

**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION  
WASHINGTON, DC 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 March 31, 2022

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: 001-39567

**C4 Therapeutics, Inc.**

(Exact Name of Registrant as Specified in its Charter)

**Delaware**  
(State or other jurisdiction of  
incorporation or organization)  
**490 Arsenal Way, Suite 200**  
**Watertown, MA**  
(Address of principal executive offices)

**47-5617627**  
(I.R.S. Employer  
Identification No.)

**02472**  
(Zip Code)

**Registrant's telephone number, including area code: (617) 231-0700**

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

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Common Stock, \$0.0001 par value per share	CCCC	The Nasdaq Global Select Market

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, 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 checked="" type="checkbox"/>	Accelerated filer	<input type="checkbox"/>
Non-accelerated filer	<input type="checkbox"/>	Smaller reporting company	<input type="checkbox"/>
		Emerging growth company	<input 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

As of April 25, 2022, the registrant had 48,797,226 shares of common stock, \$0.0001 par value per share, outstanding.

## SPECIAL NOTE REGARDING FORWARD-LOOKING STATEMENTS

This Quarterly Report on Form 10-Q, or Form 10-Q, including the section entitled “Management’s Discussion and Analysis of Financial Condition and Results of Operations,” contains express or implied forward-looking statements that are based on our management’s belief and assumptions and on information currently available to our management. Although we believe that the expectations reflected in these forward-looking statements are reasonable, these statements relate to future events or our future operational or financial performance, and involve known and unknown risks, uncertainties, and other 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 these forward-looking statements. Forward-looking statements in this Form 10-Q may include, but are not limited to, statements about:

- the initiation, timing, progress, results, safety and efficacy, and cost of our research and development programs and our current and future preclinical studies and clinical trials, including statements regarding the timing of initiation and completion of studies or trials, the period during which the results of the trials will become available, and our research and development programs;
  - the ultimate impact of the ongoing coronavirus, or COVID-19, pandemic, or any other health epidemic, on our business, manufacturing, clinical trials, research programs, supply chain, regulatory review, healthcare systems or the global economy as a whole;
  - risks related to the direct or indirect impact of the ongoing COVID-19 pandemic or any future large-scale adverse health event, such as the scope and duration of the pandemic, government actions and restrictive measures implemented in response, material delays in diagnoses, initiation or continuation of treatment for diseases that may be addressed by our development candidates and investigational medicines, or in patient enrollment in clinical trials, potential clinical trials, regulatory review or supply chain disruptions, and other potential impacts to our business, the effectiveness or timeliness of steps taken by us to mitigate the impact of the pandemic or other future large-scale adverse health event, and our ability to execute business continuity plans to address disruptions caused by the ongoing COVID-19 pandemic or future large-scale adverse health event;
  - our ability to obtain funding for our operations necessary to complete further development, manufacturing and commercialization of our product candidates;
  - our ability to obtain and maintain regulatory approval for any of our current or future product candidates;
  - the period of time over which we anticipate our existing cash and cash equivalents, and marketable securities will be sufficient to fund our operating expenses and capital expenditure requirements;
  - our ability to identify and develop product candidates for treatment of additional disease indications;
  - the potential attributes and benefits of our product candidates;
  - the rate and degree of market acceptance and clinical utility for any product candidates we may develop;
  - the pricing and reimbursement of our product candidates, if approved;
  - the effects of competition with respect to any of our current or future product candidates, as well as innovations by current and future competitors in our industry;
  - the implementation of our strategic plans for our business, any product candidates we may develop and our TORPEDO® platform;
  - the ability and willingness of our third-party strategic collaborators to continue research, development and manufacturing activities relating to our product candidates, including our ability to advance programs under our existing collaboration agreements with F. Hoffmann-La Roche Ltd and Hoffmann-La Roche Inc., or Roche, Biogen MA, Inc., or Biogen, and Calico Life Sciences LLC, or Calico, or other new collaboration agreements;
  - the scope of protection we are able to establish and maintain for intellectual property rights covering our product candidates;
  - estimates of our future expenses, revenues, capital requirements, and our needs for additional financing;
  - future agreements with third parties in connection with the manufacturing and commercialization of our product candidates, if approved;
  - the size and growth potential of the markets for our product candidates and our ability to serve those markets;
  - our financial performance;
  - regulatory developments in the United States and foreign countries;
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- our ability to contract with third-party suppliers and manufacturers and their ability to perform adequately;
- the success of competing therapies that are or may become available;
- our ability to attract and retain key scientific or management personnel;
- developments relating to our competitors and our industry; and
- other risks and uncertainties, including those discussed in Part II, Item 1A - Risk Factors in this Form 10-Q.

In some cases, forward-looking statements can be identified by terminology such as “will,” “may,” “should,” “could,” “expects,” “intends,” “plans,” “aims,” “anticipates,” “believes,” “estimates,” “predicts,” “potential,” “continue,” or the negative of these terms or other comparable terminology, although not all forward-looking statements contain these words. These statements are only predictions. You should not place undue reliance on forward-looking statements because they involve known and unknown risks, uncertainties, and other factors, which are, in some cases, beyond our control and which could materially affect results. Factors that may cause actual results to differ materially from current expectations include, among other things, those listed under the section titled “Risk Factors” and elsewhere in this Form 10-Q. If one or more of these risks or uncertainties occur, or if our underlying assumptions prove to be incorrect, actual events or results may vary significantly from those expressed or implied by the forward-looking statements. No forward-looking statement is a promise or a guarantee of future performance.

The forward-looking statements in this Form 10-Q represent our views as of the date of this Form 10-Q. We anticipate that subsequent events and developments will cause our views to change. However, while we may elect to update these forward-looking statements at some point in the future, we have no current intention of doing so except to the extent required by applicable law. You should therefore not rely on these forward-looking statements as representing our views as of any date subsequent to the date of this Form 10-Q.

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## SUMMARY OF RISK FACTORS

Our ability to implement our business strategy is subject to numerous risks that you should be aware of before making an investment decision. These risks are described more fully in Part II, Item 1A - Risk Factors in this Form 10-Q. These risks include, among others:

- We are a clinical-stage biopharmaceutical company with a limited operating history and have incurred significant losses since our inception. To date, we have not generated any revenue from product sales. We expect to continue to incur significant expenses and increasing operating losses for at least the next several years and may never achieve or maintain profitability. Our net loss was \$31.6 million and \$21.0 million for the three months ended March 31, 2022 and 2021, respectively.
  - We will need substantial additional funding to pursue our business objectives and continue our operations. If we are unable to raise capital when needed, we may be required to delay, limit, reduce or terminate our research or product development programs or future commercialization efforts.
  - Our approach to the discovery and development of product candidates based on our TORPEDO platform is unproven, which makes it difficult to predict the time, cost of development and likelihood of successfully developing any products.
  - Most of our product candidates are still in preclinical development and we have never completed a clinical trial of any of our product candidates. Our business could be harmed if we are unable to advance to clinical development, develop, obtain regulatory approval for and/or commercialize our product candidates, if we experience significant delays in doing any of these things, or if we experience significant cost increases.
  - We cannot be certain of the timely completion or outcome of our preclinical testing and clinical trials. In addition, the results of preclinical studies may not be predictive of the results of clinical trials and the results of any early-stage clinical trials we commence may not be predictive of the results of later-stage clinical trials.
  - Our preclinical studies and clinical trials may fail to demonstrate adequately the safety, potency and efficacy of any of our product candidates, which would prevent, delay, or require additional research or analysis to proceed with development, regulatory approval and commercialization of our current and future product candidates.
  - We have entered into collaboration agreements with Roche, Biogen and Calico, and may in the future seek to enter into collaborations with third parties for the development and/or commercialization of certain of our product candidates, but we may never realize the full potential benefits under these existing or potential collaboration arrangements.
  - The continuing effects of the ongoing COVID-19 pandemic, including the spread of new strains or variants of the virus, could adversely impact our business, including our preclinical studies and clinical trials.
  - We face substantial competition, which may result in others discovering, developing or commercializing products for the same indication and/or patient population before or more successfully than we do.
  - We rely, and expect to continue to rely, on third parties for the manufacture of our product candidates for preclinical and clinical testing, as well as for commercial manufacture if any of our product candidates receive marketing approval. This reliance on third parties may increase the risk that we will not have sufficient quantities of our product candidates in a timely manner, or at an acceptable cost or quality.
  - If we are unable to obtain required marketing approvals for, commercialize, manufacture, obtain and maintain patent protection for or gain market acceptance of our product candidates, or if we experience significant delays in doing so, our business will be materially harmed and our ability to generate revenue from product sales will be materially impaired.
  - If we are unable to obtain and maintain patent protection for our technology and products or if the scope of the patent protection obtained is not sufficiently broad or enforceable, our competitors could develop and commercialize technology, product candidates and products similar or identical to ours, and our ability to successfully commercialize our technology, product candidates and products may be impaired.
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#### **NOTE REGARDING COMPANY REFERENCES**

Unless the context otherwise requires, the terms “C4 Therapeutics,” “the Company,” “we,” “us,” and “our” in this Form 10-Q refer to C4 Therapeutics, Inc. and its consolidated subsidiary.

#### **NOTE REGARDING TRADEMARKS**

We own or have rights to various trademarks, service marks and trade names that are used in connection with the operation of our business, including our company name, C4 Therapeutics, our logo, the name of our TORPEDO technology platform and the names of our BIDAC and MONODAC protein degrader product candidates. This Form 10-Q may also contain trademarks, service marks and trade names of third parties, which are the property of their respective owners. Our use or display of third parties’ trademarks, service marks, trade names or products in this prospectus is not intended to and does not imply a relationship with, or endorsement or sponsorship by us. Solely for convenience, the trademarks, service marks and trade names referred to in this prospectus may appear without the ®, TM or SM symbols, but the omission of such references is not intended to indicate, in any way, that we will not assert, to the fullest extent under applicable law, our rights or the right of the applicable owner of these trademarks, service marks and trade names.

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## PART I—FINANCIAL INFORMATION

## Item 1. Financial Statements.

## C4 Therapeutics, Inc.

**Condensed Consolidated Balance Sheets**  
(in thousands, except share and per share amounts)  
(Unaudited)

	March 31, 2022	December 31, 2021
<b>Assets</b>		
Current assets:		
Cash and cash equivalents	\$ 46,004	\$ 76,124
Marketable securities, current	245,170	233,155
Accounts receivable	1,773	5,716
Prepaid expenses and other current assets	9,999	10,694
Total current assets	302,946	325,689
Marketable securities, non-current	130,478	142,200
Property and equipment, net	3,026	3,108
Right-of-use asset	74,828	31,945
Restricted cash	3,279	3,279
Other assets	928	544
Total assets	\$ 515,485	\$ 506,765
<b>Liabilities and Stockholders' Equity</b>		
Current liabilities:		
Accounts payable	\$ 3,405	\$ 4,506
Accrued expenses and other current liabilities	9,840	13,606
Deferred revenue, current	31,266	31,800
Operating lease liability, current	1,727	1,334
Long-term debt – related party, current	750	—
Total current liabilities	46,988	51,246
Deferred revenue, net of current	19,020	24,368
Operating lease liability, net of current	74,754	30,777
Long-term debt – related party, net of current	10,194	10,768
Total liabilities	150,956	117,159
Commitments and contingencies		
Stockholders' equity:		
Preferred stock, par value of \$0.0001 per share; 10,000,000 shares authorized, and no shares issued or outstanding as of March 31, 2022 and December 31, 2021, respectively	—	—
Common stock, par value of \$0.0001 per share; 150,000,000 shares authorized, and 48,751,490 and 48,688,875 shares issued and outstanding as of March 31, 2022 and December 31, 2021, respectively	5	5
Additional paid-in capital	667,509	658,091
Accumulated other comprehensive (loss) income	(3,650)	(775)
Accumulated deficit	(299,335)	(267,715)
Total stockholders' equity	364,529	389,606
Total liabilities and stockholders' equity	\$ 515,485	\$ 506,765

See accompanying notes to unaudited condensed consolidated financial statements.

C4 Therapeutics, Inc.

Condensed Consolidated Statements of Operations and Comprehensive Loss  
(in thousands, except share and per share amounts)  
(Unaudited)

	Three Months Ended March 31,	
	2022	2021
Revenue from collaboration agreements	\$ 7,654	\$ 7,426
Operating expenses:		
Research and development	26,203	20,526
General and administrative	12,820	7,409
Total operating expenses	39,023	27,935
Loss from operations	(31,369)	(20,509)
Other (expense) income, net:		
Interest expense and amortization of long-term debt – related party	(527)	(534)
Interest and other income, net	276	72
Total other (expense) income, net	(251)	(462)
Loss before income taxes	(31,620)	(20,971)
Income tax benefit	—	—
Net loss	\$ (31,620)	\$ (20,971)
Net loss per share attributable to common stockholders – basic and diluted	\$ (0.65)	\$ (0.49)
Weighted-average number of shares used in computed net loss per share – basic and diluted	\$ 48,734,827	\$ 43,084,978
Other comprehensive income (loss)		
Unrealized gain (loss) on marketable securities	(2,875)	(107)
Comprehensive loss	\$ (34,495)	\$ (21,078)

See accompanying notes to unaudited condensed consolidated financial statements.

C4 Therapeutics, Inc.

**Condensed Consolidated Statements of Stockholders' Equity (Deficit)**  
**(in thousands, except share amounts)**  
(Unaudited)

	Common Stock		Additional Paid-In Capital	Accumulated Other Comprehensive Income (Loss)	Accumulated Deficit	Total Stockholders' Equity
	Shares	Amount				
Balance as of December 31, 2021	48,688,875	\$ 5	\$ 658,091	(775)	\$ (267,715)	\$ 389,606
Exercise of stock options	52,707	—	260	—	—	260
Issuances of common stock under 2020 ESPP	8,028	—	220	—	—	220
Stock-based compensation	—	—	8,879	—	—	8,879
Unrealized loss on marketable securities	—	—	—	(2,875)	—	(2,875)
Net loss	—	—	—	—	(31,620)	(31,620)
Other	1,880	—	59	—	—	59
Balance as of March 31, 2022	<u>48,751,490</u>	<u>\$ 5</u>	<u>\$ 667,509</u>	<u>\$ (3,650)</u>	<u>\$ (299,335)</u>	<u>\$ 364,529</u>

See accompanying notes to unaudited condensed consolidated financial statements.

C4 Therapeutics, Inc.

Condensed Consolidated Statements of Stockholders' Equity (Deficit) - Continued  
(in thousands, except share amounts)  
(Unaudited)

	Common Stock		Additional Paid-In Capital	Accumulated Other Comprehensive Income (Loss)	Accumulated Deficit	Total Stockholders' Deficit
	Shares	Amount				
Balance as of December 31, 2020	43,059,632	\$ 4	\$ 464,597	\$ 13	\$ (183,823)	\$ 280,791
Exercise of stock options	49,328	—	166	—	—	166
Stock-based compensation	—	—	3,845	—	—	3,845
Unrealized loss on marketable securities	—	—	—	(107)	—	(107)
Net loss	—	—	—	—	(20,971)	(20,971)
Other	—	—	(19)	—	—	(19)
Balance as of March 31, 2021	43,108,960	\$ 4	\$ 468,589	\$ (94)	\$ (204,794)	\$ 263,705

See accompanying notes to unaudited condensed consolidated financial statements.

C4 Therapeutics, Inc.

Condensed Consolidated Statements of Cash Flows  
(in thousands)  
(Unaudited)

	Three Months Ended March 31,	
	2022	2021
Cash flows used in operating activities:		
Net loss	\$ (31,620)	\$ (20,971)
Adjustments to reconcile net loss to cash used in operating activities:		
Stock-based compensation expense	8,938	3,845
Depreciation expense	305	451
Reduction in carrying amount of right-of-use asset	1,184	320
Accretion of discount on marketable securities	664	102
Amortization of debt discount – related party	176	182
Changes in operating assets and liabilities:		
Accounts receivable	3,944	828
Prepaid expenses and other current and long-term assets	309	(40)
Accounts payable	(1,152)	(1,965)
Accrued expenses and other current liabilities	(3,766)	(3,666)
Operating lease liability	304	(251)
Deferred revenue	(5,882)	(3,769)
Net cash used in operating activities	(26,596)	(24,934)
Cash flows used in investing activities:		
Proceeds from maturities of marketable securities	72,387	114,994
Purchases of marketable securities	(76,219)	(176,303)
Purchases of property and equipment	(172)	(421)
Net cash used in investing activities	(4,004)	(61,730)
Cash flows provided by (used in) financing activities:		
Proceeds from exercises of stock options	260	166
Payment of initial public offering costs	—	(314)
Other	220	(3)
Net cash provided by (used in) financing activities	480	(151)
Net change in cash, cash equivalents and restricted cash	(30,120)	(86,815)
Cash, cash equivalents and restricted cash at beginning of period	79,403	184,304
Cash, cash equivalents and restricted cash at end of period	\$ 49,283	\$ 97,489
Reconciliation of cash, cash equivalents and restricted cash:		
Cash, cash equivalents and restricted cash at end of period	\$ 49,283	\$ 97,489
Less: restricted cash	(3,279)	(2,577)
Cash and cash equivalents at end of the period	\$ 46,004	\$ 94,912

C4 Therapeutics, Inc.

Condensed Consolidated Statements of Cash Flows - Continued  
(in thousands)  
(Unaudited)

	Three Months Ended March 31,	
	2022	2021
Supplemental disclosures of cash flow information:		
Cash paid for interest – related party	\$ 473	\$ 358
Supplemental disclosures of non-cash investing and financing activities:		
Operating lease liabilities arising from obtaining right-of-use assets	\$ 44,067	\$ —
Capital expenditures in accounts payable and accrued expenses	\$ 52	\$ 722

See accompanying notes to unaudited condensed consolidated financial statements.

## Notes to Condensed Consolidated Financial Statements

(Unaudited)

**Note 1. Nature of the business and basis of presentation**

C4 Therapeutics, Inc., or, together with its subsidiary, the Company, is a clinical-stage biopharmaceutical company dedicated to advancing targeted protein degradation science to develop a new generation of small-molecule medicines to transform how disease is treated. The Company leverages its proprietary technology platform, TORPEDO (Target ORiented ProtEIn Degradation Optimizer), to efficiently design and optimize small-molecule medicines that harness the body's natural protein recycling system to rapidly degrade disease-causing proteins, offering the potential to overcome drug resistance, drug undruggable targets and improve patient outcomes. The Company is advancing multiple targeted oncology programs to the clinic and expanding its research platform to deliver the next wave of medicines for difficult-to-treat diseases. The Company was incorporated in Delaware on October 7, 2015 and has its principal office in Watertown, Massachusetts.

***Liquidity and capital resources***

Since its inception, the Company's primary activities have been focused on research and development activities, building the Company's intellectual property, recruiting personnel and raising capital to support these activities. To date, the Company has funded its operations primarily with proceeds received from the sales of redeemable convertible preferred stock, public offerings of the Company's common stock, through its collaboration agreements, and debt financing.

The Company has incurred recurring losses since its inception, including net losses of \$31.6 million and \$21.0 million for the three months ended March 31, 2022 and 2021, respectively. In addition, as of March 31, 2022, the Company had an accumulated deficit of \$299.3 million. To date, the Company has not generated any revenue from product sales as none of its product candidates has been approved for commercialization. The Company expects to continue to generate operating losses for the foreseeable future.

The Company expects that its cash, cash equivalents and marketable securities of \$421.7 million as of March 31, 2022 will be sufficient to fund its operations for at least the next twelve months from the date of issuance of these condensed consolidated financial statements. Accordingly, the condensed consolidated financial statements have been prepared on a basis that assumes the Company will continue as a going concern and contemplates the realization of assets and satisfaction of liabilities and commitments in the ordinary course of business.

***Risks and uncertainties***

The Company is subject to risks common to other life science companies in the early development stage 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 and intellectual property, ability to raise additional financing and compliance with the Food and Drug Administration, or the FDA, and other government regulations. If the Company does not successfully advance its programs into and through human clinical trials and commercialize any of its product candidates either directly or through collaborations with other companies, the Company may be unable to produce product revenue or achieve profitability. There can be no assurance that the Company's research and development efforts will be successful, adequate protection for the Company's intellectual property will be obtained and maintained, any products developed will obtain necessary government regulatory approval, or any approved products will be commercially viable. Even if the Company's product development efforts are successful, it is uncertain when, if ever, the Company will generate significant revenue from product sales. The Company operates in an environment of rapid change in technology and substantial competition from pharmaceutical and biotechnology companies.

**Note 2. Summary of significant accounting policies*****Basis of presentation and consolidation***

The accompanying unaudited interim condensed consolidated financial statements have been prepared in accordance with generally accepted accounting principles in the United States, or U.S. GAAP, and applicable rules and regulations of the Securities and Exchange Commission, or the SEC, regarding interim financial reporting, and the instructions to Form 10-Q and Article 10 of Regulation S-X. Accordingly, they do not include all of the information and footnotes required by U.S. GAAP for complete financial statements. These condensed consolidated financial statements include the accounts of C4 Therapeutics, Inc. and its subsidiary, C4T Securities Corporation. All intercompany balances and transactions have been eliminated in consolidation.

***Unaudited interim financial information***

The accompanying condensed consolidated balance sheet as of March 31, 2022, the condensed consolidated statements of operations and comprehensive loss for the three months ended March 31, 2022 and 2021, the condensed consolidated statements of stockholders'

equity (deficit) for the three months ended March 31, 2022 and 2021, the condensed consolidated statements of cash flows for the three months ended March 31, 2022 and 2021, and the related interim disclosures are unaudited. These unaudited condensed consolidated financial statements include all adjustments necessary, consisting of only normal recurring adjustments, to fairly state the financial position and the results of the Company's operations and cash flows for interim periods in accordance with U.S. GAAP. Interim period results are not necessarily indicative of results of operations or cash flows for a full year or any subsequent interim period. The accompanying condensed consolidated financial statements should be read in conjunction with the Company's audited consolidated financial statements as of and for year ended December 31, 2021, and notes thereto, which are included in the Company's Annual Report on Form 10-K that was filed with the SEC on February 24, 2022.

### Reclassifications

Certain amounts that were previously reported have been reclassified to conform to current year presentation. The reclassifications had no effect on the reported results of operations.

### Use of estimates

The preparation of condensed 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 condensed consolidated financial statements and the reported amounts of revenues and expenses during the reporting period. The Company bases its estimates and assumptions on historical experience when available and on various factors that it believes to be reasonable under the circumstances. This process may result in actual results differing materially from those estimated amounts used in the preparation of the condensed consolidated financial statements if these results differ from historical experience or other assumptions do not turn out to be substantially accurate, even if such assumptions are reasonable when made. Significant estimates and assumptions reflected in these condensed consolidated financial statements include, but are not limited to, amounts and timing of revenues recognized under the Company's research and development collaboration arrangements, prepaid and accrued research and development expense, incremental borrowing rate used in the measurement of lease liability, and estimated volatility used in fair valuation of stock options. The Company assesses estimates on an ongoing basis; however, actual results could materially differ from those estimates.

### Significant accounting policies

The Company's significant accounting policies are disclosed in the audited consolidated financial statements for the year ended December 31, 2021, which are included in the Company's Annual Report on Form 10-K that was filed with the SEC on February 24, 2022. Since the date of those consolidated financial statements, there have been no material changes to the Company's significant accounting policies.

### Recently issued accounting standards

In March 2020, the Financial Accounting Standards Board, or FASB, issued Accounting Standards Update, or ASU, 2020-04, *Reference Rate Reform (Topic 848): Facilitation of the Effects of Reference Rate Reform on Financial Reporting*, which provides temporary optional guidance for a limited time to ease the potential accounting impacts associated with transitioning away from reference rates that are expected to be discontinued, such as the London Interbank Offered Rate, or LIBOR. Optional expedients in Topic 848 are generally available until December 31, 2022.

In January 2021, the FASB also issued ASU 2021-01, *Reference Rate Reform (Topic 848): Scope*, which extends some of Topic 848's optional expedients to derivative contracts modified as a result of rate reform, including certain derivatives that do not reference LIBOR or other reference rates that are expected to be discontinued. The amendments in this ASU affect the guidance in ASU 2020-04 and are effective in the same timeframe as ASU 2020-04. The adoptions of ASU 2020-04 and ASU 2021-01 did not have a material effect on the Company's condensed consolidated financial statements.

### Note 3. Fair value measurements

The following table presents information about the Company's financial assets measured at fair value on a recurring basis and indicates the level of the fair value hierarchy utilized to determine these fair values as of March 31, 2022 (in thousands):

	Fair Value	Level 1	Level 2	Level 3
Cash equivalents:				
Money market funds	\$ 40,248	\$ 40,248	\$ —	\$ —
Corporate debt securities	5,501	—	5,501	—
Marketable securities:				
Corporate debt securities	298,738	—	298,738	—
U.S. government debt securities	51,058	—	51,058	—
U.S. Treasury securities	25,852	—	25,852	—
Total cash equivalents and marketable securities	<u>\$ 421,397</u>	<u>\$ 40,248</u>	<u>\$ 381,149</u>	<u>\$ —</u>

There have been no transfers between fair value levels during the three months ended March 31, 2022.

The following table presents information about the Company's financial assets measured at fair value on a recurring basis and indicates the level of the fair value hierarchy utilized to determine these fair values as of December 31, 2021 (in thousands):

	Fair Value	Level 1	Level 2	Level 3
<b>Cash equivalents:</b>				
Money market funds	\$ 59,162	\$ 59,162	\$ —	\$ —
Corporate debt securities	11,649	—	11,649	—
U.S. Treasury securities	5,000	—	5,000	—
<b>Marketable securities:</b>				
Corporate debt securities	308,300	—	308,300	—
U.S. government debt securities	37,883	—	37,883	—
U.S. Treasury securities	29,172	—	29,172	—
<b>Total cash equivalents and marketable securities</b>	<b>\$ 451,166</b>	<b>\$ 59,162</b>	<b>\$ 392,004</b>	<b>\$ —</b>

The Company classifies its money market funds, which are valued based on quoted market prices in active markets, with no valuation adjustment, as Level 1 assets within the fair value hierarchy.

Marketable securities consist of U.S. Treasury securities, U.S. government debt securities, and corporate debt securities, all of which are classified as available-for-sale pursuant to Accounting Standards Codification, ASC, 320, *Investments – Debt and Equity Securities*. Marketable securities are classified within Level 2 of the fair value hierarchy because pricing inputs are other than quoted prices in active markets, which are either directly or indirectly observable as of the reporting date, and fair value is determined using models or other valuation methodologies on a recurring basis.

#### Note 4. Marketable securities

Marketable securities as of March 31, 2022 consisted of the following (in thousands):

	Amortized Cost	Gross Unrealized Gains	Gross Unrealized Losses	Fair Value
<b>Marketable securities, current:</b>				
Corporate debt securities	\$ 220,349	\$ —	\$ (1,040)	\$ 219,309
U.S. Treasury securities	16,043	—	(146)	15,897
U.S. government debt securities	10,000	—	(36)	9,964
<b>Marketable securities, non-current:</b>				
Corporate debt securities	80,816	—	(1,387)	79,429
U.S. government debt securities	41,970	—	(876)	41,094
U.S. Treasury securities	10,120	—	(165)	9,955
<b>Total marketable securities, current and non-current</b>	<b>\$ 379,298</b>	<b>\$ —</b>	<b>\$ (3,650)</b>	<b>\$ 375,648</b>

Marketable securities as of December 31, 2021 consisted of the following (in thousands):

	Amortized Cost	Gross Unrealized Gains	Gross Unrealized Losses	Fair Value
<b>Marketable securities, current:</b>				
Corporate debt securities	\$ 219,414	\$ 1	\$ (250)	\$ 219,165
U.S. government debt securities	5,006	—	—	5,006
U.S. Treasury securities	8,987	—	(3)	8,984
<b>Marketable securities, non-current:</b>				
Corporate debt securities	89,538	—	(403)	89,135
U.S. government debt securities	32,982	—	(105)	32,877
U.S. Treasury securities	20,203	—	(15)	20,188
<b>Total marketable securities, current</b>	<b>\$ 376,130</b>	<b>\$ 1</b>	<b>\$ (776)</b>	<b>\$ 375,355</b>

Marketable securities classified as current have maturities of less than one year. Marketable securities classified as non-current are those that: (i) have a maturity of greater than one year, and (ii) we do not intend to liquidate within the next twelve months, although these funds are available for use and, therefore, are classified as available-for-sale. No available-for-sale debt securities held as of March 31, 2022 or December 31, 2021 had remaining maturities greater than five years.

Marketable securities in unrealized loss positions as of March 31, 2022 consisted of the following (in thousands, except number of securities):

	Number of Securities	Fair Value	Gross Unrealized Losses
Marketable securities in continuous unrealized loss position for less than 12 months:			
Corporate debt securities	99	\$ 269,361	\$ (2,322)
U.S. government debt securities	11	51,058	(912)
U.S. Treasury securities	6	25,852	(311)
Marketable securities in continuous unrealized loss position for greater than 12 months:			
Corporate debt securities	15	26,881	(105)
<b>Total marketable securities in unrealized loss position</b>	<b>131</b>	<b>\$ 373,152</b>	<b>\$ (3,650)</b>

There were no individual securities that were in a significant unrealized loss position as of March 31, 2022, and, based on factors such as historical experience, market data, issuer-specific factors, and current economic conditions, the Company did not record an allowance for credit losses as of March 31, 2022 related to these securities. Further, given the lack of significant change in the credit risk, the Company does not consider these marketable securities to be impaired.

As of December 31, 2021, no marketable securities were in a continuous unrealized loss position for 12 months or longer.

#### Note 5. Property and equipment

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

	March 31, 2022	December 31, 2021
Property and equipment:		
Laboratory equipment	\$ 8,249	\$ 8,276
Furniture and fixtures	805	805
Leasehold improvements	548	541
Computer equipment	221	223
Office equipment	167	179
Construction in process	200	—
Total property and equipment	10,190	10,024
Less: accumulated depreciation	(7,164)	(6,916)
<b>Total property and equipment, net</b>	<b>\$ 3,026</b>	<b>\$ 3,108</b>

Depreciation expense related to property and equipment is as follows (in thousands):

	Three Months Ended March 31,	
	2022	2021
Depreciation expense	\$ 305	\$ 451

#### Note 6. Leases

In July 2017, the Company entered into a lease of office and laboratory space for its headquarters at 490 Arsenal Way, Suite 200 in Watertown, Massachusetts, or the Watertown Lease. In November 2021, the Company entered into an amendment to the Watertown Lease, or the Amended Lease. The Amended Lease serves to extend the lease term of the Company's existing leased space, or the Existing Leased Space, and provides additional office and laboratory space, or the Newly Leased Space. The lease for the Newly Leased Space commenced in January 2022 and the Company's obligation to pay rent on the Newly Leased Space commenced in March 2022, with the amount of this new rent obligation added to the Company's continuing obligation to pay rent on the Existing Leased Space. The Amended Lease terminates in March 2032, which is 10 years from the rent commencement date for the Newly Leased Space. The Amended Lease is subject to fixed rate rent escalations, provides for up to \$2.6 million in tenant improvements reimbursable to the Company, and provides an option for the Company to extend the lease term of the Amended Lease for one additional five-year period. Upon executing the Amended Lease, the Company increased the amount of its existing collateral to \$3.3 million, which is recorded as restricted cash on the accompanying condensed consolidated balance sheets as of March 31, 2022. In addition to rent, under the terms of the Amended Lease, the Company is also responsible for paying its pro rata share of costs incurred for common area maintenance, real estate taxes and property insurance related to the leased space, including both the Existing Leased Space and, as of January 2022, the Newly Leased Space, which amounts are accounted for as variable lease costs.

