balance at January 1, 2025	17,434,119	\$ 7.77	7.4 years	
Granted	4,917,140	\$ 16.00		
Exercised	(592,326)	\$ 5.95		
Forfeited	(647,096)	\$ 11.39		
Balance at June 30, 2025	<u>21,111,837</u>	\$ 9.63	7.5 years	\$ 92.6
Exercisable at June 30, 2025	<u>12,436,207</u>	\$ 7.75	6.4 years	\$ 71.3
Vested and expected to vest at June 30, 2025	<u>21,111,837</u>	\$ 9.63	7.5 years	\$ 92.6

The weighted-average grant date fair value of options granted was \$11.36 per share for the six months ended June 30, 2025. As of June 30, 2025, the total unrecognized compensation expense related to unvested stock option awards was \$69.2 million, which the Company expects to recognize over a weighted-average period of 3.2 years.

Based on the trading price of \$13.14 per ADS, which is the closing price as of June 30, 2025, the aggregate intrinsic value of outstanding options as of June 30, 2025 was \$92.6 million.

The fair value of each option was estimated on the date of grant under the Black-Scholes option pricing model using the weighted average assumptions in the table below:

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
Expected term	6.0 years	5.9 years
Expected stock price volatility	79.2 %	76.2 %
Risk-free interest rate	4.3 %	3.9 %
Expected dividend yield	0 %	0 %

**Centessa Pharmaceuticals plc**  
**Notes to the Unaudited Interim Consolidated Financial Statements**

*Restricted Share Awards and Units*

The following table summarizes ordinary share activity related to the restricted stock programs for the six months ended June 30, 2025:

	Restricted Stock Awards		Restricted Stock Units	
	Number of Shares	Weighted-Average Grant Date Fair Value Per Share	Number of Shares	Weighted-Average Grant Date Fair Value Per Share
Unvested at January 1, 2025	86,864	\$ 20.00	1,510,077	\$ 6.14
Granted	—	\$ —	108,610	\$ 16.90
Vested	(86,864)	\$ 20.00	(408,454)	\$ 5.78
Forfeited	—	\$ —	(106,850)	\$ 6.36
Unvested at June 30, 2025	—	\$ —	1,103,383	\$ 7.31
Unrecognized compensation expense at June 30, 2025 (in thousands)	\$ —		\$ 6,932	
Expected weighted average recognition period	0.0 years		2.7 years	

*Employee Share Purchase Plan*

As of June 30, 2025, no shares have been purchased or issued under the 2021 Employee Share Purchase Plan (the “ESPP”) and remaining shares reserved for the ESPP were 2,708,515.

**10. Segment Information**

Centessa Pharmaceuticals plc is a clinical-stage pharmaceutical company with a mission to discover, develop and ultimately deliver medicines that are transformational for patients. The Company’s ability to generate recurring product revenue and to become profitable will depend upon the ability to successfully develop, obtain regulatory approval and commercialize any current and future product candidates.

The Company manages the business as one segment. Its operating results are regularly reviewed by the Company’s Chief Executive Officer, who is the chief operating decision-maker (“CODM”), based on available financial information prepared on a consolidated basis. The CODM evaluates its performance based primarily on research and development efforts and the results of clinical trials. The CODM also utilizes the Company’s long-range plan as a strategic tool to allocate resources according to the Company’s strategic objectives. The consolidated net loss is used to monitor budget versus actual results in assessing segment performance and the allocation of resources. Assets provided to the CODM are consistent with those reported on the Consolidated Balance Sheets with particular emphasis on the Company’s available liquidity, including its cash, cash equivalents and investments.

The following table presents reportable segment loss, including significant expenses regularly provided to the CODM, attributable to the Company’s reportable segment for the six months ended June 30, 2025 and June 30, 2024 (in thousands):

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
License and other revenue	\$ —	\$ —	\$ 15,000	\$ —
Less:				
OX2R agonist program expenses	(32,681)	(10,300)	(52,774)	(14,952)
LockBody program expenses	(1,765)	(2,096)	(5,183)	(6,266)
Discontinued R&D program expenses	(1,192)	(16,677)	(4,109)	(28,619)
Other R&D expenses:				
Personnel expenses <sup>1</sup>	(6,077)	(4,901)	(12,379)	(10,500)
Research tax incentives	4,475	5,535	8,734	13,395
Other preclinical and clinical development expenses	(1,386)	(370)	(2,201)	(794)
General and administrative expenses <sup>1</sup>	(8,078)	(6,560)	(16,686)	(15,569)
Share-based compensation	(7,949)	(8,611)	(15,832)	(16,765)
Interest and investment income	4,380	3,240	12,270	5,831
Interest expense	(2,884)	(2,525)	(5,761)	(5,054)
Other segment items <sup>2</sup>	3,592	154	4,618	(1,383)
Income tax expense	(778)	(705)	(2,175)	(1,186)
Consolidated net loss	\$ (50,343)	\$ (43,816)	\$ (76,478)	\$ (81,862)

<sup>1</sup> Excludes share-based compensation

<sup>2</sup> Other segment items includes non-operating income (expenses), net

## Item 2. Management’s Discussion and Analysis of Financial Condition and Results of Operations

*The following discussion and analysis should be read in conjunction with the unaudited interim consolidated financial statements and related notes thereto, included elsewhere herein and the audited consolidated financial statements and notes thereto for the year ended December 31, 2024 and the related Management’s Discussion and Analysis of Financial Condition and Results of Operation, all of which are contained in our Annual Report on Form 10-K (the “2024 Annual Report”) filed with the SEC. In addition to historical financial information, some of the information contained in the following discussion and analysis contains forward-looking statements within the meaning of Section 27A of the Securities Act of 1933, as amended, and Section 21E of the Securities Exchange Act of 1934, as amended. All statements other than statements of historical facts, including statements regarding our future results of operations and financial position, business strategy, current and prospective products, product approvals, research and development costs, current and prospective collaborations, timing and likelihood of success, plans and objectives of management for future operations and future results of current and anticipated products, are forward-looking statements. These statements involve known and unknown risks, uncertainties, assumptions 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.*

### Overview and Format of Presentation

Centessa Pharmaceuticals plc is a clinical-stage pharmaceutical company with a mission to discover, develop and ultimately deliver medicines that are transformational for patients. We are pioneering a new class of potential therapies within our orexin receptor 2 (“OX2R”) agonist program intended to be orally administered for the treatment of excessive daytime sleepiness (“EDS”), impaired attention, cognitive deficits, and fatigue across neurological, neurodegenerative and neuropsychiatric disorders. We also have an early-stage immuno-oncology program focused on our novel LockBody® technology platform.

**OX2R Agonist Program:** We are developing novel, investigational, oral, highly potent and selective OX2R agonists as potential best-in-class treatments for EDS, impaired attention, cognitive deficits and fatigue in a wide spectrum of neurological, neurodegenerative and neuropsychiatric disorders. These OX2R agonists are designed to mimic the natural neuropeptide orexin (also known as ‘hypocretin’), by activating orexin receptors in the brain. The orexin system consists of two neuropeptides, orexin-A (“OXA”) and orexin-B (“OXB”) (also called hypocretin-1 and hypocretin-2) which bind to and activate the orexin receptors, Orexin Receptor-1 (“OX1R”) and Orexin Receptor-2 (“OX2R”). These receptors are abundantly expressed throughout the brain and have different distribution patterns, suggesting they have distinct physiological roles acting through different neuronal pathways. Importantly, OXA and OXB both bind to OX2R with high affinity. OX2R plays a key role in regulating wakefulness and has also been implicated in metabolism, behavioral arousal, mood and cognitive function.

**ORX750:** Our most advanced OX2R agonist candidate is ORX750, which is in an ongoing Phase 2a study of participants with narcolepsy type 1 (“NT1”), narcolepsy type 2 (“NT2”) and idiopathic hypersomnia (“IH”).

In May 2024, we announced the initiation of a Phase 1 first-in-human (“FIH”) clinical study of ORX750 in healthy volunteers. We disclosed initial data from the Phase 1 study in September 2024 and provided additional interim updates in November 2024 and January 2025. These data were also disclosed in our Annual Report on Form 10-K for the year ended December 31, 2024. In April 2025, we presented additional data from the Phase 1 study in a poster session at the American Academy of Neurology (“AAN”) Annual Meeting. Data from the Phase 1 study enabled dose selection for the Phase 2a study of ORX750 in participants with NT1, NT2, and IH. The Phase 1 study of ORX750 remains open.

In November 2024, we announced the initiation of the Phase 2a CRYSTAL-1 clinical study to evaluate the safety, tolerability, and pharmacokinetics (“PK”) of ORX750 in participants with NT1, NT2, and IH. We expect the Phase 2a study to generate clinical data for all three indications in 2025 and enable dose selection for future registrational studies.

**ORX142:** ORX142 is our second OX2R agonist development candidate. ORX142 is being advanced for the treatment of select neurological and neurodegenerative disorders, as well as other potential symptoms including impaired attention, cognitive deficits and fatigue. Following clearance of our Investigational New Drug application (“IND”) from the FDA in June 2025, we initiated a Phase 1 FIH clinical trial of ORX142 with the goal of sharing clinical data in acutely sleep-deprived healthy volunteers in 2025.

**ORX489:** ORX489 is our third OX2R agonist development candidate. ORX489 is our most potent OX2R agonist to date based on preclinical studies, and is in IND-enabling activities for the treatment of neuropsychiatric disorders.

***OX2R Agonist Pipeline:*** Our earlier OX2R agonist pipeline includes additional OX2R agonists as well as research efforts on differentiated pharmacology associated with the activation of the orexin system.

***LockBody® Technology Platform:*** Leveraging our proprietary LockBody technology, we are pioneering a novel approach that is designed to selectively drive potent effector function activity, such as CD3, into the tumor micro environment (“TME”) while avoiding systemic toxicity. Our LockBody technology platform is intended to allow for the development of LockBody constructs (a “LockBody”). A LockBody is designed to be a conditionally-active antibody drug with the potential to engage powerful immune pathways in diseased tissue, but not in non-diseased tissue or the periphery, where the drug’s action is often unwanted.

During the first quarter of 2025, we discontinued the clinical development of LB101, a conditionally tetravalent PD-L1xCD47 bispecific monoclonal antibody, and first generation LockBody candidate. This was a strategic decision based on the totality of clinical data from the Phase 1/2a FIH dose escalation study of LB101 in participants with advanced solid tumors. Although the LB101 Phase 1 results did not support further development of CD47 as a target effector, we continue to evaluate the LockBody platform with ongoing preclinical work.

In February 2025, we announced a license agreement providing Genmab access to our LockBody technology platform to research products against up to three targets during a multi-year research period, with option to take exclusive commercial licenses for worldwide development and commercialization of products against each selected target. Genmab is conducting all research and development activities under the agreement.

We own worldwide rights to all of our pipeline programs and may opportunistically evaluate and enter into strategic transactions around certain product candidates, targets, geographies, or disease areas.

### **Liquidity and Capital Resources**

As of June 30, 2025, we had cash, cash equivalents and investments of \$404.1 million. Since inception, we have devoted substantially all of our resources to acquiring and developing product and technology rights, conducting research and development in its discovery and enabling stages, in our clinical and preclinical trials, business operations and raising capital. We have incurred recurring losses and negative cash flows from operations since inception and have funded operations primarily through the sale and issuance of our equity securities and debt financing. The ability to generate product revenue sufficient to achieve profitability will depend heavily on the successful development and eventual commercialization of current or future product candidates. We expect to continue to incur significant expenses and operating losses for the foreseeable future in connection with ongoing development activities related to our pipeline programs as we advance the preclinical and clinical development of product candidates; perform research activities as we seek to discover and develop additional programs and product candidates; carry out maintenance, expansion enforcement, defense, and protection of our intellectual property portfolio; and hire additional research and development, clinical and commercial personnel. Further, inflation may affect our use of capital resources by increasing our cost of labor, research, manufacturing and clinical trial expenses. Based on our current operating model and development plans, we expect cash, cash equivalents and investments as of June 30, 2025 to fund our operations into mid-2027.

### **Components of Results of Operations**

#### ***Revenues***

While we received non-recurring revenue related to out-licensing in the past and the license of the LockBody technology platform this year, our ability to generate recurring product revenue and to become profitable will depend upon the ability to successfully develop, obtain regulatory approval and commercialize any current and future product candidates. Because of the numerous risks and uncertainties associated with product development and regulatory approval, we are unable to predict the amount or timing of product revenue.

#### ***Research and Development Expense***

Research and development expenses consist primarily of costs incurred in connection with the discovery and development of the Company’s clinical and preclinical programs, net of reimbursements. Research and development costs are expensed as incurred. These expenses include:

- expenses incurred to conduct the necessary preclinical studies and clinical trials required to obtain regulatory approval;

- milestone payments pursuant to the license agreements;
- personnel expenses, including salaries, benefits and share-based compensation expense for employees engaged in research and development functions;
- costs of funding research and development performed by third parties, including pursuant to agreements with contract research organizations (“CROs”) for active and discontinued programs, as well as investigative sites and consultants that conduct preclinical studies and clinical trials;
- expenses incurred under agreements with contract manufacturing organizations (“CMOs”), including committed costs for discontinued programs, manufacturing scale-up expenses and the cost of acquiring and manufacturing preclinical study and clinical trial materials;
- fees paid to consultants who assist with research and development activities;
- expenses related to regulatory activities, including filing fees paid to regulatory agencies; and
- allocated expenses for facility costs, including rent, utilities, depreciation and maintenance.

Research and development activities are central to our business model. Product candidates in later stages of clinical development will generally have higher development costs than those in earlier stages of clinical development, primarily due to the increased size and duration of later-stage clinical trials. We expect research and development expenses to increase significantly over the next several years due to increases in costs to conduct clinical trials for current product candidates and other clinical trials for future product candidates, costs to prepare regulatory filings for any product candidates and increases in personnel costs, including share-based compensation.

The successful development of our current or future product candidates is highly uncertain. At this time, we cannot reasonably estimate or know the nature, timing and costs of the efforts that will be necessary to complete the remainder of the development of current or future product candidates, or when, if ever, material net cash inflows may commence from product candidates. This uncertainty is due to the numerous risks and uncertainties associated with the duration and cost of clinical trials, which vary significantly over the life of a project as a result of many factors, including:

- delays in regulators or institutional review boards authorizing us or our investigators to commence our clinical trials, or in our ability to negotiate agreements with clinical trial sites or CROs;
- the ability to secure adequate supply of product candidates for trials;
- the number of clinical sites included in the trials;
- the ability and the length of time required to enroll suitable patients;
- the number of patients that ultimately participate and remain in the trials;
- the number of doses patients receive;
- any side effects associated with product candidates;
- the duration of patient follow-up;
- the results of clinical trials;
- significant and changing government regulations; and
- launching commercial sales of product candidates, if and when approved, whether alone or in collaboration with others.

Our expenditures are subject to additional uncertainties, including the terms and timing of regulatory approvals. We may never succeed in achieving regulatory approval for our product candidates.

We may obtain unexpected results from clinical trials and may elect to discontinue, delay or modify clinical trials of product candidates. A change in the outcome of any of these variables with respect to the development of a product candidate could mean a significant change in the costs and timing associated with the development of that product candidate. For example, if the European Medicines Agency (“EMA”), FDA or other comparable regulatory authorities were to require us to conduct clinical trials beyond those that are currently anticipated, or if we experience significant delays in enrollment in any clinical trials, we could be required to expend significant additional financial resources and time on the completion of clinical development. Product commercialization will take several years, and we expect to spend a significant amount in development costs.

### ***Research and Development Tax Incentives***

We participate in research tax incentive programs that are granted to companies by the United Kingdom in order to encourage technical and scientific research. Expenditures that meet the required criteria are eligible to receive a tax benefit. Estimates of the amount of the benefit expected to be received are determined at each reporting period and recorded as reductions to research and development expenses. Through December 31, 2024, we claimed relief under the Small and Medium Enterprise (“SME”) scheme. Beginning January 1, 2025, changes to the program in the UK have aligned the tax incentives earned for SME and large entities. The merged scheme provides relief for qualifying R&D expenditure. If we are loss making, a cash credit can be obtained. If we continue to meet the SME thresholds and are loss making, then a higher rate of credit may be available under the new Enhanced R&D Intensive Support (ERIS). We expect eligible R&D expenditures qualifying for the credit outside of the UK will be more limited as a result of these legislative changes and we expect to recognize less tax incentives in the United Kingdom.

### ***General and Administrative Expense***

General and administrative expense consists primarily of personnel expenses, including salaries and benefits for employees and share-based compensation. General and administrative expense also includes facility costs, including rent, utilities, depreciation and maintenance, not otherwise included in research and development expense, as well as legal fees related to intellectual property and corporate matters and fees for accounting, consulting and other professional services.

### ***Interest and Investment Income and Interest Expense***

Interest and investment income is primarily interest earned from the Company’s cash and cash equivalents and its investments and realized gains on sales of securities. Interest expense consists of interest costs related to the Loan and Security Agreement.

### ***Other Income (Expense), net***

Other income (expense), net consists primarily of foreign currency transaction gains and losses.

### ***Foreign Currency Translation***

Our financial statements are presented in U.S. dollars (“USD”), the reporting currency of the Company. The functional currency of Centessa Pharmaceuticals plc is USD and the functional currency of its subsidiaries is their respective local currency. Income and expenses have been translated into USD at average exchange rates prevailing during the period. Assets and liabilities have been translated at the rates of exchange on the balance sheets dates and equity accounts at their respective historical rates. The resulting translation gain and loss adjustments are recorded directly as a separate component of shareholders’ equity as other comprehensive income (loss). Transactions denominated in a currency other than the functional currency are remeasured based upon the exchange rate at the date of remeasurement with the resulting gain or loss included in the accompanying unaudited interim consolidated statements of operations and comprehensive loss within other non-operating income (expense), net.

## Results of Operations

### Comparison of the Three Months Ended June 30, 2025 and June 30, 2024

The following table sets forth the results of operations for the three months ended June 30, 2025 and June 30, 2024 (amounts in thousands):

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024
Operating expenses:		
Research and development	\$ 42,741	\$ 32,815
General and administrative	11,912	11,165
Loss from operations	(54,653)	(43,980)
Interest and investment income	4,380	3,240
Interest expense	(2,884)	(2,525)
Other non-operating income, net	3,592	154
Loss before income tax expense	(49,565)	(43,111)
Income tax expense	778	705
Net loss	<u>\$ (50,343)</u>	<u>\$ (43,816)</u>

### Research and Development Expenses

The following table summarizes research and development expenses by program incurred for the following periods (amounts in thousands):

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024
<b>Programs:</b>		
OX2R agonist	\$ 32,681	\$ 10,300
LockBody	1,765	2,096
<b>Discontinued programs</b>	1,192	16,677
<b>Non-asset specific costs:</b>		
Personnel expenses	10,192	8,907
Research tax incentives	(4,475)	(5,535)
Other preclinical and clinical development expenses	1,386	370
<b>Research and development expenses</b>	<u>\$ 42,741</u>	<u>\$ 32,815</u>

Research and development expenses for the three months ended June 30, 2025 were \$42.7 million, compared with \$32.8 million for the three months ended June 30, 2024. The increase in research and development expenses reflected higher development costs of \$22.4 million for our OX2R agonist program, including higher clinical study costs for ORX750 as well as developmental milestones following clearance of the IND for ORX142 by the FDA and initiation of clinical trial. The higher costs of the OX2R agonist program were partially offset by lower spending on discontinued programs. Personnel expenses in the three months ended June 30, 2025 increased \$1.3 million compared to the three months ended June 30, 2024, driven by higher headcount in 2025.

### **General and Administrative Expenses**

The following table summarizes the general and administrative expenses for the following periods (amounts in thousands):

	Three Months Ended June 30, 2025	Three Months Ended June 30, 2024
Personnel expenses	\$ 6,889	\$ 7,348
Legal and professional fees	2,776	986
Other expenses	2,247	2,831
	<u>\$ 11,912</u>	<u>\$ 11,165</u>

General and administrative expenses for the three months ended June 30, 2025 were \$11.9 million, compared to \$11.2 million for the three months ended June 30, 2024 primarily reflecting higher legal and professional fees, due in part to higher tax and public company compliance costs as well as additional intellectual property related costs. Personnel expenses in the three months ended June 30, 2025 decreased \$0.5 million compared with the three months ended June 30, 2024, reflecting lower share-based compensation expense of \$0.8 million partially offset by higher salaries expense from increased headcount.

### **Interest and Investment Income and Interest Expense**

For the three months ended June 30, 2025, interest income was \$4.4 million, which was up \$1.1 million from the three months ended June 30, 2024, largely reflecting higher interest earned from the Company's marketable securities due to a higher average cash, cash equivalent and investments balance versus a year ago.

Interest expense was \$2.9 million in the three months ended June 30, 2025, up \$0.4 million from the three months ended June 30, 2024, as a result of a higher average debt balance.

### **Other Non-Operating Income, net**

Other non-operating income (expense), net for the three months ended June 30, 2025 was a net gain of \$3.6 million, mainly reflecting foreign currency transaction gains. Other non-operating income, net for the three months ended June 30, 2024 was \$0.2 million, primarily reflecting a gain from the fair value remeasurement of the note purchase agreement with Three Peaks Capital Solutions Aggregator Fund and Cocoon SA LLC, an affiliate of Oberland Capital Management LLC ("Note Purchase Agreement") as of June 30, 2024, partially offset by foreign currency transaction losses.

### **Comparison of the Six Months Ended June 30, 2025 and June 30, 2024**

The following table sets forth the results of operations for the six months ended June 30, 2025 and June 30, 2024 (amounts in thousands):

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
License and other revenue	\$ 15,000	\$ —
Operating expenses:		
Research and development	76,184	55,467
General and administrative	24,246	24,603
Loss from operations	(85,430)	(80,070)
Interest and investment income	12,270	5,831
Interest expense	(5,761)	(5,054)
Other non-operating income (expense), net	4,618	(1,383)
Loss before income tax expense	(74,303)	(80,676)
Income tax expense	2,175	1,186
Net loss	<u>\$ (76,478)</u>	<u>\$ (81,862)</u>

### ***License and other revenue***

On February 14, 2025, the Company entered into a license agreement (the “License Agreement”) with Genmab A/S (“Genmab”). During the six months ended June 30, 2025, the Company recorded revenue of \$15.0 million related to the upfront payment received from Genmab upon execution of the License Agreement. See [Note 3 - Genmab License Agreement](#).

### ***Research and Development Expenses***

The following table summarizes research and development expenses by program incurred for the following periods (amounts in thousands):

	<b>Six Months Ended June 30, 2025</b>	<b>Six Months Ended June 30, 2024</b>
<b>Programs:</b>		
Ox2R agonist	\$ 52,774	\$ 14,952
LockBody	5,183	6,266
<b>Discontinued programs</b>	<b>4,109</b>	<b>28,619</b>
<b>Non-asset specific costs:</b>		
Personnel expenses	20,651	18,231
Research tax incentives	(8,734)	(13,395)
Other preclinical and clinical development expenses	2,201	794
<b>Research and development expenses</b>	<b>\$ 76,184</b>	<b>\$ 55,467</b>

Research and development expenses for the six months ended June 30, 2025 were \$76.2 million, compared with \$55.5 million for the six months ended June 30, 2024. The increase in research and development expenses reflected higher development costs of \$37.8 million for our Ox2R agonist program, a decline in research tax incentives and higher personnel expenses, partially offset by lower costs related to discontinued programs. The higher developmental costs for Ox2R agonist program reflected higher clinical study costs for ORX750 as well as developmental milestones related to ORX142 as a result of FDA clearance of its IND and initiation of its Phase 1 clinical trial. Personnel expenses in the first six months of 2025 increased \$2.4 million compared to the first six months ended June 30, 2024 driven by increased headcount and higher share-based compensation expense of \$0.5 million.

### ***General and Administrative Expenses***

The following table summarizes the general and administrative expenses for the following periods (amounts in thousands):

	<b>Six Months Ended June 30, 2025</b>	<b>Six Months Ended June 30, 2024</b>
Personnel expenses	\$ 13,922	\$ 14,755
Legal and professional fees	5,756	4,317
Other expenses	4,568	5,531
	<b>\$ 24,246</b>	<b>\$ 24,603</b>

General and administrative expenses for the six months ended June 30, 2025 were \$24.2 million, compared to \$24.6 million for the six months ended June 30, 2024 primarily reflecting higher legal and professional fees, driven in part by higher tax and public company compliance costs as well as additional IP related costs. The higher legal and professional fees were partially offset by lower personnel and other expenses in 2025, largely reflecting lower insurance costs. Personnel expenses in the first six months of 2025 decreased \$0.8 million compared with the first six months of 2024, reflecting lower share-based compensation expense of \$1.5 million, partially offset by higher salaries from increased headcount.

### ***Interest and Investment Income and Interest Expense***

For the six months ended June 30, 2025, interest and investment income was \$12.3 million, which was up \$6.4 million from the six months ended June 30, 2024, largely reflecting the recognition of unrealized gains of marketable securities as of

December 31, 2024 upon redemption in 2025 as well as higher interest earned from the Company's fixed investments due to a higher average cash, cash equivalent and investments balance compared to the prior year period.

Interest expense was \$5.8 million in the first six months of 2025, up \$0.7 million from the first six months of 2024, as a result of a higher average debt balance.

#### ***Other Non-Operating Income (Expense), net***

Other non-operating income (expense), net for the six months ended June 30, 2025 was a net gain of \$4.6 million, primarily reflecting foreign currency transaction gains. Other expense, net for the six months ended June 30, 2024 was \$1.4 million, mainly due to a \$0.8 million loss related to the remeasurement of the Note Purchase Agreement at fair value as of June 30, 2024, and foreign currency transaction losses of \$0.5 million.

### **Liquidity and Capital Resources**

#### ***Sources of Liquidity***

As of June 30, 2025, we had cash, cash equivalents and investments of \$404.1 million, of which \$44.2 million was classified as cash and cash equivalents, \$206.5 million was classified as short-term investments and \$153.4 million was classified as long-term investments on our Consolidated Balance Sheet. The Company invests in money market funds, U.S. Treasury securities, U.S. government agency securities, corporate notes and commercial paper. The Company's investment policy limits investments to money market funds, certain types of debt securities issued by the U.S. Government and its agencies, corporate notes and commercial paper, and places restrictions on the credit ratings, maturities and concentration by type and issuer. Securities with original maturities of three months or less when purchased are included in cash and cash equivalents. We consider investments with original maturities greater than three months and remaining maturities less than one year to be short-term investments, while remaining maturities greater than one year are classified as long-term investments.

On December 30, 2024, the Company entered into a loan and security agreement (the "Loan and Security Agreement") with Oxford Finance LLC ("Oxford"), as collateral agent and a lender, and the other lenders from time to time party thereto (collectively, the "Lenders"), pursuant to which the Lenders have agreed to lend the Company an aggregate principal amount of up to \$200.0 million in a series of term loans (the "Term Loans").

Pursuant to the Loan and Security Agreement, the Company received \$110.0 million (the "Initial Term Loan") and incurred \$1.1 million of debt issuance costs inclusive of facility and legal fees. The Company has access to up to an additional \$40.0 million of loan proceeds in an additional tranche which is available during the period commencing on the date of the occurrence of the Clinical Milestone (as defined in the Loan and Security Agreement) through the earlier of: (i) 90 days following the Clinical Milestone and (ii) June 30, 2028. An additional \$50.0 million may be made available to the Company at the Lenders' sole discretion.

The term loans are set to mature on December 1, 2029 and, following an interest-only period, will begin to amortize in equal monthly installments beginning on February 1, 2029. However, if the Extension Event as defined in the Agreement occurs, then at the Company's option, the term loans could begin to amortize in equal monthly installments beginning on February 1, 2030, and the maturity date will be extended to December 1, 2030.

On September 11, 2024, we filed an automatic shelf registration statement on Form S-3ASR ("Shelf") registering an unspecified amount of our ordinary shares, American Depositary Shares representing ordinary shares, debt securities, warrants, and/or units or any combination thereof with the SEC under the Securities Act. The Shelf automatically became effective upon filing. Under the Shelf, we may offer securities from time to time in one or more offerings, at prices and on terms to be determined by market conditions at the time of offering. The specifics of any future offerings, along with the use of proceeds of any securities offered, will be described in detail in a prospectus supplement, or other offering materials, at the time of any offering.

The Company entered into a Sales Agreement, dated January 27, 2023, by and between Centessa Pharmaceuticals plc and Leerink Partners LLC (formerly SVB Securities LLC). As sales agent, Leerink Partners LLC will provide for the issuance and sale by the Company of up to \$125.0 million of its ordinary shares represented by American Depositary Shares ("ADSs") from time to time in "at-the-market" offerings ("ATM Program"). In the six months ended June 30, 2025, the Company sold 372,538 ordinary shares under the ATM Program, resulting in net proceeds of \$6.1 million, all of which were collected during

the first quarter of 2025. Since the inception of the ATM program in 2023, as of June 30, 2025, the Company has sold 4,663,354 ordinary shares, resulting in net proceeds to us of approximately \$36.6 million.

### **Cash Flows**

The following table shows a summary of cash flows for the periods indicated (amounts in thousands):

	Six Months Ended June 30, 2025	Six Months Ended June 30, 2024
Net cash (used in) provided by:		
Operating activities	\$ (79,698)	\$ (77,866)
Investing activities	(262,342)	(38,697)
Financing activities	4,691	115,806
Exchange rate effect on cash and cash equivalents	(1,630)	99
Net decrease in cash and cash equivalents	<u>\$ (338,979)</u>	<u>\$ (658)</u>

### **Operating Activities**

During the six months ended June 30, 2025, we used \$79.7 million of net cash in operating activities. This reflected a net loss of \$76.5 million, adjusted by non-cash charges of \$14.7 million, primarily related to share-based compensation, depreciation and amortization expenses. In addition, cash used in operating activities included net working capital outflows of approximately \$20.0 million largely due to the payment of \$27.4 million in program terminations costs accrued in the fourth quarter of 2024, partially offset by increased receipts of tax incentives.

During the six months ended June 30, 2024, net cash used in operating activities was \$77.9 million. Cash used in operating activities reflected a net loss of \$81.9 million reduced by non-cash charges of \$18.0 million, which included share-based compensation, depreciation and amortization expense and changes in the fair value of debt. In addition, cash used in operating activities during the six months ended June 30, 2024 included net outflows of approximately \$15.0 million related to paying down net payables during the period.

### **Investing Activities**

During the six months ended June 30, 2025, net cash used in investing activities was \$262.3 million, largely reflecting the net investment of excess cash in marketable securities. During the six months ended June 30, 2024, net cash used in investing activities was \$38.7 million, largely reflecting the net purchase of marketable securities.

### **Financing Activities**

During the six months ended June 30, 2025, net cash provided by financing activities was \$4.7 million, primarily reflecting the proceeds from the ATM program in the first quarter and proceeds received from employee stock option exercises, partially offset by payment of employee withholding taxes related to share-based compensation. During the six months ended June 30, 2024, net cash provided by financing activities was \$115.8 million, primarily reflecting the proceeds from a share offering as well as from sales of shares under the ATM program.

### **Funding Requirements**

In future years, we expect expenses to increase as we continue further development of our product candidates and ultimately seek marketing approval for any current and future product candidates. In addition, we will begin to incur pre-commercial preparatory activities and, if marketing approval is obtained for any product candidates, we expect to incur significant commercialization expenses related to product sales, marketing, manufacturing and distribution. In addition, inflation may affect our use of capital resources by increasing our cost of labor, research and clinical trial expenses. Accordingly, there will be a need to obtain substantial additional funding in connection with the continuing operations. If we

are unable to raise capital when needed or on attractive terms, we would be forced to delay, reduce or eliminate research and development programs or future commercialization efforts.

We anticipate that our expenses will increase substantially as we:

- seek to discover and develop current and future clinical and preclinical product candidates;
- scale up clinical and regulatory capabilities;
- adapt regulatory compliance efforts to incorporate requirements applicable to marketed products;
- establish a sales, marketing and distribution infrastructure and scale up external manufacturing capabilities to commercialize any product candidates for which regulatory approval may be obtained;
- maintain, expand and protect the intellectual property portfolio;
- hire additional internal or external clinical, manufacturing and scientific personnel or consultants;
- add operational, financial and management information systems and personnel, including personnel to support product development efforts; and
- incur additional legal, accounting and other expenses in operating as a public company.

Because of the numerous risks and uncertainties associated with the research, development and commercialization of product candidates, we are unable to estimate the exact amount of our working capital requirements. Future funding requirements will depend on and could increase significantly as a result of many factors, including:

- the scope, progress, results and costs of preclinical studies and clinical trials;
- the scope, prioritization and number of research and development programs;
- the costs, timing and outcome of regulatory review of product candidates;
- the ability to establish and maintain collaborations on favorable terms, if at all;
- the costs of preparing, filing and prosecuting patent applications, maintaining and enforcing intellectual property rights and defending intellectual property-related claims;
- the costs of securing manufacturing arrangements for commercial production; and
- the costs of establishing or contracting for sales and marketing capabilities if regulatory approvals are obtained to market product candidates.

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

### **Critical Accounting Policies and Estimates**

Our management's discussion and analysis of our financial condition and results of operations is based on our consolidated financial statements, which have been prepared in accordance with generally accepted accounting principles in the United States. The preparation of these consolidated financial statements requires us to make estimates and assumptions that affect the reported amounts of assets and liabilities and the disclosure of contingent assets and liabilities at the date of the consolidated financial statements, as well as the reported expenses incurred during the reporting periods. Our estimates are based on our historical experience and on various other factors that we believe are reasonable under the circumstances, the results of which form the basis for making judgments about the carrying value of assets and liabilities that are not readily apparent from other sources. Actual results may differ from these estimates under different assumptions or conditions.

As of June 30, 2025, the Company's critical accounting policies and estimates had not changed significantly from December 31, 2024. Note 2, "Summary of Significant Accounting Policies," to the consolidated financial statements in our Annual Report on Form 10-K for the fiscal year ended December 31, 2024 describes the significant accounting policies and methods used in the preparation of the consolidated financial statements appearing in this Quarterly Report on Form 10-Q.

### **Contractual Obligations and Other Commitments**

As of June 30, 2025, other than what has been disclosed in Note 6 - Debt and Note 7 - Commitments and contingencies, there were no material changes to our contractual obligations and other commitments described under “Management’s Discussion and Analysis of Financial Condition and Results of Operations” in our Annual Report on Form 10-K for the fiscal year ended December 31, 2024.

### **Emerging Growth Company and Smaller Reporting Company Status**

We are an emerging growth company, as defined in the JOBS Act. Under the JOBS Act, emerging growth companies can delay the adoption of new or revised accounting standards issued subsequent to the enactment of the JOBS Act until such time as those standards apply to private companies. Other exemptions and reduced reporting requirements under the JOBS Act for emerging growth companies include presentation of only two years of audited financial statements in a registration statement for an initial public offering, an exemption from the requirement to provide an auditor’s report on internal controls over financial reporting pursuant to Section 404 of the Sarbanes-Oxley Act of 2002, as amended, an exemption from any requirement that may be adopted by the Public Company Accounting Oversight Board regarding mandatory audit firm rotation and less extensive disclosure about our executive compensation arrangements. We have elected to use the extended transition period for complying with new or revised accounting standards that have different effective dates for public and private companies until the earlier of the date that (i) we are no longer an emerging growth company or (ii) we affirmatively and irrevocably opt out of the extended transition period provided in the JOBS Act.

