

Dicerna Pharmaceuticals Inc  
Form 10-Q  
August 07, 2014  
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**UNITED STATES**  
**SECURITIES AND EXCHANGE COMMISSION**  
**Washington, DC 20549**

**Form 10-Q**

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

**For the quarterly period ended June 30, 2014**

**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: 001-36281**

**DICERNA PHARMACEUTICALS, INC.**

**(Exact name of registrant as specified in its charter)**

**Delaware**  
**(State or other jurisdiction of**  
**incorporation or organization)**

**20-5993609**  
**(IRS Employer**  
**Identification No.)**

**480 Arsenal Street,**  
**Building 1, Suite 120**  
**Watertown, MA 02472**

**(Address of principal executive offices and zip code)**

**(617) 621-8097**

**(Registrant's telephone number, including area code)**

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 and posted on its corporate Web site, if any, every Interactive Data File required to be submitted and posted pursuant to Rule 405 of Regulation S-T during the preceding 12 months (or for such shorter period that the registrant was required to submit and post such files). Yes  No

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

Large accelerated filer

Accelerated filer

Non-accelerated filer  (Do not check if a smaller reporting company)

Smaller reporting company

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 August 5, 2014, there were 17,768,771 shares of the registrant's common stock, par value \$0.0001 per share, outstanding.



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This Quarterly Report on Form 10-Q includes forward-looking statements within the meaning of Section 27A of the Securities Act of 1933, as amended, and Section 21E of the Securities Exchange Act of 1934, as amended. All statements other than statements of historical fact are forward-looking statements for purposes of this Quarterly Report on Form 10-Q. In some cases, you can identify forward-looking statements by terminology such as may, could, will, would, should, expect, plan, anticipate, believe, estimate, intend, predict, seek, contemporaneous, potential, ongoing or the negative of these terms or other comparable terminology. These forward-looking statements include, but are not limited to, statements about:

the initiation, timing, progress and results of our research and development programs, preclinical studies, any clinical trials and Investigational New Drug application, New Drug Application and other regulatory submissions;

our dependence on our existing collaborator, Kyowa Hakko Kirin Co., Ltd. (KHK), for developing, obtaining regulatory approval for and commercializing product candidates in the collaboration;

our receipt and timing of any milestone payments or royalties under our research collaboration and license agreement with KHK or arrangement with any future collaborator;

our ability to identify and develop product candidates for treatment of additional disease indications;

our or a collaborator's ability to obtain and maintain regulatory approval of any of our product candidates;

the rate and degree of market acceptance of any approved products candidates;

the commercialization of any approved product candidates;

our ability to establish and maintain additional collaborations and retain commercial rights for our product candidates in the collaborations;

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the implementation of our business model and strategic plans for our business, technologies and product candidates;

our estimates of our expenses, ongoing losses, future revenue and capital requirements;

our ability to obtain additional funds for our operations;

our ability to obtain and maintain intellectual property protection for our technologies and product candidates and our ability to operate our business without infringing the intellectual property rights of others;

our reliance on third parties to conduct our preclinical studies or any future clinical trials;

our reliance on third party supply and manufacturing partners to supply the materials and components for, and manufacture, our research and development, preclinical and clinical trial drug supplies;

our ability to attract and retain qualified key management and technical personnel;

our expectations regarding the time during which we will be an emerging growth company under the Jumpstart Our Business Startups Act;

our financial performance; and

developments relating to our competitors or our industry.

These statements relate to future events or to our future 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. Factors that may cause actual results to differ materially from current expectations include, among other things, those set forth in Part II, Item 1A Risk Factors below and for the reasons described elsewhere in this Quarterly Report on Form 10-Q. Any forward-looking statement in this Quarterly Report on Form 10-Q reflects our current view with respect to future events and is subject to these and other risks, uncertainties and assumptions relating to our operations, results of operations, industry and future growth. Given these uncertainties, you should not place undue reliance on these forward-looking statements. Except as required by law, we assume no obligation to update or revise these forward-looking statements for any reason, even if new information becomes available in the future.

Except where the context otherwise requires, in this Quarterly Report on Form 10-Q, we, us, our and the Company refer to Dicerna Pharmaceuticals, Inc. and, where appropriate, its consolidated subsidiary.

**Trademarks**

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This Quarterly Report on Form 10-Q includes trademarks, service marks and trade names owned by us or other companies. All trademarks, service marks and trade names included in this Quarterly Report on Form 10-Q are the property of their respective owners.

**Table of Contents****PART I. FINANCIAL INFORMATION****Item 1. Financial Statements****Dicerna Pharmaceuticals, Inc.****Condensed Consolidated Balance Sheets****(Unaudited)****(In thousands, except share and per share data)**

|  | <b>June 30,<br/>2014</b> | <b>December 31,<br/>2013</b> |
|--|--------------------------|------------------------------|
| <b>ASSETS</b>  |                          |                              |
| <b>CURRENT ASSETS:</b>                                 |                          |                              |
| Cash and cash equivalents                              | \$ 77,732                | \$ 46,595                    |
| Held-to-maturity investments                           | 22,526                   |                              |
| Prepaid expenses and other current assets              | 1,016                    | 2,058                        |
| Total current assets                                   | 101,274                  | 48,653                       |
| <b>NONCURRENT ASSETS:</b>                              |                          |                              |
| Held-to-maturity investments                           | 20,026                   |                              |
| Property and equipment net                             | 1,473                    | 877                          |
| Assets held in restriction                             | 1,380                    | 264                          |
| Total noncurrent assets                                | 22,879                   | 1,141                        |
| <b>TOTAL ASSETS</b>                                    | <b>\$ 124,153</b>        | <b>\$ 49,794</b>             |
| <b>LIABILITIES AND STOCKHOLDERS EQUITY / (DEFICIT)</b> |                          |                              |
| <b>CURRENT LIABILITIES:</b>                            |                          |                              |
| Accounts payable                                       | \$ 1,062                 | \$ 1,700                     |
| Accrued expenses and other current liabilities         | 2,808                    | 1,286                        |
| Current portion of long-term debt                      |                          | 4,587                        |
| Deferred rent  | 88                       | 105                          |
| Total current liabilities                              | 3,958                    | 7,678                        |
| <b>NONCURRENT LIABILITIES:</b>                         |                          |                              |
| Long-term debt net of current portion                  |                          | 260                          |
| Preferred stock warrant liability                      |                          | 529                          |
| Total noncurrent liabilities                           |                          | 789                          |
| <b>TOTAL LIABILITIES</b>                               | <b>3,958</b>             | <b>8,467</b>                 |



**REDEEMABLE CONVERTIBLE PREFERRED STOCK, \$0.0001 PAR VALUE NO SHARES AND 11,070,000 SHARES AUTHORIZED AT JUNE 30, 2014 AND DECEMBER 31, 2013, RESPECTIVELY; NO SHARES AND 10,589,434 SHARES ISSUED AND OUTSTANDING AT JUNE 30, 2014 AND DECEMBER 31, 2013, RESPECTIVELY:**

|  |                   |                  |
|--|-------------------|------------------|
| Series A 880,000 shares designated, 855,996 shares issued and outstanding at December 31, 2013 (aggregate liquidation preference of \$21,400 at December 31, 2013)   |                   | 21,400           |
| Series B 1,190,000 shares designated, 1,162,021 shares issued and outstanding at December 31, 2013 (aggregate liquidation preference of \$29,050 at December 31, 2013)   |                   | 29,050           |
| Series C 9,000,000 shares designated, 8,571,417 shares issued and outstanding at December 31, 2013 (aggregate liquidation preference of \$60,000 at December 31, 2013)   |                   | 59,796           |
| <b>STOCKHOLDERS EQUITY/(DEFICIT):</b>  |                   |                  |
| Preferred stock, \$0.0001 par value 5,000,000 shares and no shares authorized at June 30, 2014 and December 31, 2013, respectively; no shares issued and outstanding at June 30, 2014 and December 31, 2013, respectively                                  |                   |                  |
| Common stock, \$0.0001 par value 150,000,000 shares and 15,000,000 shares authorized at June 30, 2014 and December 31, 2013, respectively; 17,763,051 shares and 38,226 shares issued and outstanding at June 30, 2014 and December 31, 2013, respectively | 2                 | 1                |
| Additional paid-in capital   | 228,007           | 16,545           |
| Accumulated deficit  | (107,814)         | (85,465)         |
| Total stockholders equity/(deficit)  | 120,195           | (68,919)         |
| <b>TOTAL LIABILITIES AND STOCKHOLDERS EQUITY / (DEFICIT)</b>   | <b>\$ 124,153</b> | <b>\$ 49,794</b> |

The accompanying notes are an integral part of these condensed consolidated financial statements.

**Table of Contents****Dicerna Pharmaceuticals, Inc.****Condensed Consolidated Statements of Operations****(Unaudited)****(In thousands, except share and per share data)**

|  | <b>Three Months Ended June 30,</b> |             | <b>Six Months Ended</b> |             |
|--|------------------------------------|-------------|-------------------------|-------------|
|  | <b>2014</b>                        | <b>2013</b> | <b>June 30,</b>         | <b>2013</b> |
|  |                                    |             | <b>2014</b>             |             |
| Operating expenses:  |                                    |             |                         |             |
| Research and development   | \$ 6,806                           | \$ 2,513    | \$ 12,057               | \$ 4,931    |
| General and administrative   | 4,372                              | 1,142       | 7,213                   | 2,278       |
| Total operating expenses   | 11,178                             | 3,655       | 19,270                  | 7,209       |
| Loss from operations   | (11,178)                           | (3,655)     | (19,270)                | (7,209)     |
| Other income (expense):  |                                    |             |                         |             |
| Preferred stock warrant remeasurement                                    |                                    | 118         | (2,559)                 | 123         |
| Loss on extinguishment of debt   | (143)                              |             | (143)                   |             |
| Interest income  | 8                                  |             | 12                      |             |
| Interest expense   | (42)                               | (249)       | (199)                   | (519)       |
| Total other income (expense)   | (177)                              | (131)       | (2,889)                 | (396)       |
| Net loss   | \$ (11,355)                        | \$ (3,786)  | \$ (22,159)             | \$ (7,605)  |
| Less: Accretion and dividends on redeemable convertible preferred stock  |                                    | 1,022       | 204                     | 2,033       |
| Net loss attributable to common stockholders                             | (11,355)                           | (4,808)     | (22,363)                | (9,638)     |
| Net loss per share attributable to common stockholders basic and diluted | \$ (0.64)                          | \$ (171.13) | \$ (1.57)               | \$ (343.92) |
| Weighted average shares outstanding basic and diluted                    | 17,684,563                         | 28,095      | 14,272,401              | 28,024      |

The accompanying notes are an integral part of these condensed consolidated financial statements.

**Table of Contents****Dicerna Pharmaceuticals, Inc.****Condensed Consolidated Statements of Cash Flows****(Unaudited)****(In thousands, except share and per share data)**

|   | <b>Six Months Ended June 30,</b> |                |
|---|----------------------------------|----------------|
|   | <b>2014</b>                      | <b>2013</b>    |
| <b>CASH FLOWS FROM OPERATING ACTIVITIES:</b>                                |                                  |                |
| Net loss  | \$ (22,159)                      | \$ (7,605)     |
| Adjustments to reconcile net loss to net cash used in operating activities: |                                  |                |
| Depreciation  | 213                              | 271            |
| Stock-based compensation  | 4,617                            | 60             |
| Amortization of debt discount   | 48                               | 110            |
| Loss on extinguishment of debt  | 143                              |                |
| Increase (decrease) in fair value of preferred stock warrant liability      | 2,559                            | (123)          |
| Changes in operating assets and liabilities:                                |                                  |                |
| Research and license receivable   |                                  | 5,018          |
| Prepaid expenses and other current assets                                   | (944)                            | 58             |
| Accounts payable  | (272)                            | (512)          |
| Accrued expenses and other current liabilities                              | 1,542                            | 35             |
| Deferred rent   | (18)                             | (33)           |
| Net cash used in operating activities                                       | (14,271)                         | (2,721)        |
| <b>CASH FLOWS FROM INVESTING ACTIVITIES:</b>                                |                                  |                |
| Changes in assets held in restriction                                       | (1,116)                          |                |
| Purchases of property and equipment   | (808)                            | (3)            |
| Purchases of held-to-maturity investments                                   | (42,552)                         |                |
| Net cash used in investing activities                                       | (44,476)                         | (3)            |
| <b>CASH FLOWS FROM FINANCING ACTIVITIES:</b>                                |                                  |                |
| Proceeds from exercised stock options                                       | 762                              | 11             |
| Net proceeds from initial public offering                                   | 94,148                           |                |
| Payments of deferred stock issuance costs                                   |                                  | (98)           |
| Proceeds from bridge loan financing   |                                  | 3,000          |
| Repayments of long-term debt principal                                      | (5,026)                          | (2,018)        |
| Net cash provided by financing activities                                   | 89,884                           | 895            |
| <b>INCREASE (DECREASE) IN CASH AND CASH EQUIVALENTS</b>                     | <b>31,137</b>                    | <b>(1,829)</b> |
| <b>CASH AND CASH EQUIVALENTS Beginning of period</b>                        | <b>46,595</b>                    | <b>3,670</b>   |

|                           |               |    |        |    |       |
|---------------------------|---------------|----|--------|----|-------|
| CASH AND CASH EQUIVALENTS | End of period | \$ | 77,732 | \$ | 1,841 |
|---------------------------|---------------|----|--------|----|-------|

**NONCASH FINANCING ACTIVITIES:**

|                                    |  |    |       |    |  |
|------------------------------------|--|----|-------|----|--|
| Warrant conversion to common stock |  | \$ | 3,088 | \$ |  |
|------------------------------------|--|----|-------|----|--|

The accompanying notes are an integral part of these condensed consolidated financial statements.