### Accounting for the Amended Lease

As the Amended Lease extends the term of the Existing Leased Space and provides access to the Newly Leased Space, in accordance with the provisions of ASC 842, the Company accounted for the Amended Lease as two separate contracts: 1) modification of the existing lease agreement to extend the lease term of the Existing Leased Space, and 2) new lease agreement for the right-of-use of the Newly Leased Space.

As noted above, the lease for the Newly Leased Space commenced in January 2022. As a result, the Company recorded a right-of-use asset of \$44.4 million, and a corresponding lease liability of \$44.1 million for the Newly Leased Space on the accompanying condensed consolidated balance sheets. The calculation of the lease liability and the right-of-use asset of the Newly Leased Space does not include the additional five-year period option provided under the Amended Lease as the Company does not believe there is reasonable certainty that the option will be exercised. As stated above, the Amended Lease provides for up to \$2.6 million of tenant improvement allowance. As of March 31, 2022, no tenant improvement allowances have been recorded.

The elements of lease costs were as follows (in thousands):

	Three Months Ended March 31,	
	2022	2021
Lease cost:		
Operating lease cost	\$ 2,508	\$ 637
Variable lease cost	348	241
Total lease cost	<u>\$ 2,856</u>	<u>\$ 878</u>

The following table summarizes the lease term and discount rate applied in arriving at the lease liability:

	March 31, 2022	December 31, 2021
Remaining lease term	9.9 years	10.2 years
Discount rate	5.3%	5.2%

Future lease payments under non-cancelable leases as of March 31, 2022 for each of the years ending December 31 are as follows (in thousands):

Undiscounted lease payments:		
Remaining 2022	\$	6,263
2023		8,571
2024		8,828
2025		9,093
2026		9,366
Thereafter		<u>61,759</u>
Total undiscounted lease payments		103,880
Less: imputed interest		(24,757)
Less: tenant improvement allowance		(2,642)
Total operating lease liability	<u>\$</u>	<u>76,481</u>

The Company expects to receive the \$2.6 million of tenant improvement allowance from the landlord in 2022.

### Note 7. Accrued expenses and other current liabilities

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

	March 31, 2022	December 31, 2021
Accrued expenses and other current liabilities:		
Accrued research and development	\$ 6,698	\$ 6,863
Accrued compensation and benefits	1,843	5,084
Accrued professional fees	990	1,246
Other	309	413
Total accrued expenses and other current liabilities	<u>\$ 9,840</u>	<u>\$ 13,606</u>

## Note 8. Collaboration and license agreements

### *Roche Collaboration and License Agreement*

In March 2016, the Company entered into a license agreement with Roche, which was amended in June 2016 and amended further in March 2017. The Company amended and restated that agreement (as so amended) in December 2018. This amended and restated agreement is referred to as the Roche Agreement. Under the Roche Agreement, the Company and Roche agreed to collaborate in the research, development, manufacture and commercialization of target-binding degrader medicines using the Company's proprietary TORPEDO platform for the treatment of cancers and other indications. In November 2020, the Company signed a further amendment, which provided that the parties would develop up to five potential targets, with Roche maintaining its option rights to license and commercialize products directed to those targets. The November 2020 amendment also provides a mechanism through which the Company and Roche can mutually agree to terminate the Roche Agreement on a target-by-target basis by the entry into a mutual target termination agreement.

Upon signing the amendment in November 2020, the Company received an additional upfront consideration of \$40.0 million from Roche. In addition, the Company receives annual research plan payments of \$1.0 million for up to three years for each active research plan. For certain targets, Roche is required to pay the Company fees of \$2.0 million and \$3.0 million upon the progression of targets to the lead series identification achievement and good laboratory practice toxicology study phase, respectively. If Roche exercises its option right for a target, Roche is obligated to pay exercise fees ranging from \$7.0 million to \$20.0 million depending on the target. For each target option exercised by Roche, the Company is eligible to receive milestone payments ranging from \$260 million to \$275.0 million upon the achievement of certain research, development and commercial milestones with respect to corresponding products, subject to certain reductions and exclusions based on intellectual property coverage. Roche is also required to pay the Company up to \$150.0 million per target in one-time sales-based milestone payments upon the achievement of specified levels of net sales of a product directed to such target. Finally, Roche is required to pay the Company tiered royalties ranging from the mid-single digits to mid-teen percentages on net sales of products sold by Roche pursuant to its exercise of its option rights, subject to certain reductions. For sales of products for which the Company exercises its co-development right, the applicable royalty rates will be increased by a low-single digit percentage.

The collaboration is managed by a joint research committee. The Company has control over the joint research committee prior to Roche's exercise of its option rights as to a particular target, with Roche assuming control of the joint research committee thereafter, and may terminate the Roche Agreement on a target-by-target or product-by-product basis under several scenarios, upon at least 90 days' prior written notice.

### *Roche Agreement accounting*

At commencement, the Company identified twelve performance obligations within the Roche Agreement, represented by the six potential research and development targets and the option rights held by Roche for each of the six targets. A non-exclusive royalty-free license to use the Company's intellectual property to conduct research and development activities and participation on joint research committee were identified as promised services. However, the Company determined that the research and development license and research and development services were not distinct from one another, and participation on the joint research committee was determined to be quantitatively and qualitatively immaterial.

As of March 31, 2022, the total transaction price of the Roche Agreement is allocated to the performance obligations based on their relative standalone selling price.

The allocated transaction price is recognized as revenue from collaboration agreements in one of two ways:

- Research and development targets: The Company recognizes the portion of the transaction price allocated to each of the research and development performance obligations as the research and development services are provided, using an input method, in proportion to costs incurred to date for each research development target as compared to total costs incurred and expected to be incurred in the future to satisfy the underlying obligation related to said research and development target. The transfer of control occurs over this period and, in management's judgment, is the best measure of progress towards satisfying the performance obligation.
- Option rights: The transaction price allocated to the options rights, which are considered material rights, is recognized in the period that Roche elects to exercise or elects to not exercise its option right to license and commercialize the underlying research and development target.

The following table summarizes the allocation of the total transaction price to the identified performance obligations under the Roche Agreement, and the amount of the transaction price unsatisfied as of March 31, 2022 (in thousands):

	Transaction Price Allocated	Transaction Price Unsatisfied
Performance obligations:		
Research and development targets	\$ 58,851	\$ 26,109
Option rights	6,721	3,502
Total	<u>\$ 65,572</u>	<u>\$ 29,611</u>

Amounts due to the Company that have not yet been received are recorded as accounts receivable and amounts received that have not yet been recognized as revenue from collaboration agreements are recorded in deferred revenue on the Company's condensed consolidated balance sheet.

#### ***Biogen Collaboration Research and License Agreement***

In December 2018, the Company entered into a collaboration research and license agreement, or the Biogen Agreement, with Biogen MA, Inc., or Biogen, which was amended in February 2020. Pursuant to the terms of the Biogen Agreement, the Company and Biogen agreed to collaborate on research activities to develop novel treatments in the field of target protein degradation, or TPD, using the Company's degrader technology. Under the terms of the Biogen Agreement, the Company will initially develop TPD therapeutics that utilize degrader technology for up to five target proteins over a period of 54 months, ending in June 2023. On a target-by-target basis, after successful completion of a defined target evaluation period, Biogen assumes full rights and responsibility to each degrader to meet certain criteria against a target. Biogen also has the option to pay an additional \$62.5 million to extend the agreement for four additional years and select up to five additional targets for development.

In exchange for the non-exclusive research license from Biogen, as well as a \$45.0 million nonrefundable upfront payment, the Company has granted a license to develop, commercialize and manufacture products related to each of the targets (which is contingent on not cancelling the agreement), performs initial research services for drug discovery, has provided a non-exclusive research and commercial license to its intellectual property and participates on the joint steering committee, or the Biogen JSC. The Company is also obligated to participate in early research activities for other potential targets or sandbox activities, at Biogen's election up to a maximum amount; any work performed for these services is reimbursed by Biogen, and Biogen reimburses the Company for certain full-time equivalent, or FTE, costs. The Company's obligations under the sandbox activities were completed as of August 31, 2021. Biogen is also required to pay the Company up to \$35.0 million per target in development milestones and \$26.0 million per target in one-time sales-based payments for the first product to achieve certain levels of net sales. In addition, Biogen is required to pay the Company royalties on a licensed product-by-licensed product basis, on worldwide net product sales.

The collaboration is managed by the Biogen JSC, which Biogen has control over, and Biogen may terminate the Biogen Agreement on a target-by-target or product-by-product basis under several scenarios, upon at least 90 days' prior written notice.

#### ***Biogen Agreement accounting***

The Company recognizes revenue from collaboration agreements under the Biogen Agreement from two types of services: 1) research and development services, and 2) sandbox activities, which are discovery-type research services.

- **Research and development services:** The Company identified one performance obligation at the outset of the Biogen Agreement, representing a combined performance obligation consisting of (1) the licenses, (2) the research activities for the target evaluation phase for all five targets and (3) the joint research plan phase for each target. The Company determined that the licenses and research activities were not distinct from one another, as the licenses have limited value without the performance of the research activities by the Company. Participation on the Biogen JSC to oversee the research activities and the technology transfer associated with the Biogen License Agreement were determined to be quantitatively and qualitatively immaterial and therefore are excluded from performance obligations. The Company recognizes the transaction price allocated to this performance obligation as the research and development services are provided, using an input method, in proportion to costs incurred to date for each research development target as compared to total costs incurred and expected to be incurred in the future to satisfy the underlying obligation related to said research and development target. The transfer of control occurs over this period and, in management's judgment, is the best measure of progress towards satisfying the performance obligation.
- **Sandbox activities:** Biogen had the option to fund sandbox activities in exchange for consideration at market rates, whereby the Company would perform discovery-type research at Biogen's election to develop other potential targets that may be used as replacement targets for the initially nominated targets or two additional targets under the Biogen Agreement. Revenues earned under this option were recognized as services were performed and were not included in the transaction price allocated to the performance obligation described above. The Company recognized FTE reimbursement received for sandbox activities as revenue as incurred each quarter. As noted above, sandbox activities fully concluded on August 31, 2021.

As of March 31, 2022, total transaction price of the Biogen Agreement of \$55.0 million is allocated to the research and development services performance obligation and \$25.9 million of the allocated transaction price remains unsatisfied.

Amounts due to the Company that have not yet been received are recorded as accounts receivable and amounts received that have not yet been recognized as revenue from collaboration agreements are recorded in deferred revenue on the Company's condensed consolidated balance sheet.

### ***Calico Collaboration and License Agreement***

In March 2017, the Company entered into a collaboration and license agreement, or the Calico Agreement, with Calico Life Sciences LLC, or Calico, whereby the Company and Calico agreed to collaborate to develop and commercialize small molecule protein degraders for diseases of aging, including cancer, for a five-year period ending in March 2022. As further described below, in September 2021, the Company and Calico agreed to extend the research term for a certain program to March 2023.

Under the terms of the Calico Agreement, the Company will initially develop and commercialize small molecule protein degraders for up to five target proteins over the research term. On a target-by-target basis, after successful completion of a defined target evaluation period, Calico has an exclusive option to pursue further preclinical development and commercialization via a joint research plan for each target.

Under the Calico Agreement, Calico paid an upfront amount of \$5.0 million and certain annual payments totaling \$5.0 million through June 30, 2020 and pays target initiation fees and reimburses the Company for a number of FTEs, depending on the stage of the research, at specified market rates. Upon completion of the required discovery research and development services on any target, Calico is entitled to pursue commercial development of products related to that target. The Company will perform initial research services for drug discovery and preclinical development, provide a non-exclusive research and commercial license to its IP and will participate on the Calico joint research committee, or the Calico JRC. For each target, the Company is eligible to receive up to \$132.0 million in potential research, development and commercial milestone payments, on sales of all products resulting from the collaboration efforts. Calico is also required to pay the Company up to \$65.0 million in one-time sales-based payments for the first product to achieve certain levels of net sales. In addition, Calico is required to pay the Company royalties, at percentages in the mid-single digits, on a licensed product-by-licensed product basis, on worldwide net product sales.

The Calico Agreement is managed by the Calico JRC. Calico has control over the Calico JRC and may terminate the Calico Agreement on a target-by-target or product-by-product basis under several scenarios, upon prior written notice.

In August 2021, the Company provided Calico with an option to extend the research term with respect to a certain program for up to a one-year period ending in March 2023. In September 2021, Calico exercised the option resulting in a \$1.0 million extension payment to the Company. In addition, Calico will continue to reimburse the Company for a number of FTEs, depending on the stage of the research, at specified market rates.

### ***Calico Agreement accounting***

The Company identified one performance obligation at the outset of the Calico Agreement, which consists of: (1) the non-exclusive license and (2) the research activities for the target evaluation phase for five targets and the joint research plan phase for two targets. The Company determined that the license and research activities were not distinct from one another, as the license has limited value without the performance of the research activities by the Company.

The transaction price consists of the upfront amount, the committed anniversary payments, the target initiation fees related to the targets nominated at the execution of the Calico Agreement, and the extension payment upon exercise of the extension option discussed above. Initially, the Company amortized the transaction price on a straight-line basis over the initial five-year term of the Calico Agreement. Beginning in September 2021, as a result of the extension of the research term for one program and Calico's obligation to pay an additional \$1.0 million in transaction price, the Company now amortizes the revised transaction price on a straight-line basis over the six-year term of the Calico Agreement. Straight-line amortization of the transaction price was considered the best measure of progress because the customer has access to research and development services throughout the period. Incremental fees for research and development services are paid at agreed upon FTE rates and recognized in the period incurred.

As of March 31, 2022, total transaction price of the Calico Agreement of \$13.0 million is allocated to the research and development services performance obligation and \$2.2 million of the allocated transaction price remains unsatisfied.

Amounts due to the Company that have not yet been received are recorded as accounts receivable and amounts received that have not yet been recognized as revenue from collaboration agreements are recorded in deferred revenue on the Company's condensed consolidated balance sheet.

**Summary of revenue recognized from collaboration agreements**

Revenue from collaboration agreements for the three months ended March 31, 2022 and 2021 in the condensed consolidated statements of operations and comprehensive loss was as follows (in thousands):

	Three Months Ended March 31,	
	2022	2021
Revenue from collaboration agreements:		
Roche Agreement	\$ 1,123	\$ 2,193
Biogen Agreement	4,716	1,880
Calico Agreement	1,815	3,353
Total revenue from collaboration agreements	<u>\$ 7,654</u>	<u>\$ 7,426</u>

Financial information related to the collaboration and license agreements consisted of the following in the Company's condensed consolidated balance sheet as of March 31, 2022 (in thousands):

	Accounts Receivable	Deferred Revenue, Current	Deferred Revenue, Net of Current	Deferred Revenue, Total
Supplemental information:				
Roche Agreement	\$ 500	\$ 6,021	\$ 16,173	\$ 22,194
Biogen Agreement	—	23,078	2,847	25,925
Calico Agreement	1,273	2,167	—	2,167
Total	<u>\$ 1,773</u>	<u>\$ 31,266</u>	<u>\$ 19,020</u>	<u>\$ 50,286</u>

Financial information related to the collaboration and license agreements consisted of the following in the Company's condensed consolidated balance sheet as of December 31, 2021 (in thousands):

	Accounts Receivable	Deferred Revenue, Current	Deferred Revenue, Net of Current	Deferred Revenue, Total
Supplemental information:				
Roche Agreement	\$ 1,215	\$ 5,601	\$ 17,215	\$ 22,816
Biogen Agreement	3,000	24,032	6,611	30,643
Calico Agreement	1,501	2,167	542	2,709
Total	<u>\$ 5,716</u>	<u>\$ 31,800</u>	<u>\$ 24,368</u>	<u>\$ 56,168</u>

Supplemental financial information related to the collaboration and license agreements for the three months ended March 31, 2022 and 2021 are (in thousands):

	Three Months Ended March 31,	
	2022	2021
Revenue recognized that was included in the contract liability at the beginning of the period	\$ 6,267	\$ 4,519
Revenue recognized from performance obligations fully or partially satisfied in previous periods	115	—

As of March 31, 2022, the aggregate amount of the transaction price allocated to performance obligations under the Roche Agreement, the Biogen Agreement, and the Calico Agreement that are partially unsatisfied was \$57.7 million.

**Note 9. Long-term debt and warrant – related party**

On June 5, 2020, contemporaneously with the completion of its Series B Financing, the Company entered into a Credit Agreement, or the Credit Agreement, with Perceptive Credit Holdings III, LP, an affiliate of Perceptive Advisors LLC, or Perceptive, that provided for an aggregate principal borrowing amount of up to \$20.0 million, available in two tranches of \$12.5 million and \$7.5 million. Perceptive was considered a related party to the Company based on its ownership of the Company's common stock at inception of the Credit Agreement.

In June 2020, the Company drew down on the first tranche of \$12.5 million, or the Term Loan, which is outstanding as of March 31, 2022. The Company elected not to draw down the second tranche, which expired on June 30, 2021. The Term Loan bears interest at a variable rate using the greater of LIBOR or 1.75%, plus 9.50%. The interest rate was 11.25% as of March 31, 2022, and is secured by a lien on substantially all of the Company's assets. When the LIBOR interest rate is discontinued in the future, it is expected that the interest rate of the Term Loan would switch to Secured Overnight Financing Rate, or SOFR. As of March 31, 2022, the effect of switching from LIBOR to SOFR would not have been material to the Company's condensed consolidated financial statements.

The Credit Agreement requires the Company to maintain a minimum aggregate cash balance of \$3.0 million in one or more controlled accounts and contains various affirmative and negative covenants that limit its ability to engage in specified types of transactions.

The Company is required to make interest-only payments until December 5, 2022, after which point the Company will be required to make payments of principal equal to 2% of the Term Loan until maturity on June 5, 2024, or the Maturity Date. If the Company pays off the Term Loan prior to the Maturity Date, it will be required to pay a prepayment fee of \$2.3 million as of March 31, 2022.

Under the terms of the Credit Agreement, Perceptive held a warrant to purchase up to 338,784 shares of the Company's common stock at an exercise price of \$8.86 per share. As further described in Note 10, *Stockholders' equity*, in May 2021, Perceptive exercised its warrant using the net exercise method provided by the Credit Agreement.

The following table contains the anticipated future minimum payments on long-term debt as of March 31, 2022 for each of the years ending December 31 and a reconciliation to the carrying value of long-term debt on the Company's condensed consolidated balance sheets (in thousands):

Undiscounted, minimum long-term debt payments:		
Remaining 2022	\$	—
2023		3,000
2024		9,500
Total undiscounted, minimum long-term debt payments		12,500
Less: Unamortized debt issuance costs and debt discount		(1,556)
Total long-term debt—related party	\$	10,944

#### Note 10. Stockholders' equity

In October 2020, the Company authorized preferred stock issuable of 10,000,000 shares and increased its authorized common stock issuable to 150,000,000 shares, both with a \$0.0001 par value per share.

##### *Public offerings of common stock*

In June 2021, the Company completed a follow-on offering, at which time the Company issued 4,887,500 shares of its common stock, including 637,500 shares of common stock that were issued to the underwriters when they exercised in full their overallotment option. Net proceeds from the follow-on offering, including the exercise in full of the underwriters' option to purchase additional shares, were \$169.5 million, after deducting underwriting discounts and commissions, and expenses.

##### *Perceptive warrant – related party*

As described in Note 9, *Long-term debt and warrant – related party*, Perceptive held a warrant to purchase up to 338,784 shares of the Company's common stock at an exercise price of \$8.86 per share. In May 2021, Perceptive exercised its warrant using the net exercise method provided by the Credit Agreement. Under the net exercise method, Perceptive requested that the Company withhold the number of shares equivalent to the aggregate exercise price, or \$3.0 million. As a result of the net exercise, the Company issued Perceptive 256,038 shares of the Company's common stock.

##### *At-The-Market Equity Program*

In November 2021, the Company filed an automatically effective registration statement on Form S-3, or the Registration Statement, with the SEC that registers the offering, issuance and sale of an unspecified amount of common stock, preferred stock, debt securities, warrants and/or units of any combination thereof. Simultaneously, the Company entered into an equity distribution agreement with Cowen and Company, LLC, as sales agent, to provide for the issuance and sale by the Company of up to \$200.0 million of common stock from time to time in "at-the-market" offerings under the Registration Statement and related prospectus filed with the Registration Statement, or the ATM Program. As of March 31, 2022, no sales have been made under the ATM Program.

#### Note 11. Stock-based compensation

The C4 Therapeutics, Inc. 2015 Incentive Stock Option and Grant Plan, or the 2015 Plan, adopted by the Company's board of directors in December 2015 provides for the grant of incentive stock options, nonqualified stock options and restricted stock awards to eligible employees, outside directors and consultants of the Company.

In September 2020, the Company's board of directors adopted the C4 Therapeutics, Inc. 2020 Stock Option and Incentive Plan, or the 2020 Plan. Following the effectiveness of the 2020 Plan, the Company ceased making grants under the 2015 Plan. However, the 2015 Plan continues to govern the terms and conditions of the outstanding awards granted under it. Shares of common stock subject to awards granted under the 2015 Plan that cease to be subject to such awards by forfeiture or otherwise after the termination of the 2015 Plan will be available for issuance under the 2020 Plan. As of March 31, 2022, the Company had 11,056,618 shares reserved under the 2020 Plan and 2015 Plan, and 3,735,127 shares available for future issuance under the 2020 Plan.

Stock-based compensation expense for the three months ended March 31, 2022 and 2021 was classified in the Company's condensed consolidated statement of operations and comprehensive loss as follows (in thousands):

	Three Months Ended March 31,	
	2022	2021
Stock-based compensation expense:		
Research and development	\$ 3,612	\$ 1,171
General and administrative	5,326	2,674
Total stock-based compensation expense	<u>\$ 8,938</u>	<u>\$ 3,845</u>

#### Stock options

The following table summarizes the stock option activity under the Company's equity awards plans for the three months ended March 31, 2022:

	Number of Stock Options	Weighted-Average Exercise Price
Outstanding as of December 31, 2021	5,983,425	\$ 22.33
Granted	1,576,000	23.57
Exercised	(52,707)	4.96
Cancelled or forfeited	(185,227)	30.96
Outstanding as of March 31, 2022	<u>7,321,491</u>	<u>\$ 22.72</u>
Options exercisable as of March 31, 2022	<u>1,668,108</u>	<u>\$ 18.45</u>
Vested and expected to vest as of March 31, 2022	<u>7,321,491</u>	<u>\$ 22.72</u>

As of March 31, 2022, the unrecognized compensation cost related to outstanding options was \$93.0 million, which is expected to be recognized over a weighted-average period of 3.0 years.

#### Restricted stock units

During the three months ended March 31, 2022, under the 2020 Plan, the Company's Board of Directors authorized an issuance of restricted stock units to certain employees, including members of the Company's leadership team. Stock units will vest in tranches as certain discovery milestones, clinical milestones, or specified market conditions are met. Upon vesting, each stock unit automatically converts into one share of the Company's common stock.

The following table summarizes the restricted stock unit activity under the Company's equity awards plans for the three months ended March 31, 2022:

	Shares	Weighted-Average Grant Date Fair Value
Outstanding as of December 31, 2021	—	\$ —
Granted	563,500	25.47
Outstanding as of March 31, 2022	<u>563,500</u>	<u>\$ 25.47</u>

As of March 31, 2022, the unrecognized compensation cost related to outstanding restricted stock units was \$13.8 million.

#### Note 12. Income taxes

During the three months ended March 31, 2022 and 2021, the Company recorded no income tax provision or benefit. As of March 31, 2022 and December 31, 2021, the Company recorded a full valuation allowance on federal and state deferred tax assets since management does not forecast the Company to be in a profitable position in the near future.

#### Note 13. Loss per share

For periods in which the Company reports a net loss attributable to common stockholders, potentially dilutive securities have been excluded from the computation of diluted net loss per share as their effects would be anti-dilutive. Therefore, the weighted average number of common shares outstanding used to calculate both basic and diluted net loss per share attributable to common stockholders

is the same. The Company excluded the following potential common shares presented based on amounts outstanding at period end, from the computation of diluted net loss per share attributable to common stockholders for the periods indicated because including them would have had an anti-dilutive effect:

	<u>As of March 31,</u>	
	<u>2022</u>	<u>2021</u>
Anti-dilutive common stock equivalents:		
Options to purchase common stock	7,321,491	5,952,914
Warrant to purchase common stock	—	338,784
Total anti-dilutive common stock equivalents	<u>7,321,491</u>	<u>6,291,698</u>

Basic and diluted loss per share is computed by dividing net loss attributable to common stockholders by the weighted-average common shares outstanding for the three months ended March 31, 2022 and 2021 (in thousands, except share and per share data):

	<u>Three Months Ended March 31,</u>	
	<u>2022</u>	<u>2021</u>
Numerator:		
Net loss	\$ (31,620)	\$ (20,971)
Denominator:		
Weighted-average number of shares used in computed net loss per share – basic and diluted	<u>48,734,827</u>	<u>43,084,978</u>
Net loss per share attributable to common stockholders – basic and diluted	<u>\$ (0.65)</u>	<u>\$ (0.49)</u>

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

*The following discussion and analysis of our financial condition and results of operations should be read in conjunction with our unaudited interim condensed consolidated financial statements and related notes appearing elsewhere in this Quarterly Report on Form 10-Q and our audited consolidated financial statements and notes and Management's Discussion and Analysis of Financial Condition and Results of Operations, included in our Annual Report on Form 10-K for the year ended December 31, 2021 filed with the SEC on February 24, 2022. As discussed in the section titled "Special Note Regarding Forward-Looking Statements," the following discussion and analysis contains forward-looking statements that involve risks and uncertainties, as well as assumptions that, if they never materialize or prove incorrect, could cause our results to differ materially from those expressed or implied by such forward-looking statements. Factors that could cause or contribute to these differences include, but are not limited to, those identified below and those discussed in the section titled "Risk Factors" in Part II, Item 1A of this Quarterly Report on Form 10-Q and in our Annual Report on Form 10-K for the year ended December 31, 2021.*

### **Business overview**

We are a clinical-stage biopharmaceutical company dedicated to advancing targeted protein degradation science to develop a new generation of small-molecule medicines to transform how disease is treated. We leverage our proprietary technology platform, TORPEDO (Target ORiented ProtEIn Degrader Optimizer), to efficiently design and optimize small-molecule medicines that harness the body's natural protein recycling system to rapidly degrade disease-causing protein, offering the potential to overcome drug resistance, drug undruggable targets and improve patient outcomes. We are advancing multiple targeted oncology programs to the clinic and expanding our research platform to deliver the next wave of medicines for difficult-to-treat diseases.

Our most advanced product candidate, CFT7455, is an orally bioavailable MonoDAC degrader of protein targets called IKZF1 and IKZF3, currently in clinical development for multiple myeloma, or MM, and non-Hodgkin lymphomas, or NHLs, including peripheral T-cell lymphoma, or PTCL, and mantle cell lymphoma, or MCL. We initiated a first-in-human Phase 1/2 clinical trial for this product candidate in June 2021 and presented clinical data from Cohort A at the American Association for Cancer Research Annual Meeting in April 2022. We continue to enroll patients in the ongoing clinical trial. In August 2021, the FDA granted orphan drug designation to CFT7455 for the treatment of multiple myeloma.

We are also developing CFT8634, an orally bioavailable BiDAC degrader candidate targeting a protein called BRD9, for synovial sarcoma and SMARCB1-deleted solid tumors. In December 2021, the FDA cleared the IND application for CFT8634, and we expect to dose the first patient in a first-in-human Phase 1/2 clinical trial of this product candidate in the first half of 2022. In March 2022, the FDA granted orphan drug designation to CFT8634 for the treatment of soft tissue sarcoma.

Further, we are developing CFT1946, an orally bioavailable BiDAC degrader candidate specifically targeting V600X mutant BRAF to treat melanoma, non-small cell lung cancer, or NSCLC, colorectal cancer and other solid malignancies that harbor this mutation. We expect to submit an IND for this product candidate and begin a first-in human Phase 1/2 clinical trial in BRAF V600X driven cancers including melanoma, NSCLC and colorectal cancer in the second half of 2022.

Additionally, we are developing CFT8919, an orally bioavailable, allosteric, mutant-selective BiDAC degrader of epidermal growth factor receptor, or EGFR, with an L858R mutation in NSCLC. We expect to complete IND-enabling activities for this product candidate by the end of 2022.

Beyond these initial product candidates, we are further diversifying our pipeline by developing new degraders against both clinically-validated and currently undruggable targets. We have engineered degrader candidates that have successfully achieved blood-brain barrier penetration in preclinical studies, which is a key step in developing medicines with the potential to treat brain metastases in oncology as well as in therapeutic areas such neurodegenerative diseases. We also believe there are many other therapeutic areas and indications where leveraging our TORPEDO platform to develop novel degrader candidates may be advantageous.

### **Financial operations overview**

#### **Revenues**

To date, we have not generated any revenue from product sales and do not expect to generate any revenue from the sale of products for the foreseeable future. Our revenues to date have been generated through research collaboration and license agreements. We recognize revenue over the expected performance period under each agreement. We expect that our revenue for the next several years will be derived primarily from our current collaboration agreements and any additional collaborations that we may enter into in the future. To date, we have not received any royalties under any of our existing collaboration agreements.

#### *Roche Collaboration and License Agreement*

In March 2016, we entered into a license agreement, or the Original Roche Agreement, with F. Hoffmann-La Roche Ltd and Hoffmann-La Roche Inc., or Roche, whereby Roche provided us with a non-refundable upfront payment of \$15.0 million, which was creditable against our target initiation fees of either \$1.0 million or \$4.0 million, depending on the compound selected. Pursuant to the terms of the Original Roche Agreement, we collaborated on research activities to develop novel treatments in the field of targeted protein degradation using our degrader technology. Under the Original Roche Agreement, we were initially responsible for developing therapeutics that utilize degrader technology for up to ten target proteins. On a target-by-target basis, after successful completion of a

defined preclinical development phase, Roche had an exclusive option to pursue a license from us for further clinical development and commercialization.