We will remain an emerging growth company until the earliest of (i) the last day of our first fiscal year in which we have total annual gross revenues of \$1.235 billion or more, (ii) the last day of our first fiscal year following the fifth anniversary of the closing of our initial public offering, (iii) the date on which we are deemed to be a “large accelerated filer,” under the rules of the SEC, which means the market value of equity securities that is held by non-affiliates exceeds \$700.0 million as of the prior June 30th and (iv) the date on which we have issued more than \$1.0 billion in non-convertible debt securities during the prior three-year period.

We are also a “smaller reporting company” as defined in the Securities Exchange Act of 1934 (the “Exchange Act”). 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.

On the last business day of our second quarter in 2025, the aggregate market value of our shares of common stock held by non-affiliates exceeded \$700 million. As a result, as of December 31, 2025, we will be considered a “large accelerated filer” as defined in Rule 12b-2 under the Exchange Act, and we will cease to be an “emerging growth company” as defined in the JOBS Act and a “smaller reporting company” as defined in the Exchange Act. We will no longer be exempt from the auditor attestation requirements of Section 404(b) of the Sarbanes-Oxley Act, and our independent registered public accounting firm will evaluate and report on the effectiveness of internal control over financial reporting. We will also no longer be permitted to take advantage of reduced reporting requirements for smaller reporting companies.

### **Item 3. Qualitative and Quantitative Disclosures About Market Risk**

Under SEC rules and regulations, as a smaller reporting company, we are not required to provide the information required by this item.

### **Item 4. Controls and Procedures**

Our management, with the participation of our Chief Executive Officer and Chief Financial Officer (our principal executive officer and principal financial officer, respectively), evaluated the effectiveness of our disclosure controls and procedures as of June 30, 2025. The term “disclosure controls and procedures,” as defined in Rules 13a-15(e) and 15d-15(e) under the Securities Exchange Act of 1934, as amended (“Exchange Act”), means controls and other procedures of a company that are designed to ensure that information required to be disclosed by a company in the reports that it files or submits under the Exchange Act is recorded, processed, summarized and reported, within the time periods specified in the SEC’s rules and forms. Disclosure controls and procedures include, without limitation, controls and procedures designed to ensure that information required to be disclosed by us in the reports we file under the Exchange Act is accumulated and communicated to our management, including our principal executive and principal financial officers, as appropriate to allow timely decisions

regarding required disclosure. Our management recognizes that any controls and procedures, no matter how well designed and operated, can provide only reasonable assurance of achieving their objectives and our management necessarily applies its judgment in evaluating the cost-benefit relationship of possible controls and procedures. Our disclosure controls and procedures are designed to provide reasonable assurance of achieving their control objectives.

Based on the evaluation of the effectiveness of our disclosure controls and procedures as of June 30, 2025, the end of the period covered by this Quarterly Report on Form 10-Q, our Chief Executive Officer and Chief Financial Officer concluded that, as of such date, our disclosure controls and procedures were effective at the reasonable assurance level.

#### **Changes in Internal Control over Financial Reporting**

No changes in our internal control over financial reporting (as defined in Rules 13a-15(f) and 15d-15(f) under the Exchange Act) have occurred during the three months ended June 30, 2025 that have materially affected, or are reasonably likely to materially affect, our internal control over financial reporting.

## PART II - OTHER INFORMATION

### Item 1. Legal Proceedings

None

### Item 1A. Risk Factors

*Our business involves significant risks, some of which are described below. You should carefully consider the risks described below, as well as the other information in this Quarterly Report on Form 10-Q and in other documents we file with the SEC, including our financial statements and the related notes and "Management's Discussion and Analysis of Financial Condition and Results of Operations." The occurrence of any of the events or developments described below could have a material adverse effect on our business, financial condition, results of operations, growth prospects and stock price. In such an event, the market price of our ADSs could decline, and you may lose all or part of your investment. Additional risks and uncertainties not presently known to us or that we currently deem immaterial may also impair our business operations and the market price of our ADSs.*

#### **Risks Related to our Business Model and Structure**

*We may not be successful in our efforts to use our differentiated asset-centric approach to drug discovery and development to build a pipeline of product candidates with commercial value.*

A key element of Centessa's strategy is to use our differentiated asset-centric approach to drug discovery and development to develop high conviction programs, product candidates, technologies or intellectual property that we believe are novel, employ differentiated mechanisms of action, or have a combination of these attributes to ultimately deliver transformational medicines to patients. We face significant competition in sourcing high conviction programs, product candidates, technologies or intellectual property, strategic partnerships and licensing and acquisition opportunities, and the negotiation process is time-consuming, costly and complex. We may not be successful in our efforts in building a pipeline of high conviction product candidates for the treatment of various diseases and disorders through acquisitions, licensing or through internal development or in progressing these product candidates through clinical development. Although our research and development efforts to date have resulted in our identification, discovery and preclinical and clinical development of certain of our product candidates, these product candidates may not be safe or effective treatments or therapies in humans, and we may not be able to develop any other product candidates. Although we analyze whether we can replicate scientific results observed prior to our acquisition or investment in a product candidate, we may not be successful in doing so after our investment. Our asset-centric approach to drug discovery and development is evolving and may not succeed in building a pipeline of product candidates. Even if we are successful in building our pipeline of product candidates, the potential product candidates that we identify may not be suitable for clinical development or generate acceptable clinical data in humans, including as a result of unacceptable toxicity or other characteristics that indicate that they are unlikely to receive marketing approval from the FDA, or other regulatory authorities or achieve market acceptance. If we do not successfully develop and commercialize product candidates, we will not be able to generate product revenue in the future, which likely would result in significant harm to our financial position and adversely affect the price of our ADSs.

As part of our business strategy, we may expand our product candidate pipeline through in-licenses or acquisitions of discovery or development-stage assets or programs, which entails additional risk to us. While we believe our asset-centric approach offers an attractive platform for these transactions and for founder subject-matter experts and potential partners, our approach is unique and we may not be able to attract or execute transactions with founder-subject matter experts, sellers, licensors or collaborators who may choose to divest to or grant license to companies that employ more traditional licensing and collaboration approaches. Identifying, selecting, and acquiring promising product candidates requires substantial technical, financial and human resources expertise. Efforts to do so may not result in the actual acquisition or license of a successful product candidate, potentially resulting in a diversion of our management's time and the expenditure of our resources with no resulting benefit. For example, if we are unable to identify programs that ultimately result in approved products, we may spend material amounts of our capital and other resources evaluating, acquiring, and developing product candidates that ultimately do not provide a return on our investment. We may terminate programs in the future if they do not meet our criteria for advancement.

***A single or limited number of programs, developmental assets or product candidates may comprise a large proportion of our value.***

A large proportion of our value may at any time reside in a limited number of our programs and/or developmental assets or product candidates. Our consolidated financial condition and prospects may be materially diminished if the clinical development or potential commercialization prospects of one of our product candidates or programs or one or more of the intellectual property rights held by us become impaired. Furthermore, a large proportion of our consolidated revenue may at any time be derived from one, or a small number of, licensed technologies, and termination or expiration of licenses to these technologies would likely have a material adverse effect on our consolidated revenue. Any material adverse impact on the value of intellectual property rights or the clinical development of product candidates or programs, could have a material adverse effect on our consolidated business, financial condition, results of operations or prospects.

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

Because we have limited financial and managerial resources, we must focus on a limited number of research programs and product candidates and on specific indications. As a result, we may forego or delay pursuit of opportunities with other product candidates or for other indications that later prove to have greater commercial potential, or we may fail to recognize or acquire assets that may be more promising than those we acquire. 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 identification, discovery, and preclinical development programs and product candidates for specific indications may not yield any commercially viable products.

***We face challenges, risks and expenses related to our operations as well as the management of the expected growth in the scale and complexity of our operations.***

In January 2021, we acquired the ownership interests of our Centessa Subsidiaries. These Centessa Subsidiaries historically operated as independent entities with generally separate management and operational teams. As a result, we have needed to expend significant resources and efforts in integrating the operations of these Centessa Subsidiaries into our larger organization, and such integration activities may be challenging due to the number of Centessa Subsidiaries acquired and the heterogeneity of their historical operations. For example, these Centessa Subsidiaries' programs span a range of therapeutic modalities and are designed to address a variety of disease areas. In addition, the Centessa Subsidiaries have conducted their business in a variety of jurisdictions in the U.S. and Europe. All of our Centessa Subsidiaries have had historical relationships with different licensors, contract organizations and other third-party vendors.

Each Centessa Subsidiary historically had its own operational, legal, financial and management controls, reporting systems and procedures and integrating such controls, reporting systems and procedures may be challenging and we may not be successful in doing so. We believe certain synergies may be achieved by harmonizing the operational, legal, financial and management controls, reporting systems and procedures but we may not be successful in our harmonization efforts and this may result in not only being unable to take advantage of synergies but expose us to additional operational, legal and financial risks and exposures associated with several levels of disparate systems and procedures. With limited resources, historically the Centessa Subsidiaries may not have dedicated sufficient resources to ensure its operational, legal, financial and management controls, reporting systems, compliance and other procedures met required standards and this may expose us to historical non-compliance investigations and liabilities, which may have a material adverse effect on our operations.

In mid-2023, we implemented a corporate consolidation of our U.S. and UK businesses in order to simplify our administrative operations, obtain operational efficiencies, generate administrative cost savings and improve the overall control environment. This reorganization transferred the business operations and assets of our UK subsidiaries (Z Factor Limited, Centessa Pharmaceuticals (Morphogen-IX) Limited, Capella Bioscience Ltd, LockBody Therapeutics Ltd, Inexia Limited, Centessa Pharmaceuticals (Orexia) Limited and Janpix Limited) to Centessa Pharmaceuticals (UK) Limited ("CPUK"). Subsequently, on December 26, 2023, Inexia Limited, Janpix Limited, Ultrahuman Two Limited and Ultrahuman Four Limited were dissolved using the voluntary strike off procedure in sections 1003 to 1011 of the UK Companies Act 2006. On December 17, 2024, Centessa Pharmaceuticals (Morphogen-IX) Limited, Capella Bioscience Ltd, LockBody Therapeutics Ltd, and Centessa Pharmaceuticals (Orexia) Limited were dissolved using the voluntary strike off procedure in sections 1003 to 1011 of the Companies Act, and we intend to dissolve Z Factor Limited over the coming months also using the voluntary strike off procedure in sections 1003 to 1011 of the Companies Act. CPUK, a wholly owned subsidiary of Centessa Pharmaceuticals plc, was incorporated in 2020 under the laws of England and Wales with primary operations in the United Kingdom. ApcinteX Limited was incorporated in 2014 under the laws of England and Wales with primary operations in the United Kingdom.

As of August 1, 2025 we had 98 full-time equivalent employees. We may not be successful in retaining employees or finding replacements which could have a material adverse effect on our ability to develop and commercialize our programs and product candidates. As our development and commercialization plans and strategies develop, and as we refine our operations as a public company, we expect to need additional managerial, operational, sales, marketing, legal, financial and other personnel. Future growth would impose significant added responsibilities on members of management, including:

- identifying, recruiting, integrating, maintaining and motivating additional employees;
- managing our internal development efforts effectively, including the clinical and FDA review process for our product candidates, while complying with our contractual obligations to contractors and other third parties; and
- improving our operational, legal, financial and management controls, reporting systems and procedures.

Our future financial performance and our ability to commercialize any product candidates that are approved for marketing will depend, in part, on our ability to effectively manage any future growth, and our management may also have to divert a disproportionate amount of its attention away from day-to-day activities in order to devote a substantial amount of time to managing these growth activities.

We currently rely, and for the foreseeable future will continue to rely, in substantial part on certain independent organizations, advisors and consultants to provide certain services, including substantially all aspects of regulatory approval, clinical trial management and manufacturing. There can be no assurance that the services of independent organizations, advisors and consultants will continue to be available to us on a timely basis when needed, or that we can find qualified replacements. In addition, if we are unable to effectively manage our outsourced activities or if the quality or accuracy of the services provided by consultants is compromised for any reason, our clinical trials may be extended, delayed or terminated, and we may not be able to obtain regulatory approval of our product candidates or otherwise advance our business. There can be no assurance that we will be able to manage our existing consultants or find other competent outside contractors and consultants on economically reasonable terms, or at all.

If we are not able to effectively expand our organization by hiring new employees and expanding our groups of consultants and contractors, we may not be able to successfully implement the tasks necessary to further develop and potentially commercialize our product candidates and, accordingly, may not achieve our research, development and commercialization goals. We may not have sufficient funding to support our expansion.

***Our reliance on a small team of employees located in different geographies who provide services (including administrative, research and development, and other services) across our organization presents operational challenges that may adversely affect our business.***

As of August 1, 2025, we had 98 full-time equivalent employees who are located in different geographies across the U.S., UK and the European Union who provide services across our organization (including operational, administrative, research and development, and other support services). We also have consultants who we rely on for research and development, business development, and other services. While we believe this structure enables us to reduce certain infrastructure costs, the small size of our team may limit our ability to devote adequate personnel, time, and resources to support our operational, research and development activities, and the management of compliance, financial, accounting, and reporting matters. If our team fails to provide adequate operational, administrative, research and development, or other services across our entire organization, our business, financial condition, and results of operations could be harmed.

***Some of our officers currently serve, and in the future may serve, as directors or officers of our Centessa Subsidiaries, and, as a result, have and may continue to have, statutory, fiduciary and other duties to our subsidiaries causing conflicts of interest with respect to their duties to us and their duties to our subsidiaries and in determining how to devote themselves to our affairs and the affairs of our subsidiaries. Our subsidiaries' partners may also disagree with the sufficiency of resources that we provide to each Centessa Subsidiary.***

Certain of our officers, including Iqbal Hussain, our General Counsel and Gregory Weinhoff, our Chief Business Officer are directors and/or officers of certain Centessa Subsidiaries and, as a result, have fiduciary or other duties both to us and our subsidiaries. Dr. Weinhoff and Mr. Hussain do not receive any additional compensation for their service as directors of our Centessa Subsidiaries. The conflicts of interest that arise from such duties could interfere with the management of our subsidiaries and their programs and product candidates, or result in disagreements with our subsidiaries' partners. For example, an individual who is both a director of one of our subsidiaries and an officer of Centessa owes statutory and fiduciary duties to the Centessa Subsidiary and to us, and such individual may encounter circumstances in which his or her decision or action may benefit the Centessa Subsidiary while having a detrimental impact on Centessa, or vice versa, or on another Centessa

Subsidiary, including one for which he or she also serves as a director. Further, in the future, certain of our officers may serve as officers and directors of our Centessa Subsidiaries. Any such individual would need to allocate his or her time to responsibilities owed to Centessa and each of the Centessa Subsidiaries for which he or she serves as an officer or director, and would make decisions on behalf of one entity that may negatively impact others. In addition, disputes could arise between us and our Centessa Subsidiary's partners regarding a conflict of interest or perceived conflict of interest arising from the overlap between the officers and directors of the Centessa Subsidiary and those of Centessa. These partners also may disagree with the amount and quality of resources that are devoted to the Centessa Subsidiary they are invested in. Any such disputes or disagreements could distract our management, interfere with our relations with our partners, and take significant time to resolve, which could disrupt the development of our product candidates, delay our potential commercialization efforts, result in increased costs or make it less likely that other third parties will choose to partner with us in the future.

***Certain of our programs are subject to certain agreements that provide our licensors and/or collaborators with rights that could delay or impact our ability to sell assets, or enter into strategic alliances, collaborations or licensing arrangements with other third parties or the potential sale of our Centessa Subsidiaries.***

Certain of our programs are subject to licenses of intellectual property from third parties and we expect such practice to continue in the future. These third parties have certain rights that could delay collaboration, licensing or other arrangements with another third party, and the existence of these rights may adversely impact our ability to attract an acquirer or partner. These rights include rights of negotiation and fees payable upon a sale of assets or change of control of the Company or a Centessa Subsidiary that are contained in license agreements, payments upon satisfaction of milestones, royalty payments, diligence obligations and other customary terms contained in agreements for the in-license of programs and their intellectual property.

We may incorporate, form or otherwise acquire additional subsidiaries and enter into similar agreements with future counterparties, or our Centessa Subsidiaries may enter into further agreements, that in each case may contain similar provisions or other terms that are not favorable to us.

***Preclinical and clinical development is a long, expensive and uncertain process, and we may terminate one or more of our current preclinical and/or clinical development programs.***

We may determine that certain product candidates or programs (preclinical and/or clinical) do not have sufficient potential to warrant the continued allocation of resources toward them. Accordingly, we may elect to terminate our programs for and, in certain cases, our licenses to, such product candidates or programs. If we terminate programs in which we have invested significant resources, we will have expended resources on a program that will not provide a full return on our investment and missed the opportunity to have allocated those resources to potentially more productive uses. In addition, program termination may result in significant additional wind-down related costs being incurred including penalties, redundancy and severance and professional fees and may expose us to additional risks including contractual breach and employment termination claims and may divert a disproportionate amount of management time. For example, in November 2024, we announced the voluntary discontinuation of the global clinical development program for SerpinPC, a novel inhibitor of APC that was being progressed for the treatment of hemophilia B. More recently, during the first quarter of 2025, we discontinued the clinical development of LB101, a conditionally tetravalent PD-L1xCD47 bispecific monoclonal antibody, and first generation LockBody candidate. We may not be able to terminate a clinical program with an ongoing clinical trial on medical and other grounds and, to the extent we are able to terminate, such termination may expose us to additional risks including regulatory risk.

#### **Risks Related to our Financial Position, Need for Additional Capital and Growth Strategy**

***We, and our Centessa Subsidiaries, have incurred net losses since inception, and we expect to continue to incur losses for the foreseeable future and may never achieve or maintain profitability.***

We and our Centessa Subsidiaries have incurred significant net losses since inception, have not generated any revenue from product sales to date, and financed operations primarily through equity and debt financing. Centessa Pharmaceuticals plc has a limited operating history, and we expect to incur significant losses for the foreseeable future. As an organization, we have devoted substantially all of our efforts to research and development, including clinical and preclinical development of our product candidates, as well as to building out our team. We expect that it could be several years, if ever, before we have a commercialized product candidate. We expect to continue to incur significant expenses and operating losses for the foreseeable

future. The net losses we incur may fluctuate significantly from quarter to quarter each financial year. In addition, inflation could adversely impact our financial results. We anticipate that our expenses will increase substantially if, and as, we:

- continue our research and the preclinical and clinical development of our product candidates, including our ongoing and planned clinical trials;
- initiate additional clinical trials and preclinical studies for our other product candidates, including those in our pipeline that are expected to advance into the clinic in the near future; if any of our product candidates advance through and complete late-stage development, prepare and submit marketing applications with the FDA and comparable regulatory authorities;
- experience any delays or encounter any issues with any of the above, including but not limited to failed studies, complex results, safety issues or other regulatory challenges;
- seek to discover and develop additional product candidates;
- establish a sales, marketing and distribution infrastructure to commercialize any product candidates for which we may obtain marketing approval;
- maintain, expand and protect our intellectual property portfolio;
- fulfill future potential payment obligations under our incentivization agreements with each Centessa Subsidiary or program, including as a result of meeting program milestones, program divestments, Company change of control, asset sale or out-licensing; and
- acquire or in-license other product candidates and technologies.

To become and remain profitable, we must develop and eventually commercialize product candidates with significant market potential. 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 marketing approval for these product candidates, manufacturing, marketing and selling those products for which we may obtain marketing approval and satisfying any post-marketing requirements. We may never succeed in any or all of these activities and, even if we do, we may never generate revenues that are significant or large enough to achieve profitability. If we do 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 decrease the value of our company and could impair our ability to raise capital, maintain our research and development efforts and expand our business or continue our operations. A decline in the value of our company could cause you to lose all or part of your investment.

***Our limited operating history may make it difficult for investors to evaluate our business, operations and prospects.***

Our operations to date have been limited to organizing and staffing our company, business planning, developing our operating model, raising capital, acquiring our technology, identifying potential product candidates, establishing collaborations and undertaking preclinical studies and clinical trials of our most advanced product candidates. As an organization, we have not yet demonstrated a track record of completing pivotal and/or Phase 3 trials of our product candidates, obtaining marketing approvals, manufacturing a commercial-scale product or conducting sales and marketing activities necessary for successful commercialization. Consequently, any predictions made about our future success or viability may not be as accurate as they could be if we had a longer operating history.

In addition, as a new business, we may encounter unforeseen expenses, difficulties, complications, delays and other known and unknown factors. We will need to transition from a company with a research focus to a company that is also capable of supporting commercial activities. We may not be successful in such a transition.

***We have never generated revenue from product sales and may never be profitable.***

Our ability to generate revenue from product sales and achieve profitability depends on our ability, alone or with collaborative partners, to successfully complete the development of, and obtain the regulatory approvals necessary to commercialize, our product candidates. We do not anticipate generating revenues from product sales for the next several years, if ever. Our ability to generate future revenues from product sales depends heavily on our, or our collaborators', success in:

- completing research and preclinical and clinical development of our product candidates;
- seeking and obtaining regulatory and marketing approvals for product candidates for which we complete clinical trials;

- in-licensing, acquiring, discovering or otherwise expanding our pipeline of product candidates for clinical development;
- launching and commercializing product candidates for which we obtain regulatory and marketing approval by establishing a sales force, marketing and distribution infrastructure or, alternatively, collaborating with a commercialization partner;
- qualifying for adequate coverage and reimbursement by government and third-party payors for our product candidates;
- maintaining and enhancing a sustainable, scalable, reproducible and transferable manufacturing process for our product candidates;
- establishing and maintaining supply and manufacturing relationships with third parties that can provide adequate, in both amount and quality, products and services to support clinical development and the market demand for our product candidates, if approved;
- obtaining market acceptance of our product candidates as a viable treatment option;
- addressing any competing technological and market developments;
- implementing additional internal systems and infrastructure, as needed;
- negotiating favorable terms in any collaboration, licensing or other arrangements into which we may enter and performing our obligations in such collaborations;
- maintaining, protecting and expanding our portfolio of intellectual property rights, including patents, trade secrets and know-how;
- avoiding and defending against third-party interference or infringement claims; and
- attracting, hiring and retaining qualified personnel.

Even if one or more of the product candidates that we develop is approved for commercial sale, we anticipate incurring significant costs associated with commercializing any approved product candidate. Our expenses could increase beyond expectations if we are required by the FDA, the EMA, the MHRA, or other regulatory authorities to perform clinical and other studies in addition to those that we currently anticipate. For example, the FDA may require us to perform additional clinical trials of ORX750, beyond those we are currently planning to conduct, in order to support an NDA submission in due course. Even if we are able to generate revenues from the sale of any approved products, we may not become profitable and may need to obtain additional funding to continue operations.

***We will need substantial additional funds to advance development of our product candidates, and we cannot guarantee that we will have sufficient funds available in the future to develop and commercialize our current or future product candidates.***

Developing pharmaceutical products, including conducting preclinical studies and clinical trials, is a very time-consuming, expensive and uncertain process that takes years to complete. We will need substantial additional funds to expand our development, regulatory, manufacturing, marketing and sales capabilities or contract with other organizations in order to enter and advance our product candidates through preclinical studies and clinical trials. For example, in December 2024 we entered into the Oxford Finance Loan and Security Agreement (See Note 6 – “Debt” in our 2024 Annual Report on Form 10-K for more information). In January 2023, we entered into an Open Market Sale Agreement (the “2023 Sale Agreement”) with Leerink, under which Leerink is able to offer and sell, from time to time in “at-the-market” (“ATM”) offerings, shares of the Company’s ADSs having aggregate gross proceeds of up to \$125 million. In the event of a sale of Company shares under the ATM, the Company is obligated to pay to Leerink cash commissions of up to 3.0% of the gross proceeds of sales of ADSs under the 2023 Sale Agreement. Our Centessa Subsidiaries have used substantial funds in their research and development programs and will continue to expend significant resources to advance their programs and product candidates.

As of June 30, 2025, we had cash, cash equivalents and investments of \$404.1 million. Based on our current operating model and development plans, which include certain assumptions, the Company expects cash, cash equivalents and investments to fund its operations into mid-2027. Our future capital requirements and the period for which we project our existing resources to support our operations may vary significantly from what we currently expect, and changing circumstances, some of which may be beyond our control, could cause us to consume capital significantly faster than we currently anticipate, and we may need to seek additional funds sooner than planned. Our monthly spending levels vary based on new and ongoing development and corporate activities. Because the length of time and activities associated with successful development of our product

candidates is highly uncertain, we are unable to estimate the actual funds we will require for development and any approved marketing and commercialization activities.

We expect to use our cash resources to fund the continued development and pre-commercialization costs of our clinical-stage product candidates; to fund continued development of the other assets in our pipeline, including designing and conducting preclinical studies and clinical trials, as well as funding discovery, manufacturing, research and development; to fund the acquisition of and drug development activities related to new programs; although we have no material agreements, commitments or understandings with respect to any in-license or acquisition, we have and plan to continue to evaluate such opportunities and engage in related discussions with other business entities from time to time; and the remainder for working capital and other general corporate purposes.

To execute our business plan, we will need, among other things, to:

- obtain the human and financial resources necessary to develop, test, obtain regulatory approval for, manufacture and market our product candidates;
- build and maintain a strong intellectual property portfolio and avoid infringing intellectual property of third parties;
- establish and maintain successful licenses, collaborations and alliances;
- satisfy the requirements of clinical trial protocols, including patient enrollment;
- establish and demonstrate the clinical efficacy and safety of our product candidates;
- obtain regulatory approvals;
- manage our spending as costs and expenses increase due to preclinical studies and clinical trials, regulatory approvals, commercialization, legal and regulatory compliance, and increased operations;
- obtain additional capital to support and expand our operations; and
- market our products to achieve acceptance and use by the medical community in general.

We do not expect to realize revenue from product sales, milestone payments or royalties in the foreseeable future, if at all. Our revenue sources are, and will remain, extremely limited unless and until our product candidates are clinically tested, approved for commercialization and successfully marketed and/or we sell, out-license or otherwise divest certain of our assets.

We will be required to seek additional funding in the future and intend to do so through either public or private equity offerings or debt financings, credit or loan facilities or a combination of one or more of these funding sources. Attempting to secure additional financing may divert management from day-to-day activities, which may adversely affect our ability to develop our product candidates. Our ability to raise additional funds will depend on financial, economic and other factors, many of which are beyond our control. Additional funds may not be available to us on acceptable terms or at all. If we raise additional funds by issuing equity securities, our shareholders will suffer dilution and the terms of any financing may adversely affect the rights of our shareholders. In addition, as a condition to providing additional funds to us, future investors may demand, and may be granted, rights superior to those of existing shareholders. Debt financing, if available, may involve restrictive covenants limiting our flexibility in conducting future business activities, and, in the event of insolvency, debt holders would be repaid before holders of equity securities received any distribution of corporate assets.

If we raise additional 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. Certain amounts of such additional funds raised may need to be used to pay third parties in respect of obligations we owe to them including to our licensors, under Incentivization Agreements (see Contractual Obligations and Other Commitments in our 2024 Annual Report on Form 10-K) and Oxford Finance. If we are unable to raise additional funds through equity or debt financings when needed, we may be required to delay, reduce or terminate our product development or future commercialization efforts or grant rights to third parties to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

***Our loan facility and payment obligations under the Loan and Security Agreement (“LSA”), with Oxford Finance contain operating and financial covenants that restrict our business and financing activities, are subject to acceleration in specified circumstances and may adversely affect our financial position or results of operations and our ability to raise additional***

*capital which in turn may increase our vulnerability to adverse regulatory developments or economic or business downturns or which may result in Oxford Finance taking possession of our assets and disposing of any collateral.*

Our loan facility with Oxford Finance contains restrictions that limit our flexibility in operating our business. Under the terms of the LSA, we must maintain, and cause our subsidiaries to maintain, certain covenants, including with respect to limitations on new indebtedness, restrictions on the payment of dividends and maintenance of revenue levels. Our credit facility is collateralized by all of our assets including, among other things, our intellectual property.

On December 30, 2024 (the “Effective Date”), Centessa Pharmaceuticals Holdings, Inc., Centessa Biosciences, Inc. and Centessa Pharmaceuticals LLC (the “Borrowers,” together with the Company, the “Borrower Parties”) entered into a loan and security agreement (the “Loan and Security Agreement” or “LSA”) with Oxford Finance LLC (“Oxford”), as collateral agent and a lender, and the other lenders from time to time party thereto (collectively, the “Lenders”), pursuant to which the Lenders agreed to lend the Borrowers an aggregate principal amount of up to \$200.0 million in a series of term loans (the “Term Loans”). Pursuant to the Loan and Security Agreement, the Borrowers received an initial Term Loan of \$110.0 million on the Effective Date (the “Initial Term Loan”). The Borrowers have access to up to an additional \$40.0 million of loan proceeds in an additional tranche which is available during the period commencing on the date of the occurrence of the Clinical Milestone (as defined in the Loan and Security Agreement) through the earlier of: (i) 90 days following the Clinical Milestone and (ii) June 30, 2028. An additional \$50.0 million may be made available to the Borrowers at the Lenders’ sole discretion. The term loans are set to mature on December 1, 2029 and, following an interest-only period, will begin to amortize in equal monthly installments beginning on February 1, 2029. However, if a specified milestone is achieved on or after the first anniversary of the Effective Date, then the term loans will begin to amortize in equal monthly installments beginning on February 1, 2030, and the maturity date may be extended to December 1, 2030. The term loans accrue interest at a floating rate equal to (i) secured overnight financing rate for a one-month tenor from the website of the CME Group Benchmark Administration Limited, subject to a floor of 3.28%, plus (ii) an applicable margin of 5.00%. The Loan and Security Agreement provides for a minimum interest rate of 8.28% and a maximum interest rate of 10.50%. Interest on the term loans is payable monthly in arrears. The term loans once repaid or prepaid may not be reborrowed. The term loans may be prepaid in full at the option of the Borrowers. The Borrowers are required to pay a prepayment fee of 3.00% for prepayments of term loans made in the first year after funding of such term loans, 2.00% for prepayments of term loans made in the second year after funding of such term loans and 1.00% for prepayments thereafter. The Borrowers are also obligated to pay other customary fees for a loan facility of this size and type. Substantially all of the proceeds from the Initial Term Loan were used to repay in full the approximately \$110 million aggregate principal amount outstanding, accrued interest and fees related to the Company’s note purchase agreement with Three Peaks Capital Solutions Aggregator Fund and Cocoon SA LLC, an affiliate of Oberland Capital Management LLC, as well as certain fees and expenses payable to Oxford. The Borrowers’ obligations under the Loan and Security Agreement are guaranteed by the Company and certain subsidiaries of the Company and will be guaranteed by the Company’s future subsidiaries, subject to certain customary limitations pursuant to the terms of the English-law Guarantee and Indemnity (the “Guarantee”). In addition, pursuant to the terms of the LSA, the Borrowers granted Oxford, as collateral agent, a first priority security interest in substantially all of the Borrowers’ assets, including intellectual property. Furthermore, pursuant to the terms of the English law debenture entered into on the Effective Date (the “Debenture”), the Company and certain of its subsidiaries granted Oxford a first priority security interest in substantially all of the Company’s and its subsidiaries’ assets, including intellectual property. The LSA contains customary affirmative and negative covenants, including covenants limiting the ability of the Borrower Parties and their subsidiaries to, among other things, dispose of assets, incur debt, grant liens, pay dividends and distributions on their capital stock, make investments and acquisitions, and enter into transactions with affiliates, in each case subject to customary exceptions for a loan facility of this size and type. In addition, the LSA contains a minimum cash covenant commencing on October 1, 2026 and all times thereafter of: (1) 35% of the outstanding principal balance of the Term Loan; and (2) up to 80% of the outstanding principal balance of the Term Loan based on the Company’s orexin agonist program Phase 2 and Phase 3 clinical data and continued Active Development (as defined in the LSA) of its lead orexin asset programs; provided that such minimum cash covenants shall not be tested during periods when the Company’s market capitalization meets \$1.0 billion. The events of default under the LSA include, among others, payment defaults, material misrepresentations, breaches of covenants, cross defaults with certain other material indebtedness, bankruptcy and insolvency events, the occurrence of a Material Adverse Change (as defined in the LSA) and judgment defaults. The occurrence of an event of default could result in the acceleration of the Borrowers’ obligations under the LSA, the termination of the Lenders’ commitments, a 3% increase in the applicable rate of interest and the exercise by the Lender of other rights and remedies provided for under the LSA and the Debenture.

If we breach certain of our debt covenants and are unable to cure such breach within the prescribed period, or are not granted waivers in relation to such breach, it may constitute an event of default under the loan facility, giving Oxford Finance the right to require us to repay the then outstanding debt immediately, and Oxford Finance could, among other things, foreclose on the collateral granted to them to collateralize such indebtedness, if we are unable to pay the outstanding debt immediately. A

breach of the covenants contained in the credit facility documents and the acceleration of its repayment obligations by Oxford Finance could have a material adverse effect on our business, financial condition, results of operations and prospects.

The loan facility could have important negative consequences to the holders of our securities. For example, a portion of our cash flow from operations will be needed to make payments to Oxford Finance and will not be available to fund future operations. Additionally, we may have increased vulnerability to adverse general economic and industry conditions. Payment requirements under the credit facility will increase our cash outflows if and when the conditions for payment are triggered. Our future operating performance is subject to market conditions and business factors that are beyond our control. If our cash inflows and capital resources are insufficient to allow us to make required payments, we may have to reduce or delay capital expenditures, sell assets or seek additional capital. If we raise funds by selling additional equity, such sale would result in dilution to our stockholders. There is no assurance that if we are required to secure funding, we can do so on terms acceptable to us, or at all.

***If we engage in acquisitions or strategic partnerships, this may increase our capital requirements, dilute our shareholders, cause us to incur debt or assume contingent liabilities, and subject us to other risks.***

We may engage in various acquisitions and strategic partnerships in the future, including licensing or acquiring new or complementary products, intellectual property rights, technologies, or businesses. Any acquisition or strategic partnership may entail numerous risks, including:

- increased operating expenses and cash requirements;
- the assumption of indebtedness or contingent liabilities;
- the issuance of our equity securities which would result in dilution to our shareholders;
- assimilation of operations, intellectual property, products and product candidates of an acquired company, including difficulties associated with integrating new personnel;
- the diversion of our management's attention from our existing product programs and initiatives in pursuing such an acquisition or strategic partnership;
- retention of key employees, the loss of key personnel, and uncertainties in our ability to maintain key business relationships;
- risks and uncertainties associated with the other party to such a transaction, including the prospects of that party and their existing products or product candidates and regulatory approvals; and
- our inability to generate revenue from acquired intellectual property, technology and/or products sufficient to meet our objectives or even to offset the associated transaction and maintenance costs; and
- our assumption of liabilities of the acquired subsidiary or acquired assets.

In addition, if we undertake such a transaction, we may issue dilutive securities, assume or incur debt obligations, incur large one-time expenses and acquire intangible assets that could result in significant future amortization expense.