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**DICERNA PHARMACEUTICALS, INC.**

**Notes to Condensed Consolidated Financial Statements**

**(Unaudited)**

(In thousands, except share and per share data)

**1. Description of Business and Basis of Presentation**

***Nature of business***

Dicerna Pharmaceuticals, Inc. and its subsidiary (the Company) is a biopharmaceutical company focused on the discovery and development of innovative treatments for rare inherited diseases involving the liver and for cancers that are genetically defined. The Company is using its proprietary RNA interference (RNAi) technology platform to build a broad pipeline in these therapeutic areas. The Company intends to discover, develop and commercialize novel therapeutics either on its own or in collaboration with pharmaceutical partners.

The Company continues to be subject to a number of risks common to companies in similar stages of development. Principal among these risks are the uncertainties of technological innovations, which are particularly high in the field of drug discovery and development, dependence on key individuals, development of the same or similar technological innovations by the Company's competitors and protection of proprietary technology. The Company's ability to fund its planned preclinical and clinical operations, including completion of its clinical trials, is expected to depend on the amount and timing of cash receipts under its existing collaboration agreement, as well as any future collaboration or product sales and/or financing transactions.

In February 2014, the Company completed the sale of 6,900,000 shares of common stock in an initial public offering of its common stock (the IPO) at a price to the public of \$15.00 per share, resulting in proceeds to the Company of \$92,749 after deducting underwriting discounts and commissions of approximately \$7,245 and offering expenses paid by the Company of approximately \$3,506. In connection with the close of the IPO, all of the outstanding shares of Series A mandatorily redeemable, convertible preferred stock (Series A preferred stock), Series B mandatorily redeemable, convertible preferred stock (Series B preferred stock) and Series C mandatorily redeemable, convertible preferred stock (Series C preferred stock) were converted into shares of common stock on a one-for-one basis immediately prior to the closing of the IPO. Following these transactions, the Company had a total of 17,763,051 shares of common stock issued and outstanding as of June 30, 2014. The significant increase in the shares outstanding in 2014 is expected to impact the year over year comparability of the Company's net loss per share calculations for the full year.

***Basis of presentation and consolidation***

The accompanying unaudited condensed consolidated financial statements have been prepared in conformity with accounting principles generally accepted in the U.S. (GAAP) and in accordance with the rules and regulations of the Securities and Exchange Commission (SEC) for interim financial information. Accordingly, they do not include all of the information and notes required by GAAP for complete financial statements. The unaudited condensed consolidated interim financial statements have been prepared on the same basis as the annual financial statements and, in the opinion of management, reflect all adjustments, which include only normal recurring adjustments, necessary to present fairly the Company's financial position at June 30, 2014 and results of operations and cash flows for the interim periods ended June 30, 2014 and 2013. These unaudited condensed consolidated interim financial statements

should be read in conjunction with the audited consolidated financial statements and notes thereto included in the Company's Annual Report on Form 10-K for the year ended December 31, 2013. The results of the three and six month periods ended June 30, 2014 are not necessarily indicative of the results to be expected for the year ending December 31, 2014 or for any other interim period or for any other future year.

**Summary of Significant Accounting Policies** There have been no material changes to the significant accounting policies previously disclosed in the Company's Annual Report on Form 10-K, other than the held-to-maturity investments policy noted below.

#### **Held-to-Maturity Investments**

The Company accounts for its investments in marketable securities in accordance with FASB ASC 320, *Investments Debt and Equity Securities*. The Company determines the appropriate classification of investments at the time of purchase and re-evaluates such designation as of each balance sheet date. Debt securities carried at amortized cost are classified as held-to-maturity when the Company has the positive intent and ability to hold the securities to maturity. At June 30, 2014, all of the Company's investments are classified as held-to-maturity.

**Table of Contents****2. Net Loss Per Share Attributable to Common Stockholders**

The following table summarizes the computation of basic and diluted net loss per share attributable to common stockholders of the Company (in thousands, except share and per share data):

|   | Three Months Ended June 30 |            | Six Months Ended June 30, |            |
|---|----------------------------|------------|---------------------------|------------|
|   | 2014                       | 2013       | 2014                      | 2013       |
| Net loss  | \$ (11,355)                | \$ (3,786) | \$ (22,159)               | \$ (7,605) |
| Accretion of preferred stock issuance costs to redemption value |                            | (15)       | (204)                     | (31)       |
| Accrued dividends on preferred stock                            |                            | (1,007)    |                           | (2,002)    |
| Net loss attributable to common stockholders basic and diluted  | \$ (11,355)                | \$ (4,808) | \$ (22,363)               | \$ (9,638) |

The following potentially dilutive securities outstanding during the period, prior to the use of the treasury stock method or if-converted method, have been excluded from the computation of diluted weighted-average common stock outstanding, because such securities had an anti-dilutive impact since the Company has a net loss attributable to common stockholders:

|   | Three months Ended June 30, |           | Six months Ended June 30, |           |
|---|-----------------------------|-----------|---------------------------|-----------|
|   | 2014                        | 2013      | 2014                      | 2013      |
| Options to purchase common stock                            | 2,831,125                   | 25,262    | 2,159,551                 | 25,722    |
| Warrants to purchase common stock                           | 87,901                      | 2,198     | 73,424                    | 2,198     |
| Warrants to purchase redeemable convertible preferred stock |                             | 49,294    | 25,738                    | 49,294    |
| Redeemable convertible preferred stock                      |                             | 2,018,025 | 2,047,682                 | 2,018,025 |
| Unvested restricted stock                                   | 109,859                     | 10        | 87,141                    | 15        |

*Unvested Restricted Stock*

At June 30, 2014, the Company had 113,000 shares of unvested restricted stock outstanding. The unvested restricted stock resulted from the early exercise of stock options of 73,000 and the issuance of restricted stock units of 40,000 during the six-month period ended June 30, 2014.

**3. Held-to-maturity investments**

The Company invests its excess cash balances in short-term and long-term fixed-income investments. The Company determines the appropriate classification of investments at the time of purchase and re-evaluates such designation as of each balance sheet date. Debt securities carried at amortized cost are classified as held-to-maturity when the Company has the positive intent and ability to hold the securities to maturity.

The following tables provide information relating to held-to-maturity investments at June 30, 2014 (in thousands):

|  | <b>June 30, 2014</b> |                   |                   |                  |
|--|----------------------|-------------------|-------------------|------------------|
|  | <b>Amortized</b>     | <b>Gross</b>      | <b>Gross</b>      | <b>Fair</b>      |
|  | <b>Cost</b>          | <b>Unrealized</b> | <b>Unrealized</b> | <b>Value</b>     |
|  |                      | <b>Gains</b>      | <b>Losses</b>     |                  |
| <b>Held-to-maturity investments</b>            |                      |                   |                   |                  |
| U.S. Government treasury and agency securities | \$ 42,552            | \$ 7              | \$ (4)            | \$ 42,555        |
| <b>Total held-to-maturity investments</b>      | <b>\$ 42,552</b>     | <b>\$ 7</b>       | <b>\$ (4)</b>     | <b>\$ 42,555</b> |



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The amortized cost and fair value of held-to-maturity investments by contractual maturities at June 30, 2014, are as follows (in thousands):

|   | <b>Held-to-Maturity</b> |                  |
|---|-------------------------|------------------|
|   | <b>Amortized</b>        | <b>Fair</b>      |
|   | <b>Cost</b>             | <b>Value</b>     |
| Maturing in one year or less              | \$ 22,526               | \$ 22,525        |
| Maturing after one year through two years | 20,026                  | 20,030           |
| <b>Total</b>                              | <b>\$ 42,552</b>        | <b>\$ 42,555</b> |

**4. Long-term Debt**

On March 26, 2009, the Company entered into a loan and security agreement with an independent finance company, Hercules Technology II, LP (Hercules), for up to \$7,000 (Hercules loan). The Hercules loan, which has since been repaid, was collateralized by a security interest in all tangible assets. On May 28, 2010, the Company and Hercules executed an amendment to the loan and security agreement to defer \$653 in principal payments originally due in June 2010 through August 2010, and amortized such deferred principal amounts equally over the remaining payment term beginning in September 2010. On June 28, 2011, the Company and Hercules executed a second amendment to the loan and security agreement, which increased the maximum loan amount to \$12,000. Upon execution of the second amendment, the Company drew a \$7,000 advance, a portion of which the Company used to repay the outstanding balance of principal and interest under the original loan and security agreement. On December 15, 2011, the Company drew down the remaining \$5,000. Interest was payable monthly and principal was to be repaid in equal monthly installments beginning April 1, 2012 through January 2, 2015. The applicable annual interest rate was 10.15% at December 31, 2013. On April 7, 2014, the Company repaid the remaining amount of the Hercules loan in full in a total amount of \$3,590.

In connection with the Hercules loan, the Company issued warrants to Hercules for the purchase of an aggregate of 21,000 shares of Series A preferred stock and 26,400 shares of Series B preferred stock each at an exercise price of \$25.00 per share. Immediately prior to the closing of the IPO on February 4, 2014, all of the outstanding shares of Series A, Series B and Series C preferred stock were automatically converted into shares of common stock on a one-for-one basis. The fair value of the warrants was classified as a liability in the accompanying balance sheet as of December 31, 2013. After the conversion of Series A and Series B preferred stock, the fair value of the warrants was reclassified as a part of stockholders' equity. The remeasurement of the liability continued until the date of the closing of the IPO. The fair value of the outstanding Hercules warrants as of the IPO closing date was \$765 and was determined using the Black-Scholes option-pricing model with the following assumptions:

|                                 | <b>February 4,<br/>2014</b> |
|---------------------------------|-----------------------------|
| Stock price                     | \$ 32.66                    |
| Expected option term (in years) | 3.00                        |
| Expected volatility             | 62%                         |
| Risk-free interest rate         | 0.69%                       |
| Expected dividend yield         | 0.00%                       |

The estimated fair value of the outstanding Hercules warrants at December 31, 2013 was \$94. On February 11, 2014, Hercules net exercised the warrants in exchange for a total of 12,702 shares of common stock. There were no Series A and Series B warrants outstanding at June 30, 2014.

The adjustment to this preferred stock warrant liability related to the Hercules warrants was recorded in other income (expense) and amounted to \$0 and \$(671) the three and six month periods ended June 30, 2014, respectively, compared to \$0 and \$5 for the comparable periods in 2013.

## **5. Stock-Based Compensation**

### ***Stock options granted to employees and nonemployees***

Stock-based compensation expense related to stock options granted to nonemployees is recognized as the consulting services are rendered, generally on a straight-line basis. The Company believes that the fair value of the stock options is more reliably measurable than the fair value of the services received. Compensation expense is subject to re-measurement until the options vest.

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In September 2013, the Company granted stock options to purchase 132,500 shares of common stock to nonemployees with an initial fair value of \$337. During the six month period ended June 30, 2014, the services of certain nonemployees were terminated, resulting in accelerated vesting of their remaining unvested shares. Based on the terms of the nonemployee stock option agreements the Company recorded nonemployee stock-based compensation expense of \$50 and \$1,661 for the three and six month periods ended June 30, 2014. The assumptions used to estimate fair value were as follows:

|                                 | <b>Three Months Ended<br/>June 30, 2014</b> |         | <b>Six Months Ended<br/>June 30, 2014</b> |         |
|---------------------------------|---|---------|---|---------|
| Stock price                     | \$ 16.16                                    | \$22.57 | \$ 16.16                                  | \$41.12 |
| Expected option term (in years) | 6.11  | 6.40    | 0.25                                      | 6.86    |
| Expected volatility             |   | 73%     | 56%                                       | 73%     |
| Risk-free interest rate         | 2.12%                                       | 2.27%   | 0.08%                                     | 2.29%   |
| Expected dividend yield         |   | 0.00%   |   | 0.00%   |

As of June 30, 2014, there were 61,875 unvested stock options held by nonemployees.

During the three-month period ended June 30, 2014, the Company granted stock options to purchase 1,810,806 shares of common stock to employees with an initial fair value of \$19,622. The assumptions used to estimate fair value were as follows:

|                                 | <b>Three Months Ended<br/>June 30, 2014</b> |         |
|---------------------------------|---|---------|
| Stock price                     | \$ 16.30                                    | \$18.73 |
| Expected option term (in years) | 5.50  | 6.25    |
| Expected volatility             | 73%   | 74%     |
| Risk-free interest rate         | 1.92%                                       | 2.00%   |
| Expected dividend yield         |   | 0.00%   |

There were no employee options granted during the three-months ended March 31, 2014.

**6. Redeemable Convertible Preferred Stock**

The consummation of the IPO on February 4, 2014 resulted in the conversion of all of the shares of the Company's Series A, Series B and Series C preferred stock into shares of common stock. Each share of Series A, Series B and Series C preferred stock was automatically converted into common stock on a one-for-one basis. The conversion of Series A, Series B and Series C preferred stock resulted in the issuance of 10,589,434 shares of common stock.