On December 22, 2018, we amended and restated the Original Roche Agreement, or the Roche Agreement. Under the Roche Agreement, we have a more active role in the manufacturing and commercialization of products related to the targets included in the collaboration, whereby if we elect to opt into certain co-development rights, we will receive an increased royalty rate on future product sales from commercializing products directed to the target. If we opt into certain co-detailing rights, we are entitled to reimbursement of certain commercialization costs. The target structure was revised to six potential targets, three of which had been nominated as of the execution of the Roche Agreement and represent continuations of the initial preclinical research and development efforts begun under the Original Roche Agreement, and three additional targets that were not nominated as of the date of execution of the Roche Agreement. At the time of entry into the Roche Agreement, Roche maintained its option rights to license and commercialize products related to these six targets.

In November 2020, we signed a further amendment to the Roche Agreement that created a mechanism through which we and Roche can mutually agree to terminate the Roche Agreement on a target-by-target basis by the entry into a Mutual Target Termination Agreement. Upon a termination of this nature, the Roche Agreement, as amended, provides that all rights in know-how and intellectual property in support of products that use inhibition as their mode of action, referred to as the Roche Field, will revert to Roche and all rights in know-how and intellectual property in support of products that use degradation as their mode of action, referred to as the C4T Field, will revert to us. Further, this amendment states that, following the entry into a Mutual Target Termination Agreement, Roche will have rights in and responsibility for any know-how and intellectual property generated as a result of the collaboration that fits within the Roche Field and we will have rights in and responsibility for any know-how and intellectual property generated as a result of the collaboration that fits within the C4T Field. In support of this allocation of rights, under the amendment, Roche provided us, and we provided Roche, with a perpetual, irrevocable, fully paid up, exclusive (even as to party granting the license), sublicensable (including in multiple tiers) license to the patents that are allocated to a party under the mutual target termination agreement and a perpetual, irrevocable fully paid up, non-exclusive, sublicensable (including in multiple tiers) license to the know-how that is allocable to a party under the Mutual Target Termination Agreement. Through the entry into this amendment, we and Roche mutually agreed to terminate the Roche Agreement as to the target EGFR and, in November 2021, we entered into a Mutual Target Termination Agreement with Roche through which we agreed to terminate the Roche Agreement as to the target BRAF. Following this mutual target termination decision, the number of targets on which the parties shall continue to collaborate has been reduced to four, with Roche maintaining its option rights to license and commercialize products directed to those four targets. As a result, Roche is now free to pursue the targets EGFR and BRAF in the Roche Field and we are free to pursue the targets EGFR and BRAF in the C4T Field. All rights in and responsibility for know-how and intellectual property related to EGFR and BRAF in the Roche Field reverted to the Roche parties and all rights in and responsibility for know-how and intellectual property related to EGFR and BRAF in the C4T Field reverted to us. Roche is in the process of assigning the relevant patents related to BRAF in the C4T Field to us.

Under the Roche Agreement, we received additional upfront consideration of \$40.0 million from Roche. Roche will make annual research plan payments of \$1.0 million for each active research plan. Finally, adjustments were made to the option exercise fees, whereby targets that have progressed through standard good laboratory practice, or GLP, toxicology studies at the time of exercise now have option exercise fees of \$7.0 million to \$12.0 million and those progressed through Phase 1 trials have option exercise fees of \$20.0 million.

For certain targets, Roche is required to pay us fees of \$2.0 million and \$3.0 million upon the identification of a lead series and the commencement of GLP toxicology studies, respectively. For each target option exercised by Roche, we are eligible to receive up to \$275 million in research, development and commercial milestone payments per target. Roche is also required to pay us up to \$150 million per target in one-time sales-based payments if the target achieves certain levels of net sales. Roche is also required to pay us royalties, at percentages from the mid-single digits to the low double-digits, on a licensed product-by licensed product basis, on worldwide net product sales.

#### *Biogen Collaboration Research and License Agreement*

On December 28, 2018, we entered into a collaborative research and license agreement, or the Biogen Agreement, with Biogen MA, Inc., or Biogen, whereby we agreed to collaborate on research and development efforts for up to five targets to discover and develop potential new treatments for neurological conditions, such as Alzheimer's disease and Parkinson's disease. The Biogen Agreement also has an option for Biogen to nominate additional targets and extend the Biogen Agreement. In February 2020, we entered into an amendment to the Biogen Agreement that provided further clarity around Biogen's ownership of target binding moieties, which are portions of molecules, and any related intellectual property that are directed at or bind to collaboration targets. This amendment further provides that Biogen licenses to us rights to use these Biogen target binding moieties and any related intellectual property as needed in order to conduct the research and development activities contemplated under the Biogen Agreement.

We granted Biogen a non-exclusive research license under our intellectual property to perform research activities, select and optimize degraders and develop products including the degraders, as well as a commercial license to manufacture and commercialize the products related to the targets once the initial research and development work is complete. The research under the Biogen Agreement will take place over a 54-month research term, ending in June 2023, with Biogen having an option to extend the Biogen Agreement for

up to four additional years. If Biogen elects to extend the term of the Biogen Agreement, Biogen would be required to make an additional payment of \$62.5 million and would be entitled to nominate up to five additional targets.

The Biogen Agreement provides for three initial targets, with Biogen having the right to initiate up to an additional two targets and to control all post-discovery activities. Biogen paid us a nonrefundable upfront payment of \$45.0 million for access to our technology and research services through the discovery research phase. The nonrefundable upfront cash payment of \$45.0 million is not creditable against any of the target development milestone fees.

Following the achievement of development candidate criteria, prior to any IND-enabling study, for any target, Biogen will bear all costs and expenses of and will have sole discretion and decision-making authority with respect to the performance of further activities with respect to any degrader under development under the Biogen Agreement and all products that incorporate that degrader. Biogen is also required to pay us up to \$35.0 million per target in development milestones and \$26.0 million per target in one-time sales-based payments for the first product to achieve certain levels of net sales. In addition, Biogen is required to pay us royalties on a licensed product-by-licensed product basis, on worldwide net product sales, at percentages in the mid-single digits. All milestone and sales-based payments are made after we have met the defined criteria in the joint research plan for that target, at which time Biogen will have control of the products related to the targets for commercialization; the receipt of these payments is contingent on the further development of products directed to the targets to commercialization by Biogen, without any additional research and development efforts from us.

Biogen also had the option to fund additional discovery activities, referred to as sandbox activities, whereby we performed discovery-type research at Biogen's election to develop other potential targets that may be used as replacement targets for the initially nominated targets or two additional targets under the Biogen Agreement. Sandbox activities fully concluded on August 31, 2021. Revenues earned under this option, were recognized as services were performed and were not included in the transaction price at the outset of the arrangement. These research activities were reimbursed on a full-time equivalent, or FTE, basis at specified market rates. These additional discovery activities were purchased up to a maximum amount by Biogen on an à la carte basis at an amount consistent with standalone selling price. If Biogen exercised these options, we recognized revenue as those options were exercised.

#### *Calico License Agreement*

In March 2017, we entered into Collaboration and License Agreement, or the Calico Agreement, with Calico Life Sciences LLC, or Calico, whereby we agreed to collaborate to develop and commercialize a set number of targets for small molecule protein degraders for diseases of aging, including cancer, for a five-year period ending in March 2022. In August 2021, we provided Calico with an option to extend the research term with respect to a certain program for up to a one-year period ending in March 2023, such period referred to as the research term. In September 2021, Calico elected to exercise the option for a one-year extension of the research term for this specified program.

We provided Calico with a non-exclusive research license under our intellectual property to perform research activities and select and optimize degraders and develop products including the degraders. We also granted Calico a commercial license for any licensed products resulting from the development candidates supplied by us. We are required to perform research and development activities for the nominated targets over the applicable research term, with the intent to provide a development candidate for each target to Calico once the agreed-upon research is complete.

Calico is obligated to reimburse our research and development activities for each target at specified levels through the identification of a development candidate, after which Calico shall assume full responsibility for candidate development.

After the initiation of each target, the Calico Agreement does not contain any options for Calico to license the individual targets; once we complete the initial research and development activities required, Calico controls and directs the targets with no additional work required to be performed by us. There is no exercise price or incremental fee payable to us to progress the research further, though Calico is required to pay an initiation fee with the commencement of each research plan. Once Calico nominates a target and pays the applicable target initiation fee, we will commence research and development activities for that target. The Calico Agreement provides for up to five initial targets. Research activities performed are reimbursed at specified levels for the five-year term of the Calico Agreement, with the term for one research program having been extended by an additional year.

Under this agreement, Calico paid us a nonrefundable upfront amount of \$5.0 million and certain annual payments of \$5.0 million through December 31, 2020. Upon our completion of the required discovery research and development services on any target, Calico is entitled to pursue development of that target to commercialized product. For each target, we are eligible to receive potential research, development and commercial milestone payments aggregating up to \$132.0 million. Calico is also required to pay one-time sales-based payments aggregating up to \$65.0 million for the first product to achieve certain levels of net sales. In addition, Calico is required to pay us royalties, on a licensed product-by-licensed product basis, on worldwide net product sales, at percentages in the mid-single digits. All milestone and sales-based payments are made after we have met the defined criteria in the joint research plan for that target, at which time Calico will have control of the products related to targets for commercialization; the receipt of these payments by us is contingent on the further development of the targets to commercialized products by Calico, without any additional research and development efforts required by us.

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

Research and development expenses consist primarily of costs incurred for our research activities, including our discovery efforts, and the development of our product candidates, and include:

- salaries, benefits, and other related costs, including stock-based compensation expense, for personnel engaged in research and development functions;
- expenses incurred under agreements with third parties, including contract research organizations and other third parties that conduct research, preclinical, and clinical activities on our behalf as well as third parties that manufacture our product candidates for use in our preclinical and clinical trials;
- costs of outside consultants, including their fees, unit-based compensation, and related travel expenses;
- costs of laboratory supplies and acquiring materials for preclinical studies and clinical trials;
- facility-related expenses, which include direct depreciation costs of equipment and allocated expenses for rent and maintenance of facilities and other operating costs; and
- third-party licensing fees.

We expense research and development costs as incurred. Costs for external development activities are recognized based on an evaluation of the progress to completion of specific tasks using information provided to us by our vendors. Payments for these activities are based on the terms of the individual agreements, which may differ from the pattern of costs incurred, and are reflected in our condensed consolidated financial statements as prepaid or accrued research and development expenses. Nonrefundable advance payments for goods or services to be received in the future for use in research and development activities are recorded as prepaid expenses and expensed as the related goods are delivered or the services are performed.

We expect that our research and development expenses will continue to increase substantially in connection with our planned preclinical and clinical development activities.

### ***General and administrative expenses***

General and administrative expenses consist primarily of salaries and other related costs, including stock-based compensation, for personnel in our executive, finance, legal, business development and administrative functions. General and administrative expenses also include legal fees relating to intellectual property and corporate matters; professional fees for accounting, auditing, tax and consulting services; insurance costs; travel expenses; and facility-related expenses, which include direct depreciation costs and allocated expenses for rent and maintenance of facilities and other operating costs.

We expect that our general and administrative expenses will continue to increase in the future to support increased research and development activities. These increases will likely include increased costs related to the hiring of additional personnel; fees to outside consultants, lawyers and accountants; director and officer insurance costs, among other expenses. and investor and public relations costs.

### ***Other (expense) income, net***

Other (expense) income, net primarily consists of the following:

- interest expense and amortization of our long-term debt, which is discussed in greater detail in Note 9, *Long-term debt and warrant – related party*, to the unaudited condensed consolidated financial statements in this Quarterly Report on Form 10-Q; and
- interest income earned on our cash, cash equivalents, and marketable securities and accretion of discount on marketable securities.

## Results of operations

### Comparison of the three months ended March 31, 2022 and 2021

#### Revenue

Revenue from our collaboration and license agreements consisted of the following for the three months ended March 31, 2022 and 2021 (in thousands):

	Three Months Ended March 31,	
	2022	2021
Revenue from collaboration agreements:		
Roche Agreement	\$ 1,123	\$ 2,193
Biogen Agreement	4,716	1,880
Calico Agreement	1,815	3,353
Total revenue from collaboration agreements	<u>\$ 7,654</u>	<u>\$ 7,426</u>

The \$0.2 million increase in revenue in the three months ended March 31, 2022 as compared to the three months ended March 31, 2021 is primarily driven by:

- a \$2.8 million increase in revenue recognized under the Biogen Agreement, resulting from increased effort made on the nominated targets and additional revenue recognized from the Biogen milestone earned in 2021.

This was offset by:

- a \$1.5 million decrease in FTE reimbursements recognized under the Calico Agreement as a result of the extension of the research term under the Calico Agreement with respect to a specified program in September 2021; and
- a \$1.1 million decrease in revenue recognized under the Roche Agreement, resulting from the termination of the Roche Agreement as to the target BRAF in November 2021.

#### Research and development expense

The following table summarizes our research and development expense for the three months ended March 31, 2022 and 2021 (in thousands):

	Three Months Ended March 31,	
	2022	2021
Research and development expenses:		
Personnel expenses	\$ 10,911	\$ 6,934
Preclinical and development expenses	10,195	9,389
Facilities and supplies	2,273	2,297
Professional fees	1,655	1,171
Clinical expenses	809	218
Intellectual property	270	423
Other expenses	90	94
Total research and development expenses	<u>\$ 26,203</u>	<u>\$ 20,526</u>

The \$5.7 million increase in research and development expense in the three months ended March 31, 2022 as compared to the three months ended March 31, 2021 is primarily driven by:

- a \$4.0 million increase in personnel expenses, representing salary and benefit costs, including a \$2.4 million increase in stock-based compensation expense driven primarily by the equity awards issued to employees subsequent to March 31, 2021, and due to the buildout of our clinical development team.

#### General and administrative expense

The following table summarizes our general and administrative expense for the three months ended March 31, 2022 and 2021 (in thousands):

	Three Months Ended March 31,	
	2022	2021
General and administrative expenses:		
Personnel expenses	\$ 8,548	\$ 4,640
Professional fees	2,227	2,431
Facilities and other expenses	2,045	338
Total general and administrative expenses	<u>\$ 12,820</u>	<u>\$ 7,409</u>

The \$5.4 million increase in general and administrative expense in the three months ended March 31, 2022 as compared to the three months ended March 31, 2021 is primarily driven by:

- a \$3.9 million increase in personnel expenses resulting from a \$2.6 million increase in stock-based compensation expenses driven primarily by the equity awards issued to employees subsequent to March 31, 2021; and
- a \$1.6 million increase in facilities and other expenses, which is driven primarily by lease commencement in January 2022 of our additional leased space.

*Other (expense) income, net*

The following table summarizes our other (expense) income for the three months ended March 31, 2022 and 2021 (in thousands):

	<u>Three Months Ended March 31,</u>	
	<u>2022</u>	<u>2021</u>
Other (expense) income, net:		
Interest expense and amortization of long-term debt – related party	\$ (527)	\$ (534)
Interest and other income, net	276	72
Total other (expense) income, net	<u>\$ (251)</u>	<u>\$ (462)</u>

**Liquidity and capital resources**

*Sources of liquidity*

Since inception, we have incurred significant operating losses. We expect to incur significant expenses and operating losses for the foreseeable future as we advance the preclinical and, if successful, the clinical development of our programs. We do not currently have any approved products and have never generated any revenue from product sales. To date, we have financed our operations primarily through the sale of preferred stock, public offerings of our common stock, and through payments from collaboration partners. As of March 31, 2022, we had cash, cash equivalents and marketable securities of approximately \$421.7 million.

*Cash flows*

The following table summarizes our sources and uses of cash for the periods presented (in thousands):

	<u>Three Months Ended March 31,</u>	
	<u>2022</u>	<u>2021</u>
Net change in cash, cash equivalents and restricted cash:		
Net cash used in operating activities	\$ (26,596)	\$ (24,934)
Net cash used in investing activities	(4,004)	(61,730)
Net cash provided by (used in) financing activities	480	(151)
Total net change in cash, cash equivalents and restricted cash	<u>\$ (30,120)</u>	<u>\$ (86,815)</u>

*Operating activities*

Net cash used in operating activities for the three months ended March 31, 2022 was driven primarily by the following uses of cash:

- our net loss of \$31.6 million;
- a \$5.9 million change in deferred revenue due to the recognition of revenue under our collaboration agreements; and
- a \$4.9 million change in accounts payable and accrued expenses.

These were offset by non-cash expenses of \$11.3 million, which primarily consisted of stock-based compensation expense of \$8.9 million, and a \$3.9 million change in accounts receivable.

Net cash used in operating activities for the three months ended March 31, 2021 was driven primarily by the following uses of cash:

- our net loss of \$21.0 million;
- a \$5.6 million change in accounts payable and accrued expenses; and
- a \$3.8 million change in deferred revenue due to the recognition of revenue under our collaboration agreements.

These were offset by non-cash expenses of \$4.9 million, which primarily consisted of stock-based compensation expense of \$3.8 million.

#### *Investing activities*

Net cash used in investing activities for the three months ended March 31, 2022 was primarily attributable to \$3.8 million of purchases of marketable securities, net of maturities.

Net cash used in investing activities for the three months ended March 31, 2021 was primarily attributable to \$61.3 million of purchases of marketable securities, net of maturities.

#### **Funding requirements**

Since our inception, we have incurred significant operating losses and we expect to continue to incur significant expenses and increasing operating losses for the foreseeable future as we advance the preclinical and clinical development of our product candidates. In addition, we expect to continue to incur costs associated with operating as a public company.

Specifically, we anticipate that our expenses will increase substantially in the future, if and as we:

- continue our ongoing first-in-human Phase 1/2 trial and initiate and conduct planned first-in-human Phase 1/2 trials for our other lead product candidates;
- advance additional product candidates into preclinical and clinical development;
- continue to invest in our proprietary TORPEDO platform;
- advance, expand, maintain and protect our intellectual property portfolio;
- hire additional clinical, regulatory, quality and scientific personnel;
- add operational, financial and management information systems and personnel to support our ongoing research, product development, potential future commercialization efforts, operations as a public company and general and administrative roles;
- seek marketing approvals for any product candidates that successfully complete clinical trials; and
- ultimately establish a sales, marketing and distribution infrastructure and scale up external manufacturing capabilities to commercialize any products for which we may obtain marketing approval.

Because of the numerous risks and uncertainties associated with development and commercialization of our product candidates, we are unable to estimate the amounts of increased capital and operating costs associated with our current and anticipated preclinical and clinical development. Our future capital requirements will depend on many factors, including:

- the progress, costs and results of ongoing and planned first-in-human Phase 1/2 trials for our lead product candidates and any future clinical development of those lead product candidates;
- the scope, progress, costs and results of preclinical and clinical development for our other product candidates and development programs;
- the number and development requirements of other product candidates that we pursue;
- the progress and success of our collaborations with Roche, Biogen and Calico, including whether or not we receive additional research support or milestone payments from our collaboration partners upon the achievement of milestones;
- the costs, timing and outcome of regulatory review of our product candidates;
- the costs and timing of preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending any intellectual property-related claims;
- our willingness and ability to establish additional collaboration arrangements with other biotechnology or pharmaceutical companies on favorable terms, if at all, for the development or commercialization of current or additional future product candidates;
- the costs and timing of future commercialization activities, including product manufacturing, marketing, sales and distribution, for any of our product candidates for which we receive marketing approval; and
- the revenue, if any, received from commercial sales of our product candidates for which we receive marketing approval.

As a result of the anticipated expenditures described above, we will need to obtain substantial additional financing to support our continuing operations and pursue our long-term business plan. Until such time, if ever, that we can generate substantial revenue from product sales, we expect to finance our cash needs through a combination of equity offerings, debt offerings, collaborations, strategic

alliances and marketing, distribution or licensing arrangements. Although we may receive potential future milestone and royalty payments under our collaborations with Roche, Biogen and Calico, we do not have any committed external source of funds as of March 31, 2022. Adequate additional funds may not be available to us on acceptable terms, or at all. If we are unable to raise capital when needed or on attractive terms, we may be required to delay, limit, reduce or terminate our research, product development or future commercialization efforts or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

If we raise additional capital through the sale of equity securities, each investor's ownership interest will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect your rights as a common stockholder. Preferred equity financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as making acquisitions or capital expenditures or declaring dividends.

If we raise additional funds through collaborations, strategic alliances or marketing, distribution or licensing arrangements with third parties, we may have to relinquish valuable rights to our technologies, future revenue streams, research programs or product candidates or grant licenses on terms that may not be favorable to us.

#### **Contractual obligations**

We enter into contracts in the normal course of business with contract manufacturing organizations, contract research organizations and other vendors to assist in the performance of our research and development activities and other services and products for operating purposes. These contracts generally provide for termination on notice, and therefore are cancelable contracts and not included in the table of contractual obligations and commitments.

During the three months ended March 31, 2022, except for the minimum rental commitments disclosed in Note 6, *Leases*, and Note 9, *Long-term debt and warrant – related party*, to the unaudited condensed consolidated financial statements in this Quarterly Report on Form 10-Q, there were no significant changes to our contractual obligations and 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 year ended December 31, 2021.

#### **Critical accounting policies and use of estimates**

Our critical accounting policies are those policies that require the most significant judgments and estimates in the preparation of our unaudited condensed consolidated financial statements. We have determined that our most critical accounting policies are those relating to revenue recognition from collaborations, research and development expense recognition, lease liability measurement, and stock-based compensation. There have been no significant changes to our existing critical accounting policies discussed in our Annual Report on Form 10-K for the year ended December 31, 2021 filed with the SEC on February 24, 2022.

#### **Off-balance sheet arrangements**

We have not entered into any off-balance sheet arrangements, as defined under the applicable regulations of the SEC.

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

We are exposed to market risk related to changes in interest rates. Our marketable securities are subject to interest rate risk and may decrease in value if market interest rates increase. As of March 31, 2022, we had marketable securities of \$375.6 million, which consisted of corporate debt securities, U.S. government debt securities, and U.S. Treasury securities. Our marketable securities are short term in nature with a weighted-average maturity date of 0.8 years. As such, while these interest-earning instruments carry a degree of interest rate risk, historical fluctuations in interest income have not been significant for us.

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

#### **Evaluation of disclosure controls and procedures**

Our management, with the participation of our principal executive officer and our principal financial officer have evaluated the effectiveness of our disclosure controls and procedures as defined in Rules 13a-15(e) and 15d-15(e) under the Securities Exchange Act of 1934, as amended, or the Exchange Act, as of March 31, 2022. The term "disclosure controls and procedures," as defined in the 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 a company in the reports that it files or submits under the Exchange Act is accumulated and communicated to the company's management, including its principal executive and principal financial officers, or persons performing similar functions, as appropriate to allow timely decisions regarding required disclosure. Management recognizes that any controls and procedures, no matter how well designed and operated, can provide only reasonable assurance of achieving their objectives and management necessarily applies its judgment in evaluating the cost-benefit relationship of possible controls and procedures.

Based on the evaluation of our disclosure controls and procedures as of March 31, 2022, our principal executive officer and principal financial officer concluded that our disclosure controls and procedures as of such date are effective at the reasonable assurance level.

**Changes in internal control over financial reporting**

We continuously seek to improve the efficiency and effectiveness of our internal controls. This results in refinements to processes throughout the company. There were no changes in our internal control over financial reporting (as defined in Rules 13a-15(f) and 15d-15(f) under the Exchange Act) during the quarter ended March 31, 2022 that have materially affected, or are reasonably likely to materially affect, our internal control over financial reporting.

**Inherent limitation on the effectiveness over financial reporting**

The effectiveness of any system of internal control over financial reporting, including ours, is subject to inherent limitations, including the exercise of judgment in designing, implementing, operating, and evaluating the controls and procedures, and the inability to eliminate misconduct completely. Accordingly, any system of internal control over financial reporting, including ours, no matter how well designed and operated, can only provide reasonable, not absolute assurances. In addition, projections of any evaluation of effectiveness to future periods are subject to the risk that controls may become inadequate because of changes in conditions, or that the degree of compliance with the policies or procedures may deteriorate. We intend to continue to monitor and upgrade our internal controls as necessary or appropriate for our business, but there can be no assurance that such improvements will be sufficient to provide us with effective internal control over financial reporting.

## PART II—OTHER INFORMATION

### Item 1. Legal Proceedings.

From time to time, we may become involved in legal proceedings arising in the ordinary course of our business. As of the date of this quarterly report on Form 10-Q, we were not a party to any material legal matters or claims.

### Item 1A. Risk Factors.

*Investing in our common stock involves a high degree of risk. You should carefully consider the risks described below, as well as the other information in this Quarterly Report on Form 10-Q, including our financial statements and the related notes and the section of this Quarterly Report on Form 10-Q titled “Management’s Discussion and Analysis of Financial Condition and Results of Operations” before you make an investment decision. The occurrence of any of the events or developments described below could harm our business, financial condition, results of operations and growth prospects. As a result, the market price of our common stock could decline, and you may lose all or part of your investment in our common stock.*

#### **Risks related to our financial position and need for additional capital**

***We are a clinical-stage biopharmaceutical company with a limited operating history and have incurred significant losses since our inception. We expect to incur losses over at least the next several years and may never achieve or maintain profitability.***

We are a clinical-stage biopharmaceutical company with limited operating history. Our net loss was \$31.6 million and \$21.0 million for the three months ended March 31, 2022 and 2021, respectively. As of March 31, 2022, we had an accumulated deficit of \$299.3 million. To date, we have not generated any revenue from product sales and have financed our operations primarily through sales of our equity interests, including public offerings of our common stock, proceeds from our collaborations and debt financing. We are still in the early stages of development of our product candidates. As a result, we expect that it will be several years, if ever, before we have a product candidate ready for regulatory approval and commercialization. We may never succeed in these activities and, even if we do, may never generate revenues that are significant enough to achieve profitability. To become and remain profitable, we must succeed in developing, obtaining marketing approval for and commercializing products that generate significant revenue. This will require us to be successful in a range of challenging activities, including, without limitation, successfully completing preclinical studies and clinical trials of our product candidates, discovering additional product candidates, establishing arrangements with third parties for the conduct of our clinical trials, procuring clinical- and commercial-scale manufacturing, obtaining marketing approval for our product candidates, manufacturing, marketing and selling any products for which we may obtain marketing approval, identifying collaborators to develop product candidates we identify or additional uses of existing product candidates and successfully completing development of product candidates for our collaboration partners.

We expect to continue to incur significant expenses and increasing operating losses for at least the next several years. We anticipate that our expenses will increase substantially if and as we:

- initiate, conduct, and successfully complete first-in-human and later-stage clinical trials of our product candidates and as we expand the scope of our proprietary research and development portfolios;
- leverage our TORPEDO platform to identify and then advance additional product candidates into preclinical and clinical development;
- expand the capabilities of our TORPEDO platform;
- seek marketing approvals for any product candidates that successfully complete clinical trials;
- ultimately establish a sales, marketing and distribution infrastructure and scale up external manufacturing capabilities to commercialize any products for which we may obtain marketing approval;
- advance, expand, maintain and protect our intellectual property portfolio;
- hire additional personnel, including in areas such as clinical development, regulatory, quality, scientific, and general and administrative positions; and
- add operational, financial and management information systems and personnel, including personnel to support our ongoing research and development and potential future commercialization efforts.

Further, we expect to continue to incur additional costs associated with operating as a public company, including significant legal, accounting, insurance, investor relations and other expenses.

Our expenses could increase beyond our expectations if we are required by the FDA, the European Medicines Agency, or other regulatory authorities to perform trials in addition to those that we currently expect, or if we experience any delays in either

establishing appropriate manufacturing arrangements for or completing our clinical trials or the clinical development of any of our product candidates.

Because of the numerous risks and uncertainties associated with pharmaceutical product development, we are unable to accurately predict the timing or amount of increased expenses we will incur or when, if ever, we will be able to achieve profitability. Even 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 depress the value of our company and could impair our ability to raise capital, expand our business, maintain our research and development efforts, expand our business or continue operations. A decline in the value of our company, or in the value of our common stock, could also cause you to lose all or part of your investment.

If one or more of the product candidates that we develop is approved for commercial sale, we anticipate incurring significant costs associated with commercializing those approved product candidates. 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 funding to pursue our business objectives and continue our operations. If we are unable to raise capital when needed, we may be required to delay, limit, reduce or terminate our research or product development programs or future commercialization efforts.***

We expect our expenses to increase substantially in connection with our ongoing activities, particularly as we prepare for and initiate, conduct, and complete our ongoing and planned first-in-human Phase 1/2 clinical trials of our product candidates, advance our TORPEDO platform and continue research and development, expand our proprietary research and development portfolios and initiate and continue clinical trials of, and potentially seek marketing approval for, our current and future preclinical programs. In addition, if we obtain marketing approval for any of our product candidates, we expect to incur significant commercialization expenses related to product manufacturing, marketing, sales and distribution. Further, we expect to continue to incur additional costs associated with operating as a public company. Accordingly, we will need to obtain substantial additional funding in connection with our continuing operations. If we are unable to raise capital when needed or on attractive terms, we may be required to delay, limit, reduce or terminate our research, product development programs or any future commercialization efforts or grant rights to develop and market product candidates that we would otherwise prefer to develop and market ourselves.

We had cash, cash equivalents, and marketable securities of approximately \$421.7 million as of March 31, 2022. We believe that these funds, together with future payments expected to be received under existing collaboration agreements will be sufficient to fund our existing operating plan through the end of 2024. We have based this estimate on assumptions that may prove to be wrong, and we could use our capital resources sooner than we currently expect. Our future capital requirements will depend on many factors, including:

- the timing, progress, costs and results of our ongoing and planned first-in-human Phase 1/2 clinical trials for our product candidates and any future clinical development of those product candidates;
- the scope, progress, costs and results of preclinical and clinical development for CFT7455, CFT8634 and our other product candidates and development programs;
- the number and development requirements of other product candidates that we pursue;
- the success of our ongoing collaborations with Biogen, Roche and Calico;
- the costs, timing and outcomes of regulatory review of our product candidates;
- the costs and timing of future commercialization activities, including product manufacturing, marketing, sales and distribution, for any of our product candidates for which we receive marketing approval;
- the revenue, if any, received from commercial sales of our product candidates for which we receive marketing approval and the timing of the receipt of any such revenue;
- any delays or interruptions, including due to the ongoing coronavirus, or COVID-19, pandemic, that we experience in our preclinical studies, clinical trials and/or supply chain;
- the costs and timing of preparing, filing and prosecuting patent applications, maintaining and enforcing our intellectual property rights and defending any intellectual property-related claims; and
- our ability to establish collaboration arrangements with other biotechnology or pharmaceutical companies on favorable terms, if at all, for the development or commercialization of our product candidates.