***If we acquire additional assets and/or companies in the future, it could adversely affect our operating results and the value of our ADSs.***

As part of our asset-centric business model and strategy, we may acquire additional assets and/or companies. Investments in our existing and any future subsidiaries and developmental assets involve numerous risks, including, but not necessarily limited to:

- risk of conducting research and development activities in new therapeutic areas or treatment modalities in which we have little to no experience;
- diversion of financial and managerial resources from existing operations;
- successfully negotiating a proposed acquisition, in-license or investment in a timely manner and at a price or on terms and conditions favorable to us;
- successfully combining and integrating a potential acquisition into our existing business to fully realize the benefits of such acquisition;
- the impact of regulatory reviews on a proposed acquisition, in-license or investment; and

- the assumption of liabilities of acquired subsidiaries and outcome of any legal proceedings that may be instituted with respect to the proposed acquisition, in-license or investment.

If we fail to properly evaluate potential acquisitions, in-licenses, investments or other transactions associated with the creation of new research and development programs or the maintenance of existing ones, we might not achieve the anticipated benefits of any such transaction, we might incur costs in excess of what we anticipate, and management resources and attention might be diverted from other necessary or valuable activities.

### **Risks Related to Our Business and the Clinical Development, Regulatory Review and Approval**

*Our product candidates are in various stages of development, including several in discovery and preclinical stages, and may fail in development or suffer delays that materially adversely affect their commercial viability and we may fail to differentiate our molecules, including ORX750, ORX142, ORX489 and other orexin agonist molecules, LockBody molecules and/or the LockBody technology platform, from other available treatment options including other molecules in development.*

We have no products on the market and most of the product candidates in our pipeline are in the early stages of development. For example, we currently have two product candidates that are in clinical development— ORX750 and ORX142. The remainder of our programs are in preclinical development. Our ability to achieve and sustain profitability depends on obtaining regulatory approvals for, and successfully commercializing, our product candidates, either alone or with third parties. Before obtaining regulatory approval for the commercial distribution of our product candidates, we or a collaborator must conduct extensive preclinical tests and clinical trials to demonstrate the safety and efficacy of our drug product candidates and the safety, purity, and potency or efficacy, of our biologic product candidates. Preclinical testing and clinical trials are expensive, difficult to design and implement, can take many years to complete and are uncertain as to outcome. Our current or future product candidates may cause undesirable or clinically unmanageable side effects, toxicities or have other properties when used alone or in combination with other approved products or investigational new drugs that could halt their clinical development, delay or prevent their marketing approval, limit their commercial potential, result in a more restrictive label or result in significant negative consequences. Historical results of preclinical studies or clinical studies will not be fully predictive of future results in ongoing or future studies, which is particularly pertinent in the case of our orexin agonist program. For example, certain other orexin agonist programs have reported visual disturbances, drug induced liver injury (“DILI”) and other side-effects during clinical trials. There is no guarantee that ORX750 and/or ORX142 will not demonstrate the same or other serious and unexpected drug-related side effects which could materially and adversely impact our ability to develop and advance ORX750 and/or ORX142. The start or end of a clinical study is often delayed or halted due to changing regulatory requirements, manufacturing challenges, required clinical trial administrative actions, slower than anticipated patient enrollment, changing standards of care, availability or prevalence of use of a comparative drug or required prior therapy, clinical outcomes or financial constraints. For instance, delays or difficulties in patient enrollment or difficulties in retaining trial participants can result in increased costs, longer development times or termination of a clinical trial. Clinical trials of a new product candidate require the enrollment of a sufficient number of patients, including patients who are suffering from the disease the product candidate is intended to treat and who meet other eligibility criteria. Rates of patient enrollment are affected by many factors, including the size of the patient population, the eligibility criteria for the clinical trial, the age and condition of the patients, the stage and severity of disease, the nature of the protocol, the proximity of patients to clinical sites and the availability of effective treatments for the relevant disease.

A product candidate can unexpectedly fail at any stage of preclinical and clinical development. For example, in November 2024, we announced the voluntary discontinuation of the global clinical development program of SerpinPC, a novel inhibitor of APC that was being progressed for the treatment of hemophilia B. More recently, during the first quarter of 2025, we discontinued the clinical development of LB101, a conditionally tetravalent PD-L1xCD47 bispecific monoclonal antibody, and first generation LockBody candidate. The historical failure rate for product candidates is high due to scientific feasibility, safety, efficacy, changing standards of medical care and other variables. The results from preclinical testing or early clinical trials of a product candidate may not predict the results that will be obtained in later phase clinical trials of the product candidate. We, the FDA or other applicable regulatory authorities may suspend clinical trials of a product candidate at any time for various reasons, including a belief that subjects participating in such trials are being exposed to unacceptable health risks or adverse side effects. We may not be able to submit Investigational New Drug applications (“INDs”) or IND amendments to commence additional clinical trials on the timelines we expect, and even if we are able to, the FDA may not permit us to proceed. We may not have the financial resources to continue development of, or to enter into collaborations for, a product

candidate if we experience any problems or other unforeseen events that delay or prevent regulatory approval of, or our ability to commercialize, product candidates, including:

- negative or inconclusive results from our clinical trials or the clinical trials of others for product candidates similar to ours, leading to a decision or requirement to conduct additional preclinical testing or clinical trials or abandon a program;
- serious and unexpected drug-related side effects experienced by participants in our clinical trials or by individuals using drugs similar to our product candidates;
- delays in submitting INDs, Clinical Trial Applications (“CTAs”), or comparable foreign applications or delays or failure in obtaining the necessary approvals from regulators to commence a clinical trial, or a suspension or termination of a clinical trial once commenced;
- conditions imposed by the FDA or comparable foreign authorities regarding the scope or design of our clinical trials;
- delays in enrolling or our inability to enroll research subjects in clinical trials;
- high drop-out rates of research subjects;
- inadequate supply or quality of product candidate components or materials or other supplies necessary for the conduct of our clinical trials;
- greater than anticipated clinical trial costs;
- poor effectiveness of our product candidates during clinical trials;
- unfavorable FDA or other regulatory agency inspection and review of a clinical trial site;
- failure of our third-party contractors or investigators to comply with regulatory requirements or otherwise meet their contractual obligations in a timely manner, or at all;
- delays and changes in regulatory requirements, policy and guidelines, including the imposition of additional regulatory oversight around clinical testing generally or with respect to our technology in particular; or
- varying interpretations of data by the FDA and similar foreign regulatory agencies.

Some of the clinical trials performed to date were, and in the future we may conduct, open-label studies involving only a limited number of clinical sites and a limited number of patients. 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. Moreover, patients selected for early clinical studies often include the most severe sufferers and their symptoms may have been bound to improve notwithstanding the new 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.

***We may not be successful in our efforts to identify, discover, in-license or otherwise acquire additional product candidates and may fail to capitalize on programs or product candidates that may be a greater commercial opportunity or for which there is a greater likelihood of success.***

The success of our business depends upon our ability to identify, develop and commercialize product candidates. Research programs to identify new product candidates require substantial technical, financial and human resources. Although certain of our product candidates are currently in clinical or preclinical development, we may fail to identify other potential product candidates for clinical development for several reasons. For example, our research may be unsuccessful in identifying potential product candidates or our potential product candidates may be shown to have harmful side effects, may be commercially impracticable to manufacture or may have other characteristics that may make the products unmarketable or unlikely to receive marketing approval.

Additionally, because we have limited resources, we may forego or delay pursuit of opportunities with certain programs or product candidates or for indications that later prove to have greater commercial potential. Our spending on current and future research and development programs may not yield any commercially viable products. If we do not accurately evaluate the commercial potential for a particular product candidate, we may relinquish valuable rights to that product candidate through strategic collaboration, licensing or other arrangements in cases in which it would have been more advantageous for us to retain sole development and commercialization rights to such product candidate. Alternatively, we may allocate internal resources to a product candidate in a therapeutic area in which it would have been more advantageous to enter into a partnering arrangement.

If any of these events occur, we may be forced to abandon our development efforts with respect to a particular product candidate or fail to develop a potentially successful product candidate, which could have a material adverse effect on our business, financial condition, results of operations and prospects.

***Success in preclinical studies or early clinical trials may not be indicative of results obtained in later trials.***

Results from preclinical studies or previous clinical trials are not necessarily predictive of future clinical trial results, and interim results of a clinical trial are not necessarily indicative of final results. The results generated to date in preclinical studies or clinical trials for our product candidates do not ensure that later preclinical studies or clinical trials will demonstrate similar results. Our product candidates may fail to show the desired safety and efficacy in clinical development despite demonstrating positive results in preclinical studies or having successfully advanced through initial clinical trials. Product candidates in later stages of clinical trials may fail to show the desired safety and efficacy traits despite having progressed through preclinical and earlier stage clinical trials. In later-stage clinical trials, we will likely be subject to more rigorous statistical analyses than in completed earlier stage clinical trials. In some instances, there can be significant variability in safety or efficacy results between different clinical trials of the same product candidate due to numerous factors, including changes in clinical trial procedures set forth in protocols, differences in the size and type of the patient populations, adherence to the dosing regimen and other clinical trial protocols, and the rate of dropout among clinical trial participants. We cannot guarantee that any of our clinical trials will be conducted as planned or completed on schedule, or at all. Clinical trials can fail at any stage of testing and failure may result from a multitude of factors, including, among other things, flaws in study design, dose selection issues, placebo effects, patient enrollment criteria and failure to demonstrate favorable safety or efficacy traits.

There is a high failure rate for small molecule drugs and biologic products proceeding through clinical development. 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, and many companies that have believed their product candidates performed satisfactorily in preclinical studies and clinical trials have nonetheless failed to obtain marketing approval of their products. In addition, we may experience regulatory delays or rejections as a result of many factors, including due to changes in regulatory policy during the period of our product candidate development. Furthermore, the failure of any of our product candidates to demonstrate safety and efficacy in any clinical trial could negatively impact the perception of our other product candidates and/or cause the FDA or other regulatory authorities to require additional testing before approving any of our product candidates. Any such delays could materially and adversely affect our business, financial condition, results of operations and prospects.

***We may encounter substantial delays or challenges in the initiation, conduct or completion of our clinical trials, and the results of clinical development are uncertain.***

Before obtaining marketing approval from regulatory authorities for the sale of our product candidates, we must conduct extensive clinical trials to demonstrate the safety and efficacy of the product candidate for its intended indications. Clinical trials are expensive, time-consuming and uncertain as to outcome. We cannot guarantee that any clinical trials will be conducted as planned or completed on schedule, if at all. A failure of one or more clinical trials can occur at any stage of testing. Events that may prevent successful or timely completion of clinical development include, without limitation:

- delay in completing preclinical studies;
- delays in reaching a consensus with regulatory authorities on trial design;
- delays in obtaining authorizations of INDs to commence a clinical trial;
- delays in reaching agreement or failing to agree on acceptable terms with prospective CROs, and clinical trial sites, the terms of which can be subject to extensive negotiation and may vary significantly among different CROs and clinical trial sites;

- delays in obtaining or failure to obtain Institutional Review Board (“IRB”), or independent ethics committee approval at each clinical trial site;
- delays in opening or failure to open a sufficient number of clinical trial sites and recruiting an adequate number of suitable patients to participate in our clinical trials;
- imposition of a clinical hold by regulatory authorities as a result of a serious adverse event, concerns with a class of product candidates or after an inspection of our clinical trial operations or trial sites;
- delays in having patients complete participation in a trial or return for post-treatment follow-up;
- occurrence of clinical trial sites deviating from clinical trial protocol or dropping out of a clinical trial;
- obtaining sufficient product supply of product candidate for use in preclinical studies or clinical trials from third-party suppliers;
- occurrence of serious adverse events associated with the product candidate that are viewed to outweigh its potential benefits;
- changes in regulatory requirements and guidance that require amending or submitting new clinical protocols;
- failure to recruit and maintain a sufficient number of, or any, subjects in our existing and anticipated studies or clinical trials including trials of ORX750, ORX142, other orexin agonist molecules and any other LockBody candidates, and failure to meet expectations on executing our research and clinical development plans and the timing thereof; or
- geopolitical or macro factors such as the ongoing Russia-Ukraine war, the Middle East conflict(s), tensions in U.S.-China relations and the impact of changes in trade policy, including the imposition of tariffs, on our business and results of operations.

Any inability to successfully complete preclinical and clinical development could result in additional costs to us or impair our ability to generate revenue from future drug sales and regulatory and commercialization milestones. In addition, if we make manufacturing or formulation changes to our product candidates, we may need to conduct additional testing to bridge our modified product candidate to earlier versions. Clinical trial delays could also shorten any periods during which we may have the exclusive right to commercialize our product candidates, if approved, or allow our competitors to bring comparable drugs to market before we do, which could impair our ability to successfully commercialize our product candidates and may harm our business, financial condition, results of operations and prospects.

Additionally, if the results of our clinical trials are inconclusive or if there are safety concerns or serious adverse events associated with our product candidates, we may:

- be delayed in obtaining marketing approval, if at all;
- obtain approval for indications or patient populations that are not as broad as intended or desired;
- obtain approval with labeling that includes significant use or distribution restrictions or safety warnings;
- be subject to additional post-marketing testing requirements;
- be required to perform additional clinical trials to support approval or be subject to additional post-marketing testing requirements;
- have regulatory authorities withdraw, or suspend, their approval of the drug or impose restrictions on its distribution in the form of a modified risk evaluation and mitigation strategy (“REMS”) plan;
- be subject to the addition of labeling statements, such as warnings or contraindications;
- be sued; or
- experience damage to our reputation.

Our drug development costs will also increase if we experience delays in testing or obtaining marketing approvals. We do not know whether any of our preclinical studies or clinical trials will begin as planned, need to be restructured or be completed on schedule, if at all.

Further, we, the FDA or other regulatory authorities, or an IRB or ethics committee of the institutions in which our clinical trials are being conducted, or the Data Safety Monitoring Board for such trials, if any, may suspend or terminate our clinical trials. Such authorities may suspend or terminate a clinical trial at any time due to a number of factors, including if it

appears that we or our collaborators are failing to conduct a trial in accordance with regulatory requirements, including the FDA's current Good Clinical Practice ("GCP"), regulations, unforeseen safety issues or unacceptable health risks, failure to demonstrate a benefit from the product candidates, or if the FDA finds deficiencies in our INDs or the conduct of these trials. Therefore, we cannot predict with any certainty the schedule for commencement and completion of future clinical trials. 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 negatively impacted, and our ability to generate revenues from our product candidates may be delayed or eliminated entirely.

***Even if we complete the necessary preclinical studies and clinical trials, the marketing approval process is expensive, time-consuming and uncertain and may prevent us from obtaining approvals for the commercialization of our product candidates.***

Any product candidate we develop and the activities associated with its development and commercialization, including its design, testing, manufacture, safety, efficacy, recordkeeping, labeling, storage, approval, advertising, promotion, sale, and distribution, are subject to comprehensive regulation by the FDA and other regulatory authorities in the United States and by comparable authorities in other countries. Failure to obtain marketing approval for a product candidate will prevent us from commercializing the product candidate in a given jurisdiction. We have not received approval to market any product candidates from regulatory authorities in any jurisdiction and it is possible that none of the product candidates we are developing or may seek to develop in the future will ever obtain regulatory approval. We have no experience in submitting and supporting the applications necessary to gain marketing approvals and expect to rely on third-party CROs or regulatory consultants to assist us in this process. Securing regulatory approval requires the submission of extensive preclinical and clinical data and supporting information to the various regulatory authorities for each therapeutic indication to establish the product candidate's safety and efficacy. Securing regulatory approval also requires the submission of information about the product manufacturing process to, and inspection of manufacturing facilities by, the relevant regulatory authority. Any product candidates we develop may not be effective, may be only moderately effective, or may prove to have undesirable or unintended side effects, toxicities or other characteristics that may preclude its obtaining marketing approval or prevent or limit commercial use.

The process of obtaining marketing approvals is expensive, may take many years if additional clinical trials are required, if approval is obtained at all, and can vary substantially based upon a variety of factors, including the type, complexity, and novelty of the product candidates involved. Changes in marketing approval policies during the development period, changes in or the enactment of additional statutes or regulations, or changes in regulatory review for each submitted product application, may cause delays in the approval or rejection of an application. The FDA and comparable authorities in other countries have substantial discretion in the approval process and may refuse to accept any application or may decide that our data are insufficient for approval and require additional preclinical, clinical or other studies. This is particularly true for clinical trials in rare diseases, where the very small patient population makes it difficult or impossible to conduct traditional, adequate and well-controlled studies, and therefore the FDA or comparable foreign regulatory authorities are often required to exercise flexibility in approving therapies for such diseases. To the extent that the results of the trials are not satisfactory to the FDA or comparable 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. In addition, varying interpretations of the data obtained from preclinical and clinical testing could delay, limit, or prevent marketing approval of a product candidate. Any marketing approval that we may ultimately obtain could be limited or subject to restrictions or post-approval commitments that render the approved product not commercially viable.

If we experience delays in obtaining approval or if we fail to obtain approval of any product candidates we may develop, the commercial prospects for those product candidates may be harmed, and our ability to generate revenues may be materially impaired.

***We may find it difficult to enroll patients in our clinical trials, which could delay or prevent us from proceeding with clinical trials of our product candidates.***

Identifying and qualifying patients to participate in clinical trials of our product candidates is critical to our success. The timing of our clinical trials depends on our ability to recruit patients to participate in such trials as well as the completion of any required follow-up periods. Some of our product candidates are designed to target orphan indications. For example, SerpinPC was being developed for the treatment of hemophilia. Trials in orphan indications often take longer to enroll than trials for other indications due to the smaller patient population from which subjects can be recruited. We may experience delays in any of our future clinical trials. If patients are unwilling to participate in our studies because of negative publicity from adverse events related to certain modalities utilized in one or more of our product candidates, competitive clinical trials for similar patient populations or for other reasons, the timeline for recruiting patients, conducting studies and obtaining regulatory

approval of our product candidates may be delayed. These delays could result in increased costs, delays in advancing our product candidates, delays in testing the effectiveness of our product candidates or termination of the clinical trials altogether.

We may not be able to identify, recruit and enroll a sufficient number of patients, or those with required or desired characteristics, to complete our clinical trials in a timely manner. Patient enrollment and trial completion is affected by factors including:

- size of the patient population and process for identifying subjects;
- design of the trial protocol;
- eligibility and exclusion criteria;
- perceived risks and benefits of the product candidate under study;
- perceived risks and benefits of approaches utilized by one or more of our product candidates for the treatment of diseases;
- availability of competing therapies and clinical trials;
- severity of the disease under investigation;
- proximity and availability of clinical trial sites for prospective subjects;
- ability to recruit clinical trial investigators with the appropriate competencies and experience;
- ability to obtain and maintain subject consent;
- risk that enrolled subjects will drop out before completion of the trial;
- patient referral practices of physicians;
- ability to monitor subjects adequately during and after treatment; and
- factors we may not be able to control, such as current or potential pandemics that may limit patients, principal investigators or staff or clinical site availability and geopolitical conflicts such as the Russia-Ukraine war, the Middle East conflict(s), tensions in U.S.-China relations and the impact of changes in trade policy, including the imposition of tariffs, on our business and results of operations.

We plan to seek initial marketing approval in the United States and certain other major markets such as major countries in the EU, and the United Kingdom. We may not be able to initiate or continue clinical trials if we cannot enroll a sufficient number of eligible patients to participate in the clinical trials required by FDA, EMA, MHRA or other regulatory authorities. Our ability to successfully initiate, enroll and complete a clinical trial in any foreign country is subject to numerous risks unique to conducting business in foreign countries, including:

- difficulty in establishing or managing relationships with CROs, and physicians;
- difficulty in obtaining local regulatory approval to conduct clinical trials;
- different standards for the conduct of clinical trials;
- our inability to locate qualified local consultants, physicians and partners; and
- the potential burden of complying with a variety of foreign laws, medical standards and regulatory requirements, including the regulation of pharmaceutical and biotechnology products and treatment.

If we have difficulty enrolling a sufficient number of patients to conduct our clinical trials as planned, we may need to delay, limit or terminate ongoing or planned clinical trials, any of which would have an adverse effect on our business, financial condition, results of operations and prospects.

***We are dependent on third parties having accurately generated, collected, interpreted and reported data from certain preclinical studies and clinical trials that were previously conducted for our product candidates.***

We have licensed patent and other intellectual property rights from third parties and we may continue to seek and enter into similar licenses for future programs. In certain cases, we intend to rely on results of studies previously conducted by third parties to support our own development of these candidates. In such cases, we may have no involvement with or control over the preclinical and clinical development of any of such product candidates prior to obtaining the in-license. Therefore, we would be dependent on these third parties having conducted their research and development in accordance with the applicable

protocols, legal and regulatory requirements, and scientific standards; having accurately reported the results of all preclinical studies and clinical trials conducted with respect to such product candidates and having correctly collected and interpreted the data from these studies and trials. If these activities were not compliant, accurate or correct, the clinical development, regulatory approval or commercialization of our product candidates will be adversely affected.

***We may be unable to obtain U.S. or foreign regulatory approval and, as a result, unable to develop and/or commercialize our product candidates.***

Our product candidates are subject to extensive governmental regulations relating to, among other things, research, testing, development, manufacturing, safety, efficacy, approval, recordkeeping, reporting, labeling, storage, packaging, advertising and promotion, pricing, marketing and distribution of drugs. Rigorous preclinical testing and clinical trials and an extensive regulatory approval process are required to be successfully completed in the U.S. and in many foreign jurisdictions before a new drug can be marketed. Satisfaction of these and other regulatory requirements is costly, time consuming, uncertain and subject to unanticipated delays. Regulatory agencies may also impose restrictions or conditions on the development of our product candidates which may delay or adversely impact their planned development. For example, we might need to exceed the current maximum exposure limit set by FDA for our ongoing clinical trials of ORX750, and this might result in a material delay to the current planned development of ORX750, or add significant additional cost, or require that we make significant adjustments including protocol changes, or we might not be successful in having the maximum exposure limit amended or removed by the FDA at all, in each case, which may materially and adversely impact our ability to develop or commercialize ORX750 in a timely manner or at all and consequently, our business could be substantially harmed. It is possible that none of the product candidates we may develop will obtain the regulatory approvals necessary for us or our collaborators to begin selling them. Regulatory authorities may also fail to approve the facilities or processes used to manufacture a product candidate, our dosing or delivery methods.

We have very limited experience in conducting and managing the clinical trials necessary to obtain regulatory approvals, including approval by the FDA. The time required to obtain FDA and other approvals is unpredictable but typically takes many years following the commencement of clinical trials, depending upon the type, complexity and novelty of the product candidate. The standards that the FDA and its foreign counterparts use when regulating us are not always applied predictably or uniformly and can change. Any analysis we perform on data from preclinical and clinical activities is subject to confirmation and interpretation by regulatory authorities, which could delay, limit or prevent regulatory approval. We may also encounter unexpected delays or increased costs due to new government regulations, for example, from future legislation or administrative action, or from changes in FDA policy during the period of product development, clinical trials and FDA regulatory review. It is impossible to predict whether legislative changes will be enacted, or whether FDA or foreign regulations, guidance or interpretations will be changed, or what the impact of such changes, if any, may be. In addition, the U.S. Supreme Court's July 2024 decision to overturn established case law giving deference to regulatory agencies' interpretations of ambiguous statutory language has introduced uncertainty regarding the extent to which the FDA's regulations, policies and decisions may become subject to increasing legal challenges, delays, and/or changes.

In certain cases in the future, we may develop therapies that may represent a new class of drug for which the FDA and its foreign counterparts have not yet established any definitive policies, practices or guidelines in relation to these drugs. For example, we may in the future develop product candidates that we believe are regulated as new drugs under the Federal Food, Drug, and Cosmetic Act, but the FDA could decide to regulate them or other products we may develop as biologics under the Public Health Service Act. The lack of policies, practices or guidelines may hinder or slow review by the FDA of any regulatory filings that we may submit. Moreover, the FDA may respond to these submissions by defining requirements we may not have anticipated. Such responses could lead to significant delays in the clinical development of our product candidates. In addition, because there may be approved treatments for some of the diseases for which we may seek approval, in order to receive regulatory approval, we may need to demonstrate through clinical trials that the product candidates we develop to treat these diseases, if any, are not only safe and effective, but safer or more effective than existing products.

Any delay or failure in obtaining required approvals could have a material adverse effect on our ability to generate revenues from the particular product candidate for which we are seeking approval. Furthermore, any regulatory approval to market a product may be subject to limitations on the approved uses for which we may market the product or the labeling or other restrictions. In addition, the FDA has the authority to require a REMS plan as part of a new drug application ("NDA"), or biologics license application ("BLA"), or after approval, which may impose further requirements or restrictions on the distribution or use of an approved drug or biologic, such as limiting prescribing to certain physicians or medical centers that have undergone specialized training, limiting treatment to patients who meet certain safe-use criteria and requiring treated patients to enroll in a registry. These limitations and restrictions may limit the size of the market for the product and affect reimbursement by third-party payors.

We are also subject to numerous foreign regulatory requirements governing, among other things, the conduct of clinical trials, manufacturing and marketing authorization, pricing and third-party reimbursement. The foreign regulatory approval process varies among countries and may include all of the risks associated with FDA approval described above as well as risks attributable to the satisfaction of local regulations in foreign jurisdictions. Moreover, the time required to obtain approval may differ from that required to obtain FDA approval. Approval by the FDA does not ensure approval by regulatory authorities outside the U.S. and vice versa.

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

From time to time, we may publish interim, “top-line,” or preliminary data from our clinical studies. Interim data 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” data 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. Material adverse changes between preliminary, “top-line,” or interim data and final data could significantly harm our business prospects.

***We may be unable to obtain orphan drug designation or exclusivity. If our competitors are able to obtain orphan drug exclusivity for products that constitute the same drug and treat the same indications as our product candidates, we may not be able to have our product candidates approved by the applicable regulatory authority for a significant period of time.***

We may seek orphan drug designation for our product candidates where applicable, but there is no assurance the FDA or other regulatory authorities would grant orphan drug designation or, if granted, there is no assurance we can maintain orphan drug designation or obtain any benefits associated with orphan drug designation, including market exclusivity. Regulatory authorities in some jurisdictions, including the United States and the European Union, may designate drugs and biologics intended to treat relatively small patient populations as orphan drugs. Under the Orphan Drug Act of 1983, FDA may designate a product candidate as an orphan drug if it is intended to treat a rare disease or condition, which is defined as a disease or condition having a patient population of fewer than 200,000 individuals in the United States, or a patient population greater than 200,000 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 EU, the European Commission, after recommendation from the EMA’s Committee for Orphan Medicinal Products, grants orphan designation in respect of products that are intended for the diagnosis, prevention or treatment of a life-threatening or chronically debilitating condition which either affects not more than 5 in 10,000 persons in the EU when the application for orphan designation is made, or products intended for the diagnosis, prevention or treatment of a life-threatening, seriously debilitating or serious and chronic condition when, without incentives, it is unlikely that sales of the product in the EU would be sufficient to justify the necessary investment in developing the product. In each case, there must be no satisfactory method of diagnosis, prevention or treatment which is authorized for marketing in the EU, or, if such a method exists, the product would be of significant benefit to those affected by the condition.

Certain of our current product candidates, and our future potential product candidates may target patient populations that are smaller than the numbers described above. If we request orphan drug designation for our product candidates, there can be no assurances that FDA or the European Commission will grant any of our product candidates such designation. Additionally, the designation of any of our product candidates as an orphan product does not guarantee that any regulatory agency will accelerate regulatory review of, or ultimately approve, that product candidate, nor does it limit the ability of any regulatory agency to grant orphan drug designation to product candidates of other companies that treat the same indications as our product candidates prior to our product candidates receiving exclusive marketing approval.

Generally, if a product candidate with an orphan drug designation receives the first marketing approval for the indication for which it has such designation, the product is entitled to a period of marketing exclusivity, which precludes the FDA or the European Commission from approving another marketing application for a product that constitutes the same drug (or a “similar medicinal product” in the EU) treating the same indication for that marketing exclusivity period, except in limited circumstances. If another sponsor receives such approval before we do (regardless of our orphan drug designation), we will be precluded from receiving marketing approval for our product for the applicable exclusivity period. The applicable period is seven years in the United States and 10 years in the European Union. The exclusivity period in the United States can be extended by six months if the sponsor submits pediatric data that fairly respond to a written request from the FDA for such data, and the exclusivity period in the EU can be extended by two years when the results of pediatric studies are completed in accordance with a fully compliant pediatric investigation plan and reflected in the summary of product characteristics (SmPC). The exclusivity period in the European Union can be reduced to six years if a product no longer meets the criteria for orphan

drug designation or if the product is sufficiently profitable so that market exclusivity is no longer justified. Proposed amendments to EU legislation regarding orphan medicines are under consideration that, if implemented following the conclusion of the legislative process, have the potential to shorten the ten-year period of marketing exclusivity. Orphan drug exclusivity may be revoked if any regulatory agency determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantity of the product to meet the needs of patients with the rare disease or condition.

Even if we obtain orphan drug exclusivity for a product candidate, that exclusivity may not effectively protect the product candidate from competition because different drugs can be approved for the same condition. In the United States, even after an orphan drug is approved, the FDA may subsequently approve another drug for the same condition if the FDA concludes that the latter drug is not the same drug or is clinically superior in that it is shown to be safer, more effective or makes a major contribution to patient care. In the EU, a marketing authorization may be granted to a similar medicinal product for the same orphan indication if:

- the second applicant can establish in its application that its medicinal product, although similar to the orphan medicinal product already authorized, is safer, more effective or otherwise clinically superior;
- the holder of the marketing authorization for the original orphan medicinal product consents to a second orphan medicinal product application; or
- the holder of the marketing authorization for the original orphan medicinal product cannot supply sufficient quantities of orphan medicinal product.

***We face significant competition in an environment of rapid technological change and the possibility that our competitors may achieve regulatory approval before us or develop therapies that are more advanced or effective than ours, which may adversely affect our ability to successfully market or commercialize our product candidates and our financial condition.***

The biotechnology and pharmaceutical industries are characterized by rapidly changing technologies, significant competition and a strong emphasis on intellectual property. We face substantial competition from many different sources, including large and specialty pharmaceutical and biotechnology companies, academic research institutions, government agencies and public and private research institutions. In addition, we face competition from other companies that have adopted business models that are similar to ours in which they establish strategic alliances, create joint ventures or collaborations, or enter into licensing arrangements with third parties for programs, product candidates, technologies or intellectual property. We may not be able to compete effectively with such companies. See “—We may not be successful in our efforts to use our differentiated asset-centric approach to drug discovery and development to build a pipeline of product candidates with commercial value.”

Many of our potential competitors, alone or with their strategic partners, may have substantially greater financial, technical and other resources, such as larger research and development, clinical, marketing and manufacturing organizations. Mergers and acquisitions in the biotechnology and pharmaceutical industries may result in even more resources being concentrated among a smaller number of competitors. Our commercial opportunity could be reduced or eliminated if competitors develop and commercialize products that are safer, more effective, have fewer or less severe side effects, are more convenient or are less expensive than any products that we may develop. Competitors also may obtain FDA or other regulatory approval for their products more rapidly or earlier than we may obtain approval for ours, which could result in our product being prevented from being marketed for significant periods (for example, where our competitor has secured regulatory exclusivity) or our competitors establishing a strong market position before we are able to enter the market. Additionally, technologies developed by our competitors may render our potential product candidates uneconomical or obsolete, and we may not be successful in marketing our product candidates against competitors.

In addition, as a result of the expiration or successful challenge of our patent rights, we could face more litigation with respect to the validity and/or scope of patents relating to our competitors’ products. The availability of our competitors’ products could limit the demand, and the price we are able to charge, for any products that we may develop and commercialize.

***Our product candidates and the process for administering our product candidates may cause undesirable side effects or have other properties that could delay or prevent their regulatory approval, limit the commercial potential or result in significant negative consequences following any potential marketing approval.***

Our product candidates may cause undesirable side effects. Additionally, the administration process or related procedures also can cause adverse side effects. Adverse events that occur in our trials may cause us, or cause regulatory authorities or others to order us to halt, delay or amend preclinical development or clinical development of our product

candidates and could result in more restrictive labeling or the denial of regulatory approval of our product candidates for any or all targeted indications. Even if serious adverse events are unrelated to study treatment, such occurrences could affect patient enrollment or the ability of enrolled patients to complete the trial. In addition, if any of our product candidates are tested or used in combination with other drugs, these combinations may have additional side effects, which could be more severe than those caused by either therapy alone.

In June 2022, we announced our strategic decision to discontinue development of lixivaptan for ADPKD. The decision was based on a thorough reassessment of the commercial potential of lixivaptan as a potential best-in-class therapy for patients with ADPKD, and the incremental development challenges and associated costs, following a recent observation of ALT and AST elevations in one subject in the ALERT Study.

We also decided to discontinue development of ZF874 for the treatment of AATD following a report of an adverse event involving elevated liver enzymes in a PiMZ subject dosed with 5 mg/kg BID of ZF874 in our Phase 1 study. In November 2021, we reported that elevated liver enzymes were observed in a subject dosed with 15 mg/kg BID of ZF874 in the first cohort of patients within Part B of the Phase 1 study. Based on the results of the Phase 1 study observed to date, we concluded that ZF874 was unlikely to achieve the desired target product profile.

Additionally, certain of our product candidates could cause undesirable side effects in clinical trials related to on-target toxicity. If on-target toxicity is observed, or if our product candidates have characteristics that are unexpected, we may need to abandon their development or limit development to more narrow uses or subpopulations in which the undesirable side effects or other characteristics are less prevalent, less severe or more acceptable from a risk-benefit perspective. Many compounds that initially showed promise in early stage testing for treating cancer have later been found to cause side effects that prevented further development of the compound.

Furthermore, clinical trials by their nature utilize a sample of the potential patient population. With a limited number of patients and limited duration of exposure, rare and severe side effects of our product candidates or those of our competitors may only be uncovered when a significantly larger number of patients have been exposed to the drug. While we believe that our product candidates have demonstrated manageable tolerability profiles thus far in the target indications, there can be no assurance that it or any of our other product candidates will not cause more severe side effects in a greater proportion of patients. In addition, some of our product candidates are intended to address limitations in current treatment approaches by offering potentially greater tolerability. If we do not observe a favorable tolerability profile in testing of such product candidates that differentiate them from competitors in the market, we may decide to suspend or terminate development of such candidates.

In addition, certain of our product candidates target diseases that are life-threatening or are associated with significant co-morbidities. For example, some of our product candidates are designed to address cancers, an indication in which patients may undergo treatment with other therapies such as chemotherapy, radiation, and/or other high dose or myeloablative treatments in the course of treatment of their disease, and may therefore experience side effects or AEs, including death, that are unrelated to our product candidates. While these side effects or AEs may be unrelated to our product candidates, they may still affect the success of our clinical trials. The inclusion of critically ill patients in our clinical trials may also result in deaths or other adverse medical events due to underlying disease or to other therapies or medications that such patients may receive.