After the conversion of Series C preferred stock, the fair value of the warrants related to Series C preferred stock outstanding immediately prior to the closing of the IPO was reclassified as a part of stockholders' equity. The remeasurement of the Series C preferred stock warrant liability continued until the closing date of the IPO. The fair value of the Series C preferred stock warrants as of the IPO closing date was \$2,323 and was determined using the Black-Scholes option-pricing model with the following assumptions:

|                                 | <b>February 4,<br/>2014</b> |
|---------------------------------|-----------------------------|
| Stock price                     | \$ 32.66                    |
| Expected option term (in years) | 4.39                        |
| Expected volatility             | 65%                         |
| Risk-free interest rate         | 1.52%                       |
| Expected dividend yield         | 0.00%                       |

The adjustment to the Series C preferred stock warrant liability was recorded in other income (expense) and amounted to \$0 and \$(1,888) for the three and six months ended June 30, 2014, respectively, and \$118 for the three and six month periods ended June 30, 2013.

## 7. Fair Value Measurements

Fair value is an exit price, representing the amount that would be received from the sale of an asset or paid to transfer a liability in an orderly transaction between market participants. As such, fair value is a market-based measurement that should be determined based on assumptions that market participants would use in pricing an asset or liability. As a basis for considering such assumption the accounting literature establishes a three-tier value hierarchy which prioritizes the inputs used in measuring fair value as follows: (Level 1) observable inputs, such as

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quoted prices in active markets; (Level 2) inputs other than the quoted prices in active markets that are observable either directly or indirectly; and (Level 3) unobservable inputs for which there is little or no market data, which requires the Company to develop its own assumptions. This hierarchy requires the Company to use observable market data, when available, and to minimize the use of unobservable inputs when determining fair value.

The Company's cash equivalents, primarily money market accounts and assets held in restriction are classified within Level 1 of the fair value hierarchy because they are valued using quoted prices as of June 30, 2014 and December 31, 2013, respectively.

The Company's assets held in restriction bore interest at the prevailing market rates for instruments with similar characteristics and, accordingly, the carrying value of these instruments also approximated their fair value and the financial instruments were classified within Level 2 of the fair value hierarchy because the inputs to the fair value measurement are valued using observable inputs as of June 30, 2014 and December 31, 2013, respectively.

The Company's held-to-maturity investments bore interest at the prevailing market rates for instruments with similar characteristics and, accordingly, the carrying value of these instruments also approximated their fair value and the financial instruments were classified within Level 2 of the fair value hierarchy because the inputs to the fair value measurement are valued using observable inputs as of June 30, 2014. The fair value of these instruments is \$42,555, and the amortized cost basis is \$42,552.

The carrying value of other current assets, accounts payable, and accrued expenses approximate fair value, due to the short maturity of these instruments. The Company's long-term debt bore an interest at the prevailing market rates for instruments with similar characteristics and, accordingly, the carrying value for these instruments also approximated their fair value and the financial measurements were also classified within Level 2 of the fair value hierarchy. There was no debt outstanding as of June 30, 2014.

The fair value of the preferred stock warrant liability was determined using the Black-Scholes option-pricing model for the year ended December 31, 2013 and until the conversion date of February 4, 2014. After the closing of the IPO, the remaining preferred stock warrant liability was no longer subject to re-measurement as the warrants to purchase the Company's preferred stock became warrants to purchase shares of our common stock. As of the IPO closing date, the fair value of the preferred stock warrants was based significantly on the fair value of our publicly traded common stock and other observable inputs and was reclassified to Level 2. There were no preferred stock warrants outstanding as of June 30, 2014.

The following table provides a roll-forward of the Company's liabilities measured at fair value on a recurring basis using unobservable inputs (Level 3):

|   |         |
|---|---------|
| BALANCE January 1, 2014                   | \$ 529  |
| Change in fair value of warrant liability | 2,559   |
| Transfers to Level 2                      | (3,088) |
| BALANCE June 30, 2014                     | \$      |
| BALANCE January 1, 2013                   | \$ 331  |
| Issuance of preferred stock warrants      | 324     |
| Change in fair value of warrant liability | (123)   |

|         |   |    |       |
|---------|---|----|-------|
| BALANCE | June 30, 2013                             | \$ | 532   |
| BALANCE | April 1, 2013                             | \$ | 326   |
|         | Issuance of preferred stock warrants      |    | 324   |
|         | Change in fair value of warrant liability |    | (118) |

|         |               |    |     |
|---------|---------------|----|-----|
| BALANCE | June 30, 2013 | \$ | 532 |
|---------|---------------|----|-----|

## 8. Commitments and Contingencies

### *City of Hope license agreement*

In September 2007, the Company entered into a license agreement with City of Hope, an independent academic research and medical center (the Medical Center). In consideration for the right to develop, manufacture, and commercialize products based on certain of the Medical Center's intellectual property, the Company paid a one-time, non-refundable license fee and issued shares of common stock as consideration for the license.

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The Company is required to reimburse the Medical Center for patent costs incurred, and pay an amount within the range of \$5,000 to \$10,000 upon the achievement of certain milestones, and royalties on future sales, if any. The license agreement will remain in effect until the expiration of the last patents or copyrights licensed under the agreement or until all obligations under the agreement with respect to payment of milestones have terminated or expired. The Company may terminate the license agreement at any time upon 90 days written notice to the Medical Center. The Company did not record any research and development expense, related to the agreement with the Medical Center during the three and six month periods ended June 30, 2014 and 2013, respectively.

### ***Plant Bioscience Limited (PBL) license agreement***

In September 2013, the Company entered into a commercial license agreement with Plant Bioscience Limited (PBL), pursuant to which PBL has granted to the Company a license to certain of its U.S. patents and patent applications to research, discover, develop, manufacture, sell, import and export, products incorporating one or more short RNA molecules (SRMs).

The Company has paid PBL a one-time, non-refundable signature fee and will pay PBL a nomination fee for any additional SRMs nominated by the Company under the agreement. The Company is further obligated to pay PBL milestone payments upon achievement of certain clinical and regulatory milestones. The Company made a payment of \$100 to PBL upon the commencement of its clinical trial in MYC. In addition, PBL is entitled to receive royalties of any net sale revenue of any licensed product candidates sold by the Company.

### **9. Subsequent Event**

On July 11, 2014, the Company executed a non-cancelable operating lease for office and laboratory space in Cambridge, Massachusetts. The lease agreement obligates the Company to future minimum payments totaling \$9.6 million over a six-year lease term. The lease commences on the later to occur of (i) December 1, 2014 or (ii) the date on which the landlord's work is substantially complete. As part of the lease agreement, the Company established a \$1.1 million letter of credit, secured by a certificate of deposit which is included in assets held in restriction at June 30, 2014.

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

*The following discussion contains forward-looking statements that involve risks and uncertainties. Our actual results could differ materially from those discussed here. Factors that could cause or contribute to such differences include, but are not limited to, those discussed in this section as well as factors described in Part II, Item 1A Risk Factors.*

**Overview**

We are a biopharmaceutical company focused on the discovery and development of innovative treatments for rare inherited diseases involving the liver and for cancers that are genetically defined. We are using our RNA interference (RNAi) technology platform to build a broad pipeline in these therapeutic areas. In both rare diseases and oncology, we are pursuing targets that have historically been difficult to inhibit using conventional approaches, but where we believe connections between targets and diseases are well understood and documented. We aim to discover, develop and commercialize these novel therapeutics either on our own or in collaboration with pharmaceutical partners. In indications such as rare diseases in which a small sales force will suffice, we seek to retain substantially all commercial rights in key markets. In oncology and other more prevalent disease areas, we may partner our products while seeking to retain significant portions of the commercial rights in North America.

In the rare disease field, we are developing a proprietary treatment, DCR-PH1, for the rare and serious inherited disorder Primary Hyperoxaluria 1 (PH1). We seek to begin clinical trials of DCR-PH1 in 2015. We also have discovery and early development programs against a series of additional disease targets in the liver. In oncology, we are currently directing our development efforts towards our proprietary product candidate DCR-MYC for the treatment of MYC-related cancers, including hepatocellular carcinoma (HCC), which we believe to be the third leading cause of cancer deaths worldwide. We submitted an Investigational New Drug application for DCR-MYC for the treatment of solid tumors, lymphoma and multiple myeloma to the U.S. Food and Drug Administration in the first quarter of 2014 and began our clinical trials of DCR-MYC in the second quarter of 2014.

As part of our collaboration with Kyowa Hakko Kirin Co., Ltd. (KHK), a global pharmaceutical company, we are developing a product candidate that targets the oncogene KRAS, which is frequently mutated in numerous major cancers, including non-small cell lung cancer, colorectal cancer and pancreatic cancer. KHK is responsible for global development of the KRAS program, including all development expenses. For the KRAS product candidate, we retain an option to co-promote in the U.S. for an equal share of the profits from U.S. net sales. We are also developing a product candidate targeting another oncogene in collaboration with KHK. For each product candidate in our collaboration with KHK, we have the potential to receive clinical, regulatory and commercialization milestone payments of up to \$110.0 million and royalties on net sales of each such product candidate. We expect that our strategy to partner the development of product candidates will help us fund the costs of clinical development and enable us to diversify risk across a number of programs.

Since our inception in October 2006, we have devoted substantial resources to the research and development of DsiRNA molecules and drug delivery technologies and the protection and enhancement of our intellectual property estate. We have no products approved for sale and all of our revenue to date has been collaboration revenue or government grant revenue. To date, we have funded our operations primarily through the initial public offering of our common stock, previous private placements of preferred stock and convertible debt securities, from research funding, license fees, option exercise fees and preclinical payments under our research collaboration and license agreement with KHK and from a government grant. In addition, we have borrowed under a secured term loan from Hercules (Hercules loan) to fund our operations. More particularly, since our inception and through June 30, 2014, we have raised an aggregate of \$233.2 million to fund our operations, of which \$92.7 million of net proceeds was from the initial public offering of our common stock as described below, \$110.5 million was from the sale of preferred stock and convertible debt securities (including \$3.0 million from a bridge loan financing that closed in June 2013 (2013



bridge note financing), \$17.5 million was through our collaboration and license agreement with KHK, \$0.5 million was from a federal government grant for our Qualifying Therapeutic Discovery Project in November 2010 and \$12.0 million was from borrowings under the Hercules loan. As of June 30, 2014, our cash and cash equivalents and held-to-maturity investments were \$120.3 million and we also had \$1.4 million in assets held in restriction. On April 7, 2014, we repaid the remaining amount of the Hercules loan of approximately \$3.6 million.

On February 4, 2014, we completed the initial public offering of our common stock, in which we issued and sold a total of 6,900,000 shares of common stock, including 900,000 shares sold pursuant to the exercise in full by the underwriters of an option to purchase additional shares, at a public offering price of \$15.00 per share. We received net proceeds of \$92.7 million after deducting the underwriting commissions and discounts and offering expenses paid by us. All of the shares of our preferred stock were converted into shares of common stock and our warrants to purchase preferred stock became exercisable to purchase common stock immediately prior to the completion of our initial public offering.

Since inception, we have incurred significant operating losses. Our net loss was \$22.2 million and \$7.6 million for the six months ended June 30, 2014 and 2013, respectively. Substantially all of our operating losses resulted from expenses incurred in connection with our research programs and from general and administrative costs associated with our operations. No revenue was recognized for the three and six months

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ended June 30, 2014 or 2013. Our revenue to date has been generated through our research collaboration and license agreement with KHK and a government grant. We have not generated any commercial product revenue. As of June 30, 2014, we had an accumulated deficit of \$107.8 million. We expect to continue to incur significant and increasing losses in the foreseeable future. Our net losses may fluctuate significantly from quarter to quarter and year to year. We anticipate that our expenses will increase substantially as we:

advance our product candidates into preclinical development;

conduct the clinical trial of DCR-MYC and any future clinical trials of DCR-PH1 and other potential product candidates;

continue our research and development efforts, including to expand our pipeline and to enhance our technology platform;

increase research and development related activities for the discovery and development of additional product candidates;

manufacture clinical study materials and develop large-scale manufacturing capabilities;

seek regulatory approval for our product candidates that successfully complete clinical trials;

maintain, expand and protect our intellectual property portfolio;

add operational, financial and management information systems and personnel, including personnel to support our product development and planned future commercialization efforts; and

operate as a public company.

We do not expect to generate substantial revenue from product sales unless and until we successfully complete development and obtain regulatory approval for one or more of our product candidates, which is subject to significant uncertainty and which we expect will take at least seven years. If we obtain regulatory approval for any of our product candidates, we expect to incur significant commercialization expenses related to product sales, marketing, manufacturing and distribution. Until such time, if ever, that we generate product revenue, we expect to finance our cash needs through a combination of public or private equity offerings, debt financings and research collaboration and license agreements. We may be unable to raise capital or enter into such other arrangements when needed or on favorable terms. Our failure to raise capital or enter into such other arrangements as and when needed would have a negative impact on our financial condition and our ability to develop our product candidates.

***Collaboration agreement***

In December 2009, we entered into a research collaboration and license agreement with KHK for the research, development and commercialization of DsiRNA molecules and drug delivery technologies for therapeutic targets in oncology. We have granted KHK an exclusive, worldwide, royalty-bearing and sub-licensable license to our DsiRNA molecules and drug delivery technologies and intellectual property for certain selected DsiRNA-based compounds. Under the research collaboration and license agreement, KHK is responsible for activities to develop, manufacture and commercialize the selected DsiRNA-based compounds and pharmaceutical products containing such compounds. For the KRAS product candidate, we have an option to co-promote in the U.S. for an equal share of the profits from U.S. net sales. In addition, for each product candidate under the research collaboration and license agreement, we have the potential to receive clinical, regulatory and commercialization milestone payments of up to \$110.0 million and royalties on net sales of such product candidate.