Our current cash, cash equivalents, and marketable securities will not be sufficient for us to fund any of our product candidates through regulatory approval. As a result, we will need to raise substantial additional capital to complete the development and commercialization of our product candidates. Identifying potential product candidates and conducting preclinical studies and clinical trials is a time-consuming, expensive and uncertain process that takes years to complete. We may never generate the necessary data or results required to obtain marketing approval and achieve product sales. In addition, our product candidates, if approved, may not

achieve commercial success. Our commercial revenues, if any, will be derived from sales of products that we do not expect to be commercially available for several years, if at all. Adequate additional funds may not be available to us on acceptable terms, or at all. In addition, we may seek additional capital due to favorable market conditions or strategic considerations, even if we believe we have sufficient funds for our current or future operating plans.

If one or more of the product candidates that we develop is approved for commercial sale, we anticipate incurring significant costs associated with commercializing those approved product candidates. 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 remain early in the development lifecycle, which may make it difficult for you to evaluate the success of our business to date and assess our future viability.***

We commenced operations in 2015 and our operations to date have been limited to organizing and staffing our company, business planning, raising capital, conducting discovery and research activities, filing patent applications, identifying potential product candidates, developing and advancing our TORPEDO platform, undertaking preclinical studies, establishing arrangements with third parties for the manufacture of initial quantities of our product candidates, and preparing for and initiating clinical trials. While we commenced a clinical trial of CFT7455 in June 2021 and expect to commence a clinical trial for CFT8634 during the first half of 2022, all of our other product candidates are still in preclinical development or in the discovery stage. We have not yet demonstrated our ability to successfully complete any clinical trials, obtain marketing approvals, manufacture a commercial scale product directly or through a third party or conduct sales, marketing and distribution activities necessary for successful product commercialization. Consequently, any predictions you make about our future success or viability may not be as accurate as they could be if we had a longer operating history or if we had already successfully completed some or all of these types of activities in the past.

In addition, as a biopharmaceutical company, we may encounter unforeseen expenses, difficulties, complications, delays and other known and unknown challenges. We will need to transition at some point from a company with a research and development focus to a company capable of supporting commercial activities and we may not be successful in making that transition.

We expect our financial condition and operating results to continue to fluctuate significantly from quarter to quarter and year to year due to a variety of factors, many of which are beyond our control. Accordingly, you should not rely upon the results of any quarterly or annual periods as indications of future operating performance.

***The ongoing global COVID-19 pandemic could continue to adversely impact our business, including our discovery programs, preclinical studies and development programs, supply chain, business development activities, and general corporate matters.***

The COVID-19 pandemic, which began in December 2019 and remains ongoing, has spread worldwide and caused governments worldwide to implement measures to slow the spread of the pandemic through quarantines, travel restrictions, heightened border scrutiny, business shutdowns and other measures. The pandemic and government measures taken in response have also had a significant impact, both direct and indirect, on businesses and commerce, as worker shortages have occurred, supply chains have been disrupted, facilities and production have been suspended and demand for certain goods and services, such as medical services and supplies, has spiked, while demand for other goods and services, such as travel, has fallen. Since the beginning of the COVID-19 pandemic, three vaccines for COVID-19 have received Emergency Use Authorization by the FDA and two of those later received marketing approval. Additional vaccines may be authorized or approved in the future. The resultant demand for vaccines and potential for manufacturing facilities and materials to be commandeered under the Defense Production Act of 1950, or equivalent foreign legislation, may make it more difficult to obtain materials or manufacturing slots for the products needed for our clinical trials, which could lead to delays in the initiation or conduct of our trials. While these COVID-19 vaccines are now actively being distributed in the United States and around the world, the vaccines have not been universally adopted by and distributed throughout the global population and new strains of COVID-19 have accelerated and expanded the continued spread of this virus. As a result, the future progression of the pandemic, the definition and establishment of a new normal, and its effects on our business and operations remain uncertain. Current and future government shutdown orders in the U.S. and other countries may impact our discovery programs, preclinical studies and development programs, as well as our supply chain. In addition, any delays in foreign shipments coming into the United States could also impact our preclinical studies, clinical trial plans or other business operations.

We and our contract manufacturing organizations, or CMOs, and contract research organizations, or CROs, may face disruptions that could affect our ability to initiate, conduct and complete discovery or preclinical research activities, or clinical trials. These disruptions may arise from staffing shortages or difficulties in procuring items that are essential for our research and development activities, including, due to shortages arising in raw materials used in the manufacturing of our product candidates, laboratory supplies for our preclinical studies and clinical trials or animals that are used for discovery and/or preclinical testing. For example, as recently as February 2022, the FDA acknowledged the worldwide shortage of non-human primates and released guidance encouraging sponsors of clinical trials to consider the use of other non-rodent species in their non-clinical toxicity studies. However, this guidance is in effect only for the duration of the public health emergency related to COVID-19, as declared by the Department of Health and Human Services and the FDA has announced that it intends to revise and replace this guidance within sixty days following the termination of the public health emergency. As a result, if we were to elect to use an alternate species for our non-clinical toxicity studies under this guidance, it is possible that we might not be able to initiate and conduct the necessary non-clinical toxicity studies in non-human primates in a timely and cost-effective manner in the future if or when this temporary FDA guidance expires. We and our CROs and

CMOs may face disruptions related to our clinical trials arising from potential delays in IND-enabling studies, manufacturing disruptions and/or the ability to obtain necessary institutional review board, or IRB, or other necessary site approvals, as well as other delays at clinical trial sites, including delays related to site staffing. To date, we have experienced delays at certain clinical sites which, due to staffing constraints or other internal policy requirements, have not been able to complete site activation activities or enroll patients in our clinical trial as quickly as likely would have been possible absent the ongoing COVID-19 pandemic.

Further, the impact of the ongoing COVID-19 pandemic to our operations remains uncertain. For example, at various times throughout the COVID-19 pandemic, we have elected to close our office and laboratory spaces in our Watertown, Massachusetts facility or otherwise restrict access to our facility to only that subset of our employees whose work must necessarily be performed in our laboratories, transitioning our remaining employees to work from home. While the ongoing impact of this pandemic remains uncertain, we believe the redundancies we have in place between our China and India based CROs and our Watertown, Massachusetts-based laboratory staff, as well as the transition of employees to remote work arrangements as and when needed, have generally mitigated the impact of these disruptions on our discovery efforts. However, it also remains possible that we will see delays in some of our preclinical research, technical operations, clinical development or other business activities if the spread and resurgence of COVID-19 continue to arise in the greater Boston area and/or in various parts of the world, which may continue until the impacts of COVID-19, and its various variants, are better mitigated and addressed.

The response to the ongoing COVID-19 pandemic may result in the redirection of resources with respect to regulatory and intellectual property matters in a way that would adversely impact our ability to progress regulatory approvals and protect our intellectual property. For example, as of May 26, 2021, the FDA noted that it is continuing to ensure timely reviews of applications for medical products during the ongoing COVID-19 pandemic in line with its user fee performance goals and conducting mission critical domestic and foreign inspections to ensure compliance of manufacturing facilities with FDA quality standards, including for oncology product development. However, FDA may not be able to continue its current pace and review timelines could be extended, including where a pre-approval inspection or an inspection of clinical sites is required and due to the ongoing COVID-19 pandemic and travel restrictions FDA is unable to complete such required inspections during the review period. In 2020 and 2021, a number of companies announced receipt of complete response letters due to the FDA's inability to complete required inspections for their applications. In addition, we may face impediments to regulatory meetings and approvals due to measures intended to limit in-person interactions.

While we were successfully able to raise capital twice during 2020 and once during 2021, the pandemic has caused significant disruptions in the financial markets and may continue to cause these types of disruptions, which could impact our ability to raise additional funds through public offerings and may also contribute to volatility in our stock price and otherwise impact trading in our stock. Moreover, it is possible the pandemic will significantly impact economies worldwide, which could adversely affect our business prospects, financial condition and results of operations. Any significant disruption of global financial markets, reducing our ability to access capital, could negatively affect our liquidity and ability to continue operations.

COVID-19 and actions taken to reduce its spread continue to rapidly evolve. The extent to which COVID-19 may impede the development of our product candidates, reduce the productivity of our employees, disrupt our supply chains, delay our preclinical studies or clinical trials, reduce our access to capital or limit our business development activities, will depend on future developments, which are highly uncertain and cannot be predicted with confidence. To the extent the ongoing COVID-19 pandemic adversely affects our business prospects, financial condition and results of operations, it may also have the effect of heightening many of the other risks described in this "Risk Factors" section, such as those relating to the timing and results of our planned and future clinical trials and our financing needs.

***Our Credit Agreement with Perceptive Credit contains restrictions that limit our flexibility in operating our business.***

In June 2020, we entered into a credit agreement and guaranty, or the Credit Agreement, with Perceptive Credit, an affiliate of Perceptive Advisors LLC, or Perceptive Advisors, which is one of our significant stockholders. The Credit Agreement provided for a \$20.0 million senior secured delayed draw term loan facility, though we only drew down \$12.5 million of that amount and our ability to borrow the remaining \$7.5 million tranche expired in June 2021. The Credit Agreement is secured by a lien on substantially all of our and our subsidiaries' assets, including, but not limited to, shares of our subsidiaries, our current and future intellectual property, insurance, trade and intercompany receivables, inventory and equipment, and contract rights. The Credit Agreement requires us to meet specified minimum cash requirements, as described below, and contains various affirmative and negative covenants that limit our ability to engage in specified types of transactions. These covenants, which are subject to customary exceptions, limit our ability to, without Perceptive Credit's prior written consent, effect any of the following, among other things:

- sell, lease, transfer or otherwise dispose of certain assets;
- acquire another company or business or enter into a merger or similar transaction with third parties;
- incur additional indebtedness;
- make investments;
- enter into certain inbound and outbound licenses of intellectual property, subject to certain exceptions;

- encumber or permit liens on certain assets; and
- pay dividends and make other restricted payments with respect to our common stock.

Our board of directors or management team could believe that taking any one of these actions would be in our best interests and the best interests of our stockholders. If that were the case and if we were unable to complete any of these actions because Perceptive Credit does not provide its consent, that could adversely impact our business, financial condition and results of operations.

In addition, we are required to deposit into controlled accounts all cash or other payments received in respect of any and all of our accounts receivable or any other contract or right and interest and, at all times, to maintain a minimum aggregate balance of \$3.0 million in cash in one or more such controlled accounts. These accounts are required to be maintained as cash collateral accounts securing our obligations under the Credit Agreement. Until our obligations under the Credit Agreement have been discharged, our ability to use the cash amounts held in these controlled accounts in the operation of our business will be limited.

In the event of a default under the Credit Agreement, including, among other things, our failure to make any payment when due or our failure to comply with any provision of the Credit Agreement, subject to customary grace periods, Perceptive Credit could elect to declare all amounts outstanding to be immediately due and payable and terminate all commitments to extend further credit. If we are unable to repay the amounts due under the Credit Agreement, Perceptive Credit could proceed against the collateral granted to it to secure this indebtedness, which could have an adverse effect on our business, financial condition and results of operations.

Perceptive Credit's interests as a lender may not always be aligned with our interests or with Perceptive Advisor's interests as a stockholder. If our interests come into conflict with those of Perceptive Credit, including in the event of a default under the Credit Agreement, Perceptive Credit may choose to act in its self-interest, which could adversely affect the success of our current and future collaborative efforts with Perceptive Advisor.

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

Until the time, if ever, when we can generate substantial revenue from product sales, we expect to finance our cash needs through a combination of equity offerings, debt financings, collaborations, strategic alliances and marketing, distribution or licensing arrangements. Although we may receive potential future payments under our collaborations with Biogen, Roche and Calico, we do not currently have any committed external source of funds. If we raise additional capital through the sale of equity or convertible debt securities, your ownership interest will be diluted and the terms of any securities we may issue in the future may include liquidation or other preferences that adversely affect your rights as a common stockholder. Debt financing and preferred equity financing, if available, may involve agreements that include covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making acquisitions or capital expenditures or declaring dividends. Covenants in the Credit Agreement impose certain limitations and obligations on us, including restrictions on our ability to incur additional debt and to enter into certain business combinations without Perceptive Credit's prior written consent.

If we raise additional funds through collaborations, strategic alliances or marketing, distribution 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 grant licenses on terms that may not be favorable to us.

**Risks related to the discovery and development of our product candidates**

***Our approach to the discovery and development of product candidates based on our TORPEDO platform for targeted protein degradation is unproven, which makes it difficult to predict the time, cost of development and likelihood of successfully developing any products.***

Treating diseases using targeted protein degradation is a new treatment modality. Our future success depends on the successful development of this novel therapeutic approach. Very few small molecule product candidates using targeted protein degradation, such as those developed through our TORPEDO platform, have been tested in humans and none have been approved in the United States or Europe. The data underlying the feasibility of developing these types of therapeutic products is both preliminary and limited. If any adverse learnings are made by other developers of targeted protein degraders, there is a risk that development of our product candidates could be materially impacted. Discovery and development of small molecules that harness the ubiquitin proteasome pathway to degrade protein targets have been impeded largely by the complexities and limited understanding of the functions, biochemistry and structural biology of the specific components of the ubiquitin-proteasome system, including E3 ligases and their required accessory proteins involved in target protein ubiquitination, as well as by challenges of engineering compounds that promote protein-to-protein interactions.

The scientific research that forms the basis of our efforts to develop our degrader product candidates under our TORPEDO platform is ongoing and the scientific evidence to support the feasibility of developing TORPEDO platform-derived therapeutic treatments is both preliminary and limited. Further, certain cancer patients have shown inherent primary resistance to approved drugs that inhibit disease-causing proteins and other patients have developed acquired secondary resistance to these inhibitors. Although we believe our products candidates may have the ability to degrade the specific mutations that confer resistance to currently marketed inhibitors of

disease-causing enzymes, any inherent primary or acquired secondary resistance to our product candidates in patients, or if the scientific research that forms the basis of our efforts proves to be contradicted, would prevent or diminish their clinical benefit.

While we commenced a clinical trial of CFT7455 in June 2021, at this time, we have not yet completed a clinical trial of any product candidate. As a result, we are only starting to assess the safety of CFT7455 in patients and we have not yet assessed the safety of any of our other product candidates in humans. Although some of our product candidates have produced observable results in animal studies, there is a limited safety data set for their effects in animals. In addition, these product candidates may not demonstrate the same chemical and pharmacological properties in humans and may interact with human biological systems in unforeseen, ineffective or harmful ways. As a result, there could be adverse effects from treatment with any of our current or future product candidates that we cannot predict at this time.

Additionally, the regulatory approval process for novel product candidates such as ours can be more expensive and take longer than for other, better-known or extensively studied product candidates. Although other companies are also developing therapeutics based on targeted protein degradation, no regulatory authority has granted approval for any therapeutic of this nature at this time. As a result, it is more difficult for us to predict the time and cost of developing our product candidates and we cannot predict whether the application of our TORPEDO platform, or any similar or competitive protein degradation platforms, will result in the development of product candidates that make it through to marketing approval. Any development problems we experience in the future related to our TORPEDO platform or any of our research programs may cause significant delays or unanticipated costs or may prevent the development of a commercially viable product. Any of these factors may prevent us from completing our preclinical studies or any clinical trials that we may initiate, as well as from commercializing any product candidates we may develop on a timely or profitable basis, if at all.

***We are a clinical stage biotechnology company and, while we commenced a clinical trial of CFT7455 in June 2021 and plan to commence a clinical trial of CFT8634 in the first half of 2022, all of our other product candidates are still in preclinical development or in the discovery stage. If we are unable to advance to clinical development, develop, obtain regulatory approval for and commercialize our product candidates or experience significant delays in doing so, our business may be materially harmed.***

We are a clinical-stage biotechnology company and, while we commenced a clinical trial of CFT7455 in June 2021 and anticipate commencing a clinical trial of CFT8634 in the first half of 2022, all of our other product candidates are currently in preclinical development or in the discovery stage. As a result, their risk of failure is high. We have invested substantially all of our efforts and financial resources in building our TORPEDO platform and identifying and conducting preclinical development of our current product candidates, including our lead programs. Our ability to generate revenue from product sales, which we do not expect will occur for several years, if ever, will depend heavily on the successful development and eventual commercialization of one or more of our product candidates. The success of our product candidates will depend on several factors, including the following:

- sufficiency of our financial and other resources;
- successful initiations and completion of preclinical studies;
- successful submission and clearance of INDs and initiation of clinical trials;
- successful patient enrollment in, and conduct and completion of, clinical trials;
- receipt and related terms of marketing approvals from applicable regulatory authorities;
- obtaining and maintaining patent or trade secret protection and regulatory exclusivity for our product candidates;
- making suitable arrangements with third-party manufacturers for both clinical and commercial supplies of our product candidates;
- developing product candidates that achieve the therapeutic properties desired and appropriate for their intended indications;
- establishing sales, marketing and distribution capabilities and launching commercial sales of our products, if and when approved, whether alone or in collaboration with others;
- acceptance of our products, if and when approved, by patients, the medical community and third-party payors;
- obtaining and maintaining third-party coverage and adequate reimbursement;
- establishing a continued acceptable safety profile of our products and maintaining that profile following approval; and
- effectively competing with other therapies.

If we do not successfully achieve one or more of these factors in a timely manner, or at all, we could experience significant delays or an inability to successfully commercialize our product candidates, which could materially harm our business. Moreover, if we do not receive regulatory approvals, we may not be able to continue our operations.

***We have limited experience as a company in completing IND-enabling preclinical studies, submitting INDs or commencing, enrolling and conducting clinical trials.***

We have limited experience as a company in completing IND-enabling preclinical studies and, while we commenced clinical development of CFT7455 and plan to commence clinical development of CFT8634 in the first half of 2022, we have limited experience as a company in commencing, enrolling and conducting clinical trials. In part because of this lack of experience, we cannot be certain that our preclinical studies will be completed on time, that we will submit INDs in a timely manner, that any INDs we submit will be cleared by the FDA in a timely manner, if at all, or if our planned clinical trials will begin, enroll or be completed on time, if at all. Large-scale clinical trials would require significant additional financial and management resources and reliance on third-party clinical investigators and consultants. Relying on third-party clinical investigators, CROs and consultants may cause us to encounter delays that are outside of our control. In addition, relying on third parties in the conduct of our preclinical studies or clinical trials exposes us to a risk that they may not adequately adhere to study or trial protocols or comply with good laboratory practice or good clinical practice, or GCP, as required for any studies or trials we plan to submit to a regulatory authority. We may be unable to identify and contract with sufficient investigators, CROs and consultants on a timely basis or at all. For each of CFT7455 and CFT8634, we have entered into a master services agreement with a CRO to lead our first-in-human Phase 1/2 clinical trial for the applicable product candidate. There can be no assurance that we will be able to negotiate and enter into additional master services agreements with these or other CROs, if and when necessary for our other product candidates, on terms that are acceptable to us on a timely basis or at all.

***Our preclinical studies and clinical trials may fail to demonstrate adequately the safety, potency and efficacy of any of our product candidates, which would prevent or delay development, regulatory approval and commercialization.***

Before obtaining regulatory approval for the commercial sale of any of our product candidates, we must demonstrate through lengthy, complex and expensive preclinical studies and clinical trials that our product candidates are both safe and effective for use in each target indication. Preclinical and clinical testing is expensive and can take many years to complete. Further, the outcome of these activities is inherently uncertain. Failure can occur at any time during the preclinical study and clinical trial processes and, because most of our product candidates are in an early stage of development and have never been tested in humans, there is a high risk of failure. In addition, because targeted protein degraders are a relatively new class of product candidates, any failures or adverse outcomes in preclinical or clinical testing seen by other developers in this class could materially impact the success of our programs. We may never succeed in developing marketable products.

It is also possible that the results of preclinical studies and early clinical trials of our product candidates may not be predictive of the results of later-stage clinical trials. Although product candidates may demonstrate promising results in preclinical studies and early clinical trials, they may not prove to be effective or safe in subsequent clinical trials. For example, testing on animals occurs under different conditions than testing in humans and, therefore, the results of animal studies may not accurately predict human experience. There is typically an extremely high rate of attrition from the failure of product candidates proceeding through preclinical studies and clinical trials. Product candidates in later stages of clinical trials may fail to show the desired safety, potency, purity and efficacy profile despite having progressed successfully through preclinical studies and/or initial clinical trials. Likewise, early, smaller-scale clinical trials may not be predictive of eventual safety or effectiveness in large-scale pivotal clinical trials. Many companies in the biopharmaceutical industry have suffered significant setbacks in advanced clinical trials due to lack of potency or efficacy, insufficient durability of potency or efficacy or unacceptable safety issues, notwithstanding promising results in earlier trials. Most product candidates that commence preclinical studies and clinical trials are never approved as products.

Additionally, we expect that the first clinical trials for our product candidates will be open-label studies, where both the patient and investigator know whether the patient is receiving the investigational product candidate or either an existing approved drug or placebo. This is the case with the ongoing first-in-human clinical trial of CFT7455 and will be the case with our first-in-human clinical trial of CFT8634, currently planned to commence in the first half of 2022. Most typically, open-label clinical trials test only the investigational product candidate and sometimes 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. 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.

Any preclinical studies or clinical trials that we may conduct may not demonstrate the safety, potency and efficacy necessary to obtain regulatory approval to market our product candidates. If the results of our ongoing or future preclinical studies or clinical trials are inconclusive with respect to the safety, potency and efficacy of our product candidates, if we do not meet the clinical endpoints with statistical and clinically meaningful significance or if there are safety concerns associated with our product candidates, we may be prevented or delayed in obtaining marketing approval for those product candidates. In some instances, there can be significant variability in safety, potency or efficacy results between different preclinical studies and clinical trials of the same product candidate due to numerous factors, including changes in trial procedures set forth in protocols, differences in the size and type of the patient populations, changes in and adherence to the clinical trial protocols and the rate of dropout among clinical trial participants.

While we commenced a clinical trial of CFT7455 in June 2021 and are planning to commence a clinical trial of CFT8634 in the first half of 2022, we have not yet initiated clinical trials for any of our other product candidates. As is the case with all drugs, it is likely

that there may be side effects associated with the use of our product candidates related to on-target, off-target toxicity, or other mechanisms of drug toxicity including chemical-based toxicity. Results of our trials could reveal a high and unacceptable severity and prevalence of side effects of this nature. If unacceptable levels of toxicity are observed or if our product candidates have other characteristics that are unexpected, we may need to abandon their development, modify our development plans as to dose level and/or dose schedule or otherwise, 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. For example, while we commenced our ongoing Phase 1/2 clinical trial of CFT7455 with a starting dose that was informed by the results of our preclinical studies, we observed toxicity in patients enrolled in Cohort A, which caused us to modify how we dose additional patients enrolled in this clinical trial. Further, if we were to observe unacceptable levels of side effects, or if other developers of similar targeted protein degraders were to find an unacceptable severity or prevalence of side effects with their drug candidates, our trials could be suspended or terminated, and the FDA or comparable foreign regulatory authorities could order us to cease further development of or deny approval of our product candidates for any or all targeted indications. Drug-related side effects could also affect patient recruitment or the ability of enrolled patients to complete an ongoing trial or result in potential product liability claims. 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. Any of these occurrences may significantly harm our business, financial condition and prospects.

***Drug development is a lengthy and expensive process with an uncertain outcome. We may incur unexpected costs or experience delays in completing, or ultimately be unable to complete, the development and commercialization of our product candidates.***

While we commenced a clinical trial of CFT7455 in June 2021 and are planning to commence a clinical trial of CFT8634 in the first half of 2022, all of our other product candidates are still in preclinical development or in the discovery stage at this time and the risk of failure for all of our product candidates remains high. We are unable to predict when or if any of our product candidates will prove effective or safe in humans or will receive marketing approval. Before obtaining marketing approval from regulatory authorities for the sale of any product candidate, we must conduct extensive clinical trials to demonstrate the safety and efficacy of our product candidates in humans. Before we can commence clinical trials for a product candidate, we must complete extensive preclinical studies that support our planned INDs in the United States or similar applications in other jurisdictions. We cannot be certain of the timely initiation, completion or outcome of our preclinical studies and, other than in the cases of CFT7455 and CFT8634, where the FDA has cleared the IND for our planned first-in-human study of the drug candidate, we cannot predict if the FDA or similar regulatory authorities outside the United States will allow us to commence our proposed clinical trials or if the outcome of our preclinical studies ultimately will support the further development of any of our product candidates.

Clinical testing is expensive, difficult to design and implement, can take many years to enroll and complete and is uncertain as to the timing and outcome. A failure of one or more clinical trials can occur at any stage of the process. We may experience numerous unforeseen events during or as a result of clinical trials, which could delay or prevent our ability to receive marketing approval or commercialize our product candidates, including:

- delays in reaching, or the failure to reach, a consensus with regulators on clinical trial design or the inability to produce acceptable preclinical results to enable entry into human clinical trials;
- the supply or quality of our product candidates or other materials necessary to conduct clinical trials may be insufficient or inadequate, including as a result of delays in the testing, validation, manufacturing and delivery of product candidates to the clinical sites by us or by third parties with whom we have contracted to perform certain of those functions;
- delays in reaching, or the failure to reach, agreement on acceptable clinical trial contracts or clinical trial protocols with prospective trial sites or CROs;
- the failure of regulators or IRBs to authorize us or our investigators to commence a clinical trial or conduct a clinical trial at a prospective trial site;
- difficulty in designing clinical trials and in selecting endpoints for diseases that have not been well studied and for which the natural history and course of the disease is poorly understood;
- the selection of certain clinical endpoints that may require prolonged periods of clinical observation or analysis of the resulting data;
- the number of patients required for clinical trials of our product candidates may be larger than we anticipate, enrollment in these clinical trials may be slower than we anticipate, participants may drop out of these clinical trials at a higher rate than we anticipate or fail to return for post-treatment follow-up or the failure to recruit suitable patients to participate in our clinical trials;
- our product candidates may have undesirable side effects or other unexpected characteristics, causing us or our investigators, regulators or IRBs to suspend or terminate our clinical trials;
- we may have to suspend or terminate clinical trials of our product candidates for various reasons, including a finding that the participants are being exposed to unacceptable health risks;

- the third parties with whom we contract may fail to comply with regulatory requirements or meet their contractual obligations to us in a timely manner, or at all;
- the requirement from regulators or IRBs that we or our investigators suspend or terminate clinical trials for various reasons, including noncompliance with regulatory requirements or unacceptable safety risks;
- clinical trials of our product candidates may produce negative or inconclusive results and we may decide, or regulators may require us, to conduct additional clinical trials, modify our development plans as to dose level and/or dose schedule or otherwise, or abandon product development programs;
- the cost of clinical trials of our product candidates may be greater than we anticipate;
- clinical staffing shortages, including but not limited to the lack of appropriately trained or experienced clinical research associates or medical staff at the institutions where we conduct our clinical trials, may cause delays or create other challenges to the timely and efficient conduct of 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, trial sites or manufacturing facilities;
- occurrence of serious adverse events associated with the product candidate that are viewed to outweigh its potential benefits; and
- disruptions caused by the continuing spread and effects of the ongoing COVID-19 pandemic may increase the likelihood that we encounter these types of difficulties or cause other delays in initiating, enrolling, conducting or completing our planned clinical trials.

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

- be delayed in obtaining marketing approval for our product candidates, 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 required to perform additional clinical trials to support marketing approval;
- have regulatory authorities withdraw or suspend their approval, or impose restrictions on distribution of a product candidate in the form of a modified risk evaluation and mitigation strategy, or REMS;
- be subject to additional post-marketing testing requirements or changes in the way the product is administered; or
- have our product removed from the market after obtaining marketing approval.

Our product development costs also will increase if we experience delays in preclinical studies or clinical trials or in obtaining marketing approvals. While we commenced a clinical trial for CFT7455 in June 2021 and are planning to commence a clinical trial of CFT8634 in the first half of 2022, we do not know whether any of our other preclinical studies or clinical trials will begin as planned, will need to be restructured or will be completed on schedule, or at all. Significant preclinical study or clinical trial delays also could shorten any periods during which we may have the exclusive right to commercialize our product candidates or could allow our competitors to bring products to market before we do and impair our ability to successfully commercialize our product candidates, which may harm our business, results of operations, financial condition and prospects.

Further, cancer therapies sometimes are characterized as first-line, second-line or third-line. The FDA often approves new oncology therapies initially only for third-line or later use, meaning for use after two or more other treatments have failed. When cancer is detected early enough, first-line therapy, usually systemic anti-cancer therapy (e.g., chemotherapy), surgery, radiation therapy or a combination of these, is sometimes adequate to cure the cancer or prolong life without a cure. Second-line and third-line therapies are administered to patients when prior therapy has been shown to not be effective. Our ongoing clinical trial for CFT7455 and our anticipated clinical trials for CFT8634 as well as our other drug candidates will be with patients who have received one or more prior treatments and we expect that we would initially seek regulatory approval of these product candidates for second-line or third-line therapy. Subsequently, for those products that prove to be sufficiently beneficial, if any, we would expect to seek approval potentially as a first-line therapy, but any product candidates we develop, even if approved for second-line or third-line therapy, may not be approved for first-line therapy and, prior to seeking and/or receiving any approvals for first-line therapy, we may have to conduct additional clinical trials.

***Targeted protein degradation is a novel modality that continues to attract substantial interest from existing and emerging biotechnology and pharmaceutical companies. As a result, we face substantial competition, which may result in others discovering, developing or commercializing products for the same indication and/or patient population before or more successfully than we do.***

The biotechnology and pharmaceutical industries are characterized by rapidly advancing technologies, intense competition and a strong emphasis on proprietary products. We face, and will continue to face, competition from third parties that use protein degradation, antibody therapy, inhibitory nucleic acid, immunotherapy, gene editing or gene therapy development platforms and from companies focused on more traditional therapeutic modalities, such as small molecule inhibitors. The competition we face and will face is likely to come from multiple sources, including major pharmaceutical, specialty pharmaceutical and biotechnology companies, academic institutions, government agencies and public and private research institutions.