Additionally, if any of our product candidates receives marketing approval, FDA could require us to adopt a REMS to ensure that the benefits outweigh its risks, which may include, among other things, a medication guide outlining the risks of the product for distribution to patients and a communication plan to health care practitioners. Furthermore, if we or others later identify undesirable side effects caused by our product candidate, several potentially significant negative consequences could result, including:

- regulatory authorities may suspend or withdraw approvals of such product candidate;
- regulatory authorities may require additional warnings on the label;
- we may be required to change the way a product candidate is administered or conduct additional clinical trials;
- we could be sued and held liable for harm caused to patients; and
- our reputation may suffer.

Any of these events could prevent us from achieving or maintaining market acceptance of our product candidates and could significantly harm our business, prospects, financial condition and results of operations.

***We may not be able to submit INDs or IND amendments to commence additional clinical trials on the timelines we expect, and even if we are able to, the FDA may not permit us to proceed or may subject its approval to certain conditions.***

Currently, certain of the product candidates in our pipeline have not yet commenced clinical trials, and are in preclinical development. We may not be able to submit INDs for our product candidates on the timelines we expect. For example, we may experience manufacturing delays or other delays with IND-enabling studies. Moreover, we cannot be sure that submission of an IND will result in the FDA allowing further clinical trials to begin, or that, once begun, issues will not arise that suspend or terminate clinical trials. Additionally, even if such regulatory authorities agree with the design and implementation of the clinical trials set forth in an IND, we cannot guarantee that such regulatory authorities will not change their requirements in the future. These considerations also apply to new clinical trials we may submit as amendments to existing INDs or to a new IND. Any failure to file INDs on the timelines we expect or to obtain regulatory approvals for our trials may prevent us from completing our clinical trials or commercializing our products on a timely basis, if at all. The FDA may also impose restrictions or conditions on the development of our product candidates which may delay or adversely impact their planned development. For example, we might need to exceed the current maximum exposure limit set by the FDA for our ongoing clinical trials of ORX750, and this might result in a material delay to the current planned development of ORX750, or add significant additional cost, or require that we make significant adjustments including protocol changes, or we might not be successful in having the maximum exposure limit amended or removed by the FDA at all, in each case, which may materially and adversely impact our ability to conduct our ORX750 clinical trials and/or develop or commercialize ORX750 in a timely manner or at all and consequently, our business could be substantially harmed.

***We are currently conducting and plan to conduct future clinical trials for certain product candidates outside the United States, and the FDA and comparable foreign regulatory authorities may not accept data from such trials.***

We are currently conducting and plan to conduct future clinical trials for certain product candidates outside the United States, including in Europe. The acceptance of study data from clinical trials conducted outside the United States or another jurisdiction by the FDA or comparable foreign regulatory authority may be subject to certain conditions or may not be accepted at all. The FDA will generally not consider the data from a foreign clinical trial not conducted under an IND unless (i) the trial was well-designed and well-conducted in accordance with GCP requirements, including requirements for the design, conduct, performance, monitoring, auditing, recording, analysis, and reporting of clinical trials in a way that provides assurance that the data and reported results are credible and accurate and that the rights, safety, and well-being of trial subjects are protected; and (ii) the FDA is able to validate the data from the trial through an on-site inspection, if necessary. In cases where data from foreign clinical trials are intended to serve as the basis for marketing approval in the United States, the FDA will generally not approve the application on the basis of foreign data alone unless (i) the data are applicable to the U.S. population and U.S. medical practice; (ii) the trials were performed by clinical investigators of recognized competence; and (iii) the data may be considered valid without the need for an on-site inspection by the FDA or, if the FDA considers such as inspection to be necessary, the FDA is able to validate the data through an on-site inspection or other appropriate means. Additionally, the FDA's clinical trial requirements, including sufficient size of patient populations and statistical powering, must be met. Many foreign regulatory authorities have similar approval requirements. In addition, such foreign trials would be subject to the applicable local laws of the foreign jurisdictions where the trials are conducted. There can be no assurance that the FDA or any comparable foreign regulatory authority will accept data from trials conducted outside of the United States or the applicable jurisdiction. If the FDA or any comparable foreign regulatory authority does not accept such data, it would result in the need for additional trials, which could be costly and time-consuming, and which may result in product candidates that we may develop not receiving approval for commercialization in the applicable jurisdiction.

***Even if we obtain and maintain approval for our product candidates from the FDA, we may never obtain approval for our product candidates outside of the United States, which would limit our market opportunities and adversely affect our business.***

Approval of a product candidate in the United States by the FDA does not ensure approval of such product candidate by regulatory authorities in other countries or jurisdictions, and approval by one foreign regulatory authority does not ensure approval by regulatory authorities in other foreign countries or by the FDA. Sales of our product candidates outside of the United States will be subject to foreign regulatory requirements governing clinical trials and marketing approval. Even if the FDA grants marketing approval for a product candidate, comparable regulatory authorities of foreign countries also must approve the manufacturing and marketing of the product candidates in those countries. Approval procedures vary among jurisdictions and can involve requirements and administrative review periods different from, and more onerous than, those in the United States, including additional preclinical studies or clinical trials. In many countries outside the United States, a product candidate must be approved for reimbursement before it can be approved for sale in that country. In some cases, the price that we intend to charge for our products, if approved, is also subject to approval. We intend to submit a marketing authorization application to the EMA for approval of our product candidates in the European Union, but obtaining such

approval from the European Commission following the opinion of the EMA is a lengthy and expensive process. We may also submit marketing applications to regulators in other jurisdictions, such as to the MHRA in the United Kingdom. Even if a product candidate is approved, the FDA, the European Commission, the MHRA and other foreign regulatory authorities, as the case may be, may limit the indications for which the product may be marketed, require extensive warnings on the product labeling or require expensive and time-consuming additional clinical trials or reporting as conditions of approval. Regulatory authorities in countries outside of the United States and the European Union also have requirements for approval of product candidates with which we must comply prior to marketing in those countries. Obtaining foreign regulatory approvals and compliance with foreign regulatory requirements could result in significant delays, difficulties and costs for us and could delay or prevent the introduction of our product candidates in certain countries.

Further, clinical trials conducted in one country may not be accepted by regulatory authorities in other countries. Also, regulatory approval for any of our product candidates may be withdrawn. If we fail to comply with the regulatory requirements, our target market will be reduced and our ability to realize the full market potential of our product candidates will be harmed and our business, financial condition, results of operations and prospects will be adversely affected.

***We may seek Fast Track designation for any of our other current or future product candidates. This designation may not lead to a faster development or regulatory review or approval process, and does not increase the likelihood that our product candidates will receive marketing approval.***

If a drug or biologic is intended for the treatment of a serious or life-threatening condition and the product demonstrates the potential to address unmet medical needs for this condition, the product sponsor may apply for FDA Fast Track designation for a particular indication. We may seek Fast Track designation for certain of our other current and future product candidates, but there is no assurance that the FDA will grant this status to any of our proposed product candidates. The FDA has broad discretion whether or not to grant Fast Track designation, so even if we believe a particular product candidate is eligible for this designation, there can be no assurance that the FDA would decide to grant it. Even if we receive Fast Track designation for any of our product candidates, we may not experience a faster development process, regulatory review or approval compared to conventional FDA procedures, and receiving a Fast Track designation does not provide assurance of ultimate FDA approval. In addition, the FDA may withdraw Fast Track designation if it believes that the designation is no longer supported by data from our clinical development program. In addition, the FDA may withdraw any Fast Track designation at any time.

***We may seek accelerated approval for any of our current or future product candidates. Accelerated approval, even if granted, may not lead to a faster commercial launch of the product and does not increase the likelihood that our product candidates will receive marketing approval.***

We may seek approval of our product candidates, where applicable, under the FDA's accelerated approval program. A product may be eligible for accelerated approval if it treats a serious or life-threatening condition, generally provides a meaningful advantage 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. As a condition of accelerated approval, the FDA likely would require that we perform adequate and well-controlled post-marketing clinical trials to confirm the product's clinical benefit. These confirmatory trials must be completed with due diligence. Under the Food and Drug Omnibus Reform Act ("FDORA"), the FDA is permitted to require, as appropriate, that a post-approval confirmatory study or studies be underway prior to approval or within a specified time period after the date of approval for a product granted accelerated approval. FDORA also requires sponsors to send updates to the FDA every 180 days on the status of such studies, including progress toward enrollment targets, and the FDA must promptly post this information publicly. FDORA also gives the FDA increased authority to withdraw approval of a drug or biologic granted accelerated approval on an expedited basis if the sponsor fails to conduct such studies in a timely manner, send the necessary updates to the FDA, or if such post-approval studies fail to verify the drug's predicted clinical benefit. Under FDORA, the FDA is empowered to take action, such as issuing fines, against companies that fail to conduct with due diligence any post-approval confirmatory study or submit timely reports to the agency on their progress. 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. Thus, even if we seek to utilize the accelerated approval program, we may not be able to obtain accelerated approval and, even if we do, we may not experience a faster commercial launch of the product. In addition, receiving accelerated approval does not assure that the product's accelerated approval will eventually be converted to a traditional approval.

***We may seek designation for a current or future platform as a designated platform technology, but we might not receive such designation, and even if we do, such designation may not lead to a faster regulatory review or approval process.***

We may seek designation for our current or future platform as a designated platform technology. Under FDORA, a platform technology incorporated within or utilized by a drug or biological product is eligible for designation as a designated platform technology if (1) the platform technology is incorporated in, or utilized by, a drug approved under an NDA or a biologic licensed under a BLA; (2) preliminary evidence submitted by the sponsor of the approved drug or licensed biologic, or a sponsor that has been granted a right of reference to data submitted in the application for such drug or biologic, demonstrates that the platform technology has the potential to be incorporated in, or utilized by, more than one drug or biologic without an adverse effect on quality, manufacturing, or safety; and (3) data or information submitted by the applicable person indicates that incorporation or utilization of the platform technology has a reasonable likelihood to bring significant efficiencies to the drug or biologic development or manufacturing process and to the review process. A sponsor may request the FDA to designate a platform technology as a designated platform technology concurrently with, or at any time after, submission of an IND application for a drug or biologic that incorporates or utilizes the platform technology that is the subject of the request. If so designated, the FDA may expedite the development and review of any subsequent original NDA for a drug or BLA for a biologic that uses or incorporates the platform technology. Even if we believe our current or future platform technology meets the criteria for such designation, the FDA may disagree and instead determine not to grant such designation. In addition, the receipt of such designation for a platform technology does not ensure that a drug or biologic will be developed more quickly or receive FDA approval. Moreover, the FDA may revoke a designation if the FDA determines that a designated platform technology no longer meets the criteria for such designation.

***Even if we receive regulatory approval of one or more of our product candidates, we would 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 regulatory approvals that we receive for our product candidates will require surveillance to monitor the safety and efficacy of the product candidate. The FDA may also require a REMS in order to approve our product candidates, which could entail requirements for a medication guide, physician communication plans or additional elements to ensure safe use, such as restricted distribution methods, patient registries and other risk minimization tools. In addition, if the FDA or a comparable foreign regulatory authority approves our product candidates, the manufacturing processes, labeling, packaging, distribution, adverse event reporting, storage, advertising, promotion, import, export and recordkeeping for our product candidates will be subject to extensive and ongoing regulatory requirements. These requirements include submissions of safety and other post-marketing information and reports, registration, as well as continued compliance with cGMP for product manufacturing and compliance with GLP and GCP requirements for any studies that we conduct post-approval. In addition, manufacturers are required to comply with applicable product tracking and tracing requirements. Manufacturers and other parties involved in the drug supply chain for prescription drug products must also notify the FDA of counterfeit, diverted, stolen and intentionally adulterated products or products that are otherwise unfit for distribution in the United States. Later discovery of previously unknown problems with our product candidates, 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 result in, among other things:

- restrictions on the marketing or manufacturing of our product candidates, withdrawal of the product from the market or voluntary or mandatory product recalls;
- manufacturing delays and supply disruptions where regulatory inspections identify observations of noncompliance requiring remediation;
- revisions to the labeling, including limitation on approved uses or the addition of additional warnings, contraindications or other safety information, including boxed warnings;
- imposition of a REMS, which may include distribution or use restrictions;
- requirements to conduct additional post-market clinical trials to assess the safety of the product;
- fines, warning letters or holds on clinical trials;
- refusal by the FDA to approve pending applications or supplements to approved applications filed by us or suspension or revocation of approvals;
- product seizure or detention, or refusal to permit the import or export of our product candidates; and
- injunctions or the imposition of civil or criminal penalties.

Additionally, under FDORA, sponsors of approved drugs and biologics must provide 6 months' notice to the FDA of any changes in marketing status, such as the withdrawal of a drug, and failure to do so could result in the FDA placing the product on a list of discontinued products, which would revoke the product's ability to be marketed.

The FDA's and other regulatory authorities' policies may change and additional government regulations may be enacted that could prevent, limit or delay regulatory approval of our product candidates. We cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative action, either in the United States or abroad. 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 and we may not achieve or sustain profitability.

***The market opportunities for our oncology product candidates may be relatively small since the patients who may potentially be treated with our oncology product candidates are those who are ineligible for or have failed prior treatments, and our estimates of the prevalence of our target patient populations may be inaccurate.***

Cancer therapies are sometimes characterized by line of therapy (first line, second line, third line, fourth line, etc.), and the FDA often approves new therapies initially only for a particular line or lines of use. When cancer is detected early enough, first line therapy is sometimes adequate to cure the cancer or prolong life without a cure. Whenever first line therapy, usually chemotherapy, antibody drugs, tumor-targeted small molecules, hormone therapy, radiation therapy, surgery, or a combination of these, proves unsuccessful, second line therapy may be administered. Second line therapies often consist of more chemotherapy, radiation, antibody drugs, tumor-targeted small molecules, or a combination of these. Third line therapies can include chemotherapy, antibody drugs and small molecule tumor-targeted therapies, more invasive forms of surgery, and new technologies. There is no guarantee that our product candidates, even if approved as a second or third or subsequent line of therapy, would be approved for an earlier line of therapy, and, prior to any such approvals, we may have to conduct additional clinical trials.

Our projections of both the number of people who have the cancers we are targeting, who may have their tumors genetically sequenced, as well as the subset of people with these cancers in a position to receive a particular line of therapy and who have the potential to benefit from treatment with our product candidates, are based on our beliefs and estimates. These estimates have been derived from a variety of sources, including scientific literature, surveys of clinics, patient foundations or market research, and may prove to be incorrect. Further, new therapies may change the estimated incidence or prevalence of the cancers that we are targeting. Consequently, even if our product candidates are approved for a second or third line of therapy, the number of patients that may be eligible for treatment with our product candidates may turn out to be much lower than expected. In addition, we have not yet conducted market research to determine how treating physicians would expect to prescribe a product that is approved for multiple tumor types if there are different lines of approved therapies for each such tumor type.

***If we decide in the future to develop our product candidates in combination with other therapies, such strategy may expose us to additional risks.***

We may in the future develop one or more of our product candidates in combination with one or more approved or unapproved therapies. Even if any product candidate we develop were to receive marketing approval for use in combination with other approved therapies, the FDA, the EMA, the MHRA or comparable foreign regulatory authorities outside of the United States could still revoke approval of the therapy used in combination with our product. If the therapies used in combination with our product candidates are replaced as the standard of care for the indications we choose for any of our product candidates, the FDA, the EMA, the MHRA or comparable foreign regulatory authorities may require us to conduct additional clinical trials. The occurrence of any of these risks could result in our own products, if approved, being removed from the market or being less successful commercially.

Further, we will not be able to market and sell any product candidate we develop in combination with an unapproved cancer therapy for a combination indication if that unapproved therapy does not ultimately obtain marketing approval either alone or in combination with our product. In addition, unapproved cancer therapies face the same risks described with respect to our product candidates currently in development and clinical trials, including the potential for serious adverse effects, delay in their clinical trials and lack of FDA approval.

If the FDA, EMA, MHRA or comparable foreign regulatory authorities do not approve these other products or revoke their approval of, or if safety, efficacy, quality, manufacturing or supply issues arise with, the products we choose to evaluate in

combination with the product candidate we develop, we may be unable to obtain approval of or market such combination therapy.

***Certain of our product candidates are expected to be used with a drug delivery system and thus may be regulated as a combination product and may face additional challenges, risks and delays in the product development and regulatory approval process.***

Certain of our product candidates may be used with a drug delivery system and thus may be regulated as a combination product. When evaluating product candidates that utilize a specific drug delivery system or device, the FDA will evaluate the characteristics of that delivery system and its functionality, as well as the potential for undesirable interactions between the drug and the delivery system, including the potential to negatively impact the safety or effectiveness of the drug. The FDA review process can be more complicated for combination products, and may result in delays, particularly if novel delivery systems are involved. Additionally, quality or design concerns with the delivery system could delay or prevent regulatory approval and commercialization of devices.

#### **Risks Related to our Reliance on Third Parties**

***We rely, and expect to continue to rely, on third parties to conduct our preclinical studies and clinical trials and if these third parties perform in an unsatisfactory manner, our business could be substantially harmed.***

We currently conduct and expect to continue to rely on third parties such as contract development and manufacturing organizations, or CMOs, and contract research organizations, or CROs, to manufacture our products and conduct our clinical trials. We do not currently have the ability to independently conduct large-scale clinical trials, such as a Phase 3 clinical trial, without assistance of third parties.

We have relied upon and plan to continue to rely upon medical institutions, clinical investigators, contract laboratories and other third parties, such as CROs, to conduct or assist us in conducting GCP-compliant clinical trials on our product candidates properly and on time, and may not currently have all of the necessary contractual relationships in place to do so. Once we have established contractual relationships with such third-party CROs, we will have only limited control over their actual performance of these activities.

We and our CMOs, CROs and other vendors are required to comply with cGMP, GCP and GLP which are regulations and guidelines enforced by the FDA, the Competent Authorities of the Member States of the European Union and any comparable foreign regulatory authorities for all of our product candidates in preclinical and clinical development. Regulatory authorities enforce these regulations through periodic inspections of trial sponsors, principal investigators, clinical trial sites and other contractors. Although we rely on CROs to conduct any current or planned GLP-compliant preclinical studies and GCP-compliant clinical trials and have limited influence over their actual performance, we remain responsible for ensuring that each of our preclinical studies and clinical trials is conducted in accordance with its investigational plan and protocol and applicable laws and regulations, and our reliance on the CROs does not relieve us of our regulatory responsibilities. If we or any of our CROs or vendors fail to comply with applicable regulations, the data generated in our preclinical studies and clinical trials may be deemed unreliable and the FDA, EMA, MHRA or any comparable foreign regulatory agency may require us to perform additional preclinical studies and clinical trials before approving our marketing applications. We cannot assure you that upon inspection by a given regulatory agency, such regulatory agency will determine that all of our clinical trials comply with GCP regulations. In addition, our clinical trials must be conducted with products produced under cGMP requirements. Our failure to comply with these requirements may require us to repeat clinical trials, which would delay the regulatory approval process.

While we will have agreements governing their activities, our CROs will not be our employees, and we will not be able to control whether or not they devote sufficient time and resources to our future preclinical and clinical programs. These CROs may also have relationships with other commercial entities, including our competitors, for whom they may also be conducting clinical trials, or other drug development activities which could harm our business. We face the risk of potential unauthorized disclosure or misappropriation of our intellectual property by CROs, which may reduce our trade secret protection and allow our potential competitors to access and exploit our proprietary technology. CROs also may use our proprietary information and intellectual property in such a way as to result in litigation or other intellectual property-related proceedings that could jeopardize or invalidate our proprietary information and intellectual property. If our CROs do not successfully carry out their contractual duties or obligations, fail to meet expected deadlines, or if the quality or accuracy of the clinical data they obtain is compromised due to the failure to adhere to our clinical protocols or regulatory requirements or for any other reason, our clinical trials may be extended, delayed or terminated, the clinical data generated in our clinical trials may be deemed unreliable, and we may not be able to obtain regulatory approval for, or successfully commercialize any product candidate that we develop. If our CROs become subject to regulatory investigations or sanctions or are otherwise prevented or restricted from

performing their business, this may result in a material delay to our development and/or commercial activities, add significant additional cost, require that we move our non-clinical and/or clinical development activities to alternative vendors, in each case, which may materially and adversely impact our ability to conduct our clinical trials and/or develop or commercialize our products in a timely manner or at all. As a result, our financial results and the commercial prospects for any product candidate that we develop would be harmed, our costs could increase, and our ability to generate revenue could be delayed.

If our relationships with these CROs terminate, we may not be able to enter into arrangements with alternative CROs or do so on commercially reasonable terms. Switching or adding additional CROs involves substantial cost and requires management time and focus, and could delay development and commercialization of our product candidates. In addition, there is a natural transition period when a new CRO commences work. As a result, delays occur, which can negatively 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 negative impact on our business and financial condition.

***We could experience manufacturing problems that result in delays or other material disruptions in our development or commercialization of our programs or otherwise harm our business.***

The manufacturing processes our CMOs use to produce our and our affiliates' product candidates are complex. Several factors could cause production interruptions, including inability to develop novel manufacturing processes, equipment malfunctions, facility contamination, raw material shortages or contamination, natural disasters, disruption in utility services, human error or disruptions in the operations of our suppliers, including acquisition of the supplier by a third party or declaration of bankruptcy. The expertise required to manufacture these product candidates may be unique to a particular CMO, and as a result, it would be difficult and time consuming to find an alternative CMO. If our CMOs become subject to regulatory investigations or sanctions or are otherwise prevented or restricted from performing their business, this may result in material delay or other disruption to our development and / or commercial activities, add significant additional cost, require that we move our non-clinical and/or clinical development and/or manufacturing activities to alternative vendors, in each case, which may materially and adversely impact our ability to develop and manufacture our product candidates, conduct our clinical trials and/or commercialize our products in a timely manner, or at all, or result in unacceptable additional costs.

Some of our product candidates may include biologics, some of which may have physical and chemical properties that cannot be fully characterized. As a result, assays of the finished product may not be sufficient to ensure that the product is consistent from lot-to-lot or will perform in the intended manner. Accordingly, our CMOs must employ multiple steps to control the manufacturing process to assure that the process is reproducible and the product candidate is made strictly and consistently in compliance with the process. Problems with the manufacturing process, even minor deviations from the normal process, could result in product defects or manufacturing failures that result in lot failures, product recalls, product liability claims or insufficient inventory to conduct clinical trials or supply commercial markets. We may encounter problems achieving adequate quantities and quality of clinical-grade materials that meet the FDA, the EMA, the MHRA or other applicable standards or specifications with consistent and acceptable production yields and costs.

In addition, the FDA, the EMA, the MHRA and other foreign 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, the EMA, the MHRA or other foreign 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.

Our CMOs also may encounter problems hiring and retaining the experienced scientific, quality assurance, quality-control and manufacturing personnel needed to operate our manufacturing processes, which could result in delays in production or difficulties in maintaining compliance with applicable regulatory requirements.

Any problems in our CMOs' supply chain, manufacturing process or facilities could result in delays in planned clinical trials and increased costs, and could make us a less attractive collaborator for potential partners, including larger biotechnology companies and academic research institutions, which could limit access to additional attractive development programs. Problems in our manufacturing process could restrict our ability to meet potential future market demand for products.

***We currently rely and expect to rely in the future on the use of third parties to manufacture our product candidates. Our business could be harmed if the third party manufacturers experience supply chain shortages, fail to provide us with***

***sufficient quantities of our product candidates or fail to do so at acceptable quality levels or prices or deliver defective products.***

We do not currently own any facility that may be used as our clinical-scale manufacturing and processing facility and must currently rely on outside vendors to manufacture our product candidates. We will need to negotiate and maintain contractual arrangements with these outside vendors for the supply of our product candidates and we may not be able to do so on favorable terms. We have not yet caused our product candidates to be manufactured on a commercial scale and may not be able to do so for any of our product candidates.

Our reliance on a limited number of third-party manufacturers exposes us to a number of risks, including the following:

- a contract manufacturer may fail to perform its obligations, and we may be forced to enter into an agreement with a different CMO, which we may not be able to do on reasonable terms, if at all, and our clinical supply could be delayed significantly as we establish alternative supply sources;
- we may be unable to identify manufacturers on acceptable terms or at all because the number of potential manufacturers is limited and the FDA must inspect any manufacturers for current cGMP compliance as part of our marketing application;
- a new manufacturer would have to be educated in, or develop substantially equivalent processes for, the production of our product candidates and in some cases, the technical skills required to manufacture our product candidates may be unique or proprietary to the original CMO 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;
- a change in manufacturer will require us to verify that the new CMO maintains facilities and procedures that comply with quality standards and with all applicable regulations and such verification may result in material delays to our programs;
- a change in manufacturers or certain changes in manufacturing processes/procedures will require that we conduct a manufacturing comparability study to verify that any new manufacturer or manufacturing process/procedures will produce our product candidate according to the specifications previously submitted to the FDA or other regulatory authority, and such study may be unsuccessful and could require the conduct of additional clinical trials;
- our third-party manufacturers may be unable to timely manufacture our product candidates or produce the quantity and quality required to meet our clinical and commercial needs, if any;
- contract manufacturers may not be able to execute our manufacturing procedures and other logistical support requirements appropriately;
- our future contract manufacturers may not perform as agreed, may not devote sufficient resources to our product candidates or may not remain in the contract manufacturing business for the time required to supply our clinical trials or to successfully produce, store, and distribute our products, if any;
- manufacturers are subject to ongoing periodic unannounced inspection by the FDA and corresponding state agencies to ensure strict compliance with cGMP and other government regulations and corresponding foreign standards and we have no control over third-party manufacturers' compliance with these regulations and standards;
- we may not own, or may have to share, or obtain a license to, the intellectual property rights to any improvements made by our third-party manufacturers in the manufacturing process for our product candidates;
- our third-party manufacturers could breach or terminate their agreements with us;
- raw materials and components used in the manufacturing process, particularly those for which we have no other source or supplier, may not be available or may not be suitable or acceptable for use due to material or component defects;
- our contract manufacturers and critical reagent suppliers may be subject to inclement weather, as well as natural or man-made disasters; and
- our contract manufacturers may have unacceptable or inconsistent product quality success rates and yields, and we have no direct control over our contract manufacturers' ability to maintain adequate quality control, quality assurance and qualified personnel.

Each of these risks could delay or prevent the completion of our clinical trials or the approval of any of our product candidates by the FDA or any other regulatory authority, result in higher costs (including resulting from batch failures) or adversely impact commercialization of our product candidates. In addition, we will rely on third parties to perform certain

specification tests on our product candidates prior to delivery to patients. If these tests are not appropriately done and test data are not reliable, patients could be put at risk of serious harm and the FDA could place significant restrictions on our company until deficiencies are remedied. Moreover, because each of our Centessa Subsidiaries has a separate manufacturing process for their programs, we will not benefit from any synergies related to manufacturing costs. We may also face logistical problems in managing different CMOs and processes for all of our Centessa Subsidiaries.

***Certain third parties upon whom we rely for the supply of the active pharmaceutical ingredient used in our product candidates are our sole source of supply, and the loss of any of these suppliers could significantly harm our business.***

Certain of the third parties upon whom we rely for the supply of the active pharmaceutical ingredient used in our product candidates are our sole source of supply, and the loss of any of these suppliers could significantly harm our business. The active pharmaceutical ingredients (“API”) used in certain of our product candidates are supplied to us from single-source suppliers. Our ability to successfully develop our product candidates, and to ultimately supply our commercial products in quantities sufficient to meet the market demand, depends in part on our ability to obtain the API for these products in accordance with regulatory requirements and in sufficient quantities for clinical testing and commercialization. We do not currently have arrangements in place for a redundant or second-source supply of any such API in the event any of our current suppliers of such API cease their operations for any reason. We are also unable to predict how changing global economic conditions or potential global health concerns will affect our third-party suppliers and manufacturers. Any negative impact of such matters on our third-party suppliers and manufacturers may also have an adverse impact on our results of operations or financial condition. For all of our product candidates, we intend to identify and qualify additional manufacturers to provide such API prior to submission of an NDA or BLA (as applicable) to the FDA and/or EMA, MHRA or other applicable regulatory bodies. We are not certain, however, that our single-source suppliers will be able to meet our demand for their products, either because of the nature of our agreements with those suppliers, our limited experience with those suppliers or our relative importance as a customer to those suppliers. It may be difficult for us to assess their ability to timely meet our demand in the future based on past performance. While our suppliers have generally met our demand for their products on a timely basis in the past, they may subordinate our needs in the future to their other customers. Establishing additional or replacement suppliers for the API used in our product candidates, if required, may not be accomplished quickly. If we are able to find a replacement supplier, such replacement supplier would need to be qualified and may require additional regulatory inspection or approval, which could result in further delay. While we seek to maintain adequate inventory of the API used in our product candidates, any interruption or delay in the supply of components or materials, or our inability to obtain such API from alternate sources at acceptable prices in a timely manner could impede, delay, limit or prevent our development efforts, which could harm our business, results of operations, financial condition and prospects.

In the U.S., legislative proposals are pending that, if enacted, could negatively impact U.S. funding for certain biotechnology providers that have relationships with certain foreign governments or which pose a threat to national security. If these proposals become law, the potential downstream adverse impacts on entities having commercial relationships with any impacted biotechnology provider is unknown but may include supply chain disruptions or delays. If these proposals become law and any of our vendors become subject to any such laws, this may result in material delays or other disruptions to our development and/or commercial activities, add significant additional cost, require that we move our non-clinical and/or clinical development and/or manufacturing activities to alternative vendors, in each case, which may materially and adversely impact our ability to develop and manufacture our product candidates, conduct our clinical trials and/or commercialize our products in a timely manner, or at all, or result in unacceptable additional costs.

***If our third-party manufacturers use hazardous and biological materials in a manner that causes injury or violates applicable law, we may be liable for damages.***

Our research and development activities involve the controlled use of potentially hazardous substances, including chemical and biological materials, by our third-party manufacturers. Our manufacturers are subject to federal, state and local laws and regulations in the United States governing the use, manufacture, storage, handling and disposal of medical and hazardous materials. Although we believe that our manufacturers’ procedures for using, handling, storing and disposing of these materials comply with legally prescribed standards, we cannot completely eliminate the risk of contamination or injury resulting from medical or hazardous materials. As a result of any such contamination or injury, we may incur liability or local, city, state or federal authorities may curtail the use of these materials and interrupt our business operations. In the event of an accident, we could be held liable for damages or penalized with fines, and the liability could exceed our resources. We do not have any insurance for liabilities arising from medical or hazardous materials. Compliance with applicable environmental laws and regulations is expensive, and current or future environmental regulations may impair our research, development and production efforts, which could harm our business, prospects, financial condition or results of operations.

***If we are unable to obtain licenses from third parties on commercially reasonable terms or fail to comply with our obligations under such agreements, our business could be harmed.***

If it may be necessary for us to use the patented or proprietary technology of third parties to commercialize our products, in which case we would be required to obtain a license from these third parties. If we are unable to license such technology, or if we are forced to license such technology, on unfavorable terms, our business could be materially harmed. If we are unable to obtain a necessary license, we may be unable to develop or commercialize the affected product candidates, which could materially harm our business, and the third parties owning such intellectual property rights could seek either an injunction prohibiting our sales, or, with respect to our sales, an obligation on our part to pay royalties and/or other forms of compensation. Even if we are able to obtain a license, it may be non-exclusive, thereby giving our competitors access to the same technologies licensed to us.

If we fail to comply with our obligations under our license agreements, our counterparties may have the right to terminate these agreements, in which event we might not be able to develop, manufacture or market, or may be forced to cease developing, manufacturing or marketing, any product that is covered by these agreements or may face other penalties under such agreements. Such an occurrence could materially adversely affect the value of the product candidate being developed under any such agreement. Termination of these agreements or reduction or elimination of our rights under these agreements may result in our having to negotiate new or reinstated agreements with less favorable terms, cause us to lose our rights under these agreements, including our rights to important intellectual property or technology, or impede, delay or prohibit the further development or commercialization of one or more product candidates that rely on such agreements.

### **Risks Related to Intellectual Property**

***If we are unable to obtain and maintain sufficient patent and other intellectual property protection for our product candidates and technology or other product candidates that may be identified, or if the scope of the intellectual property protection obtained is not sufficiently broad, our competitors could develop and commercialize product candidates similar or identical to the product candidates, and our ability to successfully commercialize the product candidates and other product candidates that we may pursue may be impaired.***

As is the case with other pharmaceutical and biopharmaceutical companies, our success depends in significant part on our ability and the ability of our licensors and collaborators to obtain, maintain, enforce and defend patents and other intellectual property rights with respect to our product candidates and technology and to operate our business without infringing, misappropriating, or otherwise violating the intellectual property rights of others. We have and expect to continue to maintain and expand our own patent estate.

We have also licensed patent and other intellectual property rights to and from our partners. Some of these licenses give us the right to prepare, file and prosecute patent applications and maintain and enforce patents we have licensed, whereas other licenses may not give us such rights. In some circumstances, we may not have the right to control the preparation, filing and prosecution of patent applications or to maintain the patents covering technology that we license to or from our partners, and we may have to rely on our partners to fulfill these responsibilities. Consequently, these patents and applications may not be prosecuted and enforced in a manner consistent with the best interests of our business. If our current or future licensors, licensees or collaborators fail to establish, maintain or protect such patents and other intellectual property rights, such rights may be reduced or eliminated. If our licensors, licensees or collaborators are not fully cooperative or disagree with us as to the prosecution, maintenance or enforcement of any patent rights, such patent rights could be compromised.

The patent prosecution process is expensive and time-consuming. We and our current or future licensors, licensees or collaborators may not be able to prepare, file and prosecute all necessary or desirable patent applications at a reasonable cost or in a timely manner. It is also possible that we or our licensors will fail to file patent applications covering inventions made in the course of development and commercialization activities before a competitor or another third party files a patent application covering, or publishes information disclosing, a similar, independently-developed invention. Such competitor's patent application may pose obstacles to our ability to obtain or limit the scope of patent protection we may obtain. Although we enter into non-disclosure and confidentiality agreements with parties who have access to confidential or patentable aspects of our research and development output, such as our employees, collaborators, CROs, contract manufacturers, consultants, advisors and other third parties, any of these parties may breach the agreements and disclose such output before a patent application is filed, thereby jeopardizing our ability to seek patent protection. In addition, publications of discoveries in the scientific literature often lag behind the actual discoveries, and patent applications in the United States and other jurisdictions are typically not published until 18 months after filing, or in some cases not at all. Therefore, we cannot be certain that we or our licensors were the first to make the inventions claimed in our owned or licensed patents or pending patent applications, or were the first to file for patent protection of such inventions, or if such licensed patents rights may otherwise become invalid.

The patent position of biotechnology and pharmaceutical companies generally is uncertain, involves complex legal and factual questions and is the subject of much litigation. As a result, the issuance, scope, validity, enforceability and commercial value of our and our current or future licensors' patent rights are uncertain. Our and our licensors' pending and future patent applications may not result in patents being issued that protect our technology or products, in whole or in part, or which effectively exclude others from commercializing competitive technologies and products. The patent examination process may require us or our licensors to narrow the scope of the claims of our pending and future patent applications, and therefore, even if such patent applications 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. Our and our licensors' patent applications cannot be enforced against third parties practicing the technology claimed in such applications unless and until a patent issues from such applications, and then only to the extent the issued claims cover such technology. Any of the foregoing could harm our competitive position, business, financial condition, results of operations and prospects.