Since the initiation of the research collaboration and license agreement, of the various targets in the collaboration, two target programs, including the initial target KRAS, have been nominated by KHK for formal development studies. Both programs utilize our specific RNAi-inducing double-stranded DsiRNA molecules and a lipid nanoparticle drug delivery technology proprietary to KHK. We received no additional payments from KHK under the research collaboration and license agreement during the three and six month periods ended June 30, 2014.

### ***License agreement***

In September 2007, we entered into a license agreement with City of Hope (COH), an independent academic research and medical center, pursuant to which COH has granted to us an exclusive (subject to certain exceptions described below), royalty-bearing, worldwide license under certain patent rights in relation to DsiRNA, including the core DsiRNA patent (U.S. 8,084,599), to manufacture, use, offer for sale, sell and import products covered by the licensed patent rights for the prevention and treatment of any disease in humans. COH is restricted from granting any additional rights to develop, manufacture, use, offer to sell, sell or import products covered by the licensed patent rights for the prevention and treatment of any disease in humans. In addition, COH has granted to us an exclusive, royalty-bearing, worldwide license under the licensed patent rights providing certain rights for up to 20 licensed products selected by us for human diagnostic uses, provided that COH has not granted or is not negotiating a license of rights to diagnostic uses for such licensed products to a third party. The core DsiRNA patent (U.S. 8,084,599), titled methods and compositions for the specific inhibition of gene expression by double-stranded RNA, describes RNA structures having a 25 to 30 nucleotides sense strand, a blunt end at the 3' end of the sense strand and a one to four nucleotides overhang at the 3' end of the antisense strand. The expiration date of this patent is July 17, 2027.

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Pursuant to the terms of the agreement, we paid COH a one-time, non-refundable license fee and issued shares of our common stock to COH and a co-inventor of the core DsiRNA patent. COH is entitled to receive milestone payments in an aggregate amount within the range of \$5.0 million to \$10.0 million upon achievement of certain clinical and regulatory milestones. COH is further entitled to receive royalties at a low single-digit percentage of any net sale revenue of the licensed products sold by us and our sublicensees. If we sublicense the licensed patent rights to a third party, COH has the right to receive a double digit percentage of sublicense income, the percentage of which decreases after we have expended \$12.5 million in development and commercialization costs. We are also obligated to pay COH an annual license maintenance fee, which may be credited against any royalties due to COH in the same year, and reimburse COH for expenses associated with the prosecution and maintenance of the license patent rights. The license agreement will remain in effect until the expiration of the last to expire of the patents or copyrights licensed under the agreement. We have not included our obligations to make future milestone payments on our balance sheet because the achievement and timing of the related milestones is not probable and estimable.

In September 2013, we entered into a commercial license agreement with Plant Bioscience Limited (PBL), pursuant to which PBL has granted a license to us for certain of its U.S. patents and patent applications to research, discover, develop, manufacture, sell, import and export, products incorporating one or more short RNA molecules (SRMs).

We have paid PBL a one-time, non-refundable signature fee and will pay PBL a nomination fee for any additional SRMs nominated by us under the agreement. We are further obligated to pay PBL milestone payments upon achievement of certain clinical and regulatory milestones. In addition, PBL is entitled to receive royalties on any net sale revenue of any licensed product candidates sold by us. We paid PBL \$100,000 upon commencement of our MYC clinical trial.

## **Financial Operations Overview**

### ***Revenue***

Our revenue to date has been generated primarily through research funding, license fees, option exercise fees and preclinical development payments under our research collaboration and license agreement with KHK and a government grant. We have not generated any commercial product revenue. For each product candidate under our research collaboration and license agreement with KHK, we are also entitled to receive clinical, regulatory and commercialization milestone payments of up to \$110.0 million and royalties on net sales of such product candidate. We did not receive any royalty payments during the three and six month periods ended June 30, 2014 or June 30, 2013.

We did not have revenue for the three or six month periods ended June 30, 2014 or 2013.

In the future, we may generate revenue from a combination of research and development payments, license fees and other upfront payments, milestone payments, product sales and royalties in connection with our collaboration with KHK or future collaborations and licenses. We expect that any revenue we generate will fluctuate in future periods as a result of the timing of our or a collaborator's achievement of preclinical, clinical, regulatory and commercialization milestones, if at all, the timing and amount of any payments to us relating to such milestones and the extent to which any of our product candidates are approved and successfully commercialized by us or a collaborator. If we, KHK or any future collaborator fails to develop product candidates in a timely manner or obtain regulatory approval for them, our ability to generate future revenue, and our results of operations and financial position, would be materially adversely affected.

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

Research and development expenses consist of costs associated with our research activities, including discovery and development of our DsiRNA molecules and drug delivery technologies and our research activities under our research collaboration and license agreement with KHK. Our research and development expenses include:

direct research and development expenses incurred under arrangements with third parties, such as contract research organizations, contract manufacturing organizations, and consultants;

platform-related lab expenses, including lab supplies, license fees, consultants, and our scientific advisory board;

employee-related expenses, including salaries, benefits and stock-based compensation expense; and

facilities, depreciation and other allocated expenses, which include direct and allocated expenses for rent and maintenance of facilities, depreciation of leasehold improvements and equipment and laboratory and other supplies.

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We expense research and development costs as they are incurred. We account for nonrefundable advance payments for goods and services that will be used in future research and development activities as expense when the service has been performed or when the goods have been received. A significant portion of our research and development costs are not tracked by project as they benefit multiple projects or our technology platform.

In the second quarter of 2014, we initiated a multi-center, dose-escalating Phase 1 clinical study of DCR-MYC to assess the safety and tolerability of DCR-MYC in patients with solid tumors, multiple myeloma, or lymphoma who are refractory or unresponsive to standard therapies. The process of conducting preclinical studies and clinical trials necessary to obtain regulatory approval is costly and time-consuming. We, KHK or any future collaborator may never succeed in obtaining marketing approval for any of our product candidates. The probability of success for each product candidate may be affected by numerous factors, including preclinical data, clinical data, competition, manufacturing capability and commercial viability.

All of our research and development programs are at an early stage and successful development of future product candidates from these programs is highly uncertain and may not result in approved products. Completion dates and completion costs can vary significantly for each future product candidate and are difficult to predict. We anticipate we will make determinations as to which product candidates to pursue and how much funding to direct to each product candidate on an ongoing basis in response to our ability to maintain or enter into collaborations with respect to each product candidate, the scientific and clinical success of each product candidate as well as ongoing assessments as to the commercial potential of product candidates. We will need to raise additional capital and may seek additional collaborations in the future in order to advance our various product candidates. Additional private or public financings may not be available to us on acceptable terms, or at all. Our failure to raise capital as and when needed would have a material adverse effect on our financial condition and our ability to pursue our business strategy.

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

General and administrative expenses consist primarily of salaries and related benefits, including stock-based compensation, related to our executive, finance, legal, business development and support functions. Other general and administrative expenses include travel expenses, professional fees for auditing, tax and legal services and allocated facility-related costs not otherwise included in research and development expenses.

### ***Interest income***

Interest income consists of interest income earned on our cash and cash equivalents, held-to-maturity investments and assets held in restriction.

### ***Interest expense***

Interest expense consists of interest expense on the Hercules loan, which was repaid in full in April 2014.

### ***Preferred stock warrant re-measurement***

Preferred stock warrant re-measurement is associated with warrants to purchase preferred stock issued to lenders under our convertible notes and the Hercules loan. The re-measurement consists of the change in value calculated using the Black-Scholes option pricing model to estimate the fair value of the warrants. We base the estimates in the Black-Scholes option pricing model, in part, on subjective assumptions, including stock price volatility, risk-free interest rate, dividend yield and the fair value of the preferred stock underlying the warrants. The re-measurement gain or loss associated with the change in the fair value of the preferred stock warrant liability each reporting period is

recognized as a component of other income (expense). Upon the completion of the initial public offering of our common stock on February 4, 2014, the preferred stock warrant liability was reclassified as a component of equity and is no longer subject to re-measurement. The fair value of the preferred stock warrants as of the closing date of the initial public offering was \$3.1 million.

### **Critical Accounting Policies and Significant Judgments and Estimates**

Our management's discussion and analysis of financial condition and results of operations is based on our unaudited condensed consolidated financial statements, which have been prepared in accordance with accounting principles generally accepted in the U.S. (GAAP). The preparation of these condensed consolidated financial statements requires us to make estimates and assumptions that affect the reported amounts of assets and liabilities and the disclosure of contingent assets and liabilities at the date of the condensed consolidated financial statements, as well as the revenue and expenses incurred during the reported periods. On an ongoing basis, we evaluate our estimates and judgments, including those related to accrued expenses, revenue recognition, deferred revenue and stock-based compensation. We base our estimates on historical experience and on various other factors that we believe are reasonable under the circumstances, the results of which form the basis for making judgments about the carrying value of assets and liabilities that are not apparent from other sources. Changes in estimates are reflected in reported results for the period in which they become known. Actual results may differ from these estimates under different assumptions or conditions.

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The critical accounting policies that we believe impact significant judgments and estimates used in the preparation of our financial statements presented in this report are described in our Management's Discussion and Analysis of Financial Condition and Results of Operations in our Annual Report on Form 10-K filed with the SEC on March 27, 2014. There have been no material changes to our critical accounting policies during the three and six month periods ended June 30, 2014 from those discussed in Management's Discussion and Analysis of Financial Condition and Results of Operations Critical Accounting Policies and Significant Judgments and Estimates in our Annual Report on Form 10-K filed with the SEC on March 27, 2014, other than the held-to-maturity investments policy noted below.

**Held-to-Maturity Investments**

We account for our investment in marketable securities in accordance with FASB ASC 320, *Investments - Debt and Equity Securities*. We determine the appropriate classification of investments at the time of purchase and re-evaluate such designation as of each balance sheet date. Debt securities carried at amortized cost are classified as held-to-maturity when we have the positive intent and ability to hold the securities to maturity. At June 30, 2014, all of our investments are classified as held-to-maturity.

**Recent Accounting Pronouncements**

There are no recent accounting standards that would impact our results reported in the unaudited condensed consolidated interim financial statements included in this report.

**Comparison of Three and Six Month Periods Ended June 30, 2014 and 2013**

The following table summarizes the results of our operations for the periods indicated (in thousands):

|                            | THREE MONTHS ENDED JUNE 30, |            |                        | SIX MONTHS ENDED JUNE 30, |            |                        |
|----------------------------|-----------------------------|------------|------------------------|---------------------------|------------|------------------------|
|                            | 2014                        | 2013       | Increase<br>(Decrease) | 2014                      | 2013       | Increase<br>(Decrease) |
| Revenue:                   | \$                          | \$         | \$                     | \$                        | \$         | \$                     |
| Expenses:                  |                             |            |                        |                           |            |                        |
| Research and development   | 6,806                       | 2,513      | 4,293                  | 12,057                    | 4,931      | 7,126                  |
| General and administrative | 4,372                       | 1,142      | 3,230                  | 7,213                     | 2,278      | 4,935                  |
| Total expenses             | 11,178                      | 3,655      | 7,523                  | 19,270                    | 7,209      | 12,061                 |
| Loss from operations       | (11,178)                    | (3,655)    | (7,523)                | (19,270)                  | (7,209)    | (12,061)               |
| Other expense              | 177                         | 131        | 46                     | 2,889                     | 396        | 2,493                  |
| Net loss                   | \$ (11,355)                 | \$ (3,786) | \$ (7,569)             | \$ (22,159)               | \$ (7,605) | \$ (14,554)            |

**Revenue**

We did not recognize any revenue for the three or six month periods ended June 30, 2014 or 2013. We do not expect to generate any product revenue for the foreseeable future.

**Research and development expenses**



The following table summarizes our research and development expenses incurred during the periods indicated (in thousands):

|   | THREE MONTHS ENDED JUNE 30, |                 |                        | SIX MONTHS ENDED JUNE 30, |                 |                        |
|---|-----------------------------|-----------------|------------------------|---------------------------|-----------------|------------------------|
|   | 2014                        | 2013            | Increase<br>(Decrease) | 2014                      | 2013            | Increase<br>(Decrease) |
| Direct research and development expenses    | \$ 2,078                    | \$ 1,224        | \$ 854                 | \$ 2,748                  | \$ 2,140        | \$ 608                 |
| Platform-related expenses                   | 2,350                       | 411             | 1,939                  | 5,561                     | 977             | 4,584                  |
| Employee-related expenses                   | 2,287                       | 617             | 1,670                  | 3,469                     | 1,286           | 2,183                  |
| Facilities, depreciation and other expenses | 91                          | 261             | (170)                  | 279                       | 528             | (249)                  |
| <b>Total</b>                                | <b>\$ 6,806</b>             | <b>\$ 2,513</b> | <b>\$ 4,293</b>        | <b>\$ 12,057</b>          | <b>\$ 4,931</b> | <b>\$ 7,126</b>        |

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Research and development expenses were \$6.8 million and \$12.1 million for the three and six months ended June 30, 2014, respectively, compared to \$2.5 million and \$4.9 million for the three and six month periods ended June 30, 2013, respectively. On a year-to-date basis, direct research and development expenses increased by \$0.6 million for the six month period ended June 30, 2014 compared to the same period in the prior year as a result of the initiation of the clinical trial related to DCR-MYC and an increase in research activities related to DCR-PH1. Platform-related expenses increased by \$4.6 million for the six month period ended June 30, 2014 compared to the same period in the prior year primarily due to stock-based compensation of 1.6 million and an increase in expenses related to discovery and early development of future programs. Employee-related expenses increased by \$2.2 million for the six months ended June 30, 2014 compared to the six months ended June 30, 2013, primarily due to stock-based compensation of \$1.0 million and additional hiring during the period. Facilities, depreciation and other expenses have decreased by \$0.2 million for the three and six month periods ended June 30, 2014 due to reduction in occupancy costs. We expect our research and development expenses to increase in 2014 as we continue spending on our development programs and clinical trial.