Targeted protein degradation is an emerging therapeutic modality that has the potential to deliver therapies that improve outcomes for patients. As a result, a number of biotechnology and pharmaceutical companies are already working to develop degradation-based therapies and the number of companies entering this space continues to increase. We are aware of several biotechnology companies developing product candidates based on chimeric small molecules for targeted protein degradation including Arvinas, Inc., BioTheryX, Inc., Cullgen Inc., Foghorn Therapeutics, Inc., Frontier Medicines Corporation, Kymera Therapeutics, Inc., Lycia Therapeutics, Inc., Monte Rosa Therapeutics, Inc., NeoMorph Inc., Nurix Therapeutics, Inc., Proteovant Therapeutics, Inc., Treeline Biosciences, Inc., Triana Biomedicines, Inc., and Vividion Therapeutics, Inc. (a subsidiary of Bayer AG). Further, several large pharmaceutical companies have disclosed investments and research in this field, including Amgen Inc., AstraZeneca plc, Bristol-Myers Squibb Company (and its subsidiary Celgene Corporation), GlaxoSmithKline plc, Genentech, Inc. and Novartis International AG. In addition to competition from other protein degradation therapies, any products that we develop may also face competition from other types of therapies, such as small molecule, antibody, T cell or gene therapies. For example, we understand that Adaptimmune Limited, Foghorn Therapeutics, Inc. and GlaxoSmithKline plc are pursuing the development of therapies for patients with synovial sarcoma.

Many of our current or potential competitors, either alone or with their collaboration partners, have significantly greater financial resources and expertise in research and development, manufacturing, preclinical testing, conducting clinical trials, obtaining regulatory approvals and marketing approved products than we do. These competitors also compete with us in recruiting and retaining qualified scientific and management personnel and establishing clinical trial sites and patient registration for clinical trials, as well as in acquiring technologies complementary to, or necessary for, our product candidates. Mergers and acquisitions in the pharmaceutical and biotechnology industries may result in even more resources being concentrated among a smaller number of our competitors, the scale of which could be difficult to compete against. Smaller or early-stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large and established companies. Our commercial opportunity could be reduced or eliminated if our 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 product candidate that we may develop. Our competitors also may obtain FDA or other regulatory approval for their product candidates more rapidly than we may obtain approval for ours, which could result in our competitors establishing a strong market position before we are able to enter the market. In addition, our ability to compete may be affected in many cases by insurers or other third-party payors seeking to encourage the use of generic products. There are generic products currently on the market for certain of the indications that we are pursuing, and additional products are expected to become available on a generic basis over the coming years. If our product candidates are approved, we expect that they will be priced at a significant premium over competitive generic products.

***Our ability to use our net operating loss carryforwards and research and development tax credit carryforwards may be limited.***

As of December 31, 2021, we had \$152.2 million federal net operating loss carryforwards and \$216.8 million gross in United States state net operating loss carryforwards, portions of which expire at various dates through 2041. Under legislation enacted in 2017, informally titled the Tax Cuts and Jobs Act, or the TCJA, as modified by the Coronavirus Aid, Relief and Economic Security Act, or the CARES Act, federal net operating losses generated in tax years beginning after 2017, if any, will not expire and may be carried forward indefinitely, but the deductibility of such federal net operating losses in tax years beginning after December 31, 2020 will be limited to the lesser of the net operating loss carryover or 80% of the corporation's adjusted taxable income (subject to Section 382 of the Internal Revenue Code of 1986, as amended). The CARES Act temporarily allows us to carryback net operating losses arising in 2018, 2019 and 2020 to the five prior years. It is uncertain how various states will respond to the TCJA, the CARES Act or any newly enacted federal tax law. In addition, at the state level, there may be periods during which the use of net operating losses is suspended or otherwise limited, which could accelerate or permanently increase state taxes owed.

As of December 31, 2021, we also had United States federal and state research and development tax credit carryforwards of \$7.3 million and \$2.9 million, respectively, which expire at various dates through 2041. These tax credit carryforwards could expire unused and be unavailable to offset our future income tax liabilities.

In addition, under Sections 382 and 383 of the Internal Revenue Code of 1986, as amended, or the Code, and corresponding provisions of state law, if a corporation undergoes an "ownership change," which is generally defined as a greater than 50% change, by value, in its equity ownership over a three-year period, the corporation's ability to use its pre-change net operating loss carryforwards and other pre-change tax attributes to offset its post-change income or taxes may be limited. In 2021, we completed a study of ownership changes from inception through December 31, 2020, which concluded that we experienced ownership changes as

defined by Section 382 of the Code. However, there were no net operating loss carryforwards that were limited or expired unused. We have not updated the study to assess whether a change of ownership has occurred through 2022. We may have experienced additional ownership changes that have not been identified that could result in the expiration of our net operating loss and tax credit carryforwards before utilization and we may experience subsequent shifts in our stock ownership, some of which are outside our control. As a result, if we earn net taxable income and determine that an ownership change has occurred and our ability to use our historical net operating loss and tax credit carryforwards is materially limited, that will harm our future operating results by effectively increasing our future tax obligations.

***We may not be able to file INDs 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 with our planned clinical trials.***

We have limited experience as a company in preparing, submitting to and receiving clearance from the FDA on INDs. We submitted our first IND to the FDA in December 2020 for CFT7455 and, in January 2021, the FDA informed us that we are permitted to proceed with our first-in-human clinical trial for this product candidate. We submitted an IND for CFT8634 in November 2021 and, in December 2021, the FDA informed us that we are permitted to proceed with our first-in-human clinical trial for this product candidate. We plan to submit an IND application and begin a Phase 1 trial in BRAF V600X driven cancers including melanoma, colorectal and NSCLC for CFT1946 in the second half of 2022, and plan to submit an IND application and begin a Phase 1 trial for CFT8919 in EGFR L858R NSCLC in 2023. While these are our current expectations, we may not be able to file our planned INDs or INDs for other product candidates on the timelines we expect. For example, we may experience manufacturing delays or other delays with IND-enabling studies, we may determine that additional IND-enabling studies are warranted, or we may face delays due to the ongoing global COVID-19 pandemic. Moreover, we cannot be sure that submission of an IND will result in the FDA allowing us to commence clinical trials or that, once begun, issues will not arise that lead to the suspension or termination of our clinical trials. Additionally, even if the applicable regulatory authorities agree with the design and implementation of the clinical trials set forth in our INDs, we cannot guarantee that those regulatory authorities will not change their requirements in the future. These considerations apply to the INDs described above and also to new clinical trials we may submit as amendments to existing INDs or as part of new INDs in the future. Any failure to file INDs on the timelines we expect or to obtain regulatory approvals for our clinical trials may prevent us from completing our clinical trials or commercializing our products on a timely basis, if at all.

***If serious adverse events, undesirable side effects or unexpected characteristics are identified during the development of any product candidates we may develop, we may need to modify, abandon or limit our further clinical development of those product candidates.***

While we commenced a clinical trial of CFT7455 in June 2021 and are planning to commence a clinical trial of CFT8634 in the first half of 2022, all of our other product candidates are still in the preclinical or discovery stages at this time, which means that we have not yet evaluated any of our other product candidates in human clinical trials. It is impossible to predict when or if any product candidates we may develop will prove safe in humans. There can be no assurance that any of the product candidates developed through our TORPEDO platform will not cause undesirable side effects, which could arise at any time during preclinical or clinical development.

A potential risk with product candidates developed through our TORPEDO platform, or in any protein degradation product candidate, is that healthy proteins or proteins not targeted for degradation will be degraded or that the degradation of the targeted protein in and of itself could cause adverse events, undesirable side effects or unexpected characteristics. There is also the potential risk of delayed adverse events following treatment using product candidates developed through our TORPEDO platform.

If any product candidates we develop are associated with serious adverse events or undesirable side effects or have other characteristics that are unexpected, we may need to abandon their development, modify our development plans as to dose level and/or dose schedule or otherwise, or limit development to certain uses or subpopulations in which the adverse events, undesirable side effects or other characteristics are less prevalent, less severe or more acceptable from a risk-benefit perspective. The occurrence of any of these sorts of events would have an adverse effect on our business, financial condition, results of operations and prospects. Many product candidates that initially showed promise in early-stage testing for treating cancer or other diseases have later been found to cause side effects that prevented further clinical development of the product candidates or limited their competitiveness in the market. For example, single agent BRAF inhibitors can cause a secondary malignancy called keratocanthoma, which is a skin cancer caused by paradoxical activation of BRAF upon inhibitor binding.

***The results of preclinical studies may not be predictive of future results in later studies or trials. Initial success in clinical trials may not be indicative of results obtained when these trials are completed or in later stage clinical trials.***

The results of preclinical studies may not be predictive of the results of clinical trials and the results of early-stage clinical trials may not be predictive of the results of later-stage clinical trials. In addition, initial success in clinical trials may not be indicative of results obtained when those trials are completed or in later stage clinical trials. In particular, the small number of patients in our planned early clinical trials or the designs of these trials may make the results of these trials less predictive of the outcome of later clinical trials. For example, we commenced Cohort A of the ongoing Phase 1/2 clinical trial of CFT7455 with a starting dose that was based on the results of our preclinical studies; however, we have modified dosage for Cohorts B1 and C based on the results from Cohort A. Moreover, even if successful, the results of the dose escalation portion of our ongoing and planned first-in-human Phase 1/2 clinical

trials of our product candidates may not be predictive of the results of further clinical trials of these product candidates or any of our other product candidates. Moreover, preclinical and clinical data are often susceptible to varying interpretations and analyses 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. Our clinical trials may not ultimately be successful or support further clinical development of any of our product candidates. There is a high failure rate for product candidates proceeding through clinical trials. Many companies in the pharmaceutical and biotechnology industries have suffered significant setbacks in clinical development even after achieving encouraging results in earlier studies. Any setbacks of this nature in our clinical development could materially harm our business, financial condition, results of operations and prospects. In addition, we may conduct some of our clinical trials in a combination Phase 1/2 design and, if the Phase 1 portion of the trial is not successful, we will not be allowed to proceed into the Phase 2 portion of the trial.

***If we experience delays or difficulties in the enrollment of patients in our clinical trials, our timelines for submitting for and receiving necessary marketing approvals could be delayed or prevented.***

We may not be able to initiate clinical trials for our product candidates if we are unable to locate and enroll a sufficient number of eligible patients to participate in these trials, as required by the FDA or similar regulatory authorities outside of the United States. In June 2021, we advanced CFT7455 into first-in-human Phase 1/2 clinical trials in MM and NHLs, including PTCL and MCL, and anticipate advancing CFT8634 into clinical trials in the first half of 2022. In addition, we are planning to advance our other lead product candidates into first-in-human Phase 1/2 clinical trials. While we believe that we will be able to enroll a sufficient number of patients into each of these clinical trials, we cannot predict with certainty how difficult it will be to enroll patients for trials, some of which are in rare indications, either generally or specifically during the ongoing COVID-19 pandemic. Our ability to identify and enroll eligible patients for clinical trials of our product candidates may turn out to be limited or we may be slower in enrolling these trials than we anticipate. In addition, some of our competitors have ongoing clinical trials for product candidates that treat the same indications as our product candidates and, as a result, patients who would be eligible for our clinical trials may instead elect to enroll in clinical trials of our competitors' product candidates. Patient enrollment in clinical trials is also affected by other factors including:

- the severity of the disease under investigation;
- the eligibility criteria for the trial in question;
- the perceived risks and benefits of the product candidates offered in the clinical trials;
- the efforts to facilitate timely enrollment in clinical trials;
- the patient referral practices of physicians;
- the availability of suitable and sufficient staffing at clinical trial sites;
- the burden on patients due to the scope and invasiveness of required procedures under clinical trial protocols, some of which may be inconvenient and/or uncomfortable;
- the ability to monitor patients adequately during and after treatment;
- the proximity and availability of clinical trial sites for prospective patients; and
- the impact of the ongoing COVID-19 pandemic, which may affect the conduct of a clinical trial, including by slowing potential enrollment or reducing the number of eligible patients for clinical trials or by interfering with patients' ability to return to the clinical trial site for required monitoring, procedures or follow-up.

Our inability to enroll a sufficient number of patients for our planned clinical trials, or our inability to do so on a timely basis, would result in significant delays and could require us to abandon one or more clinical trials altogether. Enrollment delays in our clinical trials may also result in increased development costs for our product candidates, which would cause the value of our company to decline and limit our ability to obtain additional financing.

***The conclusions and analysis drawn from announced or published interim top-line and preliminary data from our clinical trials from time to time may change as more patient data become available. Further, all interim data that we provide remains subject to audit and verification procedures that could result in material changes in the final data.***

From time to time, we may publish interim top-line or preliminary data from our clinical trials. Interim data from clinical trials that we may conduct 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. In addition, preliminary or top-line data also remain subject to audit and verification procedures that may result in the final data being different, potentially in material ways, from the preliminary data we previously announced or published. As a result, interim and preliminary data should be viewed with caution until final data are available. Adverse differences between preliminary or interim data and final data could significantly harm our reputation, business, financial condition, results of operations and 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 focus on research programs and product candidates that we identify for 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. 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 research and development programs and product candidates for specific indications may not yield any commercially viable products. If we do not accurately evaluate the commercial potential or target market for a particular product candidate, we may relinquish valuable rights to that product candidate through collaboration, licensing or other royalty arrangements in cases in which it would have been more advantageous for us to retain sole development and commercialization rights to such product candidate.

***We may develop CFT7455 in combination with other drugs for MM. If the FDA or similar regulatory authorities outside of the United States do not approve these other drugs, revoke their approval of these other drugs or if safety, efficacy, manufacturing or supply issues arise with the drugs we choose to evaluate in combination with CFT7455, we may be unable to obtain approval of or market CFT7455.***

Once a recommended dose is identified from the dose escalation portion of our first-in-human Phase 1/2 clinical trial of CFT7455 for the treatment of MM, we plan to conduct a portion of that clinical trial in combination with a dexamethasone inhibitor. We did not develop or obtain marketing approval for, nor do we manufacture or sell, any of the currently approved drugs that we may study in combination with CFT7455. If the FDA or similar regulatory authorities outside of the United States revoke their approval of the drug or drugs we intend to deliver in combination with CFT7455, we will not be able to market CFT7455 in combination with those revoked drugs.

If safety or efficacy issues arise with any of these drugs, we could experience significant regulatory delays and the FDA or similar regulatory authorities outside of the United States may require us to redesign or terminate certain of our clinical trials. If the drugs we use are replaced as the standard of care for the indications we choose for CFT7455, the FDA or similar regulatory authorities outside of the United States may require us to conduct additional clinical trials. In addition, if manufacturing or other issues result in a shortage of supply of the drugs with which we determine to combine with CFT7455, we may not be able to complete clinical development of CFT7455 on our current timeline or at all.

Even if CFT7455 were to receive marketing approval or be commercialized for use in combination with other existing drugs, we would continue to be subject to the risks that the FDA or similar regulatory authorities outside of the United States could revoke approval of the drugs used in combination with CFT7455 or that safety, efficacy, manufacturing or supply issues could arise with these existing drugs.

Combination therapies are commonly used for the treatment of cancer and we would be subject to similar risks if we were to elect to develop any of our other product candidates for use in combination with other drugs or for indications other than cancer. This could result in our own products being removed from the market or being less successful commercially.

***We may not be successful in our efforts to identify or discover additional potential product candidates.***

While our four lead programs are focused on oncology targets, a key element of our strategy is to apply our TORPEDO platform to develop product candidates that address a broad array of targets and new therapeutic areas, such as neurodegeneration, diseases of aging and infectious disease. The therapeutic discovery activities that we are conducting may not be successful in identifying product candidates that are useful in treating cancer or other diseases. Our research programs may initially show promise in identifying potential product candidates, yet fail to yield product candidates for clinical development for a number of reasons, including:

- potential product candidates may, on further study, be shown to have harmful side effects or other characteristics that indicate that they are unlikely to be drugs that will receive marketing approval or achieve market acceptance;
- potential product candidates may not be effective in treating their targeted diseases; or
- the market size for the target indications of a potential product candidate may diminish over time due to improvements in the standard of care to the point that further development is not warranted.

Research programs to identify new product candidates require substantial technical, financial and human resources. We may choose to focus our efforts and resources on a potential product candidate that ultimately proves to be unsuccessful. If we are unable to identify suitable product candidates for preclinical and clinical development, we will not be able to obtain revenues from sale of products in future periods, which likely would result in significant harm to our financial position and adversely impact our stock price.

***If we do not achieve our projected development goals in the timeframes we announce and expect, the commercialization of our products may be delayed and, as a result, our stock price may decline.***

From time to time, we may estimate the timing of the anticipated accomplishment of various scientific, clinical, regulatory and other product development goals, which we sometimes refer to as milestones. These milestones may include the commencement or completion of preclinical studies and clinical trials and the submission of regulatory filings. From time to time, we may publicly

announce the expected timing of some of these milestones. Each of these milestones is and will be based on numerous assumptions. The actual timing of these milestones can vary dramatically compared to our estimates, in some cases for reasons beyond our control. If we do not meet these milestones as publicly announced, or at all, our revenue may be lower than expected or the commercialization of our products may be delayed or never achieved and, as a result, our stock price may decline.

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

***We expect to rely on third parties to conduct our future clinical trials and those third parties may not perform satisfactorily, including failing to meet deadlines for the completion of such trials.***

We currently rely on CROs to conduct our clinical trials, as we currently do not plan to independently conduct clinical trials of any of our product candidates. Additionally, we must contract with third-party research sites for the conduct of our clinical trials. Our agreements with these CROs and sites might terminate for a variety of reasons, including a failure to perform by the third parties. If we were ever to need to enter into alternative arrangements, we would experience delays in our product development activities.

Our reliance on CROs for research and development activities will reduce our control over these activities but will not relieve us of our responsibilities for how these activities are performed. For example, we will remain responsible for ensuring that each of our clinical trials is conducted in accordance with the general investigational plan and protocols in the applicable IND. Moreover, the FDA requires compliance with standards, commonly referred to as GCPs, for conducting, recording and reporting the results of clinical trials to assure that data and reported results are credible and accurate and that the rights, integrity and confidentiality of trial participants are protected. GCP compliance extends not only to sponsors of clinical research but also to third parties including CROs and sites involved in the conduct of clinical research.

Further, these CROs or sites may have relationships with other entities, some of which may be our peers or competitors. If the CROs or sites with whom we work do not successfully carry out their contractual duties, meet expected deadlines or conduct our clinical trials in accordance with regulatory requirements or our stated protocols, we will not be able to obtain, or may be delayed in obtaining, marketing approvals for our product candidates and will not be able to, or may be delayed in our efforts to, successfully commercialize our product candidates.

***Manufacturing pharmaceutical products is complex and subject to product delays or loss for a variety of reasons. We contract with third parties for the manufacture of our product candidates for preclinical testing and clinical trials and expect to continue to do so for commercialization. This reliance on third parties increases the risk that we will not have sufficient quantities of our product candidates or products or that we will not have the quantities we desire or require at an acceptable cost or quality or at the right time, which could delay, prevent or impair our development or commercialization efforts.***

We do not own or operate, and currently have no plans to establish, any manufacturing facilities. We rely on and expect to continue to rely on CMOs for both drug substance and finished drug product. This reliance on third parties may increase the risk that we will not have sufficient quantities of our product candidates or products or that we will not have the quantities we desire or require at an acceptable cost or quality, which could delay, prevent or impair our development or commercialization efforts, including where a pre-approval inspection or an inspection of manufacturing sites is required and due to the ongoing COVID-19 pandemic and travel restrictions, if still then in effect, FDA is unable to complete those required inspections during the review period.

We may be unable to establish agreements with CMOs or to do so on acceptable terms. Even if we are able to establish agreements with CMOs, reliance on third-party manufacturers entails additional risks, including:

- reliance on the third party for regulatory, compliance, quality assurance and manufacturing success;
- the possible breach of the manufacturing agreement by the third-party CMO;
- the possible risk that the CMO will cease offering the services we require or shut down operations altogether, either temporarily or permanently, due to a regulatory concern, financial insolvency, non-compliance with applicable law or another reason;
- the possible misappropriation of our proprietary information, including our trade secrets and know-how; and
- the possible termination or non-renewal of the agreement by the third party at a time that is costly or inconvenient for us or the inability of the CMO to provide us with a manufacturing slot when we need it.

We have only limited technology transfer agreements in place with respect to our product candidates and these existing arrangements do not extend to commercial supply. We acquire many key materials on a purchase order basis. As a result, we do not have long-term committed arrangements with respect to our product candidates and other materials. If we receive marketing approval for any of our product candidates, we will need to establish or have established an agreement for commercial manufacture with one or more third parties.

Third-party manufacturers may not be able to comply with current good manufacturing practices, or cGMP, regulations or similar regulatory requirements outside of the United States. Some of our molecules are highly potent and, in the absence of additional safety data, they receive a high occupational exposure band, or OEB. These assigned OEBs dictate the containment and other precautions

that must be taken as part of the manufacture of our product candidates and, for molecules with high OEB designations, serve to limit the number of CMOs who are qualified to manufacture our molecules. Our failure, or the failure of our CMOs, to comply with applicable regulations, including the ability of our CMOs to work with our highly potent materials and the safety protocols in connection therewith, could result in sanctions being imposed on us, including clinical holds, fines, injunctions, civil penalties, delays, suspension or withdrawal of approvals, license revocation, seizures or recalls of product candidates or products, operating restrictions and criminal prosecutions, any of which could significantly and adversely affect supplies of our products.

Our product candidates and any products that we may develop may compete with other product candidates and products for access to manufacturing facilities. As a result, we may not obtain access to these facilities on a priority basis or at all. There are a limited number of manufacturers that operate under cGMP regulations and that might be capable of manufacturing for us, particularly, in some cases, given the potency of our compounds.

Any performance failure or delay in performance on the part of our existing or future manufacturers could delay clinical development or marketing authorization. For example, in the past, our contract drug product manufacturer had a mechanical issue arise in connection with a manufacturing step for a manufacturing run for our CFT7455 product candidate. While this issue did not ultimately delay the timing of submission of our IND for CFT7455, in the future, we could experience a manufacturing issue that would have a material impact on development of our product candidates and the occurrence of an event of this nature would largely be outside of our control. We do not currently have arrangements in place for redundant supply or a second source for drug substance or drug product. If our current CMOs cannot perform as agreed, we may be required to replace them. Although we believe that there are several potential alternative manufacturers who could manufacture our product candidates, we may incur added costs and delays in identifying and qualifying any replacement manufacturers or we may not be able to reach agreement with any alternative manufacturer. While we have identified alternate vendors for some of the manufacturing work related to CFT7455 and CFT8634, switching vendors could result in significant additional costs of materials and significant delays to our operations and we may be constrained in the vendors we can select, particularly for compounds that have high OEB designations.

Our current and anticipated future dependence upon others for the manufacture of our product candidates or products may adversely affect our future profit margins and our ability to commercialize any products that receive marketing approval on a timely and competitive basis.

Additionally, we currently rely on single source suppliers for a portion of our supply chain for our preclinical and clinical trial supplies. If our current or future suppliers, whether for raw materials, drug substance or drug product, are unable to supply us with sufficient materials for our preclinical studies and clinical trials, we may experience delays in our development efforts as we locate and qualify new suppliers or manufacturers.

The third-party manufacturers on whom we rely may incorporate their own proprietary processes into our product candidate manufacturing processes. We have limited control and oversight of a third party's proprietary manufacturing processes. If a third-party manufacturer were to modify its processes, those modifications could negatively impact our manufacturing, including product loss or failure that requires additional manufacturing runs or a change in manufacturer, both of which could significantly increase the cost of and significantly delay the manufacture of our product candidates.

As our product candidates progress through preclinical studies and clinical trials towards approval and commercialization, we expect that various aspects of the product development and manufacturing process will evolve in an effort to optimize processes and results. Some of those product and manufacturing process changes may involve the use of third-party proprietary technology, which could then cause us to need to obtain a license from third parties. In addition, these types of changes may require that we make amendments to our regulatory applications, which could further delay the timeframes under which modified manufacturing processes can be used for any of our product candidates.

In addition, as we advance our product candidates into later stage clinical trials and plan for the potential commercialization of our product candidates, we may determine that it is necessary or appropriate to bring on additional suppliers of drug product and/or drug substance, which could result in changes to the manufacturing processes for our product candidates and may require us to provide additional information to regulatory authorities. If we were to bring on additional CMOs for our product candidates, we may also be required to demonstrate analytical comparability and/or conduct additional bridging studies or trials, all of which would require additional time and expense.

***We have existing collaborations with third parties under which we are engaged in the research, development and commercialization of certain product candidates. If any of these collaborations are not successful, we may not be able to capitalize on the market potential of those product candidates. In addition, these collaborations could impact our intellectual property rights.***

Previously, we entered into the following collaborations, which involve our research programs:

- a collaboration agreement with Roche in December 2015, which we amended and restated in December 2018 and further amended in November 2020 and updated as to included targets in November 2021;
- a collaboration agreement with Calico in March 2017, which was extended in respect of one program in September 2021; and
- a collaboration agreement with Biogen in December 2018, which was amended in February 2020.

Under these collaboration agreements, we are generally responsible for developing drug candidates leveraging our TORPEDO platform based on partner-selected targets. Further, these agreements provide that our collaboration partners have exclusive rights to develop degraders for their selected and reserved targets. As a result, we are not permitted to pursue a target of potential interest – either alone or with another partner – while that target is bound by these restrictions.

Further, if our collaborations do not result in the successful development and commercialization of products or if one of our collaborators terminates its agreement with us or elects not to pursue a program within a collaboration, we may not receive any future research funding or milestone or royalty payments under that collaboration or in respect of that terminated program. If that were to happen, we might decide to abandon the program or to move the program forward on our own, which would require us to devote additional resources to the program on a going-forward basis. In addition, if one of our collaborators terminates its agreement with us generally or with respect to a specific target, which they are permitted to do for convenience with between 90 and 270 days' notice or in connection with a material breach of the agreement by us that remains uncured for a specified period of time, we may find it more difficult to attract new collaborators and our development programs may be delayed or the perception of us in the business and financial communities could be adversely affected. All of the risks relating to product development, marketing approval and commercialization described in this report apply to the activities of our collaborators.

It is also possible that our collaborators may not properly obtain, maintain, enforce or defend the intellectual property or proprietary rights arising out of our licensed programs or may use our proprietary information in a way that could jeopardize or invalidate our proprietary information or expose us to potential litigation. For example, Roche, Biogen and Calico have the first right to enforce, and Roche also has the first right to defend, certain intellectual property rights under the applicable collaboration arrangement with respect to particular licensed programs and, although we may have the right to assume the enforcement and defense of these intellectual property rights if our collaborator does not, our ability to do so may be compromised by their actions. In addition, if any licensed program were later to revert to us, our ability to protect any intellectual property or other proprietary rights associated with that program would be impacted by the intellectual property filings made or other steps taken by our collaborator prior to program reversion. Further, our collaborators may own or co-own intellectual property covering our products that results from our collaborating with them and, in cases where that applies, we would not have the exclusive right to commercialize the collaboration intellectual property.

***We may form or seek collaborations or strategic alliances or enter into additional licensing arrangements in the future and we may not realize the benefits of those collaborations, alliances or licensing arrangements.***

We may in the future form or seek strategic alliances, create joint ventures or collaborations or enter into additional licensing arrangements with third parties that we believe will complement or augment our development and commercialization efforts with respect to our product candidates and any future product candidates that we may develop. Our likely collaborators in any other collaboration arrangements we may enter into include large and mid-size pharmaceutical companies and biotechnology companies. However, it is possible that we will not be able to enter into a collaboration agreement of this nature or that the terms of any potential new collaboration arrangement may not be favorable.

For example, we may seek to enter into collaboration arrangements to advance our CFT7455 product candidate in MM or other indications or we may form or seek to form collaboration arrangements to enable our development and commercialization of a product candidate in a specified geographic area. Any of these relationships may require us to incur non-recurring and other charges, increase our near and long-term expenditures, issue securities that dilute our existing stockholders or disrupt our management and business.

In addition, we face significant competition in seeking appropriate strategic partners and the negotiation process for these sorts of transactions is time-consuming, complex and expensive. Moreover, we may not be successful in our efforts to establish a strategic partnership or other alternative arrangements for our product candidates because they may be deemed to be at too early of a stage of development for collaborative effort and third parties may not view our product candidates as having the requisite potential to demonstrate safety, potency, purity and efficacy and obtain marketing approval. Additionally, our existing partners may decide to acquire or partner with other companies developing targeted protein degraders or directed at the targets or indications to which our product candidates are directed, which may have an adverse impact on our business prospects, financial condition and results of operations.

As a result, if we enter into additional collaboration agreements and strategic partnerships or license our product candidates, we may not be able to realize the benefit of those transactions if we are unable to successfully integrate them with our existing operations and company culture.

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

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

If any of our product candidates receives marketing approval, it may nonetheless fail to gain sufficient market acceptance by physicians, patients, third-party payors and others in the medical community. For example, current cancer treatments, such as chemotherapy and radiation therapy, are well-established in the medical community and doctors may continue to rely on these treatments. If our product candidates do not achieve an adequate level of acceptance, we may not generate significant revenue from

product sales and we may not become profitable. The degree of market acceptance of our product candidates, if approved for commercial sale, will depend on a number of factors, including:

- the efficacy and potential advantages compared to alternative treatments;
- the prevalence and severity of any side effects, in particular compared to alternative treatments;
- our ability to offer our products for sale at competitive prices;
- the convenience and ease of administration compared to alternative treatments;
- the willingness of the target patient population to try new therapies and of physicians treating these patients to prescribe these therapies;
- the strength of marketing, sales and distribution support;
- the availability of third-party insurance coverage and adequate reimbursement;
- the timing of any marketing approval in relation to other product approvals;
- support from patient advocacy groups; and
- any restrictions on the use of our products together with other medications.

***As a company, we currently have no marketing and sales organization and no experience in marketing products. If we are unable to establish marketing and sales capabilities or enter into agreements with third parties to market and sell our product candidates, if approved, we may not be able to generate product revenue.***

As a company, we currently have no sales, marketing or distribution capabilities and no experience in marketing products. We intend to develop an in-house marketing organization and sales force, which will require significant capital expenditures, management resources and time. We will have to compete with other pharmaceutical and biotechnology companies to recruit, hire, train and retain marketing and sales personnel.

If we are unable or decide not to establish internal sales, marketing and distribution capabilities, we will pursue arrangements with third-party sales, marketing and distribution collaborators regarding the sales and marketing of our products, if approved. However, there can be no assurance that we will be able to establish or maintain these types of arrangements on favorable terms or if at all, or if we are able to do so, that these third-party arrangements will provide effective sales forces or marketing and distribution capabilities. Any revenue we receive will depend upon the efforts of these third parties, which may not be successful. We may have little or no control over the marketing and sales efforts of these third parties and our revenue from product sales may be lower than if we had commercialized our product candidates ourselves. We also face competition in our search for third parties to assist us with the sales and marketing efforts of our product candidates.

There can be no assurance that we will be able to develop in-house sales and distribution capabilities or establish or maintain relationships with third-party collaborators to commercialize any product in the United States or overseas.