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

Our trademarks or trade names may be challenged, infringed, diluted, circumvented or declared generic or determined to be infringing on other marks. We intend to rely on both registration and common law protection for our trademarks. 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 the trademark registration process, we may receive Office Actions from the United States Patent and Trademark Office ("USPTO"), objecting to the registration of our trademark. Although we would be given an opportunity to respond to those objections, 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/or to seek the cancellation of 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 obtain a registered trademark or 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.

***If we are unable to protect the confidentiality of our trade secrets, our business and competitive position would be harmed.***

We rely on trade secrets and confidentiality agreements to protect our unpatented know-how, technology and other proprietary information and to maintain our competitive position. Trade secrets and know-how can be difficult to protect. We seek to protect these trade secrets and other proprietary technology, in part, by entering into non-disclosure and confidentiality agreements with parties who have access to them, such as our employees, collaborators, CROs, contract manufacturers, consultants, advisors and other third parties. We also enter into confidentiality and invention or patent assignment agreements with our employees and consultants. We cannot guarantee that we have entered into such agreements with each party that may have or has had access to our trade secrets or proprietary technology and processes. Despite these efforts, any of these parties may breach the agreements and disclose our proprietary information, including our trade secrets, and we may not be able to obtain adequate remedies for such breaches. These risks are heightened due to our reliance on third parties, including third party consultants, CROs and CMOs, for certain aspects of our business. The activities conducted by our third party vendors require us to share our trade secrets with them, which increases the possibility that a competitor will discover them or that our trade secrets will be misappropriated or disclosed. Enforcing a claim that a party illegally disclosed or misappropriated a trade secret is difficult, expensive and time-consuming, and the outcome is unpredictable. In addition, some courts inside and outside the United States are less willing or unwilling to protect trade secrets. If any of our trade secrets were to be lawfully obtained or independently developed by a competitor or other third party, we would have no right to prevent them 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 or other third party, our competitive position would be materially and adversely harmed.

***Third-party claims of intellectual property infringement, misappropriation or other violations may be costly and time consuming and may prevent or delay our product discovery and development efforts.***

Third parties may initiate legal proceedings alleging that we are infringing, misappropriating, or otherwise violating their intellectual property rights, the outcome of which would be uncertain and could have a material adverse effect on the success of our business. Our commercial success depends upon our ability to develop, manufacture, market and sell our current and future product candidates and use our proprietary technologies without infringing, misappropriating or otherwise violating the intellectual property rights of third parties. There is a substantial amount of litigation involving patents and other intellectual property rights in the biotechnology and pharmaceutical industries, as well as administrative proceedings for challenging patents, including derivation, interference, reexamination, inter partes review, and post grant review proceedings before the

USPTO or oppositions and other comparable proceedings in foreign jurisdictions. We or any of our future licensors or strategic partners may be party to, exposed to, or threatened with, future adversarial proceedings or litigation by third parties having patent or other intellectual property rights alleging that our current or future product candidates and/or proprietary technologies infringe, misappropriate or otherwise violate their intellectual property rights. Even if we believe third-party intellectual property claims are without merit, there is no assurance that a court would find in our favor on questions of infringement, validity, enforceability, or priority.

We cannot assure that our product candidates and other technologies that we have developed, are developing or may develop in the future do not or will not infringe, misappropriate or otherwise violate existing or future patents or other intellectual property rights owned by third parties. Numerous U.S. and foreign issued patents and pending patent applications, which 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 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, 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 may allege they have patent rights encompassing our product candidates, technologies or methods.

If a third party claims that we infringe, misappropriate or otherwise violate its intellectual property rights, we may face a number of issues, including, but not limited to:

- infringement, misappropriation and other intellectual property claims which, regardless of merit, may be expensive and time-consuming to litigate and may divert our management's attention from our core business and may impact our reputation;
- substantial damages for infringement, misappropriation or other violations, which we may have to pay if a court decides that the product candidate or technology at issue infringes, misappropriates or violates the third party's rights, and, if the court finds that the infringement was willful, we could be ordered to pay treble damages and the patent owner's attorneys' fees;
- a court prohibiting us from developing, manufacturing, marketing or selling our product candidates, or from using our proprietary technologies, unless the third party licenses its product rights to us, which it is not required to do, on commercially reasonable terms or at all;
- if a license is available from a third party, we may have to pay substantial royalties, upfront fees and other amounts, and/or grant cross-licenses to intellectual property rights for our products, or the license to us may be non-exclusive, which would permit third parties to use the same intellectual property to compete with us;
- redesigning our product candidates or processes so they do not infringe, misappropriate or violate third party intellectual property rights, which may not be possible or may require substantial monetary expenditures and time; and
- 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, it could have a substantial adverse effect on the price of our ADSs.

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. In addition, any uncertainties resulting from the initiation and continuation of any litigation could have a material adverse effect on our ability to raise the funds necessary to continue our operations or could otherwise have a material adverse effect on our business, results of operations, financial condition and prospects. The occurrence of any of the foregoing could have a material adverse effect on our business, financial condition, results of operations or prospects.

We may choose to challenge the patentability of claims in a third party's U.S. patent by requesting that the USPTO review the patent claims in an ex parte re-exam, inter partes review or post-grant review proceedings. These proceedings are expensive and may consume our time or other resources. We may choose to challenge a third party's patent in patent opposition proceedings in the European Patent Office ("EPO"), or other foreign patent office. The costs of these opposition proceedings could be substantial, and may consume our time or other resources. If we fail to obtain a favorable result at the USPTO, EPO or other patent office then we may be exposed to litigation by a third party alleging that the patent may be infringed by our product candidates or proprietary technologies.

Third parties may assert that we are employing their proprietary technology without authorization. Patents issued in the United States by law enjoy a presumption of validity that can be rebutted only with evidence that is “clear and convincing,” a heightened standard of proof. There may be issued third-party patents of which we are currently unaware with claims to compositions, formulations, methods of manufacture or methods for treatment related to the use or manufacture of our product candidates. Patent applications can take many years to issue. In addition, because some patent applications in the United States may be maintained in secrecy until the patents are issued, patent applications in the United States and many foreign jurisdictions are typically not published until 18 months after filing, and publications in the scientific literature often lag behind actual discoveries, we cannot be certain that others have not filed patent applications covering our product candidates or technology. If any such patent applications issue as patents, and if such patents have priority over our patent applications or patents we may own or in-license, we may be required to obtain rights to such patents owned by third parties which may not be available on commercially reasonable terms or at all, or may only be available on a non-exclusive basis. There may be currently pending patent applications which may later result in issued patents that our product candidates may infringe. It is also possible that patents owned by third parties of which we are aware, but which we do not believe are relevant to our product candidates or other technologies, could be found to be infringed by our product candidates or other technologies. In addition, third parties may obtain patents in the future and claim that use of our technologies infringes upon these patents. Moreover, we may fail to identify relevant patents or incorrectly conclude that a patent is invalid, not enforceable, exhausted, or not infringed by our activities. If any third-party patents were held by a court of competent jurisdiction to cover the manufacturing process of our product candidates, molecules used in or formed during the manufacturing process, or any final product itself, the holders of any such patents may be able to block our ability to commercialize the product candidate unless we obtained a license under the applicable patents, or until such patents expire or they are finally determined to be held invalid or unenforceable. Similarly, if any third-party patent were held by a court of competent jurisdiction to cover aspects of our formulations, processes for manufacture or methods of use, including combination therapy or patient selection methods, the holders of any such patent may be able to block our ability to develop and commercialize the product candidate unless we obtained a license or until such patent expires or is finally determined to be held invalid or unenforceable. In either case, such a license may not be available on commercially reasonable terms or at all. If we are unable to obtain a necessary license to a third-party patent on commercially reasonable terms, or at all, our ability to commercialize our product candidates may be impaired or delayed, which could in turn significantly harm our business. Even if we obtain a license, it may be nonexclusive, thereby giving our competitors access to the same technologies licensed to us. In addition, if the breadth or strength of protection provided by our patent applications or any patents we may own or in-license in the future is threatened, it could dissuade companies from collaborating with us to license, develop or commercialize current or future product candidates.

Parties making claims against us may seek and obtain injunctive or other equitable relief, which could effectively block our ability to further develop and commercialize our product candidates. Defense of these claims, regardless of their merit, could involve substantial litigation expense and would be a substantial diversion of employee resources from our business. In the event of a successful claim of infringement, misappropriation or other violation against us, we may have to pay substantial damages, including treble damages and attorneys’ fees for willful infringement, obtain one or more licenses from third parties, pay royalties or redesign our infringing products, which may be impossible or require substantial time and monetary expenditure. We cannot predict whether any such license would be available at all or whether it would be available on commercially reasonable terms. Furthermore, even in the absence of litigation, we may need or may choose to obtain licenses from third parties to advance our research or allow commercialization of our product candidates. We may fail to obtain any of these licenses at a reasonable cost or on reasonable terms, if at all. In that event, we would be unable to further develop and commercialize our product candidates, which could harm our business significantly.

***We may become involved in lawsuits to protect or enforce our patents or other intellectual property, which could be expensive, time-consuming and unsuccessful, and issued patents covering our technology and product candidates could be found invalid or unenforceable if challenged.***

Competitors and other third parties may infringe or otherwise violate our issued patents or other intellectual property or the patents or other intellectual property of our licensors. In addition, our patents or the patents of our licensors may become involved in inventorship or priority disputes. Our pending patent applications cannot be enforced against third parties practicing the technology claimed in such applications unless and until a patent issues from such applications. To counter infringement or other unauthorized use, we may be required to file infringement claims, which can be expensive and time-consuming. Any claims we assert against perceived infringers could provoke these parties to assert counterclaims against us alleging that we infringe their patents or that our patents or our licensed patents are invalid or unenforceable. In a patent infringement proceeding, a court may decide that a patent of ours or a licensed patent is invalid or unenforceable, in whole or in part, construe the patent’s claims narrowly or refuse to stop the other party from using the technology at issue on the grounds that our patents do not cover the technology. An adverse result in any litigation proceeding could put one or more of our owned or

licensed patents at risk of being invalidated, held unenforceable or interpreted narrowly. We may find it impractical or undesirable to enforce our intellectual property against some third parties.

If we were to initiate legal proceedings against a third party to enforce a patent directed to our product candidates, or one of our future product candidates, the defendant could counterclaim that our patent is invalid or unenforceable. In patent litigation in the United States, defendant counterclaims alleging invalidity or unenforceability are commonplace. Grounds for a validity challenge could be an alleged failure to meet any of several statutory requirements, including lack of novelty, obviousness, non-enablement or insufficient written description. Grounds for an unenforceability assertion could be an allegation that someone connected with prosecution of the patent withheld relevant information from the USPTO or made a misleading statement during prosecution. Third parties may also raise similar claims before the USPTO or an equivalent foreign body, even outside the context of litigation. Potential proceedings include re-examination, post-grant review, inter partes review, interference proceedings, derivation proceedings and equivalent proceedings in foreign jurisdictions (e.g., opposition proceedings). Such proceedings could result in the revocation of, cancellation of, or amendment to our patents in such a way that they no longer cover our technology or any product candidates that we may develop. 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 and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity or unenforceability, we would lose at least part, and perhaps all, of the patent protection on the applicable product candidates or technology covered by the patent rendered invalid or unenforceable. Such a loss of patent protection would materially harm our business, financial condition, results of operations and prospects.

Interference proceedings provoked by third parties or brought by us or declared by the USPTO may be necessary to determine the priority of inventions with respect to our patents or patent applications. An unfavorable outcome could require us to cease using the related technology or to attempt to license rights to it from the prevailing party. Our business could be materially harmed if the prevailing party does not offer us a license on commercially reasonable terms.

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.

Our competitors may be larger than we are and may have substantially greater resources. They are, therefore, likely to be able to sustain the costs of complex patent litigation or proceedings more effectively than we can because of their greater financial resources and more mature and developed intellectual property portfolios. Accordingly, despite our efforts, we may not be able to prevent third parties from infringing upon, misappropriating or otherwise violating our intellectual property. Even if resolved in our favor, litigation or other legal proceedings relating to intellectual property claims could result in substantial costs and diversion of management resources, which could harm our business. In addition, the uncertainties associated with litigation could compromise our ability to raise the funds necessary to continue our clinical trials, continue our internal research programs, or in-license needed technology or other product candidates. There could also be public announcements of the results of the hearing, motions, or other interim proceedings or developments. If securities analysts or investors perceive those results to be negative, it could cause the price of shares of our ADSs to decline. Any of the foregoing events could harm our business, financial condition, results of operation and prospects.

***Intellectual property litigation could cause us to spend substantial resources and distract our personnel from their normal responsibilities.***

Even if resolved in our favor, 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, it could have a substantial adverse effect on the price of our ADSs. Such litigation or proceedings could substantially increase our operating losses and reduce the resources available for development activities or any future sales, marketing or distribution activities. We may not have sufficient financial or other resources to conduct such litigation or proceedings adequately. Some of our competitors may be able to sustain the costs of such litigation or proceedings more effectively than we can because of their greater financial resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could compromise our ability to compete in the marketplace, including compromising our ability to raise the funds necessary to continue our clinical trials, continue our research programs, license necessary technology from third parties, or enter into development collaborations that would help us commercialize our product candidates, if approved. Any of the foregoing events would harm our business, financial condition, results of operations and prospects.

***The patent protection we obtain for our product candidates and technology may be challenged or not sufficient enough to provide us with any competitive advantage.***

Even if our owned or licensed patent applications issue as patents, the issuance of any such patents is not conclusive as to their inventorship, scope, validity, or enforceability, and such patents may be challenged, invalidated or held to be unenforceable, including in the courts or patent offices in the United States and abroad, or circumvented. We may be subject to a third party pre-issuance submission of prior art to the USPTO, or equivalent foreign bodies, or become involved in opposition, derivation, revocation, re-examination, post-grant and inter partes review, or interference proceedings challenging our patent rights or the patent rights of others.

An adverse determination in any such submission, proceeding or litigation could reduce the scope of, or invalidate, our 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. Moreover, we, or one of our licensors, may have to participate in interference proceedings declared by the USPTO to determine priority of invention or in post-grant challenge proceedings, such as oppositions in a foreign patent office, that challenge priority of invention or other features of patentability. Such proceedings and any other patent challenges may result in loss of patent rights, loss of exclusivity, loss of priority, or in patent claims being narrowed, invalidated, or held unenforceable, which could limit our ability to stop others from using or commercializing similar or identical technology and products, or limit the duration of the patent protection of our technology and product candidates. Such proceedings also may result in substantial cost and require significant time from our scientists and management, even if the eventual outcome is favorable to us. Any of the foregoing could harm our business, financial condition, results of operations and prospects.

Furthermore, 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. As a result, our owned and licensed patent portfolios may not provide us with adequate protection against third parties seeking to commercialize products similar or identical to ours. We expect to request extensions of patent terms to the extent available in countries where we obtain issued patents. In the United States, the Drug Price Competition and Patent Term Restoration Act of 1984 permits a patent term extension of up to five years beyond the expiration of the patent. However, there are no assurances that the FDA or any comparable foreign regulatory authority or national patent office will grant such extensions, in whole or in part. In such case, our competitors may launch their products earlier than might otherwise be anticipated. Moreover, some of our owned or in-licensed patents and patent applications are, and may in the future be, co-owned with third parties. If we are unable to obtain an exclusive license to any such co-owners' interest in such patents or patent applications, such co-owners may be able to license their rights to other third parties, including our competitors, and our competitors could market competing products and technology. In addition, we may need the cooperation of any such co-owners in order to enforce such patents against third parties, and such cooperation may not be provided to us.

In addition, our owned and in-licensed patents may be subject to a reservation of rights by the licensor, its affiliates and one or more third parties. When new technologies are developed with government funding, the government generally obtains certain rights in any resulting patents, including a non-exclusive license authorizing the government to use the invention for noncommercial purposes. These rights may permit the government to disclose our confidential information to third parties or allow third parties to use our licensed technology. The government can also exercise its march-in rights if it determines that action is necessary because we fail to achieve practical application of the government-funded technology, because action is necessary to alleviate health or safety needs, to meet requirements of federal regulations, or to give preference to U.S. industry. In addition, our rights in such inventions may be subject to certain requirements to manufacture products embodying such inventions in the United States. Any of the foregoing could harm our competitive position, business, financial condition, results of operations and prospects.

***We may be subject to claims by third parties asserting that we or our employees have infringed upon, misappropriated or otherwise violated their intellectual property rights, or claiming ownership of what we regard as our own intellectual property.***

Many of our employees were previously employed at other biotechnology or pharmaceutical 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 these individuals have used or disclosed intellectual property, including trade secrets or other proprietary information, of any such individual's former employer. Litigation may be necessary to defend against these claims.

In addition, we or our licensors may be subject to claims that former employees, collaborators, or other third parties have an interest in our owned or in-licensed patents or other intellectual property as an inventor or co-inventor. While it is our

policy to require our employees and contractors who may be involved in the development of intellectual property to execute agreements assigning such intellectual property to us, we may be unsuccessful in executing such an agreement with each party who in fact develops intellectual property that we regard as our own. Our and their assignment agreements may not be self-executing or may be breached, and we may be forced to bring claims against third parties, or defend claims they may bring against us, to determine the ownership of what we regard as our intellectual property.

If we fail in prosecuting or defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights or personnel. Even if we are successful in prosecuting or defending against such claims, litigation could result in substantial costs, delay development of our product candidates and be a distraction to management. Any of the foregoing events would harm our business, financial condition, results of operations and prospects.

***We may not be able to protect our intellectual property rights throughout the world.***

Filing, prosecuting, maintaining, defending and enforcing patents on our product candidates in all countries throughout the world would be prohibitively expensive, and our intellectual property rights in some countries outside the United States can be less extensive than those in the United States. In addition, the laws of some foreign countries do not protect intellectual property rights to the same extent as federal and state laws in the United States. Consequently, we may not be able to prevent third parties from practicing our inventions in all countries outside the United States, or from selling or importing products made using our inventions in and into the United States or other jurisdictions. Competitors may use our technologies in jurisdictions where we have not obtained patent protection to develop their own drugs and may export otherwise infringing drugs to territories where we have patent protection, but enforcement rights are not as strong as those in the United States. These drugs may compete with our product candidates and our patents or other intellectual property rights may not be effective or sufficient to prevent them from competing.

Many companies have encountered significant problems in protecting and defending intellectual property rights in foreign jurisdictions. The legal systems of some countries do not favor the enforcement of patents and other intellectual property protection, which could make it difficult for us to stop the infringement of our patents generally. Proceedings to enforce our patent rights in foreign jurisdictions could result in substantial costs and divert our efforts and attention from other aspects of our business, could put our patents at risk of being invalidated or interpreted narrowly and our patent applications at risk of not issuing 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.

Many countries, including major European Union countries, India, Japan and China, have compulsory licensing laws under which a patent owner may be compelled under specified circumstances to grant licenses to third parties. In addition, many countries limit the enforceability of patents against government agencies or government contractors. In those countries, we may have limited remedies if patents are infringed or if we are compelled to grant a license to a third party, which could materially diminish the value of those patents. This could limit our potential revenue opportunities. 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, which could adversely affect our business, financial condition, results of operations and prospects.

***A number of our programs and associated product candidates are heavily dependent on licensed intellectual property. If we were to lose our rights to licensed intellectual property, we may not be able to continue developing or commercializing our product candidates, if approved. If we breach any of the agreements under which we license the use, development and commercialization rights to our product candidates or technology from third parties or, in certain cases, we fail to meet certain development deadlines, we could lose license rights that are important to our business.***

We are heavily reliant upon licenses to certain patent rights and other intellectual property from third parties that are important or necessary to the development of our product candidates. We may also need to obtain additional licenses to advance the development and commercialization of other product candidates we may develop. We expect that future license agreements will impose upon us, various development, regulatory and or commercial diligence obligations, payment of milestones and/or royalties and other obligations. If we fail to comply with our obligations under these agreements, or we are subject to a bankruptcy-related event, the licensor may have the right to terminate the license, in which event we would not be able to develop, market or otherwise commercialize products covered by the license, and in some instances, may be also obligated to transfer back to licensor our developments related to the licensed product and associated regulatory rights. Our business could suffer, for example, if any current or future licenses terminate, if the licensors fail to abide by the terms of the license, if the licensed patents or other rights are found to be invalid or unenforceable, or if we are unable to enter into necessary licenses on acceptable terms.

Licensing of intellectual property is of critical importance to our business and involves complex legal, business and scientific issues and certain provisions in intellectual property license agreements may be susceptible to multiple interpretations. Disputes may 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;
- 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 transfer, assign, or sublicense patent and other rights to third parties;
- 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 ownership of inventions and know-how resulting from the joint creation or use of intellectual property by our licensors and us and our partners;
- our right to transfer or assign the license;
- the ability and effects of termination; and
- restrictive covenants that may restrict our abilities to compete or market competing products.

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 harm our business, financial condition, results of operations and prospects. Moreover, 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 product candidates.

We may enter into additional licenses to third-party intellectual property that are necessary or useful to our business. Our current licenses and any future licenses that we may enter into impose various fees, royalty payment, milestone and other obligations on us. Under some license agreements, we may not control prosecution of the licensed intellectual property, or may not have the first right to enforce the intellectual property. In those cases, we may not be able to adequately influence patent prosecution or enforcement, or prevent inadvertent lapses of coverage due to failure to pay maintenance fees. If we fail to comply with any of our obligations under a current or future license agreement, the licensor may allege that we have breached our license agreement, and may accordingly seek to terminate our license. Termination of any of our current or future licenses could result in our loss of the right to use the licensed intellectual property, which could materially adversely affect our ability to develop and commercialize a product candidate or product, if approved, as well as harm our competitive business position and our business prospects. Under some license agreements, termination may also result in the transfer or granting of rights under certain of our intellectual property and information related to the product candidate being developed under the license, such as regulatory information.

In addition, if our licensors fail to abide by the terms of the license, if the licensors fail to prevent infringement by third parties, if the licensed patents or other rights are found to be invalid or unenforceable, or if we are unable to enter into necessary licenses on acceptable terms, our business, competitive position, financial condition, results of operations and prospects could be materially harmed.

***Patent terms may be inadequate to protect our competitive position on our product candidates for an adequate amount of time. If we are not able to obtain patent term extension or non-patent exclusivity in the United States under the Hatch-Waxman Act and in foreign countries under similar legislation, thereby potentially extending the marketing exclusivity term of our product candidates, our business may be materially harmed.***

Patents have a limited lifespan. In the United States, if all maintenance fees are timely paid, the natural expiration of a patent is generally 20 years from its earliest U.S. non-provisional filing date. Various extensions may be available, but the life of a patent, and the protection it affords, is limited. Even if patents covering our product candidates are obtained, once the patent life has expired for a product candidate, we may be open to competition from competitive medications, including generic medications. Given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such product candidates might expire before or shortly after such product candidates are commercialized. As a result, our owned and licensed patent portfolio may not provide us with sufficient rights to exclude others from commercializing product candidates similar or identical to ours.

Our in-licensed patents may expire before, or soon after, our first product achieves marketing approval in the United States or foreign jurisdictions. Upon the expiration of our current patents, we may lose the right to exclude others from practicing these inventions. The expiration of these patents could also have a similar material adverse effect on our business, financial condition, prospects and results of operations.

With respect to our biologics products, we hope to take advantage of enhanced regulatory exclusivity periods, such as the 12 years of regulatory exclusivity available to biologics manufacturers under the Biologics Competition and Innovation Act of 2009. However, despite these measures, we may still lose the right to exclude others from practicing these inventions, which may negatively impact our business.

Depending upon the timing, duration and conditions of any FDA marketing approval of our product candidates, one or more of our U.S. 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. However, we may not receive an extension if we fail to exercise due diligence during the testing phase or regulatory review process, 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. Only one patent per approved product can be extended, the extension cannot extend the total patent term beyond 14 years from approval and only those claims covering the approved drug, a method for using it or a method for manufacturing it may be extended. 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 the applicable product candidate 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. Further, if this occurs, our competitors may take advantage of our investment in development and trials by referencing our clinical and preclinical data and launch their product earlier than might otherwise be the case, and our competitive position, business, financial condition, results of operations and prospects could be materially harmed.

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

Obtaining and enforcing patents in the pharmaceutical industry is inherently uncertain, due in part to ongoing changes in the patent laws. Depending on decisions by Congress, the federal courts, and the USPTO, the laws and regulations governing patents, and interpretation thereof, could change in unpredictable ways that could weaken our and our licensors' or collaborators' ability to obtain new patents or to enforce existing or future patents. For example, the 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. Therefore, there is increased uncertainty with regard to our and our licensors' or collaborators' ability to obtain patents in the future, as well as uncertainty with respect to the value of patents once obtained.

Recent patent reform legislation could increase the uncertainties and costs surrounding the prosecution of our and our licensors' or collaborators' patent applications and the enforcement or defense of our or our licensors' or collaborators' issued patents. Assuming that other requirements for patentability are met, prior to March 2013, in the United States, the first to invent the claimed invention was entitled to the patent, while outside the United States, the first to file a patent application was entitled to the patent. After March 2013, under the Leahy-Smith America Invents Act (the "Leahy-Smith Act"), enacted in September 2011, the United States transitioned to a first inventor to file system in which, assuming that other requirements for patentability are met, the first inventor to file a patent application will be entitled to the patent on an invention regardless of whether a third party was the first to invent the claimed invention. The Leahy-Smith Act also includes a number of significant changes that affect the way patent applications are prosecuted and may also affect patent litigation. These include allowing third party submission of prior art to the USPTO during patent prosecution and additional procedures to attack the validity of a patent by USPTO-administered post-grant proceedings, including post-grant review, inter partes review and derivation proceedings. The USPTO recently developed new regulations and procedures to govern administration of the Leahy-Smith Act, and many of the substantive changes to patent law associated with the Leahy-Smith Act, particularly the first inventor-to-file provisions. Accordingly, it is not clear what, if any, impact the Leahy-Smith Act will have on the operation of our business. However, the Leahy-Smith Act 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, all of which could harm our business, financial condition, results of operations and prospects.

***Intellectual property rights do not necessarily address all potential threats.***

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. For example:

- others may be able to make products that are similar to any product candidates we may develop or utilize similar technology but that are not covered by the claims of the patents that we license or may own in the future;
- we, or our current or future licensors might not have been the first to make the inventions covered by the issued patent or pending patent application that we license or may own in the future;
- we, or our current or future licensors might not have been the first to file patent applications covering certain of our or their inventions;
- others may independently develop similar or alternative technologies or duplicate any of our technologies without infringing our owned or licensed intellectual property rights;
- it is possible that our pending owned or licensed patent applications or those that we may own or license in the future will not lead to issued patents;
- issued patents that we hold rights to may be held invalid or unenforceable, including as a result of legal challenges by our competitors;
- our competitors might conduct research and development activities in countries where we do not have patent rights and then use the information learned from such activities to develop competitive products for sale in our major commercial markets;
- we may not develop additional proprietary technologies that are patentable;
- the patents of others may harm our business; and
- we may choose not to file a patent in order to maintain certain trade secrets or know-how, and a third party may subsequently file a patent covering such intellectual property.

Should any of these events occur, they could harm our business, financial condition, results of operations and prospects.

***Obtaining and maintaining our patent protection depends on compliance with various procedural, document submission, fee payment and other requirements imposed by governmental 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 governmental fees on patents and/or applications will be due to be paid to the USPTO and various governmental patent agencies outside of the United States in several stages over the lifetime of the patents and/or applications. We have systems in place to remind us to pay these fees, and we employ an outside firm and rely on our outside counsel to pay these fees due to non-U.S. patent agencies. The USPTO and various non-U.S. governmental patent agencies require compliance with a number of procedural, documentary, fee payment and other similar provisions during the patent application process. We employ reputable law firms and other professionals to help us comply, and in many cases, an inadvertent lapse can be cured by payment of a late fee or by other means in accordance with the applicable rules. However, there are situations in which non-compliance can result in abandonment or lapse of the patent or patent application, resulting in partial or complete loss of patent rights in the relevant jurisdiction. In such an event, our competitors might be able to enter the market and this circumstance would have a material adverse effect on our business.

***We engage consultants employed by academic institutions in jurisdictions that contain inventorship laws mandating that any inventions developed by such consultants while performing consultancy services automatically or otherwise shall reside in the employing institution and granting such institutions the first right to develop and/or commercialize such inventions. We may not be able to secure rights (whether through ownership or license interest) in inventions developed by such consultants during performance of consulting services for our companies.***

We enter into confidentiality and intellectual property assignment agreements with our employees, consultants, outside scientific collaborators, sponsored researchers and other advisors. These agreements generally provide that inventions conceived by the party in the course of rendering services to us will be our exclusive property. However, these agreements may not be honored and may not effectively assign intellectual property rights to us. For example, even if we have a consulting agreement in place with an academic advisor pursuant to which such academic advisor is required to assign any inventions

developed in connection with providing services to us, such academic advisor may not have the right to assign such inventions to us, as it may conflict with his or her obligations to assign their intellectual property to his or her employing institution.

Despite our undertaking of the measures listed above, we may be subject to claims challenging the inventorship or ownership of our patents and other intellectual property and may be subject to further claims in the future. Litigation may be necessary to defend against 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.

### **Risks Related to Commercialization**

***We have never commercialized a product candidate and we may lack the necessary expertise, personnel and resources to successfully commercialize any of our products that receive regulatory approval on our own or together with collaborators.***

We have never commercialized a product candidate. Our operations to date have been limited to organizing and staffing our company, business planning, raising capital, acquiring the rights to our product candidates and undertaking preclinical studies and clinical trials of our product candidates. We currently have no sales force, marketing or distribution capabilities. To achieve commercial success of our product candidates, if any are approved, we will have to develop our own sales, marketing and supply capabilities or outsource these activities to a third party.

Factors that may affect our ability to commercialize our product candidates on our own include recruiting and retaining adequate numbers of effective sales and marketing personnel, obtaining access to or persuading adequate numbers of physicians to prescribe our product candidates and other unforeseen costs associated with creating an independent sales and marketing organization. Developing a sales and marketing organization requires significant investment, is time-consuming and could delay the launch of our product candidates. We may not be able to build an effective sales and marketing organization in the United States, the European Union, the United Kingdom or other key global markets. If we are unable to build our own distribution and marketing capabilities or to find suitable partners for the commercialization of our product candidates, we may have difficulties generating revenue from them.

***The commercial success of any of our product candidates will depend upon its degree of market acceptance by physicians, patients, third-party payors and others in the medical community.***

Ethical, social and legal concerns about our product candidates could result in additional regulations restricting or prohibiting our products. Even with the requisite approvals from the FDA in the United States, the European Commission (on the recommendation of the EMA) in the European Economic Area, the MHRA in the United Kingdom and other regulatory authorities internationally, the commercial success of our product candidates will depend, in part, on the acceptance of physicians, patients and health care payors of our product candidates as medically necessary, cost-effective and safe. Any product that we commercialize may not gain acceptance by physicians, patients, health care payors and others in the medical community. If these products do not achieve an adequate level of acceptance, we may not generate significant product revenue and may not become profitable. The degree of market acceptance of our product candidates, if approved for commercial sale, will depend on several factors, including:

- the efficacy and safety of such product candidates as demonstrated in clinical trials;
- the potential and perceived advantages of product candidates over alternative treatments;
- the cost of treatment relative to alternative treatments;
- the clinical indications for which the product candidate is approved by FDA, the European Commission or the MHRA;
- patient awareness of, and willingness to seek, genotyping;
- the willingness of physicians to prescribe new therapies;
- the willingness of the target patient population to try new therapies;
- the prevalence and severity of any side effects;
- product labeling or product insert requirements of FDA, EMA, MHRA or other regulatory authorities, including any limitations or warnings contained in a product's approved labeling;

- relative convenience and ease of administration;
- the strength of marketing and distribution support;
- the timing of market introduction of competitive products;
- publicity concerning our products or competing products and treatments; and
- sufficient third-party payor coverage and reimbursement.

Even if a potential product displays a favorable efficacy and safety profile in preclinical studies and clinical trials, market acceptance of the product will not be fully known until after it is launched.

***If the market opportunities for our product candidates are smaller than we believe they are, it may not be financially viable to commercialize, and if we do commercialize, our product revenues for any therapies that are approved for commercial sale may be adversely affected and our business may suffer.***

We focus our research and product development on treatments for various diseases. Our understanding of both the number of people who have these diseases, as well as the subset of people with these diseases who have the potential to benefit from treatment with our product candidates, are based on estimates. These estimates may prove to be incorrect and new studies may reduce the estimated incidence or prevalence of these diseases. The number of patients in the United States, the European Union, the United Kingdom and elsewhere may turn out to be lower than expected, may not be otherwise amenable to treatment with our products 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.

Further, there are several factors that could contribute to making the actual number of patients who receive our potential products less than the potentially addressable market. These include the lack of widespread availability of, and limited reimbursement for, new products or therapies in many underdeveloped markets.

***If we are unable to establish sales, medical affairs and marketing capabilities or enter into agreements with third parties to market and sell our product candidates, we may be unable to generate any product revenue.***

We currently have no sales and marketing organization. To successfully commercialize any products that may result from our development programs, we will need to develop these capabilities, either on our own or with others. The establishment and development of our own commercial team or the establishment of a contract sales force to market any products we may develop will be expensive and time-consuming and could delay any product launch. Moreover, we cannot be certain that we will be able to successfully develop this capability. We may enter into collaborations regarding our product candidates with entities to utilize their established marketing and distribution capabilities, but we may be unable to enter into such agreements on favorable terms, if at all. If any current or future collaborators do not commit sufficient resources to commercialize our products, or we are unable to develop the necessary capabilities on our own, we will be unable to generate sufficient product revenue to sustain our business. We compete with many companies that currently have extensive, experienced and well-funded medical affairs, marketing and sales operations to recruit, hire, train and retain marketing and sales personnel. We also face competition in our search for third parties to assist us with the sales and marketing efforts of our product candidates. 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.

Our efforts to educate the medical community and third-party payors on the benefits of our product candidates may require significant resources and may never be successful. Such efforts may require more resources than are typically required due to the complexity and uniqueness of our potential products. If any of our product candidates is approved but fails to achieve market acceptance among physicians, patients or third-party payors, we will not be able to generate significant revenues from such product, which could have a material adverse effect on our business, financial condition, results of operations and prospects.

***Healthcare legislative reform measures may have a material adverse effect on our business and results of operations and may increase the difficulty and cost for us to obtain marketing approval of and commercialize our product candidates and affect the prices we may charge for such product candidates.***

In the United States, there have been, and continue to be, several legislative initiatives to contain healthcare costs. These enacted or proposed legislative and regulatory changes affecting the healthcare system could prevent or delay marketing approval of our product candidates, restrict or regulate post-approval activities and affect our ability to profitably sell any

product for which we obtain marketing approval. For example, in 2010, the Patient Protection and Affordable Care Act, as amended by the Health Care and Education Reconciliation Act (“ACA”), was passed, which substantially changes the way health care is financed by both the government and private insurers, and significantly impacts the U.S. pharmaceutical industry. See “Business - Government Regulation - Health Reform” in our Annual Report on Form 10-K for the year ended December 31, 2024.

Additional changes that may affect our business include those governing enrollment in federal healthcare programs, reimbursement changes, rules regarding prescription drug benefits under the health insurance exchanges and fraud and abuse and enforcement. Continued implementation of the ACA and the passage of additional laws and regulations may result in the expansion of new programs such as Medicare payment for performance initiatives, and may impact existing government healthcare programs, such as by improving the physician quality reporting system and feedback program.