*General and administrative expenses*

General and administrative expenses were \$4.4 million and \$7.2 million for the three and six month periods ended June 30, 2014, respectively, compared to \$1.1 million and \$2.3 million for the three and six month periods ended June 30, 2013, respectively. Employee related expenses were \$2.0 million and \$3.0 million for the three and six month periods ended June 30, 2014, compared to \$0.4 million and \$0.9 million in the comparable period in 2013. The increase in these costs related to stock based compensation and an increase in head count. Professional fees were \$1.4 million and \$2.6 million for the three and six month periods ended June 30, 2014, compared to \$0.4 million and \$0.7 million in the comparable period in 2013. Other expense were \$1.0 million and \$1.6 million for the three and six month periods ended June 30, 2014, compared to \$0.3 million and \$0.7 million in the comparable period in 2013. These increases were primarily due to the transition and increased costs associated with operating as a public company. We expect general and administrative expenses to increase in the future as we continue to expand our operating activities and incur additional costs associated with being a publicly-traded company. These increases will likely include legal, accounting and other professional services costs, directors and officers liability insurance premiums and costs associated with investor relations.

*Other expense*

Other expense was \$0.2 and \$2.9 million for the three and six month periods ended June 30, 2014, respectively, compared to \$0.1 million and \$0.4 million for the three and six month periods ended June 30, 2013, respectively. On a year-to-date basis, the increase of \$2.5 million was primarily due to an increase in expense related to the re-measurement of the preferred stock warrant liability of \$2.7 million, which was partially offset by a decrease in interest expense of \$0.2 million. The decrease in interest expense was due to the Hercules loan being repaid in full, in April 2014.

**Liquidity and Capital Resources**

Since our inception and through June 30, 2014, we have raised an aggregate of \$233.2 million to fund our operations, of which \$92.7 million was from the initial public offering of our common stock, which closed on February 4, 2014, \$110.5 million was from the sale of preferred stock and convertible debt securities (including \$3.0 million from the 2013 bridge note financing), \$17.5 million was through our collaboration and license agreement with KHK, \$0.5 million was from a federal government grant for our Qualifying Therapeutic Discovery Project in November 2010 and \$12.0 million was from borrowings under the Hercules loan. As of June 30, 2014, our cash and cash equivalents and held-to-maturity investments were \$120.3 million and we also had \$1.4 million in assets held in restriction.

On February 4, 2014, we closed our initial public offering, in which we issued and sold a total of 6,900,000 shares of our common stock, including 900,000 shares sold pursuant to the exercise in full by the underwriters of their option to purchase additional shares, at a public offering price of \$15.00 per share, and received net proceeds of \$92.7 million after deducting underwriting commissions and discounts and offering expenses payable by us.

In June 2011, we entered into an amendment to our original loan and security agreement with Hercules, pursuant to which we were entitled to borrow a term loan in the principal amount of up to \$12.0 million with a floating interest rate equal to the greater of (1) 10.15 percent or (2) the sum of 10.15 percent plus the prime rate published on The Wall Street Journal minus 5.75 percent, not to exceed 12.75 percent per annum, which interest was computed daily based on the actual number of days elapsed. On April 7, 2014, we repaid the remaining amount of the Hercules loan in full for a total payment of \$3.6 million. We granted Hercules a security interest in certain of our assets. In connection with the loan and security agreement, as amended, we issued to Hercules warrants to purchase 21,000 shares of Series A preferred stock and 26,400 shares of Series B preferred stock, respectively, each at an exercise price of \$25.00 per share. The warrants became exercisable to purchase our common stock immediately prior to the closing of our initial public offering. On February 11, 2014, Hercules net exercised these warrants in exchange for a total of 12,702 shares of our common stock.

In addition to our existing cash and cash equivalents, for each product candidate under our research collaboration and license agreement with KHK, we are entitled to receive clinical, regulatory and commercialization milestone payments of up to \$110.0 million and royalties on net sales of such product candidate. Our ability to earn these milestone payments and the timing of achieving these milestones is dependent upon the outcome of our research and development and regulatory activities and is uncertain at this time. Our right to receive the payment of certain milestones under our agreement with KHK is our only committed external source of funds.

**Table of Contents****Cash flows**

As of June 30, 2014, we had \$120.3 million in cash and cash equivalents and held-to-maturity investments and \$1.4 million in assets held in restriction.

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

|   | <b>SIX MONTHS ENDED</b> |                |                   |
|---|-------------------------|----------------|-------------------|
|   | <b>JUNE 30,</b>         |                |                   |
|   | <b>2014</b>             | <b>2013</b>    | <b>Difference</b> |
| Net cash used in operating activities     | \$ (14,271)             | \$ (2,721)     | \$ (11,550)       |
| Net cash used in investing activities     | (44,476)                | (3)            | (44,473)          |
| Net cash provided by financing activities | 89,884                  | 895            | 88,989            |
| <br>Increase in cash and cash equivalents | <br>\$ 31,137           | <br>\$ (1,829) | <br>\$ 32,966     |

**Operating activities**

Net cash used in operating activities for the six month periods ended June 30, 2014 and 2013 was \$14.3 million and \$2.7 million, respectively. The increase in cash used in operating activities of \$11.6 million for the six months ended June 30, 2014 was primarily due to an increase in our net loss of \$14.6 million for the six months ended June 30, 2014, which was partially offset by the change in working capital and other non-cash items of \$3.0 million, including a payment of \$5.0 million received in 2013 under a license agreement related to an option exercise fee and preclinical payments earned in December 2012.

**Investing activities**

Net cash used in investing activities for the six month periods ended June 30, 2014 and 2013 was \$44.5 million and \$0, respectively. Net cash used in investing activities for the periods presented relates to purchases of investments in held-to-maturity investments, purchases of property and equipment, primarily laboratory equipment, and an increase in assets held in restriction.

**Financing activities**

Net cash provided by financing activities for the six month periods ended June 30, 2014 and 2013 was \$89.9 million and \$0.9 million, respectively. In 2014, net proceeds from our initial public offering was \$94.1 million and proceeds from other issuance of common stock was \$0.8 million, partially offset by the repayment and principal payments related to the Hercules loan for \$5.0 million. In 2013, we had net proceeds of \$3.0 million from a bridge loan financing, offset by \$2.0 million of repayments on long-term debt.

**Funding requirements**

We expect that our primary uses of capital will continue to be third-party clinical research and development services, compensation and related expenses, laboratory and related supplies, legal and other regulatory expenses and general overhead costs. We believe that our cash and cash equivalents as of June 30, 2014, but excluding any potential option exercise fees or milestone payments, will be sufficient to meet our anticipated cash requirements through 2016.

However, we may require additional capital for the further development of our existing product candidates and may also need to raise additional funds sooner to pursue other development activities related to additional product candidates.

Our forecast of the period of time through which our financial resources will be adequate to support our operations is a forward-looking statement that involves risks and uncertainties, and actual results could vary materially as a result of a number of factors. We have based this estimate on assumptions that may prove to be wrong, and we could utilize our available capital resources sooner than we currently expect. Our future capital requirements are difficult to forecast and will depend on many factors, including:

the receipt of milestone payments under our research collaboration and license agreement with KHK;

the terms and timing of any other collaboration, licensing and other arrangements that we may establish;

the initiation, progress, timing and completion of preclinical studies and clinical trials for our potential product candidates;

the number and characteristics of product candidates that we pursue;

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the progress, costs and results of our preclinical studies and clinical trials;

the outcome, timing and cost of regulatory approvals;

delays that may be caused by changing regulatory requirements;

the cost and timing of hiring new employees to support our continued growth;

the costs involved in filing and prosecuting patent applications and enforcing and defending patent claims;

the costs of filing and prosecuting intellectual property rights and enforcing and defending any intellectual property-related claims;

the costs and timing of procuring clinical and commercial supplies of our product candidates;

the extent to which we acquire or in-license other product candidates and technologies; and

the extent to which we acquire or invest in other businesses, product candidates or technologies.

Please see the risk factors set forth in Part II, Item 1A Risk Factors in this Quarterly Report on Form 10-Q for additional risks associated with our substantial capital requirements.

Until such time, if ever, we generate product revenue, we expect to finance our cash needs through a combination of equity offerings, debt financings and research collaboration and license agreements. We may be unable to raise capital or enter into such other arrangements when needed or on favorable terms, or at all. Our failure to raise capital or enter into such other arrangements as and when needed would have a negative impact on our financial condition and our ability to develop our product candidates.

**Contractual Obligations and Commitments**

The following is a summary of our significant contractual obligations as of June 30, 2014 (in thousands).

| CONTRACTUAL OBLIGATIONS                             | TOTAL    | PAYMENTS DUE BY PERIOD |  |                   |                   |
|---|----------|------------------------|--|-------------------|-------------------|
|   |          | LESS THAN 1 YEAR       | MORE THAN 1 YEAR AND LESS THAN 5 YEARS | MORE THAN 5 YEARS | MORE THAN 5 YEARS |
| Existing operating lease obligations <sup>(1)</sup> | \$ 1,484 | \$ 594                 | \$ 890                                 |                   |                   |
| Future operating lease obligations <sup>(2)</sup>   | 9,625    | 868                    | 3,076                                  | 3,259             | 2,422             |

|       |           |          |          |          |          |
|-------|-----------|----------|----------|----------|----------|
| Total | \$ 11,109 | \$ 1,462 | \$ 3,966 | \$ 3,259 | \$ 2,422 |
|-------|-----------|----------|----------|----------|----------|

- (1) Future minimum lease payments under our existing non-cancelable operating lease for our current office and laboratory space in Watertown, Massachusetts, as amended on July 3, 2013, that expires on November 30, 2016 with an average rent of approximately \$51 per month.
- (2) Future minimum lease payments under a non-cancelable operating lease for our future office and laboratory space in Cambridge, Massachusetts, as executed on July 11, 2014 with an average rent of approximately \$134 per month.

We also have obligations to make future payments to COH, PBL and Carnegie Institution of Washington that become due and payable on the achievement of certain development, regulatory and commercial milestones. We have not included these commitments on our balance sheet or in the table above because the achievement and timing of these milestones is not probable and estimable.

### **Off-balance Sheet Arrangements**

We did not have during the periods presented, and we do not currently have, any off-balance sheet arrangements, as defined in the rules and regulations of the Securities and Exchange Commission.

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**Segment Reporting**

We view our operations and manage our business as one segment, which is the discovery, research and development of treatments based on our RNAi technology platform.

**Item 3. QUANTITATIVE AND QUALITATIVE DISCLOSURE ABOUT MARKET RISK**

The primary objectives of our investment activities are to ensure liquidity and to preserve principal while at the same time maximizing the income we receive from our marketable securities without significantly increasing risk. Some of the securities that we invest in may have market risk related to changes in interest rates. As of June 30, 2014, we had cash and cash equivalents and held-to-maturity investments of \$120.3 million. Our primary exposure to market risk is interest rate sensitivity, which is affected by changes in the general level of U.S. interest rates. Due to the short-term maturities of our cash and cash equivalents and the low risk profile of our investments, an immediate 100 basis point change in interest rates would not have a material effect on the fair market value of our cash and cash equivalents. To minimize the risk in the future, we intend to maintain our portfolio of cash and cash equivalents and short-term investments in a variety of securities, including commercial paper, money market funds, government and non-government debt securities and corporate obligations.



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**Item 4. CONTROLS AND PROCEDURES**

**Disclosure Controls and Procedures**

We maintain disclosure controls and procedures that are designed to ensure that information required to be disclosed in our periodic and current reports that we file under the Securities Exchange Act of 1934, as amended (Exchange Act), with the Securities and Exchange Commission (SEC) is recorded, processed, summarized and reported within the time periods specified in the SEC's rules and forms, and that such information is accumulated and communicated to our management, including our chief executive officer and principal financial officer, as appropriate, to allow timely decisions regarding required disclosure.

As of the end of the period covered by this Quarterly Report on Form 10-Q, we carried out an evaluation, under the supervision and with the participation of our management, including the chief executive officer and the chief financial officer, of the effectiveness of the design and operation of our disclosure controls and procedures pursuant to Exchange Act Rule 13a-15. Based upon, and as of the date of, this evaluation, the chief executive officer and the chief financial officer concluded that our disclosure controls and procedures were effective. Accordingly, management believes that the financial statements included in this report fairly present in all material respects our financial condition, results of operations and cash flows for the periods presented.

**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 was no change in our internal control over financial reporting during the quarter ended June 30, 2014, which was identified in connection with our management's evaluation required by Exchange Act Rules 13a-15(f) and 15d-15(f) that has materially affected, or is reasonably likely to materially affect, our internal control over financial reporting.