***The market opportunities for our product candidates may be relatively small as we expect that they will initially be approved only for those patients who are ineligible for other approved treatments or have failed prior treatments. In addition, our estimates of the prevalence of our target patient populations may be inaccurate.***

Our product candidates may target cancer, but cancer therapies are sometimes characterized as first-line, second-line, third-line or subsequent line and the FDA often approves new therapies initially only for a particular line 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, immunotherapy, hormone therapy, radiation therapy, surgery, other targeted therapies or a combination of these therapies – 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. We expect initially to seek approval of our product candidates in most instances as a second- or third-line therapy, for use in patients with relapsed or refractory cancer. Subsequently, for those product candidates that prove to be sufficiently safe and beneficial, if any, we would expect to seek approval as a second-line therapy and potentially as a first-line therapy, but there is no guarantee that any of our product candidates, even if approved as a second- or third- or subsequent line of therapy, would subsequently be approved for an earlier line of therapy. Further, it is possible that, prior to getting any approvals for our product candidates in earlier lines of treatment, we might 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 reasonable 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 or out of date. 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 of those tumor types.

***Even if we receive marketing approval of any of our product candidates, our products may become subject to unfavorable pricing regulations, third-party reimbursement practices or healthcare reform initiatives, any of which would harm our business.***

The regulations that govern marketing approvals, pricing, coverage and reimbursement for new drug products vary widely from country to country. Current and future legislation may significantly change the approval requirements in ways that could involve additional costs and cause delays in obtaining approvals. Some countries require approval of the sale price of a drug before it can be marketed. In many countries, the pricing review period begins after marketing or product licensing approval is granted. To obtain reimbursement or pricing approval in some countries, we may be required to conduct a clinical trial that compares the cost-effectiveness of our product candidate to other available therapies. In some foreign markets, prescription pharmaceutical pricing remains subject to continuing governmental control even after initial approval is granted. As a result, we might obtain marketing approval for a product candidate in a particular country, but then be subject to price regulations that delay our commercial launch of the product, possibly for lengthy time periods, which would negatively impact the revenues, if any, we are able to generate from the sale of the product in that country. Adverse pricing limitations may, therefore, hinder our ability to recoup our investment in one or more of our product candidates, even if our product candidates obtain marketing approval.

Our ability to commercialize any product candidates successfully also will depend in part on the extent to which coverage and adequate reimbursement for these products and related treatments will be available from government healthcare programs, private health insurers and other organizations. Government authorities and third-party payors, such as private health insurers and health maintenance organizations, decide which medications they will pay for and establish reimbursement levels. A primary trend in the U.S. healthcare industry and elsewhere is cost containment. Government authorities and third-party payors have attempted to control costs by limiting coverage and the amount of reimbursement for particular medications. Increasingly, government authorities and third-party payors are requiring that drug companies provide them with predetermined discounts from list prices and are challenging the prices charged for medical products. Coverage and reimbursement may not be available for any product that we commercialize and, even if these are available, the level of reimbursement may not be satisfactory. Reimbursement may affect the demand for, or the price of, any product candidate for which we obtain marketing approval. Obtaining and maintaining coverage and adequate reimbursement for our products may be difficult. We may be required to conduct expensive pharmacoeconomic studies to justify coverage and reimbursement or the level of reimbursement relative to other therapies. If coverage and adequate reimbursement are not available or reimbursement is available only to limited levels, we may not be able to successfully commercialize any product candidate for which we obtain marketing approval.

There may be significant delays in obtaining coverage and reimbursement for newly approved drugs. In addition, coverage may be more limited than the purposes for which the drug is approved by the FDA or similar regulatory authorities outside of the United States. Moreover, eligibility for coverage and reimbursement does not imply that a drug will be paid for in all cases or at a rate that covers our costs, including research, development, intellectual property, manufacture, sale and distribution expenses. Interim reimbursement levels for new drugs, if applicable, may also not be sufficient to cover our costs and may not be made permanent. Reimbursement rates may vary according to the use of the drug and the clinical setting in which it is used, may be based on reimbursement levels already set for lower cost drugs and may be incorporated into existing payments for other services. 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.

In the United States, no uniform policy for coverage and reimbursement for products exists among third-party payors. Therefore, coverage and reimbursement for our products can differ significantly from payor to payor. The process for determining whether a payor will provide coverage for a product may be separate from the process for setting the reimbursement rate that the payor will pay for the product. One payor's determination to provide coverage for a product does not assure that other payors will also provide coverage and reimbursement for the product. Third-party payors may also limit coverage to specific products on an approved list, or formulary, which might not include all of the FDA-approved products for a particular indication. Third-party payors often rely upon Medicare coverage policy and payment limitations in setting their own reimbursement policies. Our inability to promptly obtain coverage and adequate reimbursement rates from both government-funded and private payors for any approved products that we develop could have an adverse effect on our operating results, our ability to raise capital needed to commercialize products and our overall financial condition.

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

We face an inherent risk of product liability exposure related to the testing of our product candidates in human clinical trials and will face an even greater risk if or when we commercially sell any products that we may develop. If we cannot successfully defend

ourselves against claims that our product candidates or products caused injuries, we will incur substantial liabilities. Regardless of merit or eventual outcome, liability claims may result in:

- decreased demand for any product candidates or products that we may develop;
- termination of clinical trials;
- withdrawal of marketing approval, product recall, restriction on the approval or a “black box” warning or contraindication for an approved drug;
- withdrawal of clinical trial participants;
- significant costs to defend the related litigation and/or increased product liability insurance costs;
- substantial monetary awards to trial participants or patients;
- loss of revenue;
- injury to our reputation and significant negative media attention;
- reduced resources of our management to pursue our business strategy; and
- the inability to commercialize any products that we may develop.

We currently maintain product liability insurance coverage to support our clinical development activities. We may need to purchase additional product liability insurance coverage as we expand our clinical trials and if and when we commence commercialization of our product candidates. Insurance coverage is increasingly expensive. We may not be able to maintain insurance coverage at a reasonable cost or in an amount adequate to satisfy any liability that may arise.

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

***If we are unable to obtain and maintain patent protection for our technology, product candidates and products or if the scope of the patent protection obtained is not sufficiently broad or enforceable, our competitors could develop and commercialize technology, product candidates and products similar or identical to ours, our ability to successfully commercialize our technology, product candidates and products may be impaired or we may not be able to compete effectively in our market.***

We rely upon a combination of patents, trade secret protection and confidentiality agreements to protect our intellectual property and prevent others from exploiting our platform technologies, our pipeline drug product candidates, any future drug product candidates we may develop and their use or manufacture.

Our commercial success depends in part on our ability to obtain and maintain patents and other proprietary protection in the United States and other countries with respect to our proprietary technology, product candidates and products. We seek to protect our proprietary position by filing patent applications in the United States and abroad related to our novel technologies and product candidates. Any disclosure to or misappropriation by third parties of our confidential proprietary information could enable competitors to quickly duplicate or surpass our technological achievements, thus eroding our competitive position in our market. Moreover, the patent applications we own, co-own or license may fail to result in issued patents in the United States or in other foreign countries.

The patent prosecution process is expensive and time consuming and we may not be able to file, prosecute, and maintain all necessary or desirable patent applications at a reasonable cost or in a timely manner. It is also possible that we will fail to identify patentable aspects of our research and development output before it is too late to obtain patent protection. Moreover, in some circumstances, we do not have the right to control the preparation, filing and prosecution of patent applications, or to maintain the patents and patent applications, covering technology that we license from third parties or that we license to our collaborators. Therefore, these patents and applications may not be prosecuted and enforced in a manner consistent with the best interests of our business.

The patent position of the biopharmaceutical industry generally is highly uncertain, involves complex legal and factual questions and has been the subject of much litigation. In addition, the laws of foreign countries may not protect our rights to the same extent as the laws of the United States. 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 know with certainty whether we were the first to make the inventions claimed in our owned, co-owned or licensed patents or pending patent applications, or that we were the first inventors to file for patent protection of those inventions. As a result, the issuance, scope, validity, enforceability and commercial value of our patent rights or those of our collaborators are highly uncertain. Our pending and future patent applications may not result in patents being issued that protect our technology, product candidates or products, in whole or in part, or that effectively prevent others from commercializing competitive technologies, product candidates and products. Changes in either the patent laws or interpretation of the patent or other laws in the United States and other countries may diminish the value of our patents and potential applications, narrow the scope of our patent protection, or cause us to be required to pay royalties to third parties. In addition, if the breadth or strength of protection provided by our patents and patent applications is threatened, it could dissuade companies from collaborating with us to license, develop or commercialize current or future product candidates.

Our owned, co-owned and licensed patent estate consists principally of patent applications, many of which are at an early stage of prosecution. Even if our owned, co-owned and licensed patent applications issue as patents, they may not issue in a form that will provide us with any meaningful protection, prevent competitors from competing with us or otherwise provide us with any competitive advantage. Our competitors may be able to circumvent our owned, co-owned or licensed patents by developing similar or alternative technologies, product candidates or products in a non-infringing manner.

The issuance of a patent is not conclusive as to its inventorship, scope, validity or enforceability, and our owned, co-owned and licensed patents or patents obtained by our collaborators may be challenged in the courts or patent offices in the United States and abroad. These challenges may result in loss of exclusivity or freedom to operate or in patent claims being narrowed, invalidated or held unenforceable, in whole or in part, which could limit our ability to stop others from using or commercializing similar or identical technology, product candidates and products or limit the duration of the patent protection of our technology, product candidates and products. Given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting our drug product candidates might expire before or shortly after they are commercialized. As a result, our owned, co-owned and licensed patent portfolio, or that of our collaborators, may not provide us with sufficient rights to exclude others from commercializing products similar or identical to ours.

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

Patent reform legislation could increase the uncertainties and costs surrounding the prosecution of our patent applications and the enforcement or defense of our issued patents. On September 16, 2011, the Leahy-Smith America Invents Act, or the Leahy-Smith Act, was signed into law and made a number of significant changes to United States patent law. These changes include provisions that affect the way patent applications are prosecuted and may also affect patent litigation. The United States Patent and Trademark Office, or the USPTO, 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, including the first-inventor-to-file provisions, became effective on March 16, 2013. The Leahy-Smith Act and its implementation could increase the uncertainties and costs surrounding the prosecution of our patent applications and the enforcement or defense of our issued patents, all of which could have an adverse effect on our business and financial condition. The first-to-file provision of the Leahy-Smith Act requires us to act promptly during the period from invention to filing of a patent application, as there is always a risk that a third party could file a patent application that could be blocking to our patent filings. However, even with the intention to act promptly, circumstances could prevent us from promptly filing or prosecuting patent applications on our inventions. The Leahy-Smith Act also enlarged the scope of disclosures that qualify as prior art, which can impact our ability to receive patent protection for an invention.

The Leahy-Smith Act created, for the first time, new procedures under which third parties may challenge issued patents in the United States, including post-grant review, *inter partes* review and derivations proceedings, all of which are adversarial proceedings conducted at the USPTO. Since the effectiveness of the Leahy-Smith Act, some third parties have been using these types of actions to seek and achieve the cancellation of selected or all claims of issued patents of their competitors. Under the Leahy-Smith Act, for a patent with a priority date of March 16, 2013 or later (which is the case for all of our patent filings), a third party can file a petition for post-grant review at any time during a nine-month window commencing at the time of issuance of the patent. In addition, for a patent with a priority date of March 16, 2013 or later, a third party can file a petition for *inter partes* review after the nine-month period for filing a post-grant review petition has expired. Post-grant review proceedings can be brought on any ground of challenge, whereas *inter partes* review proceedings can only be brought to raise a challenge based on published prior art. Under applicable law, the standard of review for these types of adversarial actions at the USPTO are conducted without the presumption of validity afforded to U.S. patents, which is the standard that applies if a third party were to seek to invalidate a patent through a lawsuit filed in the U.S. federal courts. The USPTO issued a Final Rule on November 11, 2018 announcing that it will now use the same claim construction currently used in the U.S. federal courts—which is the plain and ordinary meaning of words used—to interpret patent claims in these USPTO proceedings. As a result of this regulatory landscape, if any of our patents are challenged by a third party in a USPTO proceeding of this nature, there is no guarantee that we will be successful in defending the challenged patent, which could result in our losing rights under the challenged patent in part or in whole.

As a result of this legislation, the issuance, scope, validity, enforceability and commercial value of our patent rights, or those of our collaborators, are highly uncertain, which could have an adverse effect on our business, financial condition, results of operations and prospects.

***We may become involved in lawsuits to protect or enforce our patents, the patents of our licensors or other intellectual property, which could be expensive, time-consuming and unsuccessful.***

Competitors may infringe our issued patents, the patents of our licensors or collaborators or our other intellectual property. To counter infringement or unauthorized use, we may be required to file infringement claims, which can be expensive, time-consuming and unpredictable. Any claims we assert against perceived infringers could provoke these parties to assert counterclaims against us alleging that we infringe their patents. In addition, in a patent infringement proceeding, a court may decide that a patent of ours or our licensors or collaborators 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 in question. An adverse result in any litigation proceeding could put one or more of our patents at risk of being or actually invalidated, held unenforceable or

interpreted narrowly. Even if we successfully assert our patents, a court may not award remedies that sufficiently compensate us for our losses. In addition, we may not have sufficient financial or other resources to seek to enforce our patents adequately against perceived infringers, which could have a material and adverse effect on the profitability of our products.

***We may need to license intellectual property from third parties and licenses of this nature may not be available or may not be available on commercially reasonable terms.***

A third party may hold intellectual property, including patent rights, that are important or necessary to the development or manufacture of our products or our collaborators' products. It may, therefore, be necessary for us to use the patented or proprietary technology of a third party to commercialize our own technology or products or those of our collaborators, in which case we or our collaborators would be required to obtain a license from that third party. A license to that intellectual property may not be available or may not be available on commercially reasonable terms, which could have an adverse effect on our business and financial condition.

The licensing and acquisition of third-party intellectual property rights is a competitive practice. Companies that may be more established or have greater resources than we do may also be pursuing strategies to license or acquire third-party intellectual property rights that we may consider necessary or attractive in order to commercialize our product candidates. More established companies may have a competitive advantage over us due to their larger size and cash resources or greater clinical development and commercialization capabilities. We may not be able to successfully complete such negotiations and ultimately acquire the rights to the intellectual property surrounding the additional product candidates that we may seek to acquire.

***Third parties may initiate legal proceedings alleging that we are infringing their intellectual property rights, the outcome of which would be uncertain and could have an adverse effect on the success of our business.***

Our commercial success depends upon our ability and the ability of our collaborators to develop, manufacture, market and sell our product candidates and use our proprietary technologies without infringing the proprietary rights of third parties. There is considerable intellectual property litigation in the biopharmaceutical industry, as well as administrative proceedings for challenging patents, including reexamination, post-grant review, *inter partes* review, derivation proceedings or interference proceedings before the USPTO and oppositions and other comparable proceedings in foreign jurisdictions.

We may become party to or threatened with future adversarial proceedings or litigation regarding intellectual property rights with respect to our product candidates and technology, including derivation, reexamination, post-grant review, *inter partes* review, or interference proceedings before the USPTO. Third parties may assert infringement claims against us based on existing patents or patents that may be granted in the future. As the bio-pharmaceutical industry expands 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. There may be third-party patents of which we are currently unaware with claims to materials, formulations, methods of manufacture or methods for treatment related to the use or manufacture of our drug candidates. Because patent applications can take many years to issue, there may be currently pending patent applications that may later result in issued patents that our product candidates may infringe. In addition, third parties may obtain patents in the future and claim that use of our technologies infringes upon these patents.

If we are found by a court of competent jurisdiction to infringe a third party's intellectual property rights, we could be required to obtain a license from the applicable third-party intellectual property holder to continue developing and marketing our product candidates, products and technology. However, we may not be able to obtain any required license on commercially reasonable terms or at all. Even if we were able to obtain a license, it could be non-exclusive, thereby giving our competitors access to the same technologies licensed to us. We could be forced, including by court order, to cease commercializing the infringing technology or product. In addition, we could be found liable for monetary damages, including treble damages and attorneys' fees if we are found to have willfully infringed a patent. A finding of infringement could prevent us from commercializing our product candidates or force us to cease some of our business operations, which could materially harm our business. Claims that we have misappropriated the confidential information or trade secrets of third parties could have a similar negative impact on our business.

***A number of other companies, as well as universities and other organizations, file and obtain patents in the same areas as our products, which are targeted protein degraders, or our platform technologies and these patent filings could be asserted against us or our collaborators in the future, which could have an adverse effect on the success of our business and, if successful, could lead to expensive litigation that could affect the profitability of our products and/or prohibit the sale or use of our products.***

Our MonoDAC and BiDAC product candidates are small molecule pharmaceuticals, which degrade specific proteins. A number of companies and institutions have patent applications and issued patents in this general area, such as, for example, Amgen Inc., Amphista Therapeutics, Ltd., Araxes Pharma, LLC, Arvinas, Inc., AstraZeneca PLC, Bayer AG (and its subsidiary Vividion Therapeutics, Inc.), Beigene Co. Ltd., BioTheryX, Inc., Bristol Myers Squibb Company (and its subsidiary Celgene Corporation), Captor Therapeutics SA, Cullgen, Inc., the Dana-Farber Cancer Institute and its Center for Protein Degradation, Foghorn Therapeutics, Inc., GlaxoSmithKline PLC, Janssen Biotech, Inc., Kymera Therapeutics, LLC, Nurix Therapeutics, Inc., Monte Rosa Therapeutics, AG, Novartis AG, Nurix Therapeutics, Inc., Otsuka Pharmaceuticals, Inc., Proteovant Therapeutics, Inc., Roche AG, the University of Michigan School of Medicine, Vertex Pharmaceuticals, Inc. and others. If any of these companies or institutions or others not included in this list were to assert that one of its patents is infringed by any product candidate or product we might develop or its use or manufacture, we or our collaborators may be drawn into expensive litigation, which could adversely affect our business prospects, financial condition and results of operations, require extensive time from and cause the distraction of members of our

management team and employees at large. Further, if litigation of this nature were successful, that could have a material and adverse effect on the profitability of our products or prohibit their sale. We may not be aware of patent claims that are currently or may in the future be pending that could affect our business or products. Patent applications are typically published between six and eighteen months from filing and the presentation of new claims in already pending applications can sometimes not be visible to the public, which would include us, for a period of time. In addition, even after a patent application is publicly available, we may not yet have seen that patent application and may, therefore, not be aware of the claims or scope of filed and published patent applications. As a result, we cannot provide any assurance that a third party practicing in the general area of our technology will not present or has not presented a patent claim that covers one or more of our product candidates or products or their methods of use or manufacture. If that were to occur, we or our collaborators, as applicable, may have to take steps to try to invalidate the applicable patent or application and, in a situation of that nature, we or our collaborators may either choose not to do so or our attempt may not be successful. If we determine that we require a license to a third party's patent or patent application, we may discover that a license may not be available on reasonable terms, or at all, which could prevent us or our collaborators from selling a product or using our proprietary technologies.

***Our products are subject to The Drug Price Competition and Patent Term Restoration Act of 1984, which is also referred to as the Hatch Waxman Act, in the United States, which can increase the risk of litigation with generic companies trying to sell our products and may cause us to lose patent protection.***

Because our clinical candidates are pharmaceutical molecules that will be reviewed by the Center for Drug Evaluation and Research of the FDA, after commercialization they will be subject in the United States to the patent litigation process of the Hatch-Waxman Act, as amended to date, which allows a generic company to submit an Abbreviated New Drug Application, or ANDA, to the FDA to obtain approval to sell a generic version of our drug using bioequivalence data only. Under amendments made to the Hatch-Waxman Act, we will have the opportunity to list our patents that cover our drug products or their respective methods of use in the FDA's compendium of "Approved Drug Products with Therapeutic Equivalence Evaluation," sometimes referred to as the FDA's Orange Book.

There are detailed rules and requirements regarding the patents that may be submitted to the FDA for listing in the Orange Book. We may be unable to obtain patents covering our product candidates that contain one or more claims that satisfy the requirements for listing in the Orange Book. Even if we submit a patent for listing in the Orange Book, the FDA may decline to list the patent or a generic drug manufacturer may challenge the listing. If one of our product candidates is approved and a patent covering that product candidate is not listed in the Orange Book, with respect to any unlisted patent, a generic drug manufacturer would not have to provide advance notice to us of any ANDA filed with the FDA to obtain permission to sell a generic version of that product candidate.

Currently, in the United States, the FDA may grant five years of exclusivity for new chemical entities, or NCEs, which are drugs that contain no active portion that has been approved by the FDA in any other new drug application, or NDA. We expect that all of our products will qualify as NCEs; however, the FDA will not conduct an assessment for NCE status until it is reviewing a marketing application for that drug. A generic company can submit an ANDA to the FDA four years after approval of any of our drug products designated as an NCE. The submission of an ANDA by a generic company is considered a technical act of patent infringement. The generic company can certify that it will wait until the natural expiration date of our listed patents to sell a generic version of our product or can certify that one or more of our listed patents are invalid, unenforceable or not infringed. If the generic manufacturer elects the latter, we will have 45 days to bring a patent infringement lawsuit against the generic company. If we were to do so, that would likely initiate a challenge to one or more of our Orange Book listed patents based on arguments from the generic manufacturer that our listed patents are invalid, unenforceable or not infringed. Under amendments to the Hatch-Waxman Act, if a lawsuit is brought, the FDA is prevented from issuing a final approval on the generic drug until 30 months after the end of the data exclusivity period (7.5 years) or a final decision of a court holding that our asserted patent claims are invalid, unenforceable or not infringed. If we do not properly list our relevant patents in the Orange Book or if we fail to file a lawsuit in response to a certification from a generic company under an ANDA in a timely manner, or if we do not prevail in the resulting patent litigation, we can lose our ability to benefit from a proprietary market based on patent protection covering our drug products and we may find that physicians will switch to prescribing and dispensing generic versions of our drug products. Further, even if we were to list our relevant patents in the Orange Book correctly, bring a lawsuit in a timely manner and prevail in that lawsuit, the generic litigation may come at a significant cost to us, both in terms of attorneys' fees and employee time and distraction over a long period. Further, it is common for more than one generic company to try to sell an innovator's drug at the same time and, as a result, we may face the cost and distraction of multiple lawsuits from generic manufacturers at the same time. We may also determine that it is necessary to settle these types of lawsuits in a manner that allows the generic company to enter our market prior to the expiration of our patent or otherwise in a manner that adversely affects the strength, validity or enforceability of our patents.

***A number of pharmaceutical companies have been the subject of intense review by the U.S. Federal Trade Commission or a corresponding agency in another country based on how they have conducted or settled patent litigation related to pharmaceutical products. In fact, certain reviews have led to an allegation of an anti-trust violation, sometimes resulting in a fine or loss of rights. We cannot be sure that we would not also be subject to a review of this nature or that the result of a review of this nature would be favorable to us, or that any review of this nature would not result in a fine or penalty.***

The U.S. Federal Trade Commission, or FTC, has brought a number of lawsuits in federal court in the past few years to challenge ANDA litigation settlements reached between innovator companies and generic companies as anti-competitive. As an example, the

FTC has taken an aggressive position that anything of value is a payment, whether money is paid or not. Under their approach, if an innovator, as part of a patent settlement, agrees not to launch or delay its launch of an authorized generic during the 180-day period granted to the first generic company to challenge an Orange Book listed patent covering an innovator drug, or negotiates a delay in entry without payment, the FTC may consider it an unacceptable reverse payment. Companies in the pharmaceutical industry have argued that these types of agreements are rational business decisions entered into by drug innovators as a way to address risk and that these settlements should, therefore, be immune from antitrust attack if the terms of the settlement are within the scope of the exclusionary potential of the patent. In 2013, the U.S. Supreme Court in a five-to-three decision in *FTC v. Actavis, Inc.* rejected both the pharmaceutical industry's and FTC's arguments with regard to so-called reverse payments. Instead, the Supreme Court held that whether a "reverse payment" settlement involving the exchange of consideration for a delay in entry is subject to an anti-competitive analysis depends on five considerations: (a) the potential for genuine adverse effects on competition; (b) the justification of payment; (c) the patentee's ability to bring about anti-competitive harm; (d) whether the size of the payment is a workable surrogate for the patent's weakness; and (e) that antitrust liability for large unjustified payments does not prevent litigating parties from settling their lawsuits, for example, by allowing the generic drug to enter the market before the patent expires on the branded drug without the patentee paying the generic manufacturer. Further, whether a reverse payment is justified depends upon its size, scale in relation to the patentee's anticipated future litigation costs, and independence from other services for which it might represent payment (as was the case in *Actavis*), as well as the lack of any other convincing justification. The Supreme Court instead held that reverse payment settlements can potentially violate antitrust laws and are subject to the standard antitrust rule-of-reason analysis, with the burden of proving that an agreement is unlawful on the FTC. In reaching this decision, the Supreme Court left to the lower courts the structuring of this rule of reason analysis.

If we are faced with drug patent litigation, including Hatch-Waxman litigation with a generic company, we could be faced with an FTC challenge of this nature, which challenge could impact how or whether we settle the case and, even if we strongly disagree with the FTC's position, we could face a significant expense or penalty. Any litigation settlements we enter into with generic companies under the Hatch-Waxman Act could also be challenged by third-party payors such as insurance companies, direct purchasers or others who consider themselves adversely affected by the settlement. These kinds of follow-on lawsuits, which may be class action suits, can be expensive and can continue over multiple years. If we were to face lawsuits of this nature, we may not be successful in defeating these claims and we may, therefore, be subject to large payment obligations, which we may not be able to satisfy in whole or in part.

***We may not be able to obtain patent term extension under the Drug Price Competition and Patent Term Restoration Act of 1984 in the United States and, as a result, our product candidates, if approved, may not have patent protection for a sufficient period.***

In the United States, the Drug Price Competition and Patent Term Restoration Act of 1984 permits one patent term extension of up to five years beyond the normal expiration of one patent per product, which if related to a method of treatment patent, is limited to the approved indication. The length of the patent term extension is typically calculated as one-half of the clinical trial period plus the entire period of time during the review of the NDA by the FDA, minus any time of delay by us during these periods. There is also a limit on the patent term extension to a term that is no greater than fourteen years from the date of drug approval. Therefore, if we select and are granted a patent term extension on a recently filed and issued patent, we may not receive the full benefit of a possible patent term extension, if at all. We might also not be granted a patent term extension at all, because of, for example, our failure to apply within the applicable period, failure to apply prior to the expiration of relevant patents or other failure to satisfy any of the numerous applicable requirements. In addition, the regulatory review period of an FDA-approved product may not serve as the basis for a patent term extension if the active ingredient of such product was subject to regulatory review and approval in an earlier product approved by the FDA. Moreover, the applicable authorities, including the FDA and the USPTO in the United States and any equivalent regulatory authority in other countries, may not agree with our assessment of whether extensions of this nature are available and may refuse to grant extensions to our patents or may grant more limited extensions than we request. If this occurs, our competitors may be able to obtain approval of competing products following our patent expiration by referencing our clinical and preclinical data and launch their product earlier than might otherwise be the case. If this were to occur, it could have an adverse effect on our ability to generate product revenue.

In 1997, as part of the Food & Drug Administration Modernization Act, or FDAMA, Congress enacted a law that provides incentives to drug manufacturers who conduct studies of drugs in children. The law, which provides six-months exclusivity in return for conducting pediatric studies, is referred to as the "pediatric exclusivity provision." If we were to conduct clinical trials that comply with the FDAMA, we could receive an additional six-month term added to our regulatory data exclusivity period and on the patent term extension period, if received, on our product. However, if we choose not to carry out pediatric studies that comply with the FDAMA or carry out studies that are not accepted by the FDA for this purpose, we will not receive this additional six-month exclusivity extension to our data exclusivity or our patent term extension.

In Europe, supplementary protection certificates are available to extend a patent term up to five years to compensate for patent term lost during regulatory review, and this period can be extended to five and a half years if data from clinical trials is obtained in accordance with an agreed Pediatric Investigation Plan. Although all countries in Europe must provide supplementary protection certificates, there is no unified legislation among European countries and, as a result, drug developers must apply for supplementary protection certificates on a country-by-country basis. As a result, a company may need to expend significant resources to apply for and receive these certificates in all relevant countries and may receive them in some, but not all, countries, if at all.

***Weakening patent laws and enforcement by courts in the United States and foreign countries may impact our ability to protect our markets.***

The U.S. Supreme Court has issued opinions in patent cases in the last few years that many consider may weaken patent protection in the United States, either by narrowing the scope of patent protection available in certain circumstances, holding that certain kinds of innovations are not patentable or generally otherwise making it easier to invalidate patents in court. Additionally, there have been recent proposals for additional changes to the patent laws of the United States and other countries that, if adopted, could impact our ability to obtain patent protection for our proprietary technology or our ability to enforce our proprietary technology. Depending on future actions by the U.S. Congress, the U.S. courts, the USPTO and the relevant law-making bodies in other countries, the laws and regulations governing patents could change in unpredictable ways that would weaken our ability to obtain new patents or to enforce our existing patents and patents that we might obtain in the future.

The laws of some foreign jurisdictions do not protect intellectual property rights to the same extent as in the United States and many companies have encountered significant difficulties in protecting and defending such rights in foreign jurisdictions. If we encounter such difficulties in protecting or are otherwise precluded from effectively protecting our intellectual property rights in foreign jurisdictions, our business prospects could be substantially harmed. For example, we could become a party to foreign opposition proceedings, such as at the European Patent Office, or patent litigation and other proceedings in a foreign court. If so, uncertainties resulting from the initiation and continuation of such proceedings could have an adverse effect on our ability to compete in the marketplace. The cost of foreign adversarial proceedings can also be substantial, and in many foreign jurisdictions, the losing party must pay the attorney fees of the winning party.

***We may be subject to claims by third parties asserting that we, our employees, consultants or contractors have misappropriated the applicable third party's intellectual property or claiming ownership of what we regard as our own intellectual property.***

We employ individuals who were previously employed at universities as well as other biotechnology or pharmaceutical companies, including our competitors or potential competitors. We have received confidential and proprietary information from collaborators, prospective licensees and other third parties that may be subject to contractual confidentiality and non-use obligations. Although we try to ensure that our employees do not use the proprietary information or know-how of others in their work for us, we may be subject to claims that these employees or we have used or disclosed intellectual property, including trade secrets or other proprietary information, of any such employee's former employer. We may also be subject to claims that former employers or other third parties have an ownership interest in our patents. Litigation may be necessary to defend against these claims. We may not be successful in defending these claims, and 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. Even if we are successful, litigation could result in substantial cost and reputational loss and be a distraction to our management and other employees.

In addition, while it is our policy to require our employees, consultants and contractors who may be involved in the development of intellectual property to execute agreements assigning any resulting intellectual property to us, we may be unsuccessful in executing an agreement to that effect with each party who in fact develops intellectual property that we regard as our own. Assignment agreements of this nature 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. In addition, an employee or contractor could create an invention but not inform us of it, in which case we could lose the benefit of the invention and the employee or contractor may leave to develop the invention elsewhere.