For each state that does not choose to expand its Medicaid program, there likely will be fewer insured patients overall, which could impact the sales, business and financial condition of manufacturers of branded prescription drugs. Where patients receive insurance coverage under any of the new options made available through the ACA, the possibility exists that manufacturers may be required to pay Medicaid rebates on that resulting drug utilization, a decision that could impact manufacturer revenues. The U.S. federal government also has announced delays in the implementation of key provisions of the ACA. The implications of these delays for our and our partners’ business and financial condition, if any, are not yet clear.

Moreover, increasing efforts by governmental and 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. There has been increasing legislative and enforcement interest in the United States with respect to specialty drug pricing practices. Specifically, there have been several recent U.S. Congressional inquiries and proposed and enacted federal and state legislation designed to, among other things, bring more transparency to drug pricing, reduce the cost of prescription drugs under Medicare, review the relationship between pricing and manufacturer patient programs, and reform government program reimbursement methodologies for drugs.

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.

We expect that the healthcare reform measures that have been adopted and may be adopted in the future, may result in more rigorous coverage criteria and in additional downward pressure on the price that we receive for any approved product and could seriously harm our future revenues. Any reduction in reimbursement from Medicare or other government programs may result in a similar reduction in payments from private payors. The implementation of cost containment measures or other healthcare reforms may prevent us from being able to generate revenue, attain profitability or commercialize our products.

***The insurance coverage and reimbursement status of newly-approved products is uncertain. Failure to obtain or maintain adequate coverage and reimbursement for our products, if approved, could limit our ability to market those products and decrease our ability to generate product revenue.***

We expect the cost of our product candidates and programs, to be substantial, when and if they achieve regulatory approval. We expect that coverage and reimbursement by government and private payors will be essential for most patients to be able to afford these treatments. Accordingly, sales of our product candidates will depend substantially, both domestically and abroad, on the extent to which the costs of our product candidates will be paid by health maintenance, managed care, pharmacy benefit and similar healthcare management organizations, or will be reimbursed by government authorities, private health coverage insurers and other third-party payors. See section entitled “Business - Government Regulation - Reimbursement” in our Annual Report on Form 10-K for the year ended December 31, 2024.

Obtaining coverage and reimbursement for a product from third-party payors is a time-consuming and costly process that could require us to provide to the payor supporting scientific, clinical and cost-effectiveness data. We may not be able to provide data sufficient to gain acceptance with respect to coverage and reimbursement. If coverage and reimbursement are not

available, or are available only at limited levels, we may not be able to successfully commercialize our product candidates. Even if coverage is provided, the approved reimbursement amount may not be adequate to realize a sufficient return on our investment.

There is significant uncertainty related to third-party coverage and reimbursement of newly approved products. In the United States, third-party payors, including government payors such as the Medicare and Medicaid programs, play an important role in determining the extent to which new drugs and biologics will be covered and reimbursed. The Medicare and Medicaid programs increasingly are used as models for how private payors and government payors develop their coverage and reimbursement policies. It is difficult to predict what the CMS will decide with respect to coverage and reimbursement for fundamentally novel products. Moreover, reimbursement agencies in the European Union may be more conservative than CMS. For example, several cancer drugs have been approved for reimbursement in the United States and have not been approved for reimbursement in certain European Union Member States. It is difficult to predict what third-party payors will decide with respect to the coverage and reimbursement for our product candidates.

Outside the United States, international operations generally are subject to extensive government price controls and other market regulations, and increasing emphasis on cost-containment initiatives in the European Union, Canada and other countries may put pricing pressure on us. In many countries, the prices of medical products are subject to varying price control mechanisms as part of national health systems. In general, the prices of medicines under such systems are substantially lower than in the United States. Other countries allow companies to fix their own prices for medical products, but monitor and control company profits. Additional foreign price controls or other changes in pricing regulation could restrict the amount that we are able to charge for our product candidates. Accordingly, in markets outside the United States, the reimbursement for our products may be reduced compared with the United States and may be insufficient to generate commercially reasonable product revenues.

Moreover, increasing efforts by government and 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 new products approved and, as a result, they may not cover or provide adequate payment for our product candidates. Net prices for drugs may be reduced by mandatory discounts or rebates required by government healthcare programs or private payors and by any future relaxation of laws that presently restrict imports of drugs from countries where they may be sold at lower prices than in the United States. Payors increasingly are considering new metrics as the basis for reimbursement rates, such as average sales price (“ASP”), average manufacturer price, and actual acquisition cost. The existing data for reimbursement based on some of these metrics is relatively limited, although certain states have begun to survey acquisition cost data for the purpose of setting Medicaid reimbursement rates, and CMS has begun making pharmacy National Average Drug Acquisition Cost and National Average Retail Price data publicly available on at least a monthly basis. Therefore, it may be difficult to project the impact of these evolving reimbursement metrics on the willingness of payors to cover candidate products that we or our partners are able to commercialize. In addition, many pharmaceutical manufacturers must calculate and report certain price reporting metrics to the government, such as ASP, and best price. Penalties may apply in some cases when such metrics are not submitted accurately and timely. Further, these prices for drugs may be reduced by mandatory discounts or rebates required by government healthcare programs. We expect to experience pricing pressures in connection with the sale of any of our product candidates due to the trend toward managed healthcare, the increasing influence of health maintenance organizations and additional legislative changes. The downward pressure on healthcare costs in general, particularly prescription drugs and surgical procedures and other treatments, has become intense. As a result, increasingly high barriers are being erected to the entry of new products such as ours.

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

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

- decreased demand for any product candidates or medicines that we may develop;
- injury to our reputation and significant negative media attention;
- withdrawal of clinical trial participants;
- significant costs to defend the related litigation;
- substantial monetary awards to trial participants or patients;

- loss of revenue; and
- the inability to commercialize our product candidates.

Although we maintain insurance coverage for clinical trials that we sponsor, it may not be adequate to cover all liabilities that we may incur. We anticipate that we will need to increase our insurance coverage as we commence additional clinical trials and if we successfully commercialize any product candidates. The market for insurance coverage is increasingly expensive, and the costs of insurance coverage will increase as our clinical programs increase in size. 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 Business and Industry**

***Inadequate funding for the FDA, the SEC and other government agencies, including from government shut downs, significant changes in leadership, personnel, or policies, or other disruptions to these agencies' operations, could hinder their ability to hire and retain key leadership and other personnel, prevent new products and services from being developed or commercialized in a timely manner or otherwise prevent those agencies from performing normal business functions on which the operation of our business may rely, 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, the ability to hire and retain key personnel and 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 the SEC and other government agencies on which our operations may rely, including those 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, including significant changes in leadership, personnel, or policies, may also slow the time necessary for new product candidates to be reviewed and/or approved by necessary government agencies, which would adversely affect our business. 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 business. Further, future government shutdowns could impact our ability to access the public markets and obtain necessary capital in order to properly capitalize and continue our operations.

***Adverse developments affecting the financial services industry, such as actual events or concerns involving liquidity, defaults, or non-performance by financial institutions or transactional counterparties, could adversely affect the Company's current and projected business operations and its financial condition and results of operations.***

Actual events involving limited liquidity, defaults, non-performance or other adverse developments that affect financial institutions, transactional counterparties or other companies in the financial services industry or the financial services industry generally, or concerns or rumors about any events of these kinds or other similar risks, have in the past and may in the future lead to market-wide liquidity problems. For example, in May 2023, the Federal Deposit Insurance Corporation ("FDIC") took control of First Republic Bank and JP Morgan Chase & Co. has since acquired a substantial amount of assets and certain liabilities of First Republic. If any of our lenders, including Oxford Finance, or counterparties to any such instruments were to be placed into receivership, we may be unable to access funds under the LSA. In addition, if any of our suppliers, including CROs and CMOs, or other parties with whom we conduct business are unable to access funds pursuant to such instruments or lending arrangements with such a financial institution, such parties' ability to perform their existing or future obligations to us could be adversely affected.

Even though we have not experienced any adverse impact to our liquidity or to our current and projected business operations, financial condition or results of operations, uncertainty remains over liquidity concerns in the broader financial services industry, and our business, our business partners, or industry as a whole may be adversely impacted in ways that we cannot predict at this time. Inflation and rapid increases in interest rates have led to a decline in the trading value of previously issued government securities with interest rates below current market interest rates. There is no guarantee that the U.S. Department of Treasury, FDIC and Federal Reserve Board will provide access to uninsured funds in the future in the event of the closure of other banks or financial institutions, or that they would do so in a timely fashion.

Although we assess our banking relationships as we believe necessary or appropriate, our access to funding sources in amounts adequate to finance or capitalize our current and projected future business operations could be significantly impaired by factors that affect us, the financial institutions with which we have credit agreements or arrangements directly, or the financial services industry or economy in general. These factors could include, among others, events such as liquidity constraints or failures, the ability to perform obligations under various types of financial, credit or liquidity agreements or arrangements, disruptions or instability in the financial services industry or financial markets, or concerns or negative

expectations about the prospects for companies in the financial services industry. These factors could involve financial institutions or financial services industry companies with which the Company has financial or business relationships, but could also include factors involving financial markets or the financial services industry generally.

The results of events or concerns that involve one or more of these factors could include a variety of material and adverse impacts on our current and projected business operations and our financial condition and results of operations. These could include, but may not be limited to:

- delayed access to deposits or other financial assets or the uninsured loss of deposits or other financial assets;
- delayed or lost access to, or reductions in borrowings or other working capital sources and/or delays, inability or reductions in the company's ability to refund, roll over or extend the maturity of, or enter into new credit facilities or other working capital resources;
- potential or actual breach of financial covenants in our credit agreements or credit arrangements; or
- potential or actual cross-defaults in other credit agreements, credit arrangements or operating or financing agreements.

In addition, investor concerns regarding the U.S. or international financial systems could result in less favorable commercial financing terms, including higher interest rates or costs and tighter financial and operating covenants, or systemic limitations on access to credit and liquidity sources, thereby making it more difficult for us to acquire financing on acceptable terms or at all. Any decline in available funding or access to our cash and liquidity resources could, among other risks, adversely impact our ability to meet our operating expenses, financial obligations or fulfill our other obligations, result in breaches of our financial and/or contractual obligations or result in violations of federal or state wage and hour laws. Any of these impacts, or any other impacts resulting from the factors described above or other related or similar factors not described above, could have material adverse impacts on our liquidity and our current and/or projected business operations and financial condition and results of operations.

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

Our ability to compete in the highly competitive biotechnology and pharmaceutical industries depends upon our ability to attract and retain highly qualified managerial, scientific and medical personnel. We are highly dependent on our senior management, including scientific and medical personnel and other key employees. While we expect to engage in an orderly transition process as we integrate newly appointed officers and managers, we face a variety of risks and uncertainties relating to management transition, including diversion of management attention from business concerns, failure to retain other key personnel or loss of institutional knowledge. In addition, the loss of the services of any of our executive officers, other key employees and other scientific and medical advisors, and an inability to find suitable replacements could result in delays in product development and harm our business. In particular, due to our small number of employees, the loss of one employee may have a larger impact on our business than compared to a loss at one of our peers. We currently do not maintain "key person" insurance for any members of our management team.

We may in the future expand our operations in the U.S. and other geographies, particularly in certain biotech hubs. Competition for skilled personnel in our market is intense and may limit our ability to hire and retain highly qualified personnel on acceptable terms or at all. Changes to immigration and work authorization laws and regulations, including those that restrain the flow of scientific and professional talent, can be significantly affected by political forces and levels of economic activity. Our business may be materially adversely affected if legislative or administrative changes to immigration or visa laws and regulations impair our hiring processes and goals or projects in the key jurisdictions in which we operate.

To encourage valuable employees to remain at our company, in addition to salary and cash incentives, we have provided equity awards that vest over time. The value to employees of equity awards that vest over time may be significantly affected by movements in our share price that are beyond our control, and may at any time be insufficient to counteract more lucrative offers from other companies. Despite our efforts to retain valuable employees, members of our management, scientific and development teams may terminate their employment with us at any time. Although we have employment agreements with our key employees, certain of these employment agreements provide for at-will employment, which means that any of our employees could leave our employment at any time. Our success also depends on our ability to continue to attract, retain and motivate highly skilled junior, mid-level and senior managers as well as junior, mid-level and senior scientific and medical personnel.

Additionally, we rely on our scientific founders and other scientific and clinical advisors and consultants to assist us in formulating our research, development and clinical strategies. Certain of our scientific founders, advisors and consultants are not our employees and may have commitments to, or consulting or advisory contracts with, other entities that may limit their

availability to us. If a conflict of interest arises between their work for us and their work for another entity, we may lose their services. Furthermore, our advisors may have arrangements with other companies to assist those companies in developing products or technologies that may compete with ours. In particular, if we are unable to maintain consulting relationships with our scientific founders or if they provide services to our competitors, our development and commercialization efforts will be impaired and our business will be significantly harmed.

***Our employees may engage in misconduct or other improper activities, including noncompliance with regulatory standards and requirements.***

We are exposed to the risk of employee fraud or other misconduct. Misconduct by employees could include intentional failures to comply with FDA regulations, provide accurate information to the FDA, comply with manufacturing standards we may establish, comply with federal and state healthcare fraud and abuse laws and regulations, report financial information or data accurately or disclose unauthorized activities to us. 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. Employee misconduct could also involve the improper use of information obtained in the course of clinical trials, which could result in regulatory sanctions and serious harm to our reputation. It is not always possible to identify and deter employee 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 fines or other sanctions.

***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 have a material adverse effect on the success of 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. Our operations involve the use of hazardous and flammable materials, including chemicals and biological materials. Our operations also produce hazardous waste products. We generally contract with third parties for the disposal of these materials and wastes. We cannot eliminate the risk of contamination or injury from these materials. In the event of contamination or injury resulting from our use of 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.

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

***Our internal computer systems have suffered, and our collaborators or other contractors or consultants may suffer from cybersecurity incidents or data breaches, which could result in a material disruption of our product development programs.***

In the ordinary course of our business, we may store, use, process or otherwise gain access to certain sensitive information, including proprietary information, confidential information, personal data and personal health data, intellectual property, trade secrets, and proprietary business information owned or controlled by ourselves or other parties. We may use third-party service providers and sub-processors to help us operate our business and our partners or other third parties may have access to such sensitive information or our systems or infrastructure in conjunction with our business. We may be required to expend significant resources, at significant cost, fundamentally change our business activities and practices, or modify our operations, including our clinical trial activities, or information technology in an effort to protect against security compromises or incidents, including cybersecurity incidents and data breaches and to mitigate, detect, and remediate actual or potential vulnerabilities to our information technology systems as well as harm stemming from cybersecurity threats such as cybersecurity incidents and data breaches. Our internal computer systems and infrastructure (including, without limitation, any relevant sensitive information and other assets stored therein or accessible thereby) and those of our current and any future collaborators, contractors or consultants are vulnerable to damage from a variety of evolving threats, including computer viruses, bugs, malware, unauthorized access, denial-of-service attacks, service interruptions, system malfunction (such as credential stuffing), social engineering attacks (such as phishing attacks), business email compromises, ransomware attacks, user errors or malfeasance, natural disasters, terrorism, war and telecommunication and electrical failures, as well as security

compromises, from inadvertent or intentional wrongful actions by insider employees, vendors, contractors, consultants, business partners, and/or other third parties, or from cyber-attacks by malicious third parties.

Like other companies in our industry, we, and our third party vendors, have experienced threats and cybersecurity incidents relating to our information technology systems and infrastructure. For example, in the past, Centessa Subsidiaries have experienced unauthorized access to systems through social engineering schemes. Although such past cyber-attacks did not result in material disruption to our business nor did they result in material loss, if any such material system failure, accident or cybersecurity compromises, incident, or data breach were to occur, it could result in a material disruption of our development programs and our business operations, whether due to a loss of our trade secrets or other sensitive information or other similar disruptions, as well as impact to our systems and infrastructure necessary for our business operations or necessitating that we incur significant costs to address such failure, accident or security compromises, cybersecurity incidents, or data breaches or expose us to significant liability. For example, the loss of clinical trial data from completed or future clinical trials could result in delays in our regulatory approval efforts and significantly increase our costs to recover or reproduce the data and expose us to data breach claims. In addition, failures or significant downtime of our information technology or telecommunication systems or infrastructure or those used by our third-party service providers could cause significant interruptions in our operations and adversely impact the confidentiality, integrity and availability of sensitive information. We may also be the subject of server malfunction, software or hardware failures, cyber-attacks (including supply-chain cyber-attacks), loss of data or other computer assets, and other similar issues. A significant portion of our workforce works remotely, which has increased the risk to our information technology assets and data.

To the extent that any disruption or cybersecurity compromise, cybersecurity incident, or data breach were to result in a loss of, or damage to, our data systems, infrastructure, or applications, or inappropriate disclosure, access to, or use of sensitive information, we could incur liability, our competitive position could be harmed and the further development and commercialization of our product candidates could be delayed. Relevant laws, regulations, and industry standards, as well as contractual obligations, may require us to implement specific security measures or use industry-standard or reasonable measures to protect against cybersecurity compromises, cybersecurity incidents and data breaches. Even though we have taken security measures designed to protect against cybersecurity incidents and/or data breaches, there can be no assurance that such security measures or those of our service providers, partners and other third parties will be effective in protecting against disruptions or cybersecurity compromises, cybersecurity incidents, or data breaches, or mitigating against the impact or the adverse consequences thereof. We may be unable to detect, anticipate, measure, prevent, or remediate threats or techniques used to detect or exploit vulnerabilities in our (or our third parties') information technology, services, communications or software, or to cause cybersecurity compromises, cybersecurity incidents, or data breaches, because such threats and techniques change frequently, are often sophisticated in nature, and may not be detected until after a cybersecurity incident or data breach has occurred. We cannot be certain that we will be able to address any such vulnerabilities, in whole or part, and there may be delays in developing and deploying patches and other remedial measures to adequately address vulnerabilities. Relevant laws, regulations, and industry standards, as well as contractual obligations, may also require us to notify relevant stakeholders (including affected individuals, partners, collaborators, customers, regulators, law enforcement agencies, credit reporting agencies and others) of cybersecurity incidents or data breaches. Such disclosures are costly and could also have a material adverse effect on our reputation, business, or financial condition.

Actual or perceived cybersecurity compromises, cybersecurity incidents, data breaches or other information technology system vulnerabilities, lack of appropriate information security safeguards and concerns regarding data privacy or cybersecurity may cause some of our actual or prospective customers, collaborators, partners and/or clinical trial participants to stop participating in our trials, using our products or working with us. Additionally, regulators could impose penalties and monetary fines against us for similar concerns, or we could incur other liability in connection with or resulting from litigation or governmental investigations and enforcement actions. The discontinuance of relationships with third parties, or the failure to meet the expectations of such third parties, and/or litigation, regulatory investigation or enforcement, could result in material harm to our operations, financial performance or reputation and affect our ability to grow and operate our business. Our contracts may not contain limitations of liability, and even where they do, there can be no assurance that limitations of liability in our contracts are sufficient to protect us from liabilities, damages, or claims related to our privacy and data security obligations. We cannot be sure that our insurance coverage, if any, will be adequate or otherwise protect us from or adequately mitigate liabilities arising out of such cybersecurity compromises, cybersecurity incidents, data breaches or other information technology system vulnerabilities. The successful assertion of one or more large claims against us that exceeds our available insurance coverage, or results in changes to our insurance policies (including premium increases or the imposition of large excess or deductible or co-insurance requirements), could materially and adversely affect our business.

***Our international operations may expose us to business, regulatory, legal, political, operational, financial, pricing and reimbursement risks associated with doing business across multiple jurisdictions outside of the United States.***

Our business is subject to risks associated with conducting business internationally. Our Centessa Subsidiaries, suppliers, industry partners and clinical study centers are located and/or conduct business across Europe, the United States and certain other jurisdictions. Furthermore, our business strategy incorporates potential international expansion as we seek to obtain regulatory approval for, and commercialize, our product candidates in patient populations outside the United States. If approved, we may hire sales representatives and conduct physician and patient association outreach activities across multiple jurisdictions. Doing business internationally involves a number of risks, including but not limited to:

- multiple, conflicting and changing laws, regulations, and compliance requirements such as privacy regulations, tax laws and practice, export and import restrictions, employment laws, regulatory requirements, and other governmental approvals, permits and licenses;
- failure by us to obtain and maintain regulatory approvals for the use of our products in various countries;
- rejection or qualification of foreign clinical trial data by the competent authorities of other countries;
- additional potentially relevant third-party patent rights;
- complexities and difficulties in obtaining, maintaining, protecting and enforcing our intellectual property;
- difficulties in staffing and managing foreign operations;
- complexities associated with managing multiple payor reimbursement regimes, government payors or patient self-pay systems;
- limits in our ability to penetrate international markets;
- financial risks, such as longer payment cycles, difficulty collecting accounts receivable, the impact of local and regional financial crises on demand and payment for our products and exposure to foreign currency exchange rate fluctuations;
- natural disasters, political and economic instability, including wars, terrorism and political unrest, outbreak of disease, boycotts, curtailment of trade and other business restrictions;
- certain expenses including, among others, expenses for travel, translation and insurance; and
- regulatory and compliance risks that relate to anti-corruption compliance and record-keeping that may fall within the purview of the U.S. Foreign Corrupt Practices Act and/or the UK Bribery Act of 2010, or provisions of anti-corruption or anti-bribery laws in other countries.

Any of these factors could harm our future international expansion and operations and, consequently, our results of operations.

***We may be subject, directly or indirectly, to federal and state healthcare fraud and abuse laws, false claims laws and health information privacy and security laws. If we are unable to comply, or have not fully complied, with such laws, we could face substantial penalties.***

If we obtain FDA approval for any of our product candidates and begin commercializing those products in the United States, our operations will be directly, or indirectly through our prescribers, customers and purchasers, subject to various federal and state fraud and abuse laws and regulations, including, without limitation, the federal Health Care Program Anti-Kickback Statute, the federal civil and criminal False Claims Act and Physician Payments Sunshine Act and regulations. These laws will impact, among other things, our proposed sales, marketing and educational programs. In addition, we may be subject to patient privacy laws by both the federal government and the states in which we conduct our business. See section entitled “*Business – Government Regulation – Other United States Healthcare Laws*” in our Annual Report on Form 10-K for the year ended December 31, 2024.

Because of the breadth of these laws and the narrowness of the statutory exceptions and safe harbors available, it is possible that some of our business activities could be subject to challenge under one or more of such laws. If our operations are found to be in violation of any of the laws described above or any other government regulations that apply to us, we may be subject to penalties, including civil and criminal penalties, damages, fines, exclusion from participation in government health care programs, such as Medicare and Medicaid, individual imprisonment and the curtailment or restructuring of our operations, any of which could adversely affect our ability to operate our business and our results of operations.

The provision of benefits or advantages to physicians to induce or encourage the prescription, recommendation, endorsement, purchase, supply, order or use of medicinal products is prohibited in the EU. The provision of benefits or advantages to induce or reward improper performance generally is also governed by the national anti-bribery laws of EU Member States, and the Bribery Act 2010 in the UK. Infringement of these laws could result in substantial fines and individual imprisonment.

Payments made to physicians in certain EU Member States must be publicly disclosed. Moreover, agreements with physicians often must be the subject of prior notification and approval by the physician's employer, his or her competent professional organization and/or the regulatory authorities of the individual EU Member States. These requirements are provided in the national laws, industry codes or professional codes of conduct, applicable in the EU Member States. Failure to comply with these requirements could result in reputational risk, public reprimands, administrative penalties, fines or individual imprisonment.

For further information on privacy laws, regulations and standards, as well as policies, contracts and other obligations related to data privacy and security, and the potential application thereof to our operations (including in relation to our use of health-related personal data), see the subsection immediately below this.

***We are subject to stringent and changing privacy laws, regulations and standards as well as policies, contracts and other obligations related to data privacy and security. Our actual or perceived failure to comply with such obligations could lead to government enforcement actions (that could include fines and penalties), a disruption of our clinical trials or commercialization of our products, private litigation, harm to our reputation, or other adverse effects on our business or prospects.***

The legislative and regulatory framework relating to the collection, use, retention, safeguarding, disclosure, sharing, transfer, security and other processing (collectively, "Process" or "Processing") of personal data (including health-related personal data) worldwide is rapidly evolving and is likely to remain uncertain for the foreseeable future. Globally, virtually every jurisdiction in which we operate has established its own data security and privacy frameworks with which we must comply and some of which may impose potentially conflicting obligations.

Accordingly, we are, or may become, subject to data privacy and security laws, regulations, and industry standards as well as policies, contracts and other obligations that apply to the Processing of personal data both by us and on our behalf (collectively, Data Protection Requirements). If we fail, or are perceived to have failed, to address or comply with Data Protection Requirements, this could result in government enforcement actions against us that could include investigations, fines, penalties, audits and inspections, additional reporting requirements and/or oversight, temporary or permanent bans on all or some Processing of personal data, orders to destroy or not use personal data, and imprisonment of company officials. Further, individuals or other relevant stakeholders could bring a variety of claims against us for our actual or perceived failure to comply with the Data Protection Requirements. Any of these events could have a material adverse effect on our reputation, business, or financial condition, and could lead to a loss of actual or prospective customers, collaborators or partners; interrupt or stop clinical trials; result in an inability to Process personal data or to operate in certain jurisdictions; limit our ability to develop or commercialize our products; or require us to revise or restructure our operations.

For example, in Europe, the collection and use of personal data, including health related data, is governed by the General Data Protection Regulation ("EU GDPR") which is applicable across the European Economic Area ("EEA"), and by related applicable data protection and privacy laws of the member states of the EEA. Switzerland has passed similar laws, and, following Brexit, the United Kingdom ("UK") has transposed the EU GDPR into UK domestic law with effect from January 2021 ("UK GDPR"). In this Quarterly Report on Form 10-Q, "GDPR" refers to both the UK GDPR and the EU GDPR, unless specified otherwise.

Collectively, European data protection laws (including the GDPR) are wide-ranging in scope and impose numerous, significant and complex compliance burdens in relation to the Processing of personal data, which increase our obligations (including with respect to clinical trials conducted in the EEA or the UK), such as: limiting permitted Processing of personal data to only that which is necessary for specified, explicit and legitimate purposes; requirements to conduct data protection impact assessments, requiring the establishment of a legal basis for Processing personal data; adopting a broad definition of personal data to possibly include 'pseudonymized' or key-coded data; creating obligations for controllers and processors to appoint data protection officers in certain circumstances; imposing stringent transparency obligations to data subjects, which requires more detailed notices for clinical trial subjects and investigators; introducing the obligation to carry out data protection impact assessments in certain circumstances; establishing limitations on the collection and retention of personal data through 'data minimization' and 'storage limitation' principles; establishing obligations to implement 'privacy by design'; introducing

obligations to honor increased rights for data subjects; formalizing a heightened and codified standard of data subject consent; establishing obligations to implement certain technical and organizational safeguards to protect the security and confidentiality of personal data; introducing obligations to agree to certain specific contractual terms and to take certain measures when working with third-party processors or joint controllers; imposing mandatory data breach notification requirements; and mandating the appointment of representatives in the UK and/or EU in certain circumstances. In particular, the Processing of “special category personal data” (such as personal data related to health and genetic information), which is relevant to our operations in the context of our conduct of clinical trials, imposes heightened compliance burdens under European data protection laws and is a topic of active interest among relevant regulators.

In addition, the GDPR provides that EEA member states may introduce specific or additional requirements related to the Processing of special categories of personal data such as health data that we may process in connection with clinical trials or otherwise. In the UK, the UK Data Protection Act 2018 complements the UK GDPR in this regard. This fact may lead to greater divergence on the law that applies to the Processing of such personal data across the EEA and/or UK, which may increase our costs and overall compliance risk. Such country-specific regulations could also limit our ability to Process relevant personal data in the context of our EEA and/or UK operations ultimately having an adverse impact on our business, and harming our business and financial condition.

Further, certain European data protection laws restrict transfers of personal data to countries outside Europe that are not considered by the European Commission and UK government as providing an adequate level of protection to personal data, like the United States in certain circumstances (so-called “third countries”). These transfers are prohibited unless an appropriate transfer safeguard mechanism specified by the European data protection laws is implemented, such as the Standard Contractual Clauses (“SCCs”) approved by the European Commission and/or the UK International Data Transfer Agreement/Addendum approved by the UK government, or a derogation applies. 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 European data protection laws 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. These transfer restrictions and may ultimately prevent us from transferring personal data outside Europe, which would cause significant business disruption. At present, there are few, if any, viable alternatives to the SCCs and UK IDTA. On July 10, 2023, the EU adopted an adequacy decision for a new “Data Privacy Framework,” which replaces the Privacy Shield, which the European Court of Justice invalidated in 2020 for personal data transferred from the EU to the U.S. The Framework allows for data transfers from the EU to companies who self-certified in the US and provides additional certification mechanisms to provide for data transfers from the UK. However, the long term viability of the Data Privacy Framework remains uncertain and the Framework has already been challenged in several jurisdictions.

The risks associated with such exports of personal data from locations within Europe are particularly relevant to our business as our group comprises several operating entities, many of which are located, and/or sponsor clinical trials, in Europe. We have adopted and implemented certain processes, systems and other relevant measures within our organization, and/or with our relevant collaborators, service providers, contractors or consultants, which are appropriate to address relevant requirements relating to international transfers of personal data from Europe, and to minimize the potential impacts and risks resulting from those requirements, across our organization. Failure to implement valid mechanisms for personal data transfers from Europe may result in increased exposure to regulatory actions, substantial fines and injunctions against processing personal data subject to European data protection laws. Inability to export personal data may also: restrict our activities outside Europe; limit our ability to collaborate with partners as well as other service providers, contractors and other companies outside of Europe; and/or require us to increase our Processing capabilities within Europe at significant expense or otherwise cause us to change the geographical location or segregation of our relevant systems and operations – any or all of which could adversely affect our operations or financial results. Additionally, other countries outside of Europe have enacted or are considering enacting similar cross-border data transfer restrictions and laws requiring local data residency, which could increase the cost and complexity of delivering our services and operating our business. The type of challenges we face in Europe will likely also arise in other jurisdictions that adopt laws similar in construction to the GDPR or regulatory frameworks of equivalent complexity.

European data protection laws also provide for robust regulatory enforcement and significant penalties for noncompliance, including, for example, under the GDPR, fines of up to €20 million (£17.5 million for the UK) or 4% of global annual revenue of any noncompliant organization for the preceding financial year, whichever is higher. In addition to administrative fines, a wide variety of other potential enforcement powers are available to competent supervisory authorities in respect of potential and suspected violations of the GDPR, including extensive audit and inspection rights, and powers to order temporary or permanent bans on all or some Processing of personal data carried out by noncompliant businesses – including permitting authorities to require destruction of improperly gathered or used personal data. European supervisory authorities

have shown a willingness to impose significant fines and issue orders preventing the processing of personal data on non-compliant businesses. The GDPR also confers a private right of action on data subjects and non-profit associations to lodge complaints with supervisory authorities, seek judicial remedies, and obtain compensation for damages resulting from violations of the GDPR.

Further, following the UK's departure from the EU, often referred to as Brexit, the data protection obligations of the EU GDPR continue to apply to UK-related Processing of personal data in substantially unvaried form under the UK GDPR by virtue of section 3 of the EU (Withdrawal) Act 2018, as amended.) With respect to international transfers, although the UK is regarded as a third country under the EU GDPR, the European Commission has issued an adequacy finding recognizing the UK as providing adequate protection under the EU GDPR and, therefore, transfers of personal data originating in the EEA to the UK remain unrestricted. Similarly, the UK government has confirmed that personal data transfers from the UK to the EEA remain free flowing.

However, going forward, there is increasing risk for divergence in application, interpretation and enforcement of the data protection laws as between the UK and EEA, creating additional regulatory uncertainty. For example, the UK Data (Use and Access) Act 2025, now in force, further differentiates the UK's data protection regime and could impact the adequacy finding granted to the United Kingdom by the EU Commission. An assessment by the European Commission is currently ongoing, with a decision expected by the end of 2025. These developments, along with the UK's evolving regulatory landscape, may affect our efforts to maintain a harmonized approach to processing European personal data and expose us to two parallel regimes where the UK GDPR and EU GDPR both apply with differing interpretation and enforcement approaches. This could increase our legal risk, uncertainty, complexity and compliance cost associated with the handling of European personal data, and may require us to adapt our privacy and data security compliance programs to account for legal and regulatory divergence between the UK and EEA.

If we do not designate a lead supervisory authority in an EEA member state, we are not able to benefit from the GDPR's 'one stop shop' mechanism. Amongst other things, this would mean that, in the event of a violation of the GDPR affecting data subjects across the EEA, we could be investigated by, and ultimately fined by the supervisory authority in each and every EEA member state where data subjects have been affected by such violation.

In the United States, there are a broad variety of data protection laws and regulations that may apply to our activities such as state data breach notification laws, state personal data privacy laws (for example, the California Consumer Privacy Act ("CCPA"), state health information privacy laws, and federal and state consumer protection laws. A range of enforcement agencies exist at both the state and federal levels that can enforce these laws and regulations. At the federal level, failing to take appropriate steps to keep consumers' personal information secure may constitute unfair acts or practices in or affecting commerce in violation of Section 5(a) of the Federal Trade Commission Act (the FTCA), 15 U.S.C § 45(a). The FTC expects a company's data security measures to be reasonable and appropriate in light of the sensitivity and volume of consumer information it holds, the size and complexity of its business and the cost of available tools to improve security and reduce vulnerabilities.

Regulators and legislators in the U.S. are also increasingly scrutinizing and restricting certain personal data transfers and transactions involving foreign countries. For example, the Biden Administration's executive order Preventing Access to Americans' Bulk Sensitive Personal Data and United States Government-Related Data by Countries of Concern as implemented by Department of Justice regulations issued in December 2024, which went into effect April 8, 2025, prohibits data brokerage transactions involving certain sensitive personal data categories, including health data, genetic data, and biospecimens, to countries of concern, including China. The regulations also restrict certain investment agreements, employment agreements and vendor agreements involving such data and countries of concern, absent specified cybersecurity controls. Actual or alleged violations of these regulations may be punishable by criminal and/or civil sanctions, and may result in exclusion from participation in federal and state programs.

In addition, certain state laws govern privacy and security of personal information. For example, the CCPA requires covered businesses that process personal information of California residents to disclose their data collection, use and sharing practices. Further, the CCPA provided California residents with new data privacy rights (including the ability to opt out of certain disclosures of personal data), imposed new operational requirements for covered businesses, provided for civil penalties for violations as well as a private right of action for data breaches and statutory damages (that is expected to increase data breach class action litigation and result in significant exposure to costly legal judgements and settlements). In addition, the CCPA was expanded on January 1, 2023, when the California Privacy Rights Act of 2020 ("CPRA") became operative. The amendments introduced by the CPRA, among other things, gave California residents the ability to limit use of certain sensitive personal information, further restricted the use of cross-contextual advertising, established restrictions on the retention of

personal information, expanded the types of data breaches subject to the CCPA's private right of action, provided for increased penalties for violations concerning California residents under the age of 16, established the California Privacy Protection Agency to implement and enforce the law. Although there are limited exemptions for clinical trial data and information subject to HIPAA under the CCPA, the CCPA and other similar laws could impact our business activities depending on how it is interpreted.