**Inherent Limitations on the Effectiveness of Controls**

Our management, including the chief executive officer and chief financial officer, does not expect that our disclosure controls and procedures or our internal control over financial reporting will prevent all error and all fraud. A control system, no matter how well conceived and operated, can provide only reasonable, not absolute, assurance that the objectives of the control system are met. Because of the inherent limitations in all control systems, no evaluation of controls can provide absolute assurance that all control issues and instances of fraud, if any, within the company have been detected. These inherent limitations include the realities that judgments in decision making can be faulty and that breakdowns can occur because of simple error or mistake. Additionally, controls can be circumvented by the individual acts of some persons, by collusion of two or more people or by management override of the control. The design of any system of controls also is based in part upon certain assumptions about the likelihood of future events, and there can be no assurance that any design will succeed in achieving its stated goals under all potential future conditions. Over time, controls may become inadequate because of changes in conditions, or the degree of compliance with the policies or procedures may deteriorate. Because of the inherent limitations in a cost-effective control system, misstatements due to error or fraud may occur and not be detected.

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**Part II: OTHER INFORMATION**

**Item 1. LEGAL PROCEEDINGS**

We are not currently a party to or aware of any proceedings that we believe will have, individually or in the aggregate, a material adverse effect on our business, financial condition or results of operations.

**Item 1A. RISK FACTORS**

*We are providing the following cautionary discussion of risk factors, uncertainties and assumptions that we believe are relevant to our business. These are factors that, individually or in the aggregate, we think could cause our actual results to differ materially from expected and historical results and our forward-looking statements. We note these factors for investors as permitted by Section 21E of the Securities Exchange Act of 1934, as amended (Exchange Act), and Section 27A of the Securities Act of 1933, as amended (Securities Act). You should understand that it is not possible to predict or identify all such factors. Consequently, you should not consider this section to be a complete discussion of all potential risks or uncertainties that may substantially impact our business. Moreover, we operate in a competitive and rapidly changing environment. New factors emerge from time to time and it is not possible to predict the impact of all of these factors on our business, financial condition or results of operations.*

**Risks Related to Our Business**

*We are a preclinical stage biopharmaceutical company with a history of losses, expect to continue to incur significant losses for the foreseeable future and may never achieve or maintain profitability, which could result in a decline in the market value of our common stock.*

We are a preclinical stage biopharmaceutical company with a limited operating history, focused on the discovery and development of treatments based on the emerging therapeutic modality RNA interference (RNAi), a biological process in which ribonucleic acid (RNA) molecules inhibit gene expression. Since our inception in October 2006, we have devoted our resources to the development of Dicer substrate RNA (DsiRNA) molecules and delivery technologies. We have had significant operating losses since our inception. As of June 30, 2014, we had an accumulated deficit of \$108.0 million. Our net loss was \$22.4 million for the six months ended June 30, 2014. Substantially all of our losses have resulted from expenses incurred in connection with our research programs and from general and administrative costs associated with our operations. Our technologies and product candidates are in early stages of development, and we are subject to the risks of failure inherent in the development of product candidates based on novel technologies.

To date, we have generated revenue primarily from the receipt of upfront research funding, license and option exercise fees and preclinical payments under our research collaboration and license agreement with Kyowa Hakko Kirin Co., Ltd. (KHK). We have not generated, and do not expect to generate, any product revenue for the foreseeable future, and we expect to continue to incur significant operating losses for the foreseeable future due to the cost of research and development, preclinical studies and clinical trials and the regulatory approval process for product candidates. The amount of future losses is uncertain. Our ability to achieve profitability, if ever, will depend on, among other things, us or our existing collaborator, or any future collaborators, successfully developing product candidates, obtaining regulatory approvals to market and commercialize product candidates, manufacturing any approved products on commercially reasonable terms, establishing a sales and marketing organization or suitable third party alternatives for any approved product and raising sufficient funds to finance business activities. If we or our existing collaborator, or

any future collaborators, are unable to develop and commercialize one or more of our product candidates or if sales revenue from any product candidate that receives approval is insufficient, we will not achieve profitability, which could have a material adverse effect on our business, financial condition, results of operations and prospects.

***To date, our revenue has been primarily derived from our research collaboration and license agreement with KHK, and we are dependent on KHK for the successful development of product candidates in the collaboration.***

In December 2009, we entered into a research collaboration and license agreement with KHK for the research, development and commercialization of DsiRNA molecules and drug delivery technologies for therapeutic targets, primarily in oncology. Under the research collaboration and license agreement with KHK, KHK has paid us a total of \$17.5 million. During the first two years of the collaboration, we worked together with KHK to optimize KHK's lipid nanoparticles for tumor delivery and to identify DsiRNAs optimized against oncology and KRAS targets. Based on the results of this research, KHK exercised options to advance two separate DsiRNAs into the development stage, including one with a KRAS target. For each product candidate under the research collaboration and license agreement, we have the potential to receive clinical, regulatory and commercialization milestone payments of up to \$110.0 million and royalties on net sales of such product candidate. The success of our collaboration programs with KHK depends entirely upon the efforts of KHK. Except for certain co-promotion and profit sharing rights we retain with respect to the KRAS product candidate if it is approved for marketing and commercialization in the U.S., KHK has sole discretion in determining and directing the efforts and resources, including the ability to discontinue all efforts and

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resources, it applies to the development and, if approval is obtained, commercialization and marketing of the product candidates covered by the collaboration. KHK may not be effective in obtaining approvals for the product candidates developed under the collaboration arrangement or in marketing, or arranging for necessary supply, manufacturing or distribution relationships for, any approved products. Under the research collaboration and license agreement, KHK may change its strategic focus or pursue alternative technologies in a manner that results in reduced, delayed or no revenue to us. KHK has a variety of marketed products and product candidates under collaboration with other companies, including some of our competitors, and its own corporate objectives may not be consistent with our best interests. If KHK fails to develop, obtain regulatory approval for or ultimately commercialize any product candidate under our collaboration or if KHK terminates our collaboration, our business, financial condition, results of operations and prospects could be materially and adversely affected. In addition, any dispute or litigation proceedings we may have with KHK in the future could delay development programs, create uncertainty as to ownership of intellectual property rights, distract management from other business activities and generate substantial expense.

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

We will need substantial additional funds to expand our development, regulatory, manufacturing, marketing and sales capabilities or contract with other organizations to provide these capabilities for us. We have used substantial funds to develop our product candidates and delivery technologies and will require significant funds to conduct further research and development and preclinical testing and clinical trials of our product candidates, to seek regulatory approvals for our product candidates and to manufacture and market products, if any, that are approved for commercial sale. As of June 30, 2014, we had \$120.3 million in cash and cash equivalents and held-to-maturity investments, including \$92.7 million of net proceeds, after deducting the underwriting commissions and discounts and the offering expenses, we received from the initial public offering of 6,900,000 shares of our common stock, which closed on February 4, 2014. Based on our current operating plan, we believe that our available cash, cash equivalents and short-term investments will be sufficient to fund our anticipated level of operations through 2016. Our future capital requirements and the period for which we expect our existing resources to support our operations may vary significantly from what we expect. Our monthly spending levels vary based on new and ongoing development and corporate activities. Because the length of time and activities associated with successful development of our product candidates is highly uncertain, we are unable to estimate the actual funds we will require for development and any approved marketing and commercialization activities. To execute our business plan, we will need, among other things:

to obtain the human and financial resources necessary to develop, test, obtain regulatory approval for, manufacture and market our product candidates;

to build and maintain a strong intellectual property portfolio and avoid infringing intellectual property of third parties;

to establish and maintain successful licenses, collaborations and alliances;

to satisfy the requirements of clinical trial protocols, including patient enrollment;

to establish and demonstrate the clinical efficacy and safety of our product candidates;

to obtain regulatory approvals;

to manage our spending as costs and expenses increase due to preclinical studies and clinical trials, regulatory approvals and commercialization;

to obtain additional capital to support and expand our operations; and

to market our products to achieve acceptance and use by the medical community in general.

If we are unable to obtain funding on a timely basis or on acceptable terms, we may have to delay, reduce or terminate our research and development programs and preclinical studies or clinical trials, if any, limit strategic opportunities or undergo reductions in our workforce or other corporate restructuring activities. We also could be required to seek funds through arrangements with collaborators or others that may require us to relinquish rights to some of our technologies or product candidates that we would otherwise pursue on our own. We do not expect to realize revenue from product sales, milestone payments or royalties in the foreseeable future, if at all. Our revenue sources are, and will remain, extremely limited unless and until our product candidates are clinically tested, approved for commercialization and successfully marketed. To date, we have primarily financed our operations through the sale of securities, debt financings, credit and loan facilities and payments received under our collaboration and license agreement with KHK. We will be required to seek additional funding in the future and intend to do so through either collaborations, equity offerings or debt financings, credit or loan facilities or a combination of one or more of these funding sources. Our ability to raise additional funds will depend on financial, economic and other factors, many of which are beyond our control. Additional funds may not be available to us on acceptable terms or at all. If we raise additional funds by issuing equity securities, our stockholders will suffer dilution and the terms of any financing may adversely affect the rights of our stockholders. In addition, as a condition to providing additional funds to us, future investors may demand, and may be granted, rights superior to those of existing stockholders. Debt financing, if available, may involve restrictive covenants limiting our flexibility in conducting future business activities, and, in the event of insolvency, debt holders would be repaid before holders of equity securities receive any distribution of corporate assets.

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*Our quarterly operating results may fluctuate significantly or may fall below the expectations of investors or securities analysts, each of which may cause our stock price to fluctuate or decline.*

We expect our operating results to be subject to quarterly fluctuations. Our net loss and other operating results will be affected by numerous factors, including:

variations in the level of expense related to our product candidates or future development programs;

results of clinical trials, or the addition or termination of clinical trials or funding support by us, our existing collaborator or any future collaborator or licensing partner;

the timing of the release of results from any clinical trials conducted by us or our collaborator KHK;

our execution of any collaboration, licensing or similar arrangement, and the timing of payments we may make or receive under such existing or future arrangements or the termination or modification of any such existing or future arrangements;

any intellectual property infringement lawsuit or opposition, interference or cancellation proceeding in which we may become involved;

additions and departures of key personnel;

strategic decisions by us or our competitors, such as acquisitions, divestitures, spin-offs, joint ventures, strategic investments or changes in business strategy;

if any of our product candidates receives regulatory approval, market acceptance and demand for such product candidates;

regulatory developments affecting our product candidates or those of our competitors; and

changes in general market and economic conditions.

If our quarterly operating results fall below the expectations of investors or securities analysts, the price of our common stock could decline substantially. Furthermore, any quarterly fluctuations in our operating results may, in turn, cause the price of our stock to fluctuate substantially. We believe that quarterly comparisons of our financial results are not necessarily meaningful and should not be relied upon as an indication of our future performance.

***Our approach to the discovery and development of innovative therapeutic treatments based on novel technologies is unproven and may not result in marketable products.***

We plan to develop a pipeline of product candidates using our DsiRNA molecules and delivery technologies for rare inherited diseases involving the liver and cancers that are genetically defined. We believe that product candidates identified with our drug discovery and delivery platform may offer an improved therapeutic approach to small molecules and monoclonal antibodies, as well as several advantages over earlier generation RNAi molecules. However, the scientific research that forms the basis of our efforts to develop product candidates based on the therapeutic modality RNAi and the identification and optimization of DsiRNA is relatively new. Further, the scientific evidence to support the feasibility of developing therapeutic treatments based on RNAi and DsiRNA is both preliminary and limited.

Relatively few product candidates based on RNAi have been tested in animals or humans, and a number of clinical trials conducted by other companies using RNAi technologies have not been successful. We may discover that DsiRNA does not possess certain properties required for a drug to be effective, such as the ability to remain stable in the human body for the period of time required for the drug to reach the target tissue or the ability to cross the cell wall and enter into cells within the target tissue for effective delivery. We currently have only limited data, and no conclusive evidence, to suggest that we can introduce these necessary drug-like properties into DsiRNA. We may spend substantial funds attempting to introduce these properties and may never succeed in doing so. In addition, product candidates based on DsiRNA may demonstrate different chemical and pharmacological properties in patients than they do in laboratory studies. Even if product candidates, such as DCR-PH1 and DCR-MYC, have successful results in animal studies, they 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, we may never succeed in developing a marketable product, we may not become profitable and the value of our common stock will decline.

Further, the U.S. Food and Drug Administration (FDA) has relatively limited experience with RNAi and DsiRNA based therapeutics. No regulatory authority has granted approval to any person or entity, including us, to market and commercialize therapeutics using RNAi or DsiRNA, which may increase the complexity, uncertainty and length of the regulatory approval process for our product candidates. We and our

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current collaborator, or any future collaborators, may never receive approval to market and commercialize any product candidate. Even if we or a collaborator obtain regulatory approval, the approval may be for disease indications or patient populations that are not as broad as we intended or desired or may require labeling that includes significant use or distribution restrictions or safety warnings. We or a collaborator may be required to perform additional or unanticipated clinical trials to obtain approval or be subject to post-marketing testing requirements to maintain regulatory approval. If our technologies based on DsiRNA prove to be ineffective, unsafe or commercially unviable, our entire platform and pipeline would have little, if any, value, which could have a material adverse effect on our business, financial condition, results of operations and prospects.