***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 common stock. Litigation or proceedings of this nature 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 litigation or proceedings of this nature 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.

***Obtaining and maintaining patent protection depends on compliance with various procedural, documentary, fee payment and other requirements imposed by governmental patent offices and the protection of our patents could be reduced or eliminated if we fail to comply with these requirements.***

Periodic maintenance fees on any issued patent are due to be paid to the USPTO and patent offices in foreign countries in several stages over the lifetime of a patent application and any resulting patent. The USPTO and patent offices in foreign countries require compliance with many procedural, documentary, fee payment and other requirements during the patent application process. While an inadvertent lapse can in many cases be cured by payment of a late fee or by other means in accordance with the applicable rules, there are situations in which noncompliance can result in abandonment or lapse of the patent or patent application, resulting in partial or

complete loss of a patent or patent rights in the relevant jurisdiction. Non-compliance events that could result in abandonment or lapse of a patent or patent application include, but are not limited to, failure to respond to official actions within prescribed time limits, non-payment of fees and failure to properly legalize and submit formal documents. In such an event, our competitors might be able to enter the market, which would have an adverse effect on our business.

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

In addition to seeking patents for some of our technology and product candidates, we also rely on trade secrets, including unpatented know-how, technology and other proprietary information, to maintain our competitive position. We seek to protect these trade secrets, in part, by entering into non-disclosure and confidentiality agreements with parties who have access to them, such as our employees, corporate collaborators, outside scientific collaborators, contract manufacturers, consultants, advisors and other third parties. We also enter into confidentiality and invention or patent assignment agreements with our employees and consultants. These agreements may not effectively prevent disclosure of confidential information nor result in the effective assignment to us of intellectual property and may not provide an adequate remedy in the event of unauthorized disclosure of confidential information or other breaches of the agreements. In addition, others may independently discover our trade secrets and proprietary information. In that case, we could not assert any trade secret rights against that third party. 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. Enforcing a claim that a party illegally disclosed or misappropriated a trade secret is difficult, expensive and time-consuming and the outcome of a dispute of this nature is inherently unpredictable. Costly and time-consuming litigation could be necessary to seek to enforce and determine the scope of our proprietary rights and our failure to obtain or maintain trade secret protection could adversely affect our competitive business position. In addition, some courts outside of the United States are less willing or unwilling to protect trade secrets. The Defend Trade Secrets Act of 2016 is a U.S. federal law that allows an owner of a trade secret to sue in federal court when its trade secret has been misappropriated. Congress passed this law in an attempt to strengthen the rights of trade secret owners whose valuable assets are taken without authorization. If any of our trade secrets were to be lawfully obtained or independently developed by a competitor, we would have no right to prevent them, or those to whom they communicate 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, our competitive position would be harmed.

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

Filing, prosecuting, maintaining, and defending patents covering our product candidates in all countries throughout the world would be prohibitively expensive. As a result, our intellectual property rights in some countries outside the United States can be less extensive than the protection we might have in the United States. In-licensing patents covering our product candidates in all countries throughout the world may similarly be prohibitively expensive, if these in-licensing opportunities are available to us at all. Further, in-licensing or filing, prosecuting, maintaining, and defending patents even in only those jurisdictions in which we develop or commercialize our product candidates may be prohibitively expensive or impractical. Competitors may use our and our licensors' technologies in jurisdictions where we have not obtained patent protection or licensed patents to develop their own products and, further, may export otherwise infringing products to territories where we and our licensors have patent protection, but enforcement is not as strong as that in the United States or the European Union. These products may compete with our product candidates, and our or our licensors' patents or other intellectual property rights may not be effective or sufficient to prevent them from competing.

In addition, we may decide to abandon national and regional patent applications while they are still pending. The grant proceeding of each national or regional patent is an independent proceeding that may lead to situations in which applications may be rejected by the relevant patent office, while substantively similar applications are granted by others. For example, relative to other countries, China has a heightened detailed description requirement for patentability. Further, generic drug manufacturers or other competitors may challenge the scope, validity or enforceability of our or our licensors' patents, requiring us or our licensors to engage in complex, lengthy and costly litigation or other proceedings. Generic drug manufacturers may develop, seek approval for and launch generic versions of our products. It is also quite common that depending on the country, the scope of patent protection may vary for the same product candidate or technology.

The laws of some jurisdictions do not protect intellectual property rights to the same extent as the laws or regulations in the United States and the European Union, and many companies have encountered significant difficulties in protecting and defending proprietary rights in such jurisdictions. Moreover, the legal systems of certain countries, particularly certain developing countries, do not favor the enforcement of patents, trade secrets or other forms of intellectual property, which could make it difficult for us to prevent competitors in some jurisdictions from marketing competing products in violation of our proprietary rights generally.

Proceedings to enforce our patent rights in foreign jurisdictions, whether or not successful, are likely to result in substantial costs and divert our efforts and attention from other aspects of our business and could additionally put our or our licensors' patents at risk of being invalidated or interpreted narrowly, could increase the risk of our or our licensors' patent applications not issuing or could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate, while damages or other remedies may be awarded to the adverse party, which may be commercially significant. If we prevail, damages or other remedies awarded to us, if any, may not be commercially meaningful. Accordingly, our efforts to enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we develop or license. Further, while

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

In some jurisdictions, compulsory licensing laws compel patent owners to grant licenses to third parties. In addition, some countries limit the enforceability of patents against government agencies or government contractors. In these countries, the patent owner may have limited remedies, which could materially diminish the value of such patent. If we or any of our licensors are forced to grant a license to third parties under patents relevant to our business, or if we or our licensors are prevented from enforcing patent rights against third parties, our competitive position may be substantially impaired in such jurisdictions.

#### **Risks related to regulatory matters**

***The regulatory approval processes of the FDA and foreign regulatory authorities are lengthy, time-consuming and inherently unpredictable and, if we are ultimately unable to obtain marketing approval for our product candidates, our business will be substantially harmed.***

The time required to obtain approval by the FDA and foreign regulatory authorities is unpredictable but typically takes many years following the commencement of clinical trials and depends upon numerous factors, including the substantial discretion of the regulatory authorities. In addition, approval policies, regulations or the type and amount of clinical data necessary to gain approval may change during the course of a product candidate's clinical development and may vary among jurisdictions. We have not obtained marketing approval for any product candidate, and it is possible that none of our existing product candidates, or any product candidates we may seek to develop in the future (independently or with one of our collaboration partners), will ever obtain marketing approval.

Our product candidates could fail to receive marketing approval for many reasons, including the following:

- the FDA may disagree with the design or implementation of our clinical trials;
- we may be unable to demonstrate to the satisfaction of the FDA that a product candidate is safe and effective for its proposed indication, or that it is of sufficient purity or quality in accordance with FDA standards;
- results of clinical trials may not meet the level of statistical significance required by the FDA for approval;
- we may be unable to demonstrate that a product candidate's clinical and other benefits outweigh its safety risks;
- the FDA may disagree with our interpretation of data from preclinical studies or clinical trials;
- data collected from clinical trials of our product candidates may not be sufficient to support the submission of an NDA to the FDA or other submission or to obtain marketing approval in the United States;
- the FDA may find deficiencies with or fail to approve the manufacturing processes or facilities of third-party manufacturers with which we contract for clinical and commercial supplies; and
- the approval policies or regulations of the FDA may significantly change in a manner rendering our clinical data insufficient for approval.

This lengthy approval process, as well as the unpredictability of future clinical trial results, may result in our failing to obtain regulatory approval to market any of our product candidates, which would significantly harm our business, results of operations and prospects. The FDA has substantial discretion in the approval process and determining when or whether regulatory approval will be obtained for any of our product candidates. Even if we believe the data collected from clinical trials of our product candidates are promising, that data may not be sufficient to support approval by the FDA.

In addition, even if we were to obtain approval, regulatory authorities may approve any of our product candidates for fewer or more limited indications than we request, may not approve the price we intend to charge for our products, may grant approval contingent on the performance of costly post-marketing clinical trials or may approve a product candidate with a label that does not include the labeling claims necessary or desirable for the successful commercialization of that product candidate. Any of the foregoing scenarios could materially harm the commercial prospects for our product candidates.

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

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

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

Approval processes vary among countries and can involve additional product testing and validation and additional administrative review periods. Seeking foreign regulatory approval could result in difficulties and increased costs for us and require additional preclinical studies or clinical trials, which could be costly and time consuming. Regulatory requirements can vary widely from country to country and could delay or prevent the introduction of our products in those countries. We do not have any product candidates approved for sale in any jurisdiction, including in international markets, and we do not have experience as a company in obtaining regulatory approval in international markets. If we fail to comply with regulatory requirements in international markets or to obtain and maintain required approvals, or if regulatory approvals in international markets are delayed, our target market will be reduced and our ability to realize the full market potential of any product we develop will be unrealized.

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

If any of our product candidates are approved, they will be subject to ongoing regulatory requirements for manufacturing, labeling, packaging, storage, advertising, promotion, sampling, record-keeping, conduct of post-marketing studies and submission of safety, efficacy and other post-market information, including both federal and state requirements in the United States and requirements of comparable foreign regulatory authorities. In addition, we will be subject to continued compliance with cGMP and GCP requirements for any clinical trials that we conduct post-approval.

Manufacturers and manufacturers' facilities are required to comply with extensive FDA and comparable foreign regulatory authority requirements, including ensuring that quality control and manufacturing procedures conform to cGMP regulations and applicable product tracking and tracing requirements. As such, we and our CMOs will be subject to continual review and inspections to assess compliance with cGMP and adherence to commitments made in any NDA, other marketing application and previous responses to inspection observations. Accordingly, we and others with whom we work must continue to expend time, money and effort in all areas of regulatory compliance, including manufacturing, production and quality control.

Any regulatory approvals that we receive for our product candidates may be subject to limitations on the approved indicated uses for which the product may be marketed or to the conditions of approval, or contain requirements for potentially costly post-marketing testing, including Phase 4 clinical trials and surveillance to monitor the safety and efficacy of the product candidate. The FDA may also require a REMS program as a condition of approval of our product candidates, which could entail requirements for long-term patient follow-up, 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. Comparable foreign regulatory authorities may also have programs similar to REMS. In addition, if the FDA or a comparable foreign regulatory authority approves our product candidates, we will have to comply with requirements including submissions of safety and other post-marketing information and reports and registration.

The FDA may impose consent decrees or withdraw approval if compliance with regulatory requirements and standards is not maintained or if problems occur after the product reaches the market. Later discovery of previously unknown problems with our product candidates, including adverse events of unanticipated severity or frequency, or with our third-party CMOs or manufacturing processes, or failure to comply with regulatory requirements, may result in revisions to the approved labeling to add new safety information; imposition of post-market studies or clinical trials to assess new safety risks; or imposition of distribution restrictions or other restrictions under a REMS program. Other potential consequences include, among other things:

- restrictions on the marketing or manufacturing of our products, withdrawal of the product from the market or voluntary or mandatory product recalls;
- 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 license 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.

The FDA strictly regulates marketing, labeling, advertising and promotion of products that are placed on the market. Products may be promoted only for the approved indications and in accordance with the provisions of the approved label. The FDA and other agencies actively enforce the laws and regulations prohibiting the promotion of off-label uses and a company that is found to have improperly promoted off-label uses may be subject to significant liability. However, physicians may, in their independent medical judgment, prescribe legally available products for off-label uses. The FDA does not regulate the behavior of physicians in their choice of treatments, but the FDA does restrict manufacturers' communications on the subject of off-label use of their products. The policies of

the FDA and of comparable foreign regulatory authorities 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.

***A Breakthrough Therapy designation by the FDA, even if granted for any of our product candidates, may not lead to a faster development or regulatory review or approval process and it does not increase the likelihood that our product candidates will receive marketing approval.***

We may seek Breakthrough Therapy designation for our CFT7455 and CFT8634 product candidates and some or all of our future product candidates. A Breakthrough Therapy is defined as a drug that is intended, alone or in combination with one or more other drugs, to treat a serious or life-threatening disease or condition and preliminary clinical evidence indicates that the drug may demonstrate substantial improvement over existing therapies on one or more clinically significant endpoints, such as substantial treatment effects observed early in clinical development. For product candidates that have been designated as Breakthrough Therapies, interaction and communication between the FDA and the sponsor of the trial can help to identify the most efficient path for clinical development while minimizing the number of patients placed in ineffective control regimens. Drugs designated as breakthrough therapies by the FDA may also be eligible for other expedited approval programs, including accelerated approval.

Designation as a Breakthrough Therapy is within the discretion of the FDA. Accordingly, even if we believe one of our product candidates meets the criteria for designation as a Breakthrough Therapy, the FDA may disagree and instead determine not to make such a designation. In any event, the receipt of a Breakthrough Therapy designation for a product candidate may not result in a faster development process, review or approval compared to candidate products considered for approval under non-expedited FDA review procedures and does not assure ultimate approval by the FDA of a product candidate. In addition, even if one or more of our product candidates qualify as Breakthrough Therapies, the FDA may later decide that the product no longer meets the conditions for qualification. Thus, even though we intend to seek Breakthrough Therapy designation for our lead product candidates and some or all of our future product candidates for the treatment of various cancers, there can be no assurance that we will receive Breakthrough Therapy designations.

***A Fast Track designation by the FDA, even if granted for one or all of our lead product candidates, or any of our other current or future product candidates, 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.***

At various times, we may seek Fast Track designation for one or more of our product candidates. If a drug is intended for the treatment of a serious or life-threatening condition and the drug demonstrates the potential to address unmet medical needs for this condition, the drug sponsor may apply for FDA Fast Track designation for a particular indication. We may seek Fast Track designation for one or all of our lead product candidates and/or certain of our future product candidates, but there is no assurance that the FDA will grant this status to any of our proposed product candidates and we might only be successful in receiving a Fast Track designation from the FDA for a product candidate after applying on more than one occasion. Marketing applications filed by sponsors of products in Fast Track development may qualify for priority review under the policies and procedures offered by the FDA, but the receipt of a Fast Track designation does not assure any such qualification or ultimate marketing approval by the FDA. The FDA has broad discretion whether or not to grant a 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 do receive a Fast Track designation, we may not experience a faster development process, 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 a 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 have obtained Orphan Drug Designation for CFT7455 and CFT8634 and if we decide to seek Orphan Drug Designation for any other current or future product candidates, we may be unsuccessful or may be unable to maintain the benefits associated with Orphan Drug Designation, including the potential for supplemental market exclusivity.***

In August 2021, the FDA granted Orphan Drug Designation to CFT7455 for the treatment of MM, and in March 2022, the FDA granted Orphan Drug Designation to CFT8634 for the treatment of soft tissue sarcoma. We may seek Orphan Drug Designation for one or more of our other current or future product candidates. Regulatory authorities in some jurisdictions, including the United States and Europe, may designate drugs for relatively small patient populations as orphan drugs. Under the Orphan Drug Act, the FDA may grant an Orphan Drug Designation to a drug intended to treat a rare disease or condition, defined as a disease or condition with a patient population of fewer than 200,000 in the United States, or a patient population greater than 200,000 in the United States when there is no reasonable expectation that the cost of developing and making available the drug in the United States will be recovered from sales in the United States for that drug. In the United States, receipt of an Orphan Drug Designation entitles a party to financial incentives such as opportunities for grant funding towards clinical trial costs, tax advantages and user-fee waivers. After the FDA grants Orphan Drug Designation, the generic identity of the drug and its potential orphan use are disclosed publicly by the FDA. Orphan Drug Designation does not convey any advantage in, or shorten the duration of, the regulatory review and approval process.

If a product that has an Orphan Drug Designation subsequently receives the first FDA approval for a particular active ingredient for the disease for which it has such designation, the product is entitled to orphan product exclusivity, which means that the FDA may not approve any other applications, including an NDA, to market the same drug for the same indication for seven years, except in limited circumstances such as a showing of clinical superiority to the product with orphan drug exclusivity or if the FDA finds that the holder of the orphan drug exclusivity has not shown that it can assure the availability of sufficient quantities of the orphan drug to meet the needs of patients with the disease or condition for which the drug was designated. As a result, even if one of our product candidates receives orphan drug exclusivity, the FDA can still approve other drugs that have a different active ingredient for use in treating the same indication or disease. Further, the FDA can waive orphan drug exclusivity if we are unable to manufacture sufficient supply of our product.

We may also seek Orphan Drug Designations for our other lead programs and/or some or all of our other current or future product candidates in additional orphan indications in which there is a medically plausible basis for the use of these product candidates. Even when we obtain an Orphan Drug Designation, exclusive marketing rights in the United States may be limited if we seek approval for an indication broader than the orphan designated indication and may be lost if the FDA later determines that the request for designation was materially defective or if we, through our manufacturer, are unable to assure sufficient quantities of the product to meet the needs of patients with the rare disease or condition. In addition, even if we seek Orphan Drug Designation for other product candidates, we may never receive these designations. For example, the FDA has expressed concerns regarding the regulatory considerations for Orphan Drug Designation as applied to tissue agnostic therapies and the FDA may interpret the Federal Food, Drug and Cosmetic Act, and regulations promulgated thereunder, in a way that limits or blocks our ability to obtain an Orphan Drug Designation or orphan drug exclusivity, if our product candidates are approved, for our targeted indications.

***We may seek approval of our product candidates, where applicable under the FDA's accelerated approval pathway. This pathway may not lead to a faster development or regulatory review or approval process and it does not increase the likelihood that our product candidates will receive marketing approval.***

We plan to seek accelerated approval of our lead product candidates and may seek approval of future product candidates, where applicable, using the FDA's accelerated approval pathway. A product may be eligible for accelerated approval if it treats a serious or life-threatening condition and generally provides a meaningful advantage over available therapies. In addition, it must demonstrate 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. In addition, the FDA generally requires 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 pathway for any of our product candidates, we may not be able to obtain accelerated approval, and even if we do, that product may not experience a faster development or regulatory review or approval process. In addition, receiving accelerated approval does not assure the product's accelerated approval will eventually be converted to a traditional approval.

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

Healthcare providers and third-party payors in the United States and elsewhere will play a primary role in the recommendation and prescription of any product candidates for which we obtain marketing approval. Our current and future arrangements with healthcare professionals, principal investigators, consultants, customers and third-party payors subject us to various federal and state fraud and abuse laws and other healthcare laws that may constrain the business or financial arrangements and relationships through which we research, sell, market and distribute our product candidates, if we obtain marketing approval. In particular, the research of our product candidates, as well as the promotion, sales and marketing of healthcare items and services, as well as certain business arrangements in the healthcare industry, are subject to extensive laws designed to: (i) prevent fraud, kickbacks, self-dealing and other abusive practices, (ii) guarantee the security and privacy of health information, and (iii) increase transparency around the financial relationships between physicians, teaching hospitals and manufacturers of drugs, medical devices and biologics. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, structuring and commission(s), certain customer incentive programs and other business or financial arrangements. See the sections entitled "Business — Other Healthcare Laws" and "Business — Healthcare Reform" in our 2021 Annual Report.

Ensuring that our business arrangements and practices with third parties comply with applicable healthcare laws and regulations will likely be costly. It is possible that governmental authorities will conclude that our business practices may not comply with current or future statutes, regulations or case law involving applicable fraud and abuse or other healthcare laws and regulations. If our operations are found to be in violation of any of these laws or any other governmental regulations that may apply to us, we may be subject to significant civil, criminal and administrative penalties, damages, fines, disgorgement, imprisonment, exclusion from participating in government funded healthcare programs such as Medicare and Medicaid, additional reporting requirements and oversight if we become subject to a corporate integrity agreement or similar agreement to resolve allegations of non-compliance with these laws,

contractual damages, reputational harm and the curtailment or restructuring of our operations. The shifting compliance environment and the need to build and maintain robust and expandable systems to comply with multiple jurisdictions with different compliance or reporting requirements increases the possibility that a healthcare company may run afoul of one or more of the requirements.

If the physicians or other providers or entities with whom we expect to do business are found not to comply with applicable laws, they may be subject to significant criminal, civil or administrative sanctions, including exclusions from government funded healthcare programs. Even if resolved in our favor, litigation or other legal proceedings relating to healthcare laws and regulations 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. If securities analysts or investors perceive these results to be negative, it could have a substantial adverse effect on the price of our common stock. Litigation or proceedings of this nature could substantially increase our operating losses and reduce the resources available for development, manufacturing, sales, marketing or distribution activities. Uncertainties resulting from the initiation and continuation of litigation or other proceedings relating to applicable healthcare laws and regulations could have an adverse effect on our ability to compete in the marketplace.

***The successful commercialization of our product candidates in the United States and abroad will depend in part on the extent to which third-party payors, including governmental authorities and private health insurers, provide coverage and adequate reimbursement levels, as well as implement pricing policies favorable for our product candidates. Failure to obtain or maintain coverage and adequate reimbursement for our product candidates, if approved, could limit our ability to market those products and decrease our ability to generate revenue.***

Significant uncertainty exists as to the coverage and reimbursement status of any products for which we may obtain regulatory approval. In the United States and in other countries, patients who are provided medical treatment for their conditions generally rely on third-party payors to reimburse all or part of the costs associated with their treatment. The availability of coverage and adequacy of reimbursement for our products by third-party payors, including government health care programs (e.g., Medicare, Medicaid or TRICARE in the United States), managed care providers, private health insurers, health maintenance organizations and other organizations is essential for most patients to be able to afford medical services and pharmaceutical products such as our product candidates. Third-party payors decide which medications they will pay for and establish reimbursement levels. See the section entitled “Coverage and Reimbursement” in our 2021 Annual Report.

Our ability to successfully commercialize our product candidates will depend in part on the extent to which coverage and adequate reimbursement for our products and related treatments will be available from third-party payors. Moreover, a payor’s decision to provide coverage for a product does not imply that an adequate reimbursement rate will be approved. If coverage and adequate reimbursement is not available, or is available only to 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 high enough to allow us to establish or maintain pricing sufficient to realize a sufficient return on our investment. A decision by a third-party payor not to cover or not to separately reimburse for our medical products or therapies using our products could reduce physician utilization of our products once approved.

We cannot be sure that coverage and reimbursement in the United States and other countries will be available for our current or future product candidates or for any procedures using our current or future product candidates, and any reimbursement that may become available may not be adequate or may be decreased or eliminated in the future.

In the United States, no uniform policy for coverage and reimbursement for products exists among third-party payors. Therefore, coverage and reimbursement for our products can differ significantly from payor to payor. The process for determining whether a payor will provide coverage for a product may be separate from the process for setting the reimbursement rate that the payor will pay for the product. One payor’s determination to provide coverage for a product does not assure that other payors will also provide coverage and reimbursement for the product. Third-party payors may also limit coverage to specific products on an approved list, or formulary, which might not include all of the FDA-approved products for a particular indication. The principal decisions about reimbursement for new medicines in the United States are typically made by the Centers for Medicare & Medicaid Services, or CMS, an agency within the U.S. Department of Health and Human Services, or HHS. CMS will decide whether and to what extent our products will be covered and reimbursed under Medicare and private payors tend to follow CMS to a substantial degree. Factors considered by payors in determining reimbursement are based on whether the product is:

- a covered benefit under its health plan;
- safe, effective and medically necessary;
- appropriate for the specific patient;
- cost-effective; and
- neither experimental nor investigational.

We cannot be sure that coverage and reimbursement will be available for or accurately estimate the potential revenue from our product candidates or assure that coverage and reimbursement will be available for any product that we may develop.

Further, increasing efforts by third-party payors in the United States and abroad to cap or reduce healthcare costs may cause payor organizations to limit both coverage and the level of reimbursement for newly approved products and, as a result, they may not cover or provide adequate payment for our product candidates. In order to secure coverage and reimbursement for any product that might be approved for sale, we may need to conduct expensive pharmacoeconomic studies in order to demonstrate the medical necessity and cost-effectiveness of our products, in addition to the costs required to obtain FDA or comparable regulatory approvals. Additionally, we may also need to provide discounts to purchasers, private health plans or government healthcare programs. Our product candidates may nonetheless not be considered medically necessary or cost-effective. If third-party payors do not consider a product to be cost-effective compared to other available therapies, they may not cover the product after approval as a benefit under their plans or, if they do, the level of payment may not be sufficient to allow a company to sell its products at a profit. We expect to experience pricing pressures from third-party payors in connection with the potential sale of any of our product candidates.

Lastly, in some foreign countries, the proposed pricing for a drug must be approved before it may be lawfully marketed. The requirements governing drug pricing vary widely from country to country. For example, countries in the European Union, or EU, Member States can restrict the range of medicinal products for which their national health insurance systems provide reimbursement and they can control the prices of medicinal products for human use. To obtain reimbursement or pricing approval, some of these countries may require the completion of clinical trials that compare the cost effectiveness of a particular product candidate to currently available therapies. An EU Member State may approve a specific price for the medicinal product, or it may instead adopt a system of direct or indirect controls on the profitability of the company placing the medicinal product on the market. Approaches between EU Member States are diverging. For example, in France, effective market access will be supported by agreements with hospitals and products may be reimbursed by the Social Security Fund. The price of medicines is negotiated with the Economic Committee for Health Products. There can be no assurance that any country that has price controls or reimbursement limitations for pharmaceutical products will allow favorable reimbursement and pricing arrangements for any of our product candidates. Historically, products launched in the European Union do not follow price structures of the United States and generally prices in the European Union tend to be significantly lower than prices in the United States.

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

In the United States and other jurisdictions, there have been, and we expect there will continue to be, a number of legislative and regulatory changes and proposed changes to the healthcare system that could affect our future results of operations. In particular, there have been and continue to be a number of initiatives at the U.S. federal and state levels that seek to reduce healthcare costs and improve the quality of healthcare. Individual states in the United States have also increasingly passed legislation and implemented regulations designed to control pharmaceutical and biological product pricing, including price or patient reimbursement constraints, discounts, restrictions on certain product access and marketing cost disclosure and transparency measures and, in some cases, designed to encourage importation from other countries and bulk purchasing. See the section entitled “*Business — Healthcare Reform*” in our 2021 Annual Report.

We expect that additional state and federal healthcare reform measures will be adopted in the future, any of which could limit the extent to which state and federal governments cover particular healthcare products and services and could limit the amounts that the federal and state governments will pay for healthcare products and services. This could result in reduced demand for any product candidate we develop or could result in additional pricing pressures.

In markets outside of the United States, reimbursement and healthcare payment systems vary significantly by country and many countries have instituted price ceilings on specific products and therapies. The price control regulations outside of the United States can have a significant impact on the profitability of a given market, and further uncertainty is introduced if and when these laws change.

We cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative action in the United States or any other jurisdiction. It is possible that additional governmental action will be taken in response to the ongoing COVID-19 pandemic. If we or any third parties we may engage are slow or unable to adapt to changes in existing requirements or the adoption of new requirements or policies, or if we or these third parties are not able to maintain regulatory compliance, our product candidates may lose any regulatory approval that may have been obtained and we may not achieve or sustain profitability.

***We may face potential liability under applicable privacy laws if we obtain identifiable patient health information from clinical trials sponsored by us.***

Most healthcare providers, including certain research institutions from which we may obtain patient health information, are subject to privacy and security regulations promulgated under the Health Insurance Portability and Accountability Act of 1966, or HIPAA, as amended by the Health Information Technology for Economic and Clinical Health Act. Depending on the facts and circumstances, we could be subject to civil, criminal, and administrative penalties if we obtain, use, or disclose individually identifiable health information maintained by a HIPAA-covered entity in a manner that is not authorized or permitted by HIPAA. In addition, we may be subject to state laws requiring notification of affected individuals and state regulators in the event of a breach of personal information, which is a broader class of information than the health information protected by HIPAA.

The global data protection landscape is rapidly evolving, and we may be or become subject to or affected by numerous federal, state and foreign laws and regulations, as well as regulatory guidance, governing the collection, use, disclosure, transfer, security and processing of personal data, such as information that we collect about participants and healthcare providers in connection with clinical trials. Implementation standards and enforcement practices are likely to remain uncertain for the foreseeable future, which may create uncertainty in our business, affect our or our service providers' ability to operate in certain jurisdictions or to collect, store, transfer use and share personal data, result in liability or impose additional compliance or other costs on us. Any failure or perceived failure by us to comply with federal, state or foreign laws or self-regulatory standards could result in negative publicity, diversion of management time and effort and proceedings against us by governmental entities or others. Further, the California Consumer Privacy Act of 2018, or the CCPA, went into effect in January 2020. The CCPA provides new data privacy rights for consumers and new operational requirements for companies, including placing increased privacy and security obligations on entities handling certain personal data of consumers or households. These requirements could increase our compliance costs and potential liability. The CCPA gives California residents expanded rights to access and delete their personal information, opt out of certain personal information sharing and receive detailed information about how their personal information is used. The CCPA provides for civil penalties for violations, as well as a private right of action for data breaches that is expected to increase data breach litigation. While there is currently an exception for protected health information that is subject to HIPAA and clinical trial regulations, as currently written, the CCPA may impact certain of our business activities. The new California law may lead to similar laws in other U.S. states or at a national level, which could increase our potential liability and adversely affect our business, which exemplifies the vulnerability of our business to the evolving regulatory environment related to personal data and protected health information.

We expect that there will continue to be new laws, regulations and industry standards concerning privacy, data protection and information security proposed and enacted in various international jurisdictions. For example, European legislators adopted the EU General Data Protection Regulation, or GDPR, which became effective on May 25, 2018. The GDPR further implemented through binding guidance by the European Data Protection Board, or EDPB, (and supplemented by national laws in individual EU member states), imposes more stringent data protection compliance requirements and provides for more significant penalties for noncompliance in Europe. The GDPR creates new compliance obligations that may be applicable to our business, which could cause us to change our business practices, and increases financial penalties for noncompliance (including possible fines of up to the greater of €20 million and 4% of our global annual turnover for the preceding financial year for the most serious violations, as well as the right to compensation for financial or non-financial damages claimed by any individuals under Article 82 of the GDPR).

The GDPR also confers a private right of action on data subjects and consumer associations to lodge complaints with supervisory authorities, seek judicial remedies and obtain compensation for damages resulting from violations of the GDPR. In addition, the GDPR includes restrictions on cross-border data transfers. The GDPR may increase our responsibility and liability in relation to personal data that we process where that processing is subject to the GDPR. In addition, we may be required to put in place additional mechanisms to ensure compliance with the GDPR, including GDPR requirements as implemented by individual countries.

We are taking steps to comply with the GDPR as appropriate and as and when applicable to us, but this is an ongoing compliance process. Compliance with the GDPR will be a rigorous and time-intensive process that may increase our cost of doing business or require us to change our business practices. If our efforts to comply with GDPR or other applicable EU laws and regulations are not successful, or are perceived to be unsuccessful, it could adversely affect our business in the EU.