The CCPA marks the beginning of a trend toward more stringent privacy legislation in the United States, which could increase our potential liability and adversely affect our business. Already, in the United States, we have witnessed significant developments at the state level with numerous other states passing comprehensive privacy laws that incorporate many similar concepts of the CCPA. In addition, a number of other states have proposed new privacy laws, some of which are similar to the above discussed recently passed laws. Such proposed legislation, if enacted, may add additional complexity, variation in requirements, restrictions and potential legal risk, require additional investment of resources in compliance programs, impact strategies and the availability of previously useful data and could result in increased compliance costs and/or changes in business practices and policies. There are also states that are specifically regulating health information. For example, Washington state recently passed a health privacy law, effective March 31, 2024, that regulates the collection and sharing of health information, and the law also has a private right of action, which further increases the relevant compliance risk. Connecticut and Nevada have also passed similar laws regulating consumer health data. In addition, other states have proposed and/or passed legislation that regulates the privacy and/or security of certain specific types of information. For example, a small number of states have passed laws that specifically regulate biometric data. These various privacy and security laws may impact our business activities, including our identification of research subjects, relationships with business partners and ultimately the marketing and distribution of our products. State laws are changing rapidly and there have also been discussions in the U.S. Congress of new comprehensive federal data privacy laws to which we may likely become subject, if enacted.

In other foreign jurisdictions in which we operate or have operated (including sponsoring past, present or future clinical trials), such as, without limitation, Canada and Georgia, we may also be subject to stringent Data Protection Requirements. In Canada, for instance, Quebec's new comprehensive data protection law recently entered into force and is expected to have far-reaching effects.

Generally, these laws exemplify the vulnerability of our business to the evolving regulatory environment related to personal data and may require us to modify our Processing practices at substantial costs and expenses in an effort to comply.

Additionally, regulations promulgated pursuant to HIPAA, as amended, establish privacy and security standards that limit the use and disclosure of individually identifiable health information, or protected health information, and require the implementation of administrative, physical and technological safeguards designed to protect the privacy, confidentiality, integrity and availability of protected health information. These provisions may be applicable to our business or that of our collaborators, service providers, contractors or consultants.

Determining whether protected health information has been handled in compliance with applicable Data Protection Requirements can be complex and may be subject to changing interpretation. If we are unable to properly protect the privacy and security of protected health information, we could be found to have violated these privacy and security laws and/or breached certain contracts with our business partners (including as a business associate). Further, if we fail to comply with applicable Data Protection Requirements, such as, to the extent applicable, HIPAA privacy and security standards, we could face significant civil and criminal penalties. In the United States, the Department of Health and Human Services' and state attorneys general enforcement activity can result in financial liability and reputational harm, and responses to such enforcement activity can consume significant internal resources. In addition, state attorneys general are authorized to bring civil actions seeking either injunctions or damages in response to violations that threaten the privacy of state residents. We cannot be sure how these regulations will be interpreted, enforced or applied to our operations. In addition to the risks associated with enforcement activities and potential contractual liabilities, our ongoing efforts to comply with evolving laws and regulations at the federal and state level may be costly and require ongoing modifications to our policies, procedures and systems.

Given the breadth and evolving nature of Data Protection Requirements, preparing for and complying with these requirements is rigorous, time-intensive and requires significant resources and a review of our technologies, systems and practices, as well as those of any third-party collaborators, service providers, contractors or consultants that Process personal data on our behalf.

We may publish privacy policies and other documentation regarding our Processing of personal data and/or other confidential, proprietary or sensitive information. Although we endeavor to comply with our published policies and other documentation, we may at times fail to do so or may be perceived to have failed to do so. Moreover, despite our efforts, we may

not be successful in achieving compliance if our employees, third-party collaborators, service providers, contractors or consultants fail to comply with our policies and documentation. Such failures can subject us to potential foreign, local, state and federal action if they are found to be deceptive, unfair, or misrepresentative of our actual practices. Moreover, subjects about whom we or our partners obtain information, as well as the providers who share this information with us, may contractually limit our ability to use and disclose the information. Claims that we have violated individuals' privacy rights or failed to comply with data protection laws or applicable privacy notices even if we are not found liable, could be expensive and time-consuming to defend and could result in adverse publicity that could harm our business or otherwise materially and negatively impact our business.

***We are subject to U.S. and certain foreign export and import controls, sanctions, embargoes, anti-corruption laws, and anti-money laundering laws and regulations. Compliance with these legal standards could impair our ability to compete in domestic and international markets. We can face criminal liability and other serious consequences for violations, which can harm our business.***

We are subject to export control and import laws and regulations, including the U.S. Export Administration Regulations, U.S. Customs regulations, various economic and trade sanctions regulations administered by the U.S. Treasury Department's Office of Foreign Assets Controls, the U.S. Foreign Corrupt Practices Act of 1977, as amended, the U.S. domestic bribery statute contained in 18 U.S.C. § 201, the U.S. Travel Act, the USA PATRIOT Act, and other state and national anti-bribery and anti-money laundering laws in the countries in which we conduct activities. Anti-corruption laws are interpreted broadly and prohibit companies and their employees, agents, contractors, and other collaborators from authorizing, promising, offering, or providing, directly or indirectly, improper payments or anything else of value to recipients in the public or private sector. We may engage third parties to sell our products outside the United States, to conduct clinical trials, and/or to obtain necessary permits, licenses, patent registrations, and other regulatory approvals. We have direct or indirect interactions with officials and employees of government agencies or government-affiliated hospitals, universities, and other organizations. We can be held liable for the corrupt or other illegal activities of our employees, agents, contractors, and other collaborators, even if we do not explicitly authorize or have actual knowledge of such activities. Any violations of the laws and regulations described above may result in substantial civil and criminal fines and penalties, imprisonment, the loss of export or import privileges, debarment, tax reassessments, breach of contract and fraud litigation, reputational harm, and other consequences.

### **Risks Related to Ownership of Our Securities**

***We are an emerging growth company and a smaller reporting company, and we cannot be certain if the reduced reporting requirements applicable to emerging growth companies and smaller reporting companies will make our ADSs less attractive to investors.***

We are an emerging growth company, as defined in the Jumpstart Our Business Startups Act ("JOBS Act"), enacted in April 2012. For as long as we continue to be an emerging growth company, we may take advantage of exemptions from various 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 of Section 404 of the Sarbanes-Oxley Act of 2002, as amended ("Sarbanes-Oxley Act"), reduced disclosure obligations regarding executive compensation in our periodic reports and proxy statements, and exemptions from the requirements of holding nonbinding advisory votes on executive compensation and shareholder approval of any golden parachute payments not previously approved. We could be an emerging growth company for up to five years following 2021, the year in which we completed our initial public offering, although circumstances could cause us to lose that status earlier. We will remain an emerging growth company until the earlier of (1) the last day of the fiscal year (a) following the fifth anniversary of the closing of our initial public offering, (b) in which we have total annual gross revenue of at least \$1.235 billion or (c) in which we are deemed to be a large accelerated filer, which requires the market value of our ordinary shares that is held by non-affiliates to exceed \$700 million as of the prior June 30th after we have been subject to the SEC's periodic reporting requirements for at least twelve calendar months and have filed at least one annual report, and (2) the date on which we have issued more than \$1 billion in non-convertible debt during the prior three-year period.

Under the JOBS Act, emerging growth companies can also delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have elected to not "opt out" of this exemption from complying with new or revised accounting standards and, therefore, we will adopt new or revised accounting standards at the time private companies adopt the new or revised accounting 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.

On the last business day of our second quarter in 2025, the aggregate market value of our shares of common stock held by non-affiliates exceeded \$700 million. As a result, as of December 31, 2025, we will be considered a "large accelerated filer" as defined in Rule 12b-2 under the Exchange Act, and we will cease to be an "emerging growth company" as defined in the

JOBS Act and a “smaller reporting company” as defined in the Exchange Act. We will no longer be exempt from the auditor attestation requirements of Section 404(b) of the Sarbanes-Oxley Act, and our independent registered public accounting firm will evaluate and report on the effectiveness of internal control over financial reporting. We will also no longer be permitted to take advantage of reduced reporting requirements for smaller reporting companies.

We cannot predict if investors will find our ADSs less attractive because we may rely on these exemptions. If some investors find our ADSs less attractive as a result, there may be a less active trading market for our ADSs and our ADS price may be more volatile.

***Our articles of association provide that the courts of England and Wales will be the exclusive forum for the resolution of all shareholder complaints other than complaints asserting a cause of action arising under the Securities Act or the Exchange Act, and that the United States District Court for the Southern District of New York will be the exclusive forum for the resolution of any shareholder complaint asserting a cause of action arising under the Securities Act or the Exchange Act.***

Our articles of association provide that, unless we consent by ordinary resolution to the selection of an alternative forum, the courts of England and Wales shall, to the fullest extent permitted by law, be the exclusive forum for resolving: (a) any derivative action or proceeding brought on our behalf; (b) any action or proceeding asserting a claim of breach of fiduciary duty owed by any of our directors, officers or other employees to us; (c) any action or proceeding asserting a claim arising out of any provision of the Companies Act, or our articles of association (as may be amended from time to time); or (d) any action or proceeding asserting a claim or otherwise related to our affairs, or the England and Wales Forum Provision. The England and Wales Forum Provision will not apply to any causes of action arising under the Securities Act or the Exchange Act. Our articles of association further provide that unless we consent by ordinary resolution to the selection of an alternative forum, the United States District Court for the Southern District of New York shall be the exclusive forum for resolving any complaint asserting a cause of action arising under the Securities Act or the Exchange Act, or the U.S. Federal Forum Provision. In addition, our articles of association provide that any person or entity purchasing or otherwise acquiring any interest in our shares is deemed to have notice of and consented to the England and Wales Forum Provision and the U.S. Federal Forum Provision; provided, however, that our shareholders cannot and will not be deemed to have waived our compliance with the U.S. federal securities laws and the rules and regulations thereunder.

The England and Wales Forum Provision and the U.S. Federal Forum Provision in our articles of association may impose additional litigation costs on our shareholders in pursuing any such claims. Additionally, the forum selection clauses in our articles of association may limit the ability of our shareholders to bring a claim in a judicial forum that they find favorable for disputes with us or our directors, officers or employees, which may discourage the filing of lawsuits against us and our directors, officers and employees, even though an action, if successful, might benefit our shareholders. In addition, while the Delaware Supreme Court ruled in March 2020 that federal forum selection provisions purporting to require claims under the Securities Act be brought in federal court are “facially valid” under Delaware law, there is uncertainty as to whether other courts, including the courts of England and Wales and other courts within the U.S., will enforce our U.S. Federal Forum Provision. If the U.S. Federal Forum Provision is found to be inapplicable or unenforceable in an action, we may incur additional costs associated with resolving such action in other jurisdictions, which could adversely affect our results of operations and financial condition. The U.S. Federal Forum Provision may also impose additional litigation costs on our shareholders who assert that the provision is not enforceable or invalid. The courts of England and Wales and the United States District Court for the Southern District of New York may also reach different judgments or results than would other courts, including courts where a shareholder considering an action may be located or would otherwise choose to bring the action, and such judgments may be more or less favorable to us than our shareholders.

***The price of our ADSs may be volatile, and you could lose all or part of your investment.***

The trading price of our ADSs is likely to be highly volatile and could be subject to wide fluctuations in response to various factors, some of which are beyond our control, including limited trading volume. In addition to the factors discussed in this “Risk Factors” section, these factors include:

- the results of our ongoing, planned or any future preclinical studies, clinical trials or clinical development programs;
- the commencement, enrollment, or results of clinical trials of our product candidates or any future clinical trials we may conduct, or changes in the development status of our product candidates;
- adverse results or delays in preclinical studies and clinical trials;
- our decision to initiate a clinical trial, not to initiate a clinical trial, or to terminate an existing clinical trial;

- any delay in our regulatory filings or any adverse regulatory decisions, including failure to receive regulatory approval of our product candidates;
- changes in laws or regulations applicable to our products, including but not limited to clinical trial requirements for approvals;
- adverse developments concerning our manufacturers or our manufacturing plans;
- our inability to obtain adequate product supply for any licensed product or inability to do so at acceptable prices;
- our inability to establish collaborations if needed;
- our failure to commercialize our product candidates;
- changes in the structure of healthcare payment systems;
- additions or departures of key scientific or management personnel;
- unanticipated serious safety concerns related to the use of our product candidates;
- introduction of new products or services offered by us or our competitors;
- announcements of significant acquisitions, strategic partnerships, joint ventures or capital commitments by us or our competitors;
- our ability to effectively manage our growth;
- the size and growth of our initial cancer target markets;
- our ability to successfully treat additional types of cancers or at different stages;
- actual or anticipated variations in quarterly operating results;
- our cash position;
- our failure to meet the estimates and projections of the investment community or that we may otherwise provide to the public;
- publication of research reports about us or our industry, or immunotherapy in particular, or positive or negative recommendations or withdrawal of research coverage by securities analysts;
- changes in the market valuations of similar companies;
- overall performance of the equity markets;
- sales of our ADSs by us or holders of our ADSs in the future;
- trading volume of our ADSs;
- changes in accounting practices;
- ineffectiveness of our internal controls;
- disputes or other developments relating to intellectual property or proprietary rights, including patents, litigation matters and our ability to obtain patent protection for our technologies;
- significant lawsuits, including intellectual property or shareholder litigation;
- general political and economic conditions; and
- other events or factors, many of which are beyond our control.

In addition, the stock market in general, and the market for biopharmaceutical companies in particular, have experienced extreme price and volume fluctuations that have often been unrelated or disproportionate to the operating performance of these companies. Broad market and industry factors may negatively affect the market price of our ADSs, regardless of our actual operating performance. If the market price of our ADSs does not exceed the price at which you purchased them, you may not realize any return on your investment in us and may lose some or all of your investment. In the past, securities class action litigation has often been instituted against companies following periods of volatility in the market price of a company's securities. On September 28, 2022, the Company and certain of its current and former officers were named as defendants in a proposed class-action lawsuit. On August 27, 2024, the United States District Court for the Southern District of New York dismissed the plaintiff's complaint with prejudice and ordered the case closed.

***Sales of a substantial number of securities by shareholders in the public market could cause our ADS price to fall.***

If our shareholders sell, or indicate an intention to sell, substantial amounts of our ADSs in the public market after the lockup and other legal restrictions on resale lapse, the trading price of our ADSs could decline. For example, ordinary shares that are either subject to outstanding options or reserved for future issuance under equity incentive plans will become eligible for sale in the public market to the extent permitted by the provisions of various vesting schedules, the lock-up agreements and Rule 144 and Rule 701 under the Securities Act of 1933, as amended, or the Securities Act. If these additional ordinary shares are sold, or if it is perceived that they will be sold, in the public market, the trading price of our ADSs could decline.

As of June 30, 2025, the holders of 50,034,030 ordinary shares (or ordinary shares converted to ADSs) are entitled to rights with respect to the registration of their shares under the Securities Act. Registration of these shares under the Securities Act would result in such shares becoming freely tradable without restriction under the Securities Act, except for shares held by affiliates, as defined in Rule 144 under the Securities Act. Any sales of securities by these shareholders could have a material adverse effect on the trading price of our ADSs.

***We do not know whether an active, liquid and orderly trading market will develop for our ADSs or what the market price of our ADSs will be and, as a result, it may be difficult for you to sell your ADSs.***

Although our ADSs are listed on The Nasdaq Global Select Market, an active trading market for our ADSs may never develop or be sustained. You may not be able to sell your ADSs quickly or at the market price if trading in shares of our ADSs is not active. As a result of these and other factors, you may be unable to resell your ADSs at or above the price at which you purchased them. Further, an inactive market may also impair our ability to raise capital by selling additional ADSs and may impair our ability to enter into strategic partnerships or acquire companies or products by using our ADSs as consideration.

***If securities or industry analysts do not maintain research coverage of our company or publish inaccurate or unfavorable research about our business, the price of our ADSs and trading volume could decline.***

The trading market for our ADSs will depend in part on the research and reports that securities or industry analysts publish about us or our business. If one or more of the analysts who cover us downgrades our ADSs or publishes inaccurate or unfavorable research about our business, our ADS price may decline. If one or more of these analysts ceases coverage of our company or fails to publish reports on us regularly, demand for our ADSs could decrease, which might cause our ADS price and trading volume to decline.

***Our principal shareholders and management own a significant percentage of our voting shares and will be able to exert significant influence over matters subject to shareholders' approval.***

Our executive officers, directors, and 5.0% shareholders beneficially owned approximately 30.2% of our voting shares as of June 30, 2025. Therefore, these shareholders will have the ability to influence us through this ownership position. These shareholders may be able to determine matters requiring shareholder approval. For example, these shareholders may be able to control elections, re-elections and removal of directors, amendments of our articles of association, or approval of any merger, scheme of arrangement, sale of assets, or other major corporate transaction. This may prevent or discourage unsolicited acquisition proposals or offers for our ADSs that you may feel are in your best interest as a holder of our ADSs.

In addition, some of these persons or entities may have interests different than yours. For example, because many of these shareholders purchased their ordinary shares at prices substantially below the price at which you may have purchased our ADSs and have held their ordinary 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 shareholders.

***Future sales and issuances of our ADSs or rights to purchase ordinary shares, including pursuant to our equity incentive plans, could result in additional dilution of the percentage ownership of our shareholders and could cause the price of our ADSs to fall.***

We expect that significant additional capital may be needed in the future to continue our planned operations, including conducting clinical trials, expanded research and development activities, and costs associated with operating as a public company. To raise capital, we may sell ADSs, ordinary shares, convertible securities, or other equity securities in one or more transactions at prices and in a manner we determine from time to time. If we sell ADSs, ordinary shares, convertible securities, or other equity securities, investors may be materially diluted by subsequent sales, and new investors could gain rights, preferences, and privileges senior to the holders of our ADSs. Pursuant to our 2021 Plan, our management is authorized to grant share options to our employees, directors, and consultants. In September 2024, we filed a registration statement on Form

S-3ASR relating to the registration of our ordinary shares, each of which may be represented by one ADS; senior or subordinated debt securities; warrants to purchase any securities that may be sold under the prospectus; units or any combination thereof. In January 2023, we entered into an “at-the-market” offering program, which provides for the offering, issuance and sale by us of shares of our ordinary shares, represented by ADSs from time to time for aggregate gross proceeds of up to \$125.0 million in sales deemed to be “at-the-market offerings” as defined by the Securities Act of 1933, as amended. Any sale or issuance of securities pursuant to this registration statement or otherwise may result in dilution to our shareholders and may cause the market price of our ADSs to decline. Furthermore, new investors purchasing securities that we may issue and sell in the future could obtain rights superior to the rights of our existing shareholders.

As of June 30, 2025, the aggregate number of ordinary shares that may be issued pursuant to future share awards under the 2021 Plan is 11,665,913 ordinary shares. The number of ordinary shares reserved for issuance under the 2021 Plan shall be cumulatively increased on January 1, 2023 and each January 1 thereafter by up to 5.0% of the total number of ordinary shares outstanding on December 31 of the preceding calendar year or a lesser number of ordinary shares determined by our board of directors. Unless our board of directors elects not to increase the number of ordinary shares available for future grant each year, our shareholders may experience additional dilution, which could cause the price of our ADSs to fall.

***We have broad discretion in the use of our cash resources and may not use them effectively.***

Our management will have broad discretion in the application of our cash resources, and you will not have the opportunity as part of your investment decision to assess whether such resources are being used appropriately. Because of the number and variability of factors that will determine our use of our cash resources, their ultimate use may vary substantially from their currently intended use. Our management might not apply our cash resources in ways that ultimately increase or maintain the value of your investment. Pending their use, we may invest our cash resources in short-term, investment-grade, interest-bearing securities. These investments may not yield a favorable return to our shareholders.

***We do not intend to pay dividends on our ordinary shares, so any returns will be limited to the value of our ordinary shares or ADSs.***

We currently anticipate that we will retain future earnings for the development, operation, and expansion of our business and do not anticipate declaring or paying any cash dividends for the foreseeable future. In addition, we may enter into agreements that prohibit us from paying cash dividends without prior written consent from our contracting parties, or which other terms prohibiting or limiting the amount of dividends that may be declared or paid on our ADS. Furthermore, under the Companies Act, a company’s accumulated realized profits, so far as not previously utilized by distribution or capitalization, must exceed its accumulated realized losses so far as not previously written off in a reduction or reorganization of capital duly made (on a non-consolidated basis), before dividends can be paid. In the future, were our dividend policy to change, a dividend or distribution may still be restricted from being declared and paid. In addition, under the Companies Act, a public company may only effect a buyback of shares out of distributable profits or a fresh issue of shares and cannot do so out of capital. For these reasons, any return to shareholders may therefore be limited to the appreciation of their shares, which may never occur.

***As a public company, we may be at an increased risk of securities class action litigation, which is expensive and could divert management attention.***

The market price of our securities may be volatile and, in the past, companies that have experienced volatility in the market price of their stock have been subject to securities class action litigation. Historically, 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 biotechnology and pharmaceutical companies have experienced significant share price volatility in recent years. If we were to be sued, it could result in substantial costs and a diversion of management’s attention and resources, which could harm our business.

***We incur significant increased costs as a result of operating as a public company, and our management is required to devote substantial time to compliance initiatives.***

As a public company, we incur significant legal, accounting, and other expenses that we had not historically incurred as a private company. We are subject to the reporting requirements of the Securities Exchange Act of 1934, as amended, or the Exchange Act, which will require, among other things, that we file with the Securities and Exchange Commission (“SEC”), annual, quarterly, and current reports with respect to our business and financial condition. In addition, the Sarbanes-Oxley Act, as well as rules subsequently adopted by the SEC and The Nasdaq Global Select Market to implement provisions of the Sarbanes-Oxley Act, impose significant requirements on public companies, including requiring establishment and maintenance

of effective disclosure and financial controls and changes in corporate governance practices. Further, in July 2010, the Dodd-Frank Wall Street Reform and Consumer Protection Act (the “Dodd-Frank Act”), was enacted. There are significant corporate governance and executive compensation related provisions in the Dodd-Frank Act that require the SEC to adopt additional rules and regulations in these areas, such as “say on pay” and proxy access. Emerging growth companies are permitted to implement many of these requirements over a longer period and up to five years from the pricing of our initial public offering. We have taken advantage of this ; however, as of December 31, 2025, we will be considered a “large accelerated filer” as defined in Rule 12b-2 under the Exchange Act, and therefore we will cease to be a “smaller reporting company” and an “emerging growth company” as defined in the JOBS Act and will be required to implement these additional requirements, which could result in us incurring additional expenses. Shareholder activism, the current political environment, and government intervention and regulatory reform may lead to substantial new regulations and disclosure obligations, which may lead to additional compliance costs and impact the manner in which we operate our business in ways we cannot currently anticipate.

The rules and regulations applicable to public companies result in substantial legal and financial compliance costs and make some activities more time-consuming and costly than would otherwise be the case. As a result of the increased disclosure and compliance obligations we will become subject to as of December 31, 2025, including the requirement to obtain an auditor attestation of our internal control over financial reporting pursuant to Section 404(b) of the Sarbanes-Oxley Act of 2002, we will incur additional expenses in connection with compliance with these regulations and our management will need to devote additional time and effort to implement and comply with such requirements. If these requirements divert the attention of our management and personnel from other business concerns, they could have a material adverse effect on our business, financial condition, and results of operations. The increased costs will decrease our net income or increase our net loss and may require us to reduce costs in other areas of our business or increase the prices of our products or services. We cannot predict or estimate the amount or timing of additional costs we may incur to respond to these requirements. The impact of these requirements could also make it more difficult for us to attract and retain qualified persons to serve on our board of directors, our board committees, or as executive officers.

***We previously had material weaknesses in our internal control systems over financial reporting, which have been remediated; however we may identify additional or new material weaknesses in the future that may cause us to fail to meet our reporting obligations, result in material misstatements in our financial statements or fail to prevent fraud. We will need to continue to invest time and resources in the design, implementation and maintenance of internal controls.***

Our management is responsible for establishing and maintaining internal control over financial reporting, disclosure controls, and compliance with the other requirements of the Sarbanes-Oxley Act and the rules promulgated by the SEC thereunder. Internal control over financial reporting is a process designed to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements in accordance with international financial reporting standards. A material weakness is a deficiency, or combination of deficiencies, in internal control over financial reporting, such that there is a reasonable possibility that a material misstatement of the financial statements will not be prevented or detected on a timely basis.

We previously had material weaknesses identified related to inadequate controls that address segregation of certain accounting duties and reconciliation and analysis of certain key accounts. We concluded that these material weaknesses arose because, as a pre-revenue private company recently formed, neither we and nor our subsidiaries had the necessary personnel to design effective components of internal control including risk assessment control activities information/communication and monitoring to satisfy the accounting and financial reporting requirements of a public company.

Management remediated the material weaknesses described above through hiring additional qualified accounting and financial reporting personnel, and designing and implementing financial reporting systems, processes, policies and internal controls. However, management must continually evaluate the internal control environment and make enhancements to people, processes and systems which will require the investment of significant resources. For example, in March 2023, management implemented a company-wide ERP system to upgrade certain existing business, operational, and financial processes to enhance the overall control environment of the Company. There is no guarantee that new or additional material weaknesses will not be identified in the future. If material weaknesses arise in the future, our financial statements could contain material misstatements that, when discovered in the future, could cause us to fail to meet our future reporting obligations and cause the price of our ADSs to decline.

***If we fail to develop or maintain an effective system of disclosure controls and internal control over financial reporting, our ability to produce timely and accurate financial statements or comply with applicable regulations could be impaired.***

Effective internal controls over financial reporting are necessary for us to provide reliable financial reports and, together with adequate disclosure controls and procedures, are designed to prevent fraud. Any failure to implement required new or improved controls, or difficulties encountered in their implementation could cause us to fail to meet our reporting obligations. In addition, any testing by us conducted in connection with Section 404 of the Sarbanes-Oxley Act, or any subsequent testing by our independent registered public accounting firm, may reveal deficiencies in our internal controls over financial reporting that are deemed to be material weaknesses or that may require prospective or retroactive changes to our financial statements or identify other areas for further attention or improvement. Inferior internal controls could also cause investors to lose confidence in our reported financial information, which could have a negative effect on the trading price of our stock.

We are required to disclose changes made in our internal controls and procedures on a quarterly basis and our management is required to assess the effectiveness of these controls annually. An independent assessment of the effectiveness of our internal controls over financial reporting could detect problems that our management's assessment might not. Undetected material weaknesses in our internal controls over financial reporting could lead to restatements of our financial statements and require us to incur the expense of remediation.

As a public company, we are required to develop and maintain internal control over financial reporting and to report any material weaknesses in such internal controls. The Sarbanes-Oxley Act requires that we evaluate and determine the effectiveness of our internal control over financial reporting and provide a management report on internal control over financial reporting. In addition, once we are no longer an emerging growth company, as will be the case as of December 31, 2025, we will be required to include an attestation report on internal control over financial reporting issued by our independent registered public accounting firm beginning with our Annual Report on Form 10-K for the year ending December 31, 2025.

Any failure to develop or maintain effective controls, or any difficulties encountered in their implementation or improvement, could harm our results of operations, cause us to fail to meet our reporting obligations, result in a restatement of our financial statements for prior periods, or adversely affect the results of management evaluations and independent registered public accounting firm audits of our internal control over financial reporting that we will eventually be required to include in our periodic reports that will be filed with the SEC. In addition, to the extent we acquire or establish additional consolidated subsidiaries, the financial statements of such entities may not be initially prepared by us, and we will not have direct control over their financial statement preparation. As a result, we will, for our financial reporting, depend on what these entities report to us, which could result in our adding monitoring and audit processes, and increase the difficulty of implementing and maintaining adequate controls over our financial processes and reporting in the future, which could lead to delays in our external reporting. In particular, this may occur where we are establishing such entities with partners that do not have sophisticated financial accounting processes in place, or where we are entering into new relationships at a rapid pace, straining our integration capacity. Additionally, if we do not receive the information from the consolidated subsidiaries on a timely basis, it could cause delays in our external reporting. Ineffective disclosure controls and procedures and internal controls over financial reporting could also cause investors to lose confidence in our reported financial and other information, which would likely have a negative effect on the trading price of our ADSs.

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

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.

***Holder of ADSs are not treated as holders of our ordinary shares.***

By investing in our company, you are a holder of ADSs with underlying ordinary shares in a company incorporated under English law. Holders of ADSs are not treated as holders of our ordinary shares, unless they withdraw the ordinary shares underlying their ADSs in accordance with the deposit agreement and applicable laws and regulations. The depositary is the

holder of the ordinary shares underlying the ADSs. Holders of ADSs therefore do not have any rights as holders of our ordinary shares, other than the rights that they have pursuant to the deposit agreement.

***Holders of ADSs may be subject to limitations on the transfer of their ADSs and the withdrawal of the underlying ordinary shares.***

ADSs are transferable on the books of the depository. However, the depository may close its books at any time or from time to time when it deems expedient in connection with the performance of its duties. The depository may refuse to deliver, transfer or register transfers of ADSs generally when our books or the books of the depository are closed, or at any time if we or the depository think it is advisable to do so because of any requirement of law, government or governmental body, or under any provision of the deposit agreement, or for any other reason, subject to the right of ADS holders to cancel their ADSs and withdraw the underlying ordinary shares. Temporary delays in the cancellation of your ADSs and withdrawal of the underlying ordinary shares may arise because the depository has closed its transfer books or we have closed our transfer books, the transfer of ordinary shares is blocked to permit voting at a shareholders' meeting or we are paying a dividend on our ordinary shares. In addition, ADS holders may not be able to cancel their ADSs and withdraw the underlying ordinary shares when they owe money for fees, taxes and similar charges and when it is necessary to prohibit withdrawals in order to comply with any laws or governmental regulations that apply to ADSs or to the withdrawal of ordinary shares or other deposited securities.

***We are entitled to amend the deposit agreement and to change the rights of ADS holders under the terms of such agreement, or to terminate the deposit agreement, without the prior consent of the ADS holders.***

We are entitled to amend the deposit agreement and to change the rights of the ADS holders under the terms of such agreement, without the prior consent of the ADS holders. We and the depository may agree to amend the deposit agreement in any way we decide is necessary or advantageous to us or to the depository. Amendments may reflect, among other things, operational changes in the ADS program, legal developments affecting ADSs or changes in the terms of our business relationship with the depository. In the event that the terms of an amendment are materially disadvantageous to ADS holders, ADS holders will only receive 30 days' advance notice of the amendment, and no prior consent of the ADS holders is required under the deposit agreement. Furthermore, we may decide to direct the depository to terminate the ADS facility at any time for any reason. For example, terminations may occur when we decide to list our ordinary shares on a non-U.S. securities exchange and determine not to continue to sponsor an ADS facility or when we become the subject of a takeover or a going-private transaction. If the ADS facility will terminate, ADS holders will receive at least 30 days' prior notice, but no prior consent is required from them. Under the circumstances that we decide to make an amendment to the deposit agreement that is disadvantageous to ADS holders or terminate the deposit agreement, the ADS holders may choose to sell their ADSs or surrender their ADSs and become direct holders of the underlying ordinary shares, but will have no right to any compensation whatsoever.

***ADS holders may not be entitled to a jury trial with respect to claims arising under the deposit agreement, which could result in less favorable outcomes to the plaintiff(s) in any such action.***

The deposit agreement governing the ADSs representing our ordinary shares provides that, to the fullest extent permitted by law, holders and beneficial owners of ADSs irrevocably waive the right to a jury trial of any claim they may have against us or the depository arising out of or relating to the ADSs or the deposit agreement.

If this jury trial waiver provision is not permitted by applicable law, an action could proceed under the terms of the deposit agreement with a jury trial. If we or the depository opposed a jury trial demand based on the waiver, the court would determine whether the waiver was enforceable based on the facts and circumstances of that case in accordance with the applicable state and federal law. To our knowledge, the enforceability of a contractual pre-dispute jury trial waiver in connection with claims arising under the federal securities laws has not been finally adjudicated by the United States Supreme Court. However, we believe that a contractual pre-dispute jury trial waiver provision is generally enforceable, including under the laws of the State of New York, which govern the deposit agreement, by a federal or state court in the City of New York, which has non-exclusive jurisdiction over matters arising under the deposit agreement. In determining whether to enforce a contractual pre-dispute jury trial waiver provision, courts will generally consider whether a party knowingly, intelligently and voluntarily waived the right to a jury trial. We believe that this is the case with respect to the deposit agreement and the ADSs. It is advisable that you consult legal counsel regarding the jury waiver provision before entering into the deposit agreement.

If ADS holders or beneficial owners of ADSs bring a claim against us or the depository in connection with matters arising under the deposit agreement or the ADSs, including claims under federal securities laws, the ADS holder or beneficial owner may not be entitled to a jury trial with respect to such claims, which may have the effect of limiting and discouraging

lawsuits against us and/or the depository. If a lawsuit is brought against us and/or the depository under the deposit agreement, it may be heard only by a judge or justice of the applicable trial court, which would be conducted according to different civil procedures and may result in different outcomes than a trial by jury would have had, including results that could be less favorable to the plaintiff(s) in any such action, depending on, among other things, the nature of the claims, the judge or justice hearing such claims, and the venue of the hearing.

No condition, stipulation or provision of the deposit agreement or ADSs serves as a waiver by any holder or beneficial owner of ADSs or by us or the depository of compliance with the U.S. federal securities laws and the rules and regulations promulgated thereunder.

Moreover, as the jury trial waiver relates to claims arising out of or relating to the ADSs or the deposit agreement, we believe that, as a matter of construction of the clause, the waiver would likely to continue to apply to ADS holders who withdraw the ordinary shares from the ADS facility with respect to claims arising before the cancellation of the ADSs and the withdrawal of the ordinary shares, and the waiver would most likely not apply to ADS holders who subsequently withdraw the ordinary shares represented by ADSs from the ADS facility with respect to claims arising after the withdrawal. However, to our knowledge, there has been no case law on the applicability of the jury trial waiver to ADS holders who withdraw the ordinary shares represented by the ADSs from the ADS facility.

***ADS holders will not have the same voting rights as the holders of our ordinary shares and may not receive voting materials in time to be able to exercise their right to vote.***

Except as described in the deposit agreement, holders of the ADSs will not be able to exercise voting rights attaching to the ordinary shares represented by the ADSs. Under the terms of the deposit agreement, holders of the ADSs may instruct the depository to vote the ordinary shares underlying their ADSs. Otherwise, holders of ADSs will not be able to exercise their right to vote unless they withdraw the ordinary shares underlying their ADSs to vote them in person or by proxy in accordance with applicable laws and regulations and our articles of association. Even so, ADS holders may not know about a meeting far enough in advance to withdraw those ordinary shares. If we ask for the instructions of holders of the ADSs, the depository, upon timely notice from us, will notify ADS holders of the upcoming vote and arrange to deliver our voting materials to them. Upon our request, the depository will mail to holders a shareholder meeting notice that contains, among other things, a statement as to the manner in which voting instructions may be given. We cannot guarantee that ADS holders will receive the voting materials in time to ensure that they can instruct the depository to vote the ordinary shares underlying their ADSs. A shareholder is only entitled to participate in, and vote at, the meeting of shareholders, provided that it holds our ordinary shares as of the record date set for such meeting and otherwise complies with our articles of association. In addition, the depository's liability to ADS holders for failing to execute voting instructions or for the manner of executing voting instructions is limited by the deposit agreement. As a result, holders of ADSs may not be able to exercise their right to give voting instructions or to vote in person or by proxy and they may not have any recourse against the depository or us if their ordinary shares are not voted as they have requested or if their shares cannot be voted.