***The market may not be receptive to our product candidates based on a novel therapeutic modality, and we may not generate any future revenue from the sale or licensing of product candidates.***

Even if approval is obtained for a product candidate, we may not generate or sustain revenue from sales of the product due to factors such as whether the product can be sold at a competitive cost and otherwise accepted in the market. The product candidates that we are developing are based on new technologies and therapeutic approaches. Market participants with significant influence over acceptance of new treatments, such as physicians and third-party payors, may not adopt a treatment based on DsiRNA technology, and we may not be able to convince the medical community and third-party payors to accept and use, or to provide favorable reimbursement for, any product candidates developed by us or our existing collaborator or any future collaborators. Market acceptance of our product candidates will depend on, among other factors:

the timing of our receipt of any marketing and commercialization approvals;

the terms of any approvals and the countries in which approvals are obtained;

the safety and efficacy of our product candidates;

the prevalence and severity of any adverse side effects associated with our product candidates;

limitations or warnings contained in any labeling approved by the FDA or other regulatory authority;

relative convenience and ease of administration of our product candidates;

the willingness of patients to accept any new methods of administration;

the success of our physician education programs;

the availability of adequate government and third-party payor reimbursement;



the pricing of our products, particularly as compared to alternative treatments; and

availability of alternative effective treatments for the disease indications our product candidates are intended to treat and the relative risks, benefits and costs of those treatments.

With our focus on the emerging therapeutic modality RNAi, these risks may increase to the extent the space becomes more competitive or less favored in the commercial marketplace. Additional risks apply in relation to any disease indications we pursue which are classified as rare diseases and allow for orphan drug designation by regulatory agencies in major commercial markets, such as the U.S., Europe and Japan. For instance, we are in the preliminary stages of developing a treatment for the rare genetic disorder Primary Hyperoxaluria 1 (PH1) with the gene encoding the liver metabolic enzyme glycolate oxidase as our target. Because of the small patient population for a rare disease, if pricing is not approved or accepted in the market at an appropriate level for an approved product with orphan drug designation, such drug may not generate enough revenue to offset costs of development, manufacturing, marketing and commercialization despite any benefits received from the orphan drug designation, such as market exclusivity, assistance in clinical trial design or a reduction in user fees or tax credits related to development expense. Market size is also a variable in disease indications not classified as rare. Our estimates regarding potential market size for any indication may be materially different from what we discover to exist at the time we commence commercialization, if any, for a product, which could result in significant changes in our business plan and have a material adverse effect on our business, financial condition, results of operations and prospects.

***Our product candidates are in early stages of development and may fail in development or suffer delays that materially adversely affect their commercial viability.***

We have no products on the market and all of our product candidates are in early stages of development. Our ability to achieve and sustain profitability depends on obtaining regulatory approvals, including institutional review board (IRB) approval, for and successfully commercializing our product candidates, either alone or with third parties, such as our collaborator KHK. Before obtaining regulatory approval for the commercial distribution of our product candidates, we or a collaborator must conduct extensive preclinical tests and clinical trials to demonstrate the safety and efficacy in humans of our product candidates. Preclinical testing and clinical trials are expensive, difficult to design

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and implement, can take many years to complete and are uncertain as to outcome. The start or end of a clinical study is often delayed or halted due to changing regulatory requirements, manufacturing challenges, required clinical trial administrative actions, slower than anticipated patient enrollment, changing standards of care, availability or prevalence of use of a comparative drug or required prior therapy, clinical outcomes or financial constraints. For instance, delays or difficulties in patient enrollment or difficulties in retaining trial participants can result in increased costs, longer development times or termination of a clinical trial. Clinical trials of a new product candidate require the enrollment of a sufficient number of patients, including patients who are suffering from the disease the product candidate is intended to treat and who meet other eligibility criteria. Rates of patient enrollment are affected by many factors, including the size of the patient population, the eligibility criteria for the clinical trial, the age and condition of the patients, the stage and severity of disease, the nature of the protocol, the proximity of patients to clinical sites and the availability of effective treatments for the relevant disease.

A product candidate can unexpectedly fail at any stage of preclinical and clinical development. The historical failure rate for product candidates is high due to scientific feasibility, safety, efficacy, changing standards of medical care and other variables. The results from preclinical testing or early clinical trials of a product candidate may not predict the results that will be obtained in later phase clinical trials of the product candidate. We, the FDA or other applicable regulatory authorities may suspend clinical trials of a product candidate at any time for various reasons, including a belief that subjects participating in such trials are being exposed to unacceptable health risks or adverse side effects. We may not have the financial resources to continue development of, or to enter into collaborations for, a product candidate if we experience any problems or other unforeseen events that delay or prevent regulatory approval of, or our ability to commercialize, product candidates, including:

negative or inconclusive results from our clinical trials or the clinical trials of others for product candidates similar to ours, leading to a decision or requirement to conduct additional preclinical testing or clinical trials or abandon a program;

serious and unexpected drug-related side effects experienced by participants in our clinical trials or by individuals using drugs similar to our product candidates;

delays in submitting Investigational New Drug applications (INDs) or comparable foreign applications or delays or failure in obtaining the necessary approvals from regulators to commence a clinical trial, or a suspension or termination of a clinical trial once commenced;

conditions imposed by the FDA or comparable foreign authorities regarding the scope or design of our clinical trials;

delays in enrolling research subjects in clinical trials;

high drop-out rates of research subjects;

inadequate supply or quality of product candidate components or materials or other supplies necessary for the conduct of our clinical trials;

greater than anticipated clinical trial costs;

poor effectiveness of our product candidates during clinical trials;

unfavorable FDA or other regulatory agency inspection and review of a clinical trial site;

failure of our third-party contractors or investigators to comply with regulatory requirements or otherwise meet their contractual obligations in a timely manner, or at all;

delays and changes in regulatory requirements, policy and guidelines, including the imposition of additional regulatory oversight around clinical testing generally or with respect to our technology in particular; or

varying interpretations of data by the FDA and similar foreign regulatory agencies.

***If third parties on which we depend to conduct our preclinical studies, or any future clinical trials, do not perform as contractually required, fail to satisfy regulatory or legal requirements or miss expected deadlines, our development program could be delayed with materially adverse effects on our business, financial condition, results of operations and prospects.***

We rely on third party clinical investigators, contract research organizations (CROs), clinical data management organizations and consultants to design, conduct, supervise and monitor preclinical studies of our product candidates and will do the same for any clinical trials. Because we rely on third parties and do not have the ability to conduct preclinical studies or clinical trials independently, we have less control over the timing, quality and other aspects of preclinical studies and clinical trials than we would if we conducted them on our own. These investigators, CROs and consultants are not our employees and we have limited control over the amount of time and resources that they dedicate to our programs. These third parties may have contractual relationships with other entities, some of which may be our competitors, which may draw time and resources from our programs. The third parties with which we contract might not be diligent, careful or timely in conducting our preclinical studies or clinical trials, resulting in the preclinical studies or clinical trials being delayed or unsuccessful.

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If we cannot contract with acceptable third parties on commercially reasonable terms, or at all, or if these third parties do not carry out their contractual duties, satisfy legal and regulatory requirements for the conduct of preclinical studies or clinical trials or meet expected deadlines, our clinical development programs could be delayed and otherwise adversely affected. In all events, we are responsible for ensuring that each of our preclinical studies and clinical trials is conducted in accordance with the general investigational plan and protocols for the trial. The FDA requires clinical trials to be conducted in accordance with good clinical practices, including for conducting, recording and reporting the results of preclinical studies and clinical trials to assure that data and reported results are credible and accurate and that the rights, integrity and confidentiality of clinical trial participants are protected. Our reliance on third parties that we do not control does not relieve us of these responsibilities and requirements. Any such event could have a material adverse effect on our business, financial condition, results of operations and prospects.

***Because we rely on third party manufacturing and supply partners, our supply of research and development, preclinical and clinical development materials may become limited or interrupted or may not be of satisfactory quantity or quality.***

We rely on third party supply and manufacturing partners to supply the materials and components for, and manufacture, our research and development, preclinical and clinical trial drug supplies. We do not own manufacturing facilities or supply sources for such components and materials. Our manufacturing requirements include lipid nanoparticle components and nucleic acid, each of which we procure from a single source supplier on a purchase order basis. In addition, we currently contract with only one drug product formulation manufacturer for the encapsulation of the oligonucleotide in a lipid particle. There can be no assurance that our supply of research and development, preclinical and clinical development drugs and other materials will not be limited, interrupted, restricted in certain geographic regions or of satisfactory quality or continue to be available at acceptable prices. In particular, any replacement of our drug product formulation manufacturer could require significant effort and expertise because there may be a limited number of qualified replacements.

The manufacturing process for a product candidate is subject to FDA and foreign regulatory authority review. Suppliers and manufacturers must meet applicable manufacturing requirements and undergo rigorous facility and process validation tests required by regulatory authorities in order to comply with regulatory standards, such as current Good Manufacturing Practices (cGMP). In the event that any of our suppliers or manufacturers fails to comply with such requirements or to perform its obligations to us in relation to quality, timing or otherwise, or if our supply of components or other materials becomes limited or interrupted for other reasons, we may be forced to manufacture the materials ourselves, for which we currently do not have the capabilities or resources, or enter into an agreement with another third party, which we may not be able to do on reasonable terms, if at all. In some cases, the technical skills or technology required to manufacture our product candidates may be unique or proprietary to the original manufacturer and we may have difficulty, or there may be contractual restrictions prohibiting us from, transferring such skills or technology to another third party and a feasible alternative may not exist. These factors would increase our reliance on such manufacturer or require us to obtain a license from such manufacturer in order to have another third party manufacture our product candidates. If we are required to change manufacturers for any reason, we will be required to verify that the new manufacturer maintains facilities and procedures that comply with quality standards and with all applicable regulations and guidelines. The delays associated with the verification of a new manufacturer could negatively affect our ability to develop product candidates in a timely manner or within budget.

We expect to continue to rely on third party manufacturers if we receive regulatory approval for any product candidate. To the extent that we have existing, or enter into future, manufacturing arrangements with third parties, we will depend on these third parties to perform their obligations in a timely manner consistent with contractual and regulatory requirements, including those related to quality control and assurance. If we are unable to obtain or maintain third-party manufacturing for product candidates, or to do so on commercially reasonable terms, we may not

be able to develop and commercialize our product candidates successfully. Our or a third party's failure to execute on our manufacturing requirements could adversely affect our business in a number of ways, including:

an inability to initiate or continue clinical trials of product candidates under development;

delay in submitting regulatory applications, or receiving regulatory approvals, for product candidates;

loss of the cooperation of a collaborator;

subjecting our product candidates to additional inspections by regulatory authorities;

requirements to cease distribution or to recall batches of our product candidates; and

in the event of approval to market and commercialize a product candidate, an inability to meet commercial demands for our products.

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***We may not successfully engage in strategic transactions, including any additional collaborations we seek, which could adversely affect our ability to develop and commercialize product candidates, impact our cash position, increase our expense and present significant distractions to our management.***

From time to time, we may consider strategic transactions, such as collaborations, acquisitions of companies, asset purchases and out- or in-licensing of product candidates or technologies. In particular, in addition to our current arrangement with KHK, we will evaluate and, if strategically attractive, seek to enter into additional collaborations, including with major biotechnology or pharmaceutical companies. The competition for collaborators is intense, and the negotiation process is time-consuming and complex. Any new collaboration may be on terms that are not optimal for us, and we may be unable to maintain any new or existing collaboration if, for example, development or approval of a product candidate is delayed, sales of an approved product candidate do not meet expectations or the collaborator terminates the collaboration. Any such collaboration, or other strategic transaction, may require us to incur non-recurring or other charges, increase our near- and long-term expenditures and pose significant integration or implementation challenges or disrupt our management or business. These transactions would entail numerous operational and financial risks, including exposure to unknown liabilities, disruption of our business and diversion of our management's time and attention in order to manage a collaboration or develop acquired products, product candidates or technologies, incurrence of substantial debt or dilutive issuances of equity securities to pay transaction consideration or costs, higher than expected collaboration, acquisition or integration costs, write-downs of assets or goodwill or impairment charges, increased amortization expenses, difficulty and cost in facilitating the collaboration or combining the operations and personnel of any acquired business, impairment of relationships with key suppliers, manufacturers or customers of any acquired business due to changes in management and ownership and the inability to retain key employees of any acquired business. Accordingly, although there can be no assurance that we will undertake or successfully complete any transactions of the nature described above, any transactions that we do complete may be subject to the foregoing or other risks and have a material adverse effect on our business, results of operations, financial condition and prospects. Conversely, any failure to enter any collaboration or other strategic transaction that would be beneficial to us could delay the development and potential commercialization of our product candidates and have a negative impact on the competitiveness of any product candidate that reaches market.

***We face competition from entities that have developed or may develop product candidates for our target disease indications, including companies developing novel treatments and technology platforms based on modalities and technology similar to ours. If these companies develop technologies or product candidates more rapidly than we do or their technologies, including delivery technologies, are more effective, our ability to develop and successfully commercialize product candidates may be adversely affected.***

The development and commercialization of drugs is highly competitive. We compete with a variety of multinational pharmaceutical companies and specialized biotechnology companies, as well as technology being developed at universities and other research institutions. Our competitors have developed, are developing or will develop product candidates and processes competitive with our product candidates. Competitive therapeutic treatments include those that have already been approved and accepted by the medical community and any new treatments that enter the market. We believe that a significant number of products are currently under development, and may become commercially available in the future, for the treatment of conditions for which we may try to develop product candidates. We are aware of multiple companies that are working in the field of RNAi therapeutics, including major pharmaceutical companies such as Novartis International AG, Takeda Pharmaceutical Company Limited and biopharmaceutical companies such as Alnylam Pharmaceuticals, Inc. (Alnylam), which in March 2014 acquired Sirna Therapeutics, Inc. from Merck & Co., Inc., Tekmira Pharmaceuticals Corporation (Tekmira), Arrowhead Research Corporation (Arrowhead), Silence Therapeutics plc, RXi Pharmaceuticals Corporation, Quark Pharmaceuticals, Inc. and Marina Biotech, Inc. In particular, Arrowhead holds a non-exclusive license to the same patent rights of City of Hope (COH) and Integrated Data Technologies, Inc. (IDT) as we are licensed under our license agreement with COH.