European data protection laws also generally prohibit the transfer of personal data from Europe, including the European Economic Area, United Kingdom and Switzerland, to the United States and most other countries unless the parties to the transfer have implemented specific safeguards to protect the transferred personal data. One of the primary safeguards used for transfers of personal data from the European Union to the United States, namely, the Privacy Shield framework administered by the U.S. Department of Commerce, was recently invalidated by a decision of the European Union's highest court. The same decision also cast doubt on the ability to use one of the primary alternatives to the Privacy Shield, namely, the European Commission's Standard Contractual Clauses, to lawfully transfer personal data from Europe to the United States and most other countries. At present, there are few if any viable alternatives to the Privacy Shield and the Standard Contractual Clauses. To the extent that we were to rely on the EU-U.S. Privacy Shield Framework or the Standard Contractual Clauses, we may not be able to do so in the future, which could increase our costs and limit our ability to process personal data from the EU.

Further, the United Kingdom's departure from the EU, often referred to as Brexit, has created uncertainty with regard to data protection regulation in the United Kingdom and how data transfers to and from the United Kingdom will be regulated. We may, however, incur liabilities, expenses, costs and other operational losses under the GDPR and applicable EU Member States and the United Kingdom privacy laws in connection with any measures we take to comply with them.

Further, certain health privacy laws, data breach notification laws, consumer protection laws and genetic testing laws may apply directly to our operations and/or those of our collaborators and may impose restrictions on our collection, use and dissemination of individuals' health information. Patients about whom we or our collaborators may obtain health information, as well as the providers who may share this information with us, may have statutory or contractual rights that limit our ability to use and disclose the information. We may be required to expend significant capital and other resources to ensure ongoing compliance with applicable privacy and data security laws. Claims that we have violated individuals' privacy rights or breached our contractual obligations, 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.

If we or third-party CMOs, CROs or other contractors or consultants fail to comply with applicable federal, state/provincial or local regulatory requirements, we could be subject to a range of regulatory actions that could affect our or our contractors' ability to develop and commercialize our therapeutic candidates and could harm or prevent sales of any affected therapeutics that we are able to commercialize, or could substantially increase the costs and expenses of developing, commercializing and marketing our therapeutics. Any threatened or actual government enforcement action could also generate adverse publicity and require that we devote substantial resources that could otherwise be used in other aspects of our business. Increasing use of social media could give rise to liability, breaches of data security or reputational damage.

Additionally, we are subject to state and foreign equivalents of each of the healthcare laws described above, among others, some of which may be broader in scope and may apply regardless of the payor.

***If we or our third-party manufacturers and suppliers 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 an 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 research and development activities involve the use of biological and hazardous materials and 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, which could cause an interruption of our commercialization efforts, research and development efforts and business operations, environmental damage resulting in costly clean-up and liabilities under applicable laws and regulations governing the use, storage, handling and disposal of these materials and specified waste products. Although we believe that the safety procedures utilized by our third-party CMOs for handling and disposing of these materials generally comply with the standards prescribed by these laws and regulations, we cannot guarantee that this is the case or eliminate the risk of accidental contamination or injury from these materials. Upon an event of this nature, we may be held liable for any resulting damages and such liability could exceed our resources and state or federal or other applicable authorities may curtail our use of certain materials and/or interrupt our business operations. Further, environmental laws and regulations are complex, change frequently and have tended to become more stringent. We cannot predict the impact of any changes of this nature and cannot be certain of our future compliance. 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. Failure to comply with these laws and regulations also may result in substantial fines, penalties, or other sanctions.

Although we maintain workers' compensation insurance to cover us for costs and expenses, we may incur due to injuries to our employees resulting from the use of hazardous materials or other work-related injuries, this insurance may not provide adequate coverage against potential liabilities. We do not carry specific biological waste or hazardous waste insurance coverage, workers compensation or property and casualty and general liability insurance policies that include coverage for damages and fines arising from biological or hazardous waste exposure or contamination.

***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 of 2001 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. In the future, we may engage third parties for clinical trials outside of the United States, to sell our products abroad once we enter a commercialization phase and/or to obtain necessary permits, licenses, patent registrations and other regulatory approvals. We may also 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 these 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 employee matters, managing growth and operational matters**

*We are highly dependent on our key personnel and anticipate hiring new key personnel. If we are not successful in attracting and retaining highly qualified personnel, we may not be able to successfully implement our business strategy.*

Our ability to compete in the highly competitive biotechnology and pharmaceutical industries depends upon our ability to attract and retain highly qualified managerial, scientific, medical personnel, sales and marketing and other personnel. We are highly dependent on our management, scientific and medical personnel, including our President and Chief Executive Officer, Chief Scientific Officer, Chief Medical Officer, Chief Financial Officer, Chief Legal Officer, Chief People Officer, and Chief Business Officer. 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.

We conduct our operations at our facilities in Watertown, Massachusetts. The Massachusetts region is headquarters to many other biopharmaceutical companies and many academic and research institutions. 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 U.S. 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 involving personnel who are not U.S. citizens.

To encourage valuable employees to remain at our company, in addition to salary and cash incentives, we have provided stock options that vest over time. The value to our employees of stock options that vest over time may be significantly affected by movements in our stock 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 on short notice. Although we have employment agreements with our executive employees, these employment agreements provide for at-will employment, which means that any of our executive employees could leave our employment at any time, with or without notice. 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, medical, and general and administrative personnel.

In addition, we have scientific and clinical advisors who assist us in formulating our development and clinical strategies. These advisors are not our employees and may have commitments to, or consulting or advisory contracts with, other entities that may limit their availability to us. In addition, our advisors may have arrangements with other companies to assist those companies in developing products or technologies that may compete with ours.

*We will need to grow the size of our organization and we may experience difficulties in managing this growth, which could disrupt our operations.*

We expect to experience significant growth in the number of our employees and the scope of our operations, particularly in the areas of drug development, manufacturing, regulatory affairs and, if any of our product candidates receives marketing approval, sales, marketing, and distribution. In addition, in connection with our transition to being a publicly traded company, we expect to increase the size of our general and administrative teams to support the growth of our business and the requirements of being a publicly traded company. To manage our anticipated future growth, we must continue to implement and improve our managerial, operational, and financial systems, expand our facilities, and continue to recruit and train additional qualified personnel. 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 lead product candidates and any other product candidates we develop, while complying with our contractual obligations to contractors and other third parties; and
- improving our operational, financial and management controls, reporting systems and procedures.

Our future financial performance and our ability to advance into clinical development and, if approved, commercialize our lead product candidates and any of our other product candidates we develop will depend, in part, on our ability to effectively manage any future growth. Due to our limited financial resources and the limited experience of our management team in managing a company with this type of anticipated growth, we may not be able to effectively manage the expansion of our operations or recruit and train additional qualified personnel. The expansion of our operations may lead to significant costs and may divert our management and business development resources. Any inability to manage growth could delay the execution of our business plans or disrupt our operations. Further, research at our Chinese and Indian CROs also exposes us to various risks, including regulatory, economic, and political instability, potentially unfavorable tax, import and export policies, fluctuations in foreign exchange and inflation rates, international and civil hostilities, terrorism, natural disasters and pandemics.

***Our internal computer systems, or those of any of our collaborators, contractors, or consultants, may fail or suffer security breaches, which could result in a material disruption of our product development programs.***

Our internal computer systems and those of any collaborators, contractors or consultants are vulnerable to damage from computer viruses, unauthorized access, natural disasters, terrorism, war and telecommunication and electrical failures. While we have not experienced any material system failure, accidents, or security breaches of this nature to date, if an event of this nature were to occur and cause interruptions in our operations, 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 proprietary information or other similar disruptions. For example, the loss of clinical trial data from completed or future clinical trials could result in delays in our marketing approval efforts and significantly increase our costs to recover or reproduce the data. To the extent that any disruption or security breach were to result in a loss of or damage to our data or applications or the inappropriate disclosure of confidential or proprietary information, we could incur liability, our competitive position could be harmed, and the further development and commercialization of our product candidates could be delayed. Additionally, we may have data security obligations with respect to the information of third parties that we store. Unauthorized access or use of any third-party data or information of this nature could result in fines or other penalties that may impact our relationships with these third parties and our operations.

***Our employees, independent contractors, vendors, principal investigators, CROs and consultants may engage in misconduct or other improper activities, including non-compliance with regulatory standards and requirements and insider trading laws.***

We are exposed to the risk that our employees, independent contractors, vendors, principal investigators, CROs, CMOs and consultants may engage in fraudulent conduct or other illegal activity. Misconduct by these parties could include, among other things:

- intentional, reckless, or negligent conduct or disclosure of unauthorized activities that violate study and trial protocols or the regulations of the FDA or similar foreign regulatory authorities;
- violations of healthcare fraud and abuse laws and regulations in the United States and abroad;
- violations of United States federal securities laws relating to trading in our common stock; and
- failures to report financial information or data accurately.

In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, misconduct, kickbacks, self-dealing and other abusive practices. These laws and regulations regulate a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Other forms of misconduct could involve the improper use of information obtained in the course of clinical trials or creating fraudulent data in our preclinical studies or clinical trials, which could result in regulatory sanctions and cause serious harm to our reputation. We have adopted a code of business conduct and ethics and other corporate governance and compliance documents, policies and charters applicable to all of our employees. However, it is not always possible to identify and deter misconduct by employees and other third parties. Further, the precautions we take to detect and prevent this type of 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 comply with these laws or regulations. Additionally, we are subject to the risk that a person could allege fraud or other misconduct, even if none occurred. If any actions of this nature 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 civil, criminal and administrative penalties, damages, monetary fines, possible exclusion from participation in Medicare, Medicaid and other federal healthcare programs, contractual damages, reputational harm, diminished profits and future earnings and curtailment of our operations, any of which could adversely affect our business prospects, financial condition and results of operations.

#### **Risks related to our common stock**

***If we were to determine to raise additional capital in the future, you would suffer dilution of your investment.***

We may choose to raise additional capital in the future through the sale of shares or other securities convertible into shares, depending on market conditions, strategic considerations and operational requirements. To the extent we raise additional capital in this manner, our stockholders will be diluted. Future issuances of our common stock or other equity securities, or the perception that sales of this nature may occur, could adversely affect the trading price of our common stock and impair our ability to raise capital through future offerings of shares or equity securities. No prediction can be made as to the effect, if any, that future sales of common stock or the availability of common stock for future sales will have on the trading price of our common stock.

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

Prior to our initial public offering, there was no public trading market for shares of our common stock. Although our common stock is listed on The Nasdaq Global Select Market, an active trading market for our shares may not be sustained. You may not be able to sell your shares quickly or at the market price if trading in shares of our common stock is not active. Further, an inactive market may also impair our ability to raise capital by selling shares of our common stock and may impair our ability to enter into strategic partnerships or acquire companies or products by using our shares of common stock as consideration.

***If securities or industry analysts do not publish or cease publishing research or reports about our business, or if they issue an adverse or misleading opinion regarding our stock, our stock price and trading volume could decline.***

The trading market for our common stock is and will continue to be influenced by the research and reports that industry or securities analysts publish about us, our business or the targeted protein degradation space. We do not have control over these analysts. There can be no assurance that existing analysts will continue to provide research coverage or that new analysts will begin to provide coverage. Although we have obtained analyst coverage, if any of the analysts who cover us were to issue an adverse or misleading opinion regarding us, our business model, our intellectual property or our stock performance, or if our preclinical studies and future clinical trials and results of operations fail to meet the expectations of any of these analysts, our stock price would likely decline. If one or more of these covering analysts were to cease coverage of us or fail to publish reports on us regularly, we could lose visibility in the financial markets, which in turn could cause a decline in our stock price or trading volume.

***The price of our common stock may be volatile and fluctuate substantially, which could result in substantial losses for purchasers of our common stock.***

The trading price of shares of our common stock may be volatile and subject to wide fluctuations in response to various factors, some of which we cannot control. The stock market in general, and the market for smaller biopharmaceutical companies in particular, have experienced extreme volatility that has often been unrelated to the operating performance of particular companies. As a result of this volatility, you may not be able to sell your common stock at or above the price at which you acquired it. The market price for our common stock may be influenced by many factors, including:

- the degree of success of competitive products or technologies or changes in standard of care regimens;
- results of preclinical studies and clinical trials of our product candidates or those of our competitors;
- the timing and progress of our clinical development activities;
- regulatory or legal developments in the United States and other countries;
- developments or disputes concerning patent applications, issued patents or other proprietary rights;
- the recruitment or departure of key personnel;
- the level of expenses related to any of our product candidates or clinical development programs;
- the results of our efforts to discover, develop, acquire or in-license additional technologies or product candidates;
- actual or anticipated changes in estimates as to financial results, development timelines or recommendations by securities analysts;
- variations in our financial results or those of companies that are perceived to be similar to us;
- changes in the structure of healthcare payment systems;
- market conditions in the pharmaceutical and biotechnology sectors;
- effects of public health crises, pandemics and epidemics, such as COVID-19;
- general economic, industry and market conditions; and
- the other factors described in this “Risk Factors” section.

If any of the foregoing matters were to occur or if our operating results fall below the expectations of investors or securities analysts, the price of our common stock could decline substantially. In the past, following periods of volatility in the market price of a company’s securities, securities class-action litigation often has been instituted against that company. Litigation of this nature, if instituted against us, could cause us to incur substantial costs to defend these claims and divert management’s attention and resources, which could seriously harm our business, financial condition, results of operations and prospects. Further, our director and officer liability insurance cost may increase as a result of litigation of this nature and our insurance deductible may be significant before our insurers are required to provide any coverage to us.

***We have broad discretion in the use of the capital we have raised and may not use our capital effectively.***

Our management has broad discretion in the application of the net proceeds from our prior financings, including our initial and follow-on public offerings, and could spend the proceeds in ways that do not improve our results of operations or enhance the value of our common stock. The failure by our management to apply these funds effectively could result in financial losses that could have an adverse effect on our business, cause the price of our common stock to decline and delay the development of our product candidates. Pending their use, we may invest the net proceeds from our financing activities in a manner that does not produce income or that loses value.

***Our executive officers, directors and principal stockholders will have the ability to control or significantly influence matters submitted to stockholders for approval.***

Our executive officers and directors, combined with our stockholders who have reported through filings made with the Securities and Exchange Commission that they own more than 5% of our outstanding common stock, in the aggregate, beneficially own a significant percentage of our shares. As a result, our executive officers and directors, combined with our greater than 5% stockholders, have the ability to control us through this ownership position. These stockholders, if acting together, will consequently continue to control matters submitted to our stockholders for approval, as well as our management and affairs. For example, these persons, if they choose to act together, would control the election of directors and approval of any merger, consolidation or sale of all or substantially all of our assets. This concentration of ownership control may:

- delay, defer or prevent a change in control;
- entrench our management and the board of directors; or
- impede a merger, consolidation, takeover or other business combination involving us that other stockholders may desire.

***Anti-takeover provisions under our charter documents and Delaware law could delay or prevent a change of control, which could limit the market price of our common stock and may prevent or frustrate attempts by our stockholders to replace or remove our current management.***

Our amended and restated certificate of incorporation and amended and restated bylaws contain provisions that could delay or prevent a change of control of our company or changes in our board of directors that our stockholders might consider favorable. Some of these provisions include:

- a board of directors divided into three classes serving staggered three-year terms, the result of which is that not all members of the board will be elected at one time;
- a prohibition on stockholder action through written consent, the result of which is that all stockholder actions will have to be taken at a meeting of our stockholders;
- a requirement that special meetings of stockholders be called only by the board of directors acting pursuant to a resolution approved by the affirmative vote of a majority of the directors then in office;
- advance notice requirements for stockholder proposals and nominations for election to our board of directors;
- a requirement that no member of our board of directors may be removed from office by our stockholders except for cause and, in addition to any other vote required by law, upon the approval of not less than two-thirds of all outstanding shares of our voting stock then entitled to vote in the election of directors;
- a requirement of approval of not less than two-thirds of all outstanding shares of our voting stock to amend any bylaws by stockholder action or to amend specific provisions of our certificate of incorporation; and
- the authority of the board of directors to issue preferred stock on terms determined by the board of directors without stockholder approval and which preferred stock may include rights superior to the rights of the holders of common stock.

In addition, because we are incorporated in Delaware, we are governed by the provisions of Section 203 of the Delaware General Corporate Law, which may prohibit certain business combinations with stockholders owning 15% or more of our outstanding voting stock. These antitakeover provisions and other provisions in our amended and restated certificate of incorporation and amended and restated bylaws could make it more difficult for stockholders or potential acquirers to obtain control of our board of directors or initiate actions that are opposed by the then-current board of directors and could also delay or impede a merger, tender offer or proxy contest involving our company. These provisions could also discourage proxy contests and make it more difficult for you and other stockholders to elect directors of your choosing or cause us to take other corporate actions you desire. Any delay or prevention of a change of control transaction or changes in our board of directors could cause the market price of our common stock to decline.

***We will continue to incur additional costs as a result of operating as a public company and our management will be required to devote substantial time to new compliance initiatives and corporate governance practices.***

As a public company, we will continue to incur significant legal, accounting and other expenses that we did not incur as a private company. The Sarbanes-Oxley Act of 2002, or Sarbanes-Oxley, the Dodd-Frank Wall Street Reform and Consumer Protection Act, the listing requirements of The Nasdaq Stock Market LLC and other applicable securities rules and regulations impose various requirements on public companies, including establishment and maintenance of effective disclosure and financial controls and corporate governance practices. Our management and other personnel devote a substantial amount of time to these compliance initiatives. Moreover, these rules and regulations increase our legal and financial compliance and insurance costs and make some activities more time-consuming and costly. For example, these rules and regulations make it more difficult and more expensive for us to obtain director and officer liability insurance, which in turn could make it more difficult for us to attract and retain qualified members of our board of directors.

We continually evaluate these rules and regulations and cannot always predict or estimate the amount of additional costs we may incur or the timing of these costs. These rules and regulations are also often subject to varying interpretations, in many cases due to their lack of specificity and, as a result, their application in practice may evolve over time as new guidance is provided by regulatory and governing bodies. This could result in continuing uncertainty regarding compliance matters and higher costs necessitated by ongoing revisions to disclosure and governance practices.

Pursuant to Section 404 of Sarbanes-Oxley, or Section 404, we are required to furnish a report by our management on our internal control over financial reporting, including an attestation report on internal control over financial reporting issued by our independent registered public accounting firm. However, while we were an “emerging growth company” as defined in the Jumpstart Our Business Startups Act, our auditors were not required to formally attest to the effectiveness of our internal control over financial reporting. As of the end of our fiscal year ended December 31, 2021, we qualified as a “large accelerated filer” as defined in the Securities Exchange Act of 1934, as amended, or the Exchange Act and, as a result, ceased to qualify as an emerging growth company. Our compliance with Section 404 necessitates that we incur substantial accounting expense and expend significant management efforts.

We will need to continue to dedicate internal resources, potentially engage outside consultants and adopt a detailed work plan to assess and document the adequacy of internal control over financial reporting, continue steps to improve control processes as appropriate, validate through testing that controls are functioning as documented and implement a continuous reporting and improvement process for internal control over financial reporting. Despite our efforts, there is a risk that neither we nor our independent registered public accounting firm will be able to conclude, within the prescribed timeframe or at all, that our internal control over financial reporting is effective as required by Section 404. Further, we cannot assure you that the measures we have taken in the past or will take in the future will prevent the occurrence of future material weaknesses or significant deficiencies in our internal control over financial reporting. If we identify one or more material weaknesses in the future, it could result in an adverse reaction in the financial markets and restrict our future access to the capital markets due to a loss of confidence in the reliability of our consolidated financial statements.

***Our amended and restated bylaws designate specific courts as the exclusive forum for certain litigation that may be initiated by our stockholders, which could limit our stockholders’ ability to obtain a favorable judicial forum for disputes with us.***

Pursuant to our amended and restated bylaws, unless we consent in writing to the selection of an alternative forum, the Court of Chancery of the State of Delaware is the sole and exclusive forum for state law claims for (1) any derivative action or proceeding brought on our behalf; (2) any action asserting a claim of breach of a fiduciary duty owed by any of our directors, officers, or other employees to us or our stockholders; (3) any action asserting a claim arising pursuant to any provision of the Delaware General Corporation Law or our certificate of incorporation or bylaws; (4) any action to interpret, apply, enforce or determine the validity of our certificate of incorporation or bylaws; or (5) any action asserting a claim governed by the internal affairs doctrine. We refer to this provision in our bylaws as the Delaware Forum Provision. The Delaware Forum Provision will not apply to any causes of action arising under the Securities Act of 1933, as amended, the Securities Act, or the Exchange Act of 1934, as amended, or the Exchange Act. Our amended and restated bylaws further provide that unless we consent in writing to the selection of an alternative forum, the United States District Court for the District of Massachusetts shall be the sole and exclusive forum for resolving any complaint asserting a cause of action arising under the Securities Act, as our headquarters are located in Watertown, Massachusetts. We refer to this provision in our bylaws as the Federal Forum Provision. In addition, our amended and restated bylaws provide that any person or entity purchasing or otherwise acquiring any interest in shares of our capital stock is deemed to have notice of and consented to the Delaware Forum Provision and the Federal Forum Provision; provided, however, that stockholders 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 Delaware Forum Provision and the Federal Forum Provision in our bylaws may impose additional litigation costs on stockholders in pursuing any such claims. Additionally, these forum selection clauses may limit our stockholders’ ability 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 stockholders. 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 will enforce our Federal Forum Provision. If the Federal Forum Provision is found to be unenforceable, we may incur additional costs associated with resolving such matters. The Federal Forum Provision may also impose additional litigation costs on stockholders who assert that the provision is not enforceable or invalid. The Court of Chancery of the State of Delaware and the United States District Court for the District of Massachusetts may also reach different judgments or results than would other courts, including courts where a stockholder 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 stockholders.

***Because we do not anticipate paying any cash dividends on our capital stock in the foreseeable future, capital appreciation, if any, will be your sole source of gain.***

We have never declared or paid cash dividends on our capital stock. We currently intend to retain all of our future earnings, if any, to finance the growth and development of our business. In addition, the terms of our Credit Agreement with Perceptive Credit also preclude us from paying dividends. As a result, capital appreciation, if any, of our common stock will be your sole source of gain for the foreseeable future.

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

Our results of operations could be adversely affected by general conditions in the global economy and in the global financial markets. For example, the global financial crisis caused extreme volatility and disruptions in the capital and credit markets. Similarly, the recent significant volatility associated with the ongoing COVID-19 pandemic has caused significant instability and disruptions in the capital and credit markets. Our operations could be adversely affected by economic and political changes in the markets, including inflation rates, supply chain disruptions, recessions, trade restrictions, tariff increases or potential new tariffs, and economic embargoes imposed by the U.S. A severe or prolonged economic downturn could result in a variety of risks to our business, including weakened demand for our product candidates, and could also impact our ability to raise additional capital when needed on acceptable terms, if at all. Our general business strategy may be adversely affected by any economic downturn of this nature, volatile business environment or continued unpredictable and unstable market conditions. If the current equity and credit markets deteriorate, or do not improve, it may make any necessary debt or equity financing more difficult, costly and dilutive.

Failure to secure any necessary financing in a timely manner and on favorable terms could have an adverse effect on our growth strategy, financial performance and stock price and could require us to delay, modify, or abandon clinical development plans. In addition, there is a risk that one or more of our current service providers, manufacturers and other partners may not survive these difficult economic times, which could directly affect our ability to attain our operating goals on schedule and on budget.

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 stock 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 adversely affect our business prospects, financial condition and results of operations.

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

***Recent Sales of Unregistered Securities***

Not applicable.

***Use of Proceeds from our Initial Public Offering of Common Stock***

In October 2020, our Registration Statement on Form S-1 (No. 333-248719) was declared effective by the SEC pursuant to which we issued and sold an aggregate of 11,040,000 shares of common stock (inclusive of shares of sold pursuant to the underwriters' exercise of their over-allotment option) at a public offering price of \$19.00 per share for aggregate net cash proceeds of \$191.2 million, after deducting underwriting discounts, commissions and offering costs. No payments for such expenses were made directly or indirectly to (i) any of our officers or directors or their associates, (ii) any persons owning 10% or more of any class of our equity securities or (iii) any of our affiliates.

The sale and issuance of 11,040,000 shares closed on October 6, 2020. Jefferies LLC, Evercore Group L.L.C., BMO Capital Markets Corp. and UBS Securities LLC acted as joint book-running managers for the offering.

There has been no material change in the planned use of proceeds from our initial public offering from that described in the final prospectus filed with the SEC pursuant to Rule 424(b) relating to our Registration Statement on Form S-1.

***Repurchase of Shares of Company Equity Securities***

Not applicable.

**Item 3. Defaults Upon Senior Securities.**

Not applicable.

**Item 4. Mine Safety Disclosures.**

Not applicable.

**Item 5. Other Information.**

None.

**Item 6. Exhibits.**

<b>Exhibit Number</b>	<b>Description</b>	<b>Form</b>	<b>File Number</b>	<b>Date of Filing</b>	<b>Exhibit Number</b>	<b>Filed Herewith</b>
3.1	<a href="#">Fifth Amended and Restated Certificate of Incorporation of the Registrant</a>	8-K	001-39567	10/06/2020	3.3	
3.2	<a href="#">Second Amended and Restated Bylaws of the Registrant</a>	S-1	333-248719	09/10/2020	3.5	
4.1	<a href="#">Amended and Restated Investors' Rights Agreement among the Registrant, its warrant holder and certain of its stockholders</a>	S-1	333-248719	09/10/2020	4.2	
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.</a>					X
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.</a>					X
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>					X
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>					X
101.INS	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 Calculation Linkbase Document					
101.DEF	Inline XBRL Taxonomy Extension Definition Linkbase Document					
101.LAB	Inline XBRL Taxonomy Label Linkbase Document					
101.PRE	Inline XBRL Taxonomy Presentation Linkbase Document					
104	Cover Page Interactive Data File (embedded within the Inline XBRL document)					

\* Exhibits 32.1 and 32.2 are being furnished and shall not be deemed to be "filed" for purposes of Section 18 of the Exchange Act, or otherwise subject to the liability of that section, nor shall such exhibit be deemed to be incorporated by reference in any registration statement or other document filed under the Securities Act, or the Exchange Act, except as otherwise stated in 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.

**C4 THERAPEUTICS, INC.**

Date: May 5, 2022

By:

/s/ Andrew J. Hirsch

**Andrew J. Hirsch**

*President and Chief Executive Officer  
(Principal Executive Officer)*

Date: May 5, 2022

By:

/s/ Lauren A. White

**Lauren A. White**

*Chief Financial Officer and Treasurer  
(Principal Financial and Principal  
Accounting Officer)*

**CERTIFICATION 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**

I, Andrew J. Hirsch, certify that:

1. I have reviewed this Quarterly Report on Form 10-Q of C4 Therapeutics, Inc.;
2. Based on my knowledge, this report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this report;
3. Based on my knowledge, the financial statements, and other financial information included in this report, fairly present in all material respects the financial condition, results of operations and cash flows of the registrant as of, and for, the periods presented in this report;
4. The registrant's other certifying officer(s) and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) and internal control over financial reporting (as defined in Exchange Act Rules 13a-15(f) and 15d-15(f)) for the registrant and have:
  - (a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under our supervision, to ensure that material information relating to the registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this report is being prepared;
  - (b) Designed such internal control over financial reporting, or caused such internal control over financial reporting to be designed under our supervision, to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles;
  - (c) Evaluated the effectiveness of the registrant's disclosure controls and procedures and presented in this report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this report based on such evaluation; and
  - (d) Disclosed in this report any change in the registrant's internal control over financial reporting that occurred during the registrant's most recent fiscal quarter (the registrant's fourth fiscal quarter in the case of an annual report) that has materially affected, or is reasonably likely to materially affect, the registrant's internal control over financial reporting; and
5. The registrant's other certifying officer(s) and I have disclosed, based on our most recent evaluation of internal control over financial reporting, to the registrant's auditors and the audit committee of the registrant's board of directors (or persons performing the equivalent functions):
  - (a) All significant deficiencies and material weaknesses in the design or operation of internal control over financial reporting which are reasonably likely to adversely affect the registrant's ability to record, process, summarize and report financial information; and
  - (b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the registrant's internal control over financial reporting.

Date: May 5, 2022

By: /s/ Andrew J. Hirsch

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**Andrew J. Hirsch**  
**Chief Executive Officer**

**CERTIFICATION 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**

I, Lauren A. White, certify that:

1. I have reviewed this Quarterly Report on Form 10-Q of C4 Therapeutics, Inc.;
2. Based on my knowledge, this report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this report;
3. Based on my knowledge, the financial statements, and other financial information included in this report, fairly present in all material respects the financial condition, results of operations and cash flows of the registrant as of, and for, the periods presented in this report;
4. The registrant's other certifying officer(s) and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) and internal control over financial reporting (as defined in Exchange Act Rules 13a-15(f) and 15d-15(f)) for the registrant and have:
  - (a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under our supervision, to ensure that material information relating to the registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this report is being prepared;
  - (b) Designed such internal control over financial reporting, or caused such internal control over financial reporting to be designed under our supervision, to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles;
  - (c) Evaluated the effectiveness of the registrant's disclosure controls and procedures and presented in this report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this report based on such evaluation; and
  - (d) Disclosed in this report any change in the registrant's internal control over financial reporting that occurred during the registrant's most recent fiscal quarter (the registrant's fourth fiscal quarter in the case of an annual report) that has materially affected, or is reasonably likely to materially affect, the registrant's internal control over financial reporting; and
5. The registrant's other certifying officer(s) and I have disclosed, based on our most recent evaluation of internal control over financial reporting, to the registrant's auditors and the audit committee of the registrant's board of directors (or persons performing the equivalent functions):
  - (a) All significant deficiencies and material weaknesses in the design or operation of internal control over financial reporting which are reasonably likely to adversely affect the registrant's ability to record, process, summarize and report financial information; and
  - (b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the registrant's internal control over financial reporting.

Date: May 5, 2022

By: /s/ Lauren A. White

**Lauren A. White**  
**Chief Financial Officer and Treasurer**

**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 C4 Therapeutics, Inc. (the "Company") on Form 10-Q for the period ending March 31, 2022, 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:

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

Date: May 5, 2022

By: /s/ Andrew J. Hirsch

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**Andrew J. Hirsch**  
**Chief Executive 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 C4 Therapeutics, Inc. (the "Company") on Form 10-Q for the period ending March 31, 2022, 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:

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

Date: May 5, 2022

By: /s/ Lauren A. White

**Lauren A. White**

**Chief Financial Officer and Treasurer**