***ADS holders may not receive distributions on our ordinary shares represented by the ADSs or any value for them if it is illegal or impractical to make them available to holders of ADSs.***

The depository for the ADSs has agreed to pay to you the cash dividends or other distributions it or the custodian receives on our ordinary shares or other deposited securities after deducting its fees and expenses. You will receive these distributions in proportion to the number of our ordinary shares your ADSs represent. However, in accordance with the limitations set forth in the deposit agreement, it may be unlawful or impractical to make a distribution available to holders of ADSs. We have no obligation to take any other action to permit distribution on the ADSs, ordinary shares, rights or anything else to holders of the ADSs. This means that ADS holders may not receive the distributions we make on our ordinary shares or any value from them if it is unlawful or impractical to make them available. These restrictions may have an adverse effect on the value of your ADSs.

***Claims of U.S. civil liabilities may not be enforceable against us.***

We are incorporated under English law and have our registered office in England. Certain members of our board of directors and senior management are non-residents of the United States, and all or a substantial portion of our assets and the assets of such persons are located outside the United States. As a result, it may not be possible to serve process on such persons or us in the United States or to enforce judgments obtained in U.S. courts against them or us based on civil liability provisions of the securities laws of the United States.

The United States and England and Wales do not currently have a treaty providing for recognition and enforcement of judgments (other than arbitration awards) in civil and commercial matters. Consequently, a final judgment for payment given by a court in the United States, whether or not predicated solely upon U.S. securities laws, would not automatically be recognized or enforceable in England and Wales. In addition, uncertainty exists as to whether the courts of England and Wales would entertain original actions brought in England and Wales against us or our directors or senior management predicated upon the securities laws of the United States or any state in the United States. Any final and conclusive monetary judgment for a definite sum obtained against us in U.S. courts would be treated by the courts of England and Wales as a cause of action in itself and sued upon as a debt at common law so that no retrial of the issues would be necessary, provided that certain requirements are met. Whether these requirements are met in respect of a judgment based upon the civil liability provisions of the U.S. securities laws, including whether the award of monetary damages under such laws would constitute a penalty, is an issue for the court making such decision. If the courts of England and Wales give judgment for the sum payable under a U.S. judgment, the judgment of the English and Welsh court will be enforceable by methods generally available for this purpose. These methods generally permit the courts of England and Wales discretion to prescribe the manner of enforcement.

As a result, U.S. investors may not be able to enforce against us or our senior management, board of directors or certain experts named herein who are residents of England and Wales or countries other than the United States any judgments obtained in U.S. courts in civil and commercial matters, including judgments under the U.S. federal securities laws.

***ADS holders' right to participate in any future rights offerings may be limited, which may cause dilution to their holdings.***

We may from time to time distribute rights to our shareholders, including rights to acquire our securities. However, we cannot make rights available to ADS holders in the United States unless we register the rights and the securities to which the rights relate under the Securities Act or an exemption from the registration requirements is available. Also, under the deposit agreement, the depository bank will not make rights available to ADS holders unless either both the rights and any related securities are registered under the Securities Act, or the distribution of them to ADS holders is exempted from registration under the Securities Act. We are under no obligation to file a registration statement with respect to any such rights or securities or to endeavor to cause such a registration statement to be declared effective. Moreover, we may not be able to establish an exemption from registration under the Securities Act. If the depository does not distribute the rights, it may, under the deposit agreement, either sell them, if possible, or allow them to lapse. Accordingly, ADS holders may be unable to participate in our rights offerings and may experience dilution in your holdings.

***If we are a controlled foreign corporation, there could be material adverse U.S. federal income tax consequences to certain U.S. Holders.***

Each "Ten Percent Shareholder" (as defined below) in a non-U.S. corporation that is classified as a "controlled foreign corporation," or a CFC, for U.S. federal income tax purposes generally is required to include in income for U.S. federal tax purposes such Ten Percent Shareholder's pro rata share of the CFC's "Subpart F income" and investment of earnings in U.S. property, even if the CFC has made no distributions to its shareholders. Subpart F income generally includes dividends, interest, rents, royalties, "global intangible low-taxed income," gains from the sale of securities and income from certain transactions with related parties. In addition, a Ten Percent Shareholder that realizes gain from the sale or exchange of shares in a CFC may be required to classify a portion of such gain as dividend income rather than capital gain. A non-U.S. corporation generally will be classified as a CFC for U.S. federal income tax purposes if Ten Percent Shareholders own, directly or indirectly, more than 50% of either the total combined voting power of all classes of stock of such corporation entitled to vote or of the total value of the stock of such corporation. A "Ten Percent Shareholder" is a United States person (as defined by the Code) who owns or is considered to own 10% or more of the total combined voting power of all classes of stock entitled to vote or 10% or more of the total value of all classes of stock of such corporation.

We do not expect to be a CFC in the current taxable year; however, it is possible that we may become a CFC in a subsequent taxable year. The determination of CFC status is complex and includes attribution rules, the application of which is not certain. In addition, as a result of attribution rules in the Code applicable to taxable years prior to January 1, 2026, the stock of our non-U.S. subsidiaries is attributed to our U.S. subsidiary, which results in our non-U.S. subsidiaries being treated as CFCs and could result in certain United States persons being treated as Ten Percent Shareholders of such non-U.S. subsidiary CFCs. On July 4, 2025, the One Big Beautiful Bill Act (the "OBBA") was signed into law. As a result of the OBBA, certain rules applicable to Ten Percent Shareholders and CFCs will change for tax years beginning on or after January 1, 2026, including new rules applicable to "foreign controlled foreign corporations" and "foreign controlled U.S. shareholders" as defined under new Code Section 951B. We cannot provide any assurances that we will assist holders of our ordinary shares or ADSs in determining whether we are treated as a CFC or whether any holder of ordinary shares or ADSs is treated as a Ten

Percent Shareholder with respect to any such CFC or furnish to any Ten Percent Shareholders information that may be necessary to comply with the aforementioned reporting and tax paying obligations.

U.S. Holders should consult their own tax advisors with respect to the potential material adverse U.S. tax consequences of becoming a Ten Percent Shareholder in a CFC, including the possibility and consequences of becoming a Ten Percent Shareholder in our non-U.S. subsidiaries that are treated as CFCs due to the changes to the attribution rules. If we are classified as both a CFC and a PFIC (as defined below), we generally will not be treated as a PFIC with respect to those U.S. Holders that meet the definition of a Ten Percent Shareholder during the period in which we are a CFC.

***There is substantial uncertainty as to whether we are or will be a Passive Foreign Investment Company (“PFIC”). If we are a PFIC, there could be material adverse U.S. federal income tax consequences to U.S. holders.***

Under the Code, we will be a PFIC, for any taxable year in which (1) 75% or more of our gross income consists of passive income or (2) 50% or more of the average quarterly value of our assets consists of assets that produce, or are held for the production of, passive income. For purposes of these tests, passive income includes dividends, interest, gains from the sale or exchange of investment property and certain rents and royalties. In addition, for purposes of the above calculations, a non-U.S. corporation that directly or indirectly owns at least 25% by value of the shares of another corporation is treated as holding and receiving directly its proportionate share of assets and income of such corporation. If we are a PFIC for any taxable year during which a U.S. Holder holds our ordinary shares or ADSs, the U.S. Holder may be subject to material adverse tax consequences regardless of whether we continue to qualify as a PFIC, including ineligibility for any preferred tax rates on capital gains or on actual or deemed dividends, interest charges on certain taxes treated as deferred and additional reporting requirements.

While we believe we were not a PFIC for 2024, it is uncertain whether we or any of our Centessa Subsidiaries were, are, or will be treated as a PFIC for U.S. federal income tax purposes for any past, current or subsequent tax year. The determination of whether we are a PFIC is a fact-intensive determination made on an annual basis applying principles and methodologies that in some circumstances are unclear and subject to varying interpretation. The value of our assets would also be determined differently for the purposes of this determination if we were treated as a CFC, as discussed above. Under the income test described above, our status as a PFIC depends on the composition of our income which will depend on the transactions we enter into in the future and our corporate structure. The composition of our income and assets is also affected by the spending of the cash we raise in any offering, including in our initial public offering. Because PFIC status is based on our income, assets, and activities for the entire taxable year, our PFIC status may change from year to year. In addition, the total value of our assets for PFIC testing purposes may be determined in part by reference to the market price of our ordinary shares or ADSs from time to time, which may fluctuate considerably. Until we generate sufficient revenue from active licensing and other non-passive sources, there is a risk that we could be classified as a PFIC under the income test as our current operations generate limited amounts of non-passive income.

In certain circumstances, a U.S. Holder of shares in a PFIC may alleviate some of the adverse tax consequences described above by making either a “qualified electing fund” (“QEF”), election or a mark-to-market election (if our ordinary shares or ADSs constitute “marketable” securities under the Code). However, a U.S. Holder may make a QEF election with respect to our ordinary shares or ADSs only if we agree to furnish such U.S. Holder annually with required information. If we determine that we are a PFIC for this taxable year or any future taxable year, we currently expect that we would make available the information necessary for U.S. Holders to make a QEF Election. However, there is also no assurance that we will have timely knowledge of our status as a PFIC in the future or of the required information to be provided.

If we are a PFIC and, at any time, have a foreign subsidiary that is classified as a PFIC, U.S. Holders generally would be deemed to own a portion of the shares of such lower-tier PFIC, and generally could incur liability for the deferred tax and interest charge described above if we receive a distribution from, or dispose of all or part of our interest in, the lower-tier PFIC or the U.S. Holders otherwise were deemed to have disposed of an interest in the lower-tier PFIC. If we determine that we are a PFIC, to the extent appropriate, we currently expect that we will cause any lower-tier PFIC that we control to provide to a U.S. Holder the information necessary for U.S. Holders to make or maintain a QEF election with respect to the lower-tier PFIC. However, in the future, we may not hold a controlling interest in any such lower-tier PFIC and thus there can be no assurance that we will be able to cause the lower-tier PFIC to provide such required information. A mark-to-market election generally would not be available with respect to such lower-tier PFIC. U.S. Holders are urged to consult their tax advisors regarding the tax issues raised by lower-tier PFICs.

U.S. Holders should consult their own tax advisors with respect to the potential material adverse U.S. tax consequences if we or any of our Centessa Subsidiaries are or were to become a PFIC.

***Future changes to tax laws could materially adversely affect our company and reduce net returns to our shareholders.***

We conduct business globally. The tax treatment of the company or any of the group companies is subject to changes in tax laws, regulations and treaties, or the interpretation thereof, tax policy initiatives and reforms under consideration and the practices of tax authorities in jurisdictions in which we operate, as well as international tax policy initiatives and reforms including those related to the Organization for Economic Co-Operation and Development's ("OECD"), Base Erosion and Profit Shifting ("BEPS"), Project, the European Commission's state aid investigations and other initiatives. Such changes may include (but are not limited to) the taxation of operating income, investment income, dividends received or (in the specific context of withholding tax) dividends paid.

We are unable to predict what tax reform may be proposed or enacted in the future or what effect such changes would have on our business, but such changes, to the extent they are brought into tax legislation, regulations, policies or practices, could affect our financial position, future results of operations, cash flows in a particular period and overall or effective tax rates in the future in countries where we have operations, reduce post-tax returns to our shareholders, and increase the complexity, burden and cost of tax compliance.

***Tax authorities may disagree with our positions and conclusions regarding certain tax positions, resulting in unanticipated costs, taxes or non-realization of expected benefits.***

We operate through various Centessa Subsidiaries in the U.S. and UK. Consequently, we are subject to tax laws, treaties, and regulations in the countries in which we operate, and these laws and treaties are subject to interpretation. Our effective tax rate is influenced by many factors including changes in our operating structure, changes in the mix of our earnings among countries, our allocation of profits and losses among our subsidiaries, our intercompany transfer pricing agreements and rules relating to transfer pricing, the availability of U.S. research and development tax credits, and future changes in tax laws and regulations in the U.S. and foreign countries. Significant judgment is required in determining our tax liabilities including management's judgment for uncertain tax positions. We have taken, and will continue to take, tax positions based on our interpretation of such tax laws. A tax authority may disagree with tax positions that we have taken, which could result in increased tax liabilities. For example, HM Revenue & Customs ("HMRC"), the Internal Revenue Service or another tax authority could challenge our allocation of income by tax jurisdiction and the amounts paid between our affiliated companies pursuant to our intercompany arrangements and transfer pricing policies, including amounts paid with respect to our intellectual property development. There can be no assurance that a taxing authority will not have a different interpretation of applicable law and assess us with additional taxes. Similarly, a tax authority could assert that we are subject to tax in a jurisdiction where we believe we have not established a taxable connection, often referred to as a "permanent establishment" under international tax treaties, and such an assertion, if successful, could increase our expected tax liability in one or more jurisdictions. If we are assessed with additional taxes, this may result in a material adverse effect on our results of operations and/or financial condition.

A tax authority may take the position that material income tax liabilities, interest and penalties are payable by us, for example where there has been a technical violation of contradictory laws and regulations that are relatively new and have not been subject to extensive review or interpretation, in which case, we expect that we might contest such assessment. Contesting such an assessment may be lengthy and costly and if we were unsuccessful in disputing the assessment, the implications could increase our anticipated effective tax rate, where applicable, or result in other liabilities.

***We may be unable to use UK net operating loss and tax credit carryforwards and certain built-in losses to reduce future tax payments or benefit from favorable UK tax legislation.***

As a UK incorporated and tax resident entity, we are subject to UK corporate taxation on tax-adjusted trading profits. Due to the nature of our business, we have generated losses since inception and have not paid any UK corporation tax. We therefore have accumulated carryforward tax losses. Subject to numerous utilization criteria and restrictions (including those that limit the percentage of profits that can be reduced by carried forward losses and those that can restrict the use of carried forward losses where there is a change of ownership of more than half the ordinary shares of the Company and a major change in the nature, conduct or scale of the trade), we expect these to be eligible for carry forward and utilization against future operating profits. The use of loss carryforwards in relation to UK profits incurred on or after April 1, 2017 is generally limited each year to £5.0 million plus an incremental 50% of UK taxable profits. In addition, if we were to have a major change in the nature of the conduct of our trade, loss carryforwards may be restricted or extinguished.

As a company that carries out extensive research and development activities, we seek to benefit from the UK research and development ("R&D") tax relief program. For accounting periods prior to April 1, 2024 this consisted of two programs: the Small and Medium-sized Enterprises R&D tax relief program, or SME Program, and, to the extent that our projects are grant

funded or relate to work subcontracted to us by third parties, the Research and Development Expenditure Credit program, (“RDEC Program”). For accounting periods beginning on or after April 1, 2024 these two regimes have been merged into one program for all companies, R&D expenditure credit, (“RDEC”), alongside the introduction of a new enhanced R&D intensive support program, or ERIS, specifically for loss-making R&D intensive SMEs. In addition, unless limited exceptions apply, for accounting periods beginning on or after April 1, 2024, restrictions were introduced which apply to the tax relief that can be claimed for expenditure incurred on sub-contracted R&D activities or externally provided workers (where such sub-contracted activities are not carried out in the UK or such workers are not subject to UK payroll taxes). These changes, including the rate of deduction for qualifying R&D expenditures and activities for which relief may be claimed, may have a material impact on the quantum of R&D relief that we are able to claim in the future. Further, the UK R&D tax relief program’s rules are complex, and if a tax authority were to challenge or seek to disallow our claims (in whole or in part), for example by asserting that we do not (or the relevant expenditure does not) meet the technical conditions to be granted tax credits (or cash rebates), then such challenge or disallowance, if successful, could have a material impact on our cash-flow and financial performance. In addition, future changes to the UK R&D tax credit program may mean that we no longer qualify for it or have a material impact on the extent to which we can make claims (or benefit from them).

We may benefit in the future from the United Kingdom’s “patent box” regime, which allows certain profits attributable to revenues from patented products (and other qualifying income) to be taxed at an effective rate of 10%. We are the exclusive licensee or owner of several patent applications which, if granted, would cover our product candidates, and accordingly, future upfront fees, milestone fees, product revenues and royalties could be taxed at this lower tax rate. When taken in combination with the enhanced relief available on our research and development expenditures, we expect a long-term lower rate of corporation tax to apply to us. If, however, there are unexpected adverse changes to the UK research and development tax credit regime or the “patent box” regime, or for any reason we are unable to qualify for such advantageous tax legislation, or we are unable to use net operating loss and tax credit carryforwards and certain built-in losses to reduce future tax payments then our business, results of operations and financial condition may be adversely affected.

***Shareholder protections found in provisions under the UK City Code on Takeovers and Mergers, or the Takeover Code, will not apply if: (i) our place of central management and control remains outside of the United Kingdom (or the Channel Islands or the Isle of Man); and (ii) our securities are not admitted to trading on a UK-regulated market.***

We are classified as a “transition company” under the revised scope of the Takeover Code that became effective on February 3, 2025. As a result, certain provisions of the Takeover Code may apply to any takeover attempt we may be subject to from February 3, 2025 until February 2, 2027 (the “transition period”) if we either become UK quoted or UK resident at any time during the transition period and remain so at the time of any such transaction. We believe that our place of central management and control is not in the United Kingdom (or the Channel Islands or the Isle of Man) for the purposes of the jurisdictional criteria of the Takeover Code, and therefore we are not a UK resident for the purposes of the Takeover Code, and our securities are not admitted to trading on a UK-regulated market. Accordingly, we believe that any takeover attempt we may be subject to would not be subject to the Takeover Code and, as a result, our shareholders would not be entitled to the benefit of certain takeover offer protections provided under the Takeover Code, including the rules regarding mandatory takeover bids.

We will cease to be classified as a transition company at the end of the transition period and therefore from February 3, 2027 will no longer be subject to these transitional arrangements under the Takeover Code. Following the transition period, we will only be subject to the Takeover Code in the event that we become a UK quoted company. In the event that this changes, or if the interpretation and application of the Takeover Code by the Panel on Takeovers and Mergers (“Takeover Panel”), changes (including changes to the way in which the Takeover Panel assesses the application of the Takeover Code to English companies whose shares are listed outside of the United Kingdom), the Takeover Code may apply to us in the future.

The Takeover Code provides a framework within which takeovers of companies which are subject to the Takeover Code are regulated and conducted. The following is a brief summary of some of the most important rules of the Takeover Code:

- in connection with a potential offer, if following an approach by or on behalf of a potential bidder, the company is “the subject of rumor or speculation” or there is an “untoward movement” in the company’s share price, there is a requirement for the potential bidder to make a public announcement about a potential offer for the company, or for the company to make a public announcement about its review of a potential offer;
- when any person, or group of persons acting in concert, acquires, whether by a series of transactions over a period of time or not, an interest in shares which (taken together with shares already held by that person and an interest in shares held or acquired by persons acting in concert with him or her) carry 30% or more of the voting rights of a company that is subject to the Takeover Code, that person is generally required to make a mandatory offer to all

the holders of any class of equity share capital or other class of transferable securities carrying voting rights in that company to acquire the balance of their interests in the company;

- when any person who, together with persons acting in concert with him or her, is interested in shares representing not less than 30% but does not hold more than 50% of the voting rights of a company that is subject to the Takeover Code, and such person, or any person acting in concert with him or her, acquires an additional interest in shares which increases the percentage of shares carrying voting rights in which he or she is interested, then such person is generally required to make a mandatory offer to all the holders of any class of equity share capital or other class of transferable securities carrying voting rights of that company to acquire the balance of their interests in the company;
- a mandatory offer triggered in the circumstances described in the two paragraphs above must be in cash (or be accompanied by a cash alternative) and at not less than the highest price paid within the preceding 12 months to acquire any interest in shares in the company by the person required to make the offer or any person acting in concert with him or her;
- in relation to a voluntary offer (i.e. any offer which is not a mandatory offer), when interests in shares representing 10% or more of the shares of a class have been acquired for cash by an offeror (i.e., a bidder) and any person acting in concert with it in the offer period and the previous 12 months, the offer must be in cash or include a cash alternative for all shareholders of that class at not less than the highest price paid for any interest in shares of that class by the offeror and by any person acting in concert with it in that period. Further, if an offeror, or any person acting in concert with them, acquires for cash any interest in shares during the offer period, a cash alternative must be made available at not less than the highest price paid for any interest in the shares of that class;
- if, after making an offer for a company, the offeror or any person acting in concert with them acquires an interest in shares in an offeree company (i.e., a target) at a price higher than the value of the offer, the offer must be increased to not less than the highest price paid for the interest in shares so acquired;
- the offeree company must appoint a competent independent adviser whose advice on the financial terms of the offer must be made known to all the shareholders, together with the opinion of the board of directors of the offeree company;
- special or favorable deals for selected shareholders are not permitted, except in certain circumstances where independent shareholder approval is given and the arrangements are regarded as fair and reasonable in the opinion of the financial adviser to the offeree;
- all shareholders must be given the same information;
- each document published in connection with an offer by or on behalf of the offeror or offeree must state that the directors of the offeror or the offeree, as the case may be, accept responsibility for the information contained therein;
- profit forecasts, quantified financial benefits statements and asset valuations must be made to specified standards and must be reported on by professional advisers;
- misleading, inaccurate or unsubstantiated statements made in documents or to the media must be publicly corrected immediately;
- actions during the course of an offer by the offeree company, which might frustrate the offer are generally prohibited unless shareholders approve these plans. Frustrating actions would include, for example, lengthening the notice period for directors under their service contract or agreeing to sell off material parts of the target group;
- stringent requirements are laid down for the disclosure of dealings in relevant securities during an offer, including the prompt disclosure of positions and dealing in relevant securities by the parties to an offer and any person who is interested (directly or indirectly) in 1% or more of any class of relevant securities; and
- employees of both the offeror and the offeree company and the trustees of the offeree company's pension scheme must be informed about an offer. In addition, the offeree company's employee representatives and pension scheme trustees have the right to have a separate opinion on the effects of the offer on employment appended to the offeree board of directors' circular or published on a website.

***The rights of our shareholders may differ from the rights typically offered to shareholders of a U.S. corporation.***

We are incorporated under the laws of England and Wales. The rights of holders of ordinary shares and, therefore, certain of the rights of holders of ADS, are governed by English law, including the provisions of the Companies Act, and by our articles of association. These rights differ in certain respects from the rights of shareholders in typical U.S. corporations.

The principal differences include the following:

- under English law and our articles of association, each shareholder present at a meeting has only one vote unless demand is made for a vote on a poll, in which case each holder gets one vote per share owned. Under U.S. law, each shareholder typically is entitled to one vote per share at all meetings;
- under English law, it is only on a poll that the number of shares determines the number of votes a holder may cast. You should be aware, however, that the voting rights of ADS are also governed by the provisions of a deposit agreement with our depository bank;
- under English law, subject to certain exceptions and disapplications, each shareholder generally has preemptive rights to subscribe on a proportionate basis to any issuance of ordinary shares or rights to subscribe for, or to convert securities into, ordinary shares for cash. Under U.S. law, shareholders generally do not have preemptive rights unless specifically granted in the certificate of incorporation or otherwise;
- under English law and our articles of association, certain matters require the approval of 75% of the shareholders who vote (in person or by proxy) on the relevant resolution (or on a poll of shareholders representing 75% of the ordinary shares voting (in person or by proxy)), including amendments to the articles of association. This may make it more difficult for us to complete corporate transactions deemed advisable by our board of directors. Under U.S. law, generally only majority shareholder approval is required to amend the certificate of incorporation or to approve other significant transactions;
- in the United Kingdom, takeovers may be structured as takeover offers or as schemes of arrangement. Under English law, a bidder seeking to acquire us by means of a takeover offer would need to make an offer for all of our outstanding ordinary shares/ADS. If acceptances are not received for 90% or more of the ordinary shares/ADS under the offer, under English law, the bidder cannot complete a “squeeze out” to obtain 100% control of us. Accordingly, acceptances of 90% of our outstanding ordinary shares/ADSs will likely be a condition in any takeover offer to acquire us, not 50% as is more common in tender offers for corporations organized under Delaware law. By contrast, a scheme of arrangement, the successful completion of which would result in a bidder obtaining 100% control of us, requires the approval of a majority of shareholders voting at the meeting and representing 75% of the ordinary shares voting for approval; and
- under English law and our articles of association, shareholders and other persons whom we know or have reasonable cause to believe are, or have been, interested in our shares may be required to disclose information regarding their interests in our shares upon our request, and the failure to provide the required information could result in the loss or restriction of rights attaching to the shares, including prohibitions on certain transfers of the shares, withholding of dividends and loss of voting rights. Comparable provisions generally do not exist under U.S. law.

***As an English public limited company, certain capital structure decisions will require shareholder approval, which may limit our flexibility to manage our capital structure.***

English law provides that a board of directors may only allot shares (or grant rights to subscribe for or to convert any security into shares) with the prior authorization of shareholders, either pursuant to an ordinary resolution or as set out in the articles of association. This authorization must state the aggregate nominal amount of shares that it covers, can be valid up to a maximum period of five years and can be varied, renewed or revoked by shareholders. Such authority from our shareholders to allot additional shares for a period of five years from 2024 was included in the ordinary resolution passed by our shareholders on June 25, 2024, which authorization will need to be renewed upon expiration (i.e., at least every five years) but may be sought more frequently for additional five-year terms (or any shorter period).

English law also generally provides shareholders with preemptive rights when new shares are issued for cash. However, it is possible for the articles of association, or for shareholders to pass a special resolution at a general meeting, being a resolution passed by at least 75% of the votes cast, to disapply preemptive rights. Such a disapplication of preemptive rights may be for a maximum period of up to five years from the date of adoption of the articles of association, if the disapplication is contained in the articles of association, but not longer than the duration of the authority to allot shares to which this disapplication relates or from the date of the shareholder special resolution, if the disapplication is by shareholder special

resolution. In either case, this disapplication would need to be renewed by our shareholders upon its expiration (i.e., at least every five years). Such authority from our shareholders to disapply preemptive rights for a period of five years was included as a special resolution passed by our shareholders on June 25, 2024, which disapplication will need to be renewed upon expiration (i.e., at least every five years) to remain effective, but may be sought more frequently for additional five-year terms (or any shorter period).

English law also generally prohibits a public company from repurchasing its own shares without the prior approval of its shareholders by ordinary resolution, being a resolution passed by a simple majority of votes cast, and other formalities. Such approval may be provided for a maximum period of up to five years. In addition, a public company can only affect a buyback of shares out of distributable profits or a fresh issue of shares and cannot do so out of capital.

## **General Risk Factors**

***Business interruptions resulting from the Russia-Ukraine war, the Middle East conflict(s), tensions in U.S.-China relations, changes in trade policy, including the imposition of tariffs, or similar geo-political conflicts could cause a disruption to our business activities including the development of our product candidates and the conduct of clinical trials thereby adversely impacting our business.***

Geo-political conflicts including the Russia-Ukraine war, the Middle East conflict(s), tensions in U.S.-China relations and the impact of changes in trade policy, including the imposition of tariffs, may impact our CROs, clinical data management organizations, and clinical investigators' ability to conduct certain of our trials in the applicable countries, and may prevent us from obtaining data on patients already enrolled at sites in these countries. This could negatively impact the completion of our clinical trials and/or analyses of clinical results, which may increase our product development costs, elongate clinical trial timeframes and materially harm our business.

The global economy has been impacted by geopolitical tensions. The U.S. and other governments have imposed, and propose to impose additional, export controls and tariffs on certain products, and financial and economic sanctions on certain industry sectors and parties. These geopolitical tensions could result in, among other things, cyberattacks, supply chain disruptions, higher energy and other commodity costs, lower consumer demand, and changes to foreign exchange rates and financial markets, and tariffs and trade restrictions may result in increased production costs and product pricing, further supply chain disruptions, limited access to end markets, lower profitability, and uncertainty related to planning long-term investments and strategies, and may have other competitive effects. Any of the foregoing could have a material adverse effect on our business, financial condition, and results of operations.

***We or the third parties upon whom we depend may be adversely affected by earthquakes, outbreak of disease, or other natural disasters and our business continuity and disaster recovery plans may not adequately protect us from a serious disaster.***

Earthquakes, outbreak of disease, or other natural disasters could severely disrupt our operations, and have a material adverse effect on our business, results of operations, financial condition and prospects. If a natural disaster, power outage or other event occurred that prevented us from using all or a significant portion of our headquarters, that damaged critical infrastructure, such as the manufacturing facilities of our third-party CMOs, or that otherwise disrupted operations, it may be difficult or, in certain cases, impossible for us to continue our business for a substantial period of time. The disaster recovery and business continuity plans we have in place currently are limited and are unlikely to prove adequate in the event of a serious disaster or similar event. We may incur substantial expenses as a result of the limited nature of our disaster recovery and business continuity plans, which, particularly when taken together with our lack of earthquake insurance, could have a material adverse effect on our business.

***Unfavorable global economic conditions could adversely affect our business, financial condition or results of operations.***

Our results of operations could be adversely affected by general conditions in the global economy and in the global financial markets. For example, rising interest rates, high inflation and trade wars, including the imposition of tariffs, may cause our cost of doing business to materially increase and may adversely impact our ability to operate or may adversely impact other parties upon whom we rely for research and development capabilities to operate. The most recent global financial crisis caused extreme volatility and disruptions in the capital and credit markets. A severe or prolonged economic downturn, such as the most recent global financial crisis, could result in a variety of risks to our business, including weakened demand for our product candidates and our ability to raise additional capital when needed on acceptable terms, if at all. A weak or declining economy could strain our suppliers, possibly resulting in supply disruption, or cause delays in payments for our services by third-party

payors or our collaborators. Any of the foregoing could harm our business and we cannot anticipate all of the ways in which the current economic climate and financial market conditions could adversely impact our business.

**Item 2. Unregistered Sales of Equity Securities and Use of Proceeds.**

**(a) Recent Sales of Unregistered Equity Securities**

None.

**(b) Use of Proceeds from Initial Public Offering**

Not applicable.

**(c) Issuer Purchases of Equity Securities**

None.

**Item 3. Defaults Upon Senior Securities**

Not applicable.

**Item 4. Mine Safety Disclosures**

Not applicable.

**Item 5. Other Information**

***Rule 10b5-1 Trading Plans***

Our policy governing transactions in our securities by our directors, officers and employees permits our officers, directors and certain other persons to enter into trading plans complying with Rule 10b5-1 under the Exchange Act. Consistent with such regulation, our policy permits such plans to be entered into only when that person confirms they are not in possession of material non-public information. Our policy also requires a waiting period after a trading plan is created before shares can be traded under the plan. Our open trading windows are established in consultation with legal counsel. A number of our directors, officers and employees have entered into trading plans in accordance with Rule 10b5-1 and our policy governing transactions in our securities.

During the three months ended June 30, 2025, none of our directors or officers (as defined in Rule 16a-1(f) of the Securities Exchange Act of 1934, as amended) adopted, terminated or modified a Rule 10b5-1 trading arrangement or any “non-Rule 10b5-1 trading arrangement” (as defined in Item 408(c) of Regulation S-K).

**Item 6. Exhibits**

(a) Exhibits:

Exhibit number	Description of exhibit
3.1	<a href="#">Articles of Association of the registrant, as currently in effect (incorporated by reference to Exhibit 3.1 to the Registrant's Current Report on Form 8-K filed on October 6, 2022 (File No. 001-40445)).</a>
4.1	<a href="#">Form of Deposit Agreement (incorporated by reference to Exhibit 4.1 to the Registrant's Registration Statement on Form S-1 (File No. 333-255393)).</a>
4.2	<a href="#">Form of American Depositary Receipt (included in Exhibit 4.1) (incorporated by reference to Exhibit 4.2 to the Registrant's Registration Statement on Form S-1 (File No. 333-255393)).</a>
31.1	<a href="#">Certification of Principal Executive Officer Pursuant to Rules 13a-14(a) and 15d-14(a) under the Securities Exchange Act of 1934, as Adopted Pursuant to Section 302 of the Sarbanes-Oxley Act of 2002. (1)</a>
31.2	<a href="#">Certification of Principal Financial Officer Pursuant to Rules 13a-14(a) and 15d-14(a) under the Securities Exchange Act of 1934, as Adopted Pursuant to Section 302 of the Sarbanes-Oxley Act of 2002. (1)</a>
32.1*	<a href="#">Certification of Principal Executive Officer Pursuant to 18 U.S.C. Section 1350, as Adopted Pursuant to Section 906 of the Sarbanes-Oxley Act of 2002.</a>
32.2*	<a href="#">Certification of Principal Financial Officer Pursuant to 18 U.S.C. Section 1350, as Adopted Pursuant to Section 906 of the Sarbanes-Oxley Act of 2002.</a>
101 INS	Inline XBRL Instance Document - the instance document does not appear in the interactive data file because its XBRL tags are embedded within the inline XBRL document
101 SCH	Inline XBRL Taxonomy Extension Schema Document
101 CAL	Inline XBRL Taxonomy Extension Calculation Document
101 DEF	Inline XBRL Taxonomy Extension Definition Linkbase Document
101 LAB	Inline XBRL Taxonomy Extension Labels Linkbase Document
101 PRE	Inline XBRL Taxonomy Extension Presentation Link Document
104	Cover Page Interactive Data File - the cover page interactive data is embedded within the Inline XBRL document or included within Exhibit 101 attachments.

(1) Filed herewith

\* This certification will not be deemed "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), or otherwise subject to the liability of that section. Such certification will not be deemed to be incorporated by reference into any filing under the Securities Act of 1933, as amended, or the Exchange Act, except to the extent specifically incorporated by reference into such filing.

**SIGNATURES**

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned thereunto duly authorized.

**CENTESSA PHARMACEUTICALS PLC**

Date: August 12, 2025

By: /s/ Saurabh Saha, M.D., Ph.D.

Name: Saurabh Saha, M.D., Ph.D.

Title: *Chief Executive Officer (Principal Executive Officer)*

Date: August 12, 2025

By: /s/ John Crowley

Name: John Crowley

Title: *Chief Financial Officer (Principal Financial Officer)*





**CERTIFICATION PURSUANT TO  
18 U.S.C. SECTION 1350, AS ADOPTED PURSUANT TO  
SECTION 906 OF THE SARBANES-OXLEY ACT OF 2002**

In connection with the Quarterly Report of Centessa Pharmaceuticals plc (the "Company") on Form 10-Q for the period ended June 30, 2025 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I certify, pursuant to 18 U.S.C. § 1350, as adopted pursuant to § 906 of the Sarbanes-Oxley Act of 2002, that to the best of my knowledge:

- (1) The Report fully complies with the requirements of section 13(a) or 15(d) of the Securities Exchange Act of 1934, as amended; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and results of operations of the Company.

Date: August 12, 2025

By:

/s/ Saurabh Saha

**Saurabh Saha**

**Chief Executive Officer**

**Principal Executive Officer**