As a result, we cannot rely on those patent rights to prevent Arrowhead or third parties working with Arrowhead from developing, marketing and selling products that compete directly with our product candidates.

We also compete with companies working to develop antisense and other RNA-based drugs. Like RNAi therapeutics, antisense drugs target messenger RNA (mRNA) with the objective of suppressing the activity of specific genes. The development of antisense drugs is more advanced than that of RNAi therapeutics, and antisense technology may become the preferred technology for products that target mRNAs. Significant competition also exists from companies such as Tekmira and Arrowhead to discover and develop safe and effective means to deliver therapeutic RNAi molecules, such as DsiRNAs, to the relevant cell and tissue types.

If our lead product candidates are approved for the indications we are currently pursuing, they will compete with a range of therapeutic treatments that are either in development or currently marketed. For example, Nexavar, marketed by Amgen Inc. and Bayer AG, is currently in use for the treatment of hepatocellular carcinoma (HCC). Given the high unmet medical need and the commercial success of Nexavar, numerous targeted therapies for the treatment of HCC are under development. Targeted therapies represent the largest proportion of the HCC pipeline. There are also a number of pharmaceuticals and biologics that are marketed or in clinical development for the treatment of solid tumors. The most common treatments for solid tumors are various chemotherapeutic agents, radiation therapy and certain targeted therapies, including monoclonal antibodies such as Avastin, Erbitux, Herceptin and Vectibix. Small molecules, such as Nexavar, Sutent and Tareeva, are also indicated for the treatment of solid tumors. In addition, we believe that Kadmon Corporation, LLC is evaluating salirasib (KD032) in clinical trials for the treatment of KRAS-specific non-small cell lung cancer, pancreatic cancer and other solid tumors.

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Many of our competitors have significantly greater financial, technical, manufacturing, marketing, sales and supply resources or experience than we have. If we successfully obtain approval for any product candidate, we will face competition based on many different factors, including the safety and effectiveness of our products, the ease with which our products can be administered and the extent to which patients accept relatively new routes of administration, the timing and scope of regulatory approvals for these products, the availability and cost of manufacturing, marketing and sales capabilities, price, reimbursement coverage and patent position. Competing products could present superior treatment alternatives, including by being more effective, safer, less expensive or marketed and sold more effectively than any products we may develop. Competitive products may make any products we develop obsolete or noncompetitive before we recover the expense of developing and commercializing our product candidates. Such competitors could also recruit our employees, which could negatively impact our level of expertise and our ability to execute our business plan.

***Any inability to attract and retain qualified key management and technical personnel would impair our ability to implement our business plan.***

Our success largely depends on the continued service of key management and other specialized personnel, including Douglas M. Fambrough, III, Ph.D., our chief executive officer, Pankaj Bhargava, M.D., our chief medical officer, Bob D. Brown, Ph.D., our chief scientific officer, James E. Dentzer, our chief financial officer, and James B. Weissman, our chief business officer. The loss of one or more members of our management team or other key employees or advisors could delay our research and development programs and materially harm our business, financial condition, results of operations and prospects. The relationships that our key managers have cultivated within our industry make us particularly dependent upon their continued employment with us. We are dependent on the continued service of our technical personnel because of the highly technical nature of our product candidates and technologies and the specialized nature of the regulatory approval process. Because our management team and key employees are not obligated to provide us with continued service, they could terminate their employment with us at any time without penalty. We do not maintain key person life insurance policies on any of our management team members or key employees. Our future success will depend in large part on our continued ability to attract and retain other highly qualified scientific, technical and management personnel, as well as personnel with expertise in clinical testing, manufacturing, governmental regulation and commercialization. We face competition for personnel from other companies, universities, public and private research institutions, government entities and other organizations.

***If our product candidates advance into clinical trials, we may experience difficulties in managing our growth and expanding our operations.***

We have limited experience in drug development and have only recently begun our first clinical trial of our product candidates. As our product candidates enter and advance through preclinical studies and any clinical trials, we will need to expand our development, regulatory and manufacturing capabilities or contract with other organizations to provide these capabilities for us. In the future, we expect to have to manage additional relationships with collaborators or partners, suppliers and other organizations. Our ability to manage our operations and future growth will require us to continue to improve our operational, financial and management controls, reporting systems and procedures. We may not be able to implement improvements to our management information and control systems in an efficient or timely manner and may discover deficiencies in existing systems and controls.

***If any of our product candidates are approved for marketing and commercialization and we are unable to develop sales, marketing and distribution capabilities on our own or enter into agreements with third parties to perform these functions on acceptable terms, we will be unable to commercialize successfully any such future products.***



We currently have no sales, marketing or distribution capabilities or experience. If any of our product candidates is approved, we will need to develop internal sales, marketing and distribution capabilities to commercialize such products, which would be expensive and time-consuming, or enter into collaborations with third parties to perform these services. If we decide to market our products directly, we will need to commit significant financial and managerial resources to develop a marketing and sales force with technical expertise and supporting distribution, administration and compliance capabilities. If we rely on third parties with such capabilities to market our products or decide to co-promote products with collaborators, we will need to establish and maintain marketing and distribution arrangements with third parties, and there can be no assurance that we will be able to enter into such arrangements on acceptable terms or at all. In entering into third-party marketing or distribution arrangements, any revenue we receive will depend upon the efforts of the third parties and there can be no assurance that such third parties will establish adequate sales and distribution capabilities or be successful in gaining market acceptance of any approved product. If we are not successful in commercializing any product approved in the future, either on our own or through third parties, our business, financial condition, results of operations and prospects could be materially adversely affected.

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***If we fail to comply with U.S. and foreign regulatory requirements, regulatory authorities could limit or withdraw any marketing or commercialization approvals we may receive and subject us to other penalties that could materially harm our business.***

Even if we receive marketing and commercialization approval of a product candidate, we will be subject to continuing regulatory review, including in relation to adverse patient experiences with the product and clinical results that are reported after a product is made commercially available, both in the U.S. and any foreign jurisdiction in which we seek regulatory approval. The FDA has significant post-market authority, including the authority to require labeling changes based on new safety information and to require post-market studies or clinical trials to evaluate safety risks related to the use of a product or to require withdrawal of the product from the market. The FDA also has the authority to require a risk evaluation and mitigation strategies (REMS) plan after approval, which may impose further requirements or restrictions on the distribution or use of an approved drug. The manufacturer and manufacturing facilities we use to make a future product, if any, will also be subject to periodic review and inspection by the FDA and other regulatory agencies, including for continued compliance with cGMP requirements. The discovery of any new or previously unknown problems with our third-party manufacturers, manufacturing processes or facilities may result in restrictions on the product, manufacturer or facility, including withdrawal of the product from the market. If we rely on third-party manufacturers, we will not have control over compliance with applicable rules and regulations by such manufacturers. Any product promotion and advertising will also be subject to regulatory requirements and continuing regulatory review. If we or our collaborators, manufacturers or service providers fail to comply with applicable continuing regulatory requirements in the U.S. or foreign jurisdictions in which we seek to market our products, we or they may be subject to, among other things, fines, warning letters, holds on clinical trials, refusal by the FDA to approve pending applications or supplements to approved applications, suspension or withdrawal of regulatory approval, product recalls and seizures, refusal to permit the import or export of products, operating restrictions, injunction, civil penalties and criminal prosecution.

***Price controls imposed in foreign markets may adversely affect our future profitability.***

In some countries, particularly member states of the European Union, the pricing of prescription drugs is subject to governmental control. In these countries, pricing negotiations with governmental authorities can take considerable time after receipt of marketing approval for a product. In addition, there can be considerable pressure by governments and other stakeholders on prices and reimbursement levels, including as part of cost containment measures. Political, economic and regulatory developments may further complicate pricing negotiations, and pricing negotiations may continue after reimbursement has been obtained. Reference pricing used by various European Union member states and parallel distribution, or arbitrage between low-priced and high-priced member states, can further reduce prices. In some countries, we or our collaborators may be required to conduct a clinical trial or other studies that compare the cost-effectiveness of our RNAi therapeutic candidates to other available therapies in order to obtain or maintain reimbursement or pricing approval. Publication of discounts by third-party payors or authorities may lead to further pressure on the prices or reimbursement levels within the country of publication and other countries. If reimbursement of any product candidate approved for marketing is unavailable or limited in scope or amount, or if pricing is set at unsatisfactory levels, our business, financial condition, results of operations or prospects could be adversely affected.

***Our business entails a significant risk of product liability and our ability to obtain sufficient insurance coverage could have a material effect on our business, financial condition, results of operations or prospects.***

Our business exposes us to significant product liability risks inherent in the development, testing, manufacturing and marketing of therapeutic treatments. Product liability claims could delay or prevent completion of our development programs. If we succeed in marketing products, such claims could result in an FDA investigation of the safety and effectiveness of our products, our manufacturing processes and facilities or our marketing programs and potentially a

recall of our products or more serious enforcement action, limitations on the approved indications for which they may be used or suspension or withdrawal of approvals. Regardless of the merits or eventual outcome, liability claims may also result in decreased demand for our products, injury to our reputation, costs to defend the related litigation, a diversion of management's time and our resources, substantial monetary awards to trial participants or patients and a decline in our stock price. We currently have product liability insurance that we believe is appropriate for our stage of development and may need to obtain higher levels prior to marketing any of our product candidates. Any insurance we have or may obtain may not provide sufficient coverage against potential liabilities. Furthermore, clinical trial and product liability insurance is becoming increasingly expensive. As a result, we may be unable to obtain sufficient insurance at a reasonable cost to protect us against losses caused by product liability claims that could have a material adverse effect on our business.

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

We are exposed to the risk of employee fraud or other misconduct. Misconduct by employees could include intentional failures to comply with FDA regulations, provide accurate information to the FDA, comply with manufacturing standards we may establish, comply with federal and state healthcare fraud and abuse laws and regulations, report financial information or data accurately or disclose unauthorized activities to us. In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, kickbacks, self-dealing and other abusive practices. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Employee misconduct could also involve the improper use of information obtained in the course of clinical trials, which could result in regulatory sanctions and serious harm to our reputation. It is not always possible to identify and deter employee misconduct, and the precautions we take

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to detect and prevent this activity may not be effective in controlling unknown or unmanaged risks or losses or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to be in compliance with such laws or regulations. If any such actions are instituted against us, and we are not successful in defending ourselves or asserting our rights, those actions could have a significant impact on our business, including the imposition of significant fines or other sanctions.

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

Despite the implementation of security measures, our internal computer systems and those of our CROs and other contractors and consultants are vulnerable to damage from computer viruses, unauthorized access, natural disasters, terrorism, war and telecommunication and electrical failures. Such events could cause interruptions of our operations. For instance, the loss of preclinical data or data from any future clinical trial involving our product candidates could result in delays in our development and regulatory filing efforts and significantly increase our costs. To the extent that any disruption or security breach were to result in a loss of, or damage to, our data, or inappropriate disclosure of confidential or proprietary information, we could incur liability and the development of our product candidates could be delayed.

***If we do not comply with laws regulating the protection of the environment and health and human safety, our business could be adversely affected.***

Our research, development and manufacturing involves the use of hazardous materials and various chemicals. We maintain quantities of various flammable and toxic chemicals in our facilities in Watertown that are required for our research, development and manufacturing activities. We are subject to federal, state and local laws and regulations governing the use, manufacture, storage, handling and disposal of these hazardous materials. We believe our procedures for storing, handling and disposing these materials in our Watertown facilities comply with the relevant guidelines of Watertown, the Commonwealth of Massachusetts and the Occupational Safety and Health Administration of the U.S. Department of Labor. Although we believe that our safety procedures for handling and disposing of these materials comply with the standards mandated by applicable regulations, the risk of accidental contamination or injury from these materials cannot be eliminated. If an accident occurs, we could be held liable for resulting damages, which could be substantial. We are also subject to numerous environmental, health and workplace safety laws and regulations, including those governing laboratory procedures, exposure to blood-borne pathogens and the handling of animals and biohazardous materials. 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 these materials, this insurance may not provide adequate coverage against potential liabilities. We do not maintain insurance for environmental liability or toxic tort claims that may be asserted against us in connection with our storage or disposal of biological or hazardous materials. Additional federal, state and local laws and regulations affecting our operations may be adopted in the future. We may incur substantial costs to comply with, and substantial fines or penalties if we violate, any of these laws or regulations.

***Our information technology systems could face serious disruptions that could adversely affect our business.***

Our information technology and other internal infrastructure systems, including corporate firewalls, servers, leased lines and connection to the Internet, face the risk of systemic failure that could disrupt our operations. A significant disruption in the availability of our information technology and other internal infrastructure systems could cause interruptions in our collaborations with our partners and delays in our research and development work.

***Our current operations are concentrated in one location and any events affecting this location may have material adverse consequences.***

Our current operations are located in our facilities situated in Watertown, Massachusetts. Any unplanned event, such as flood, fire, explosion, earthquake, extreme weather condition, medical epidemics, power shortage, telecommunication failure or other natural or manmade accidents or incidents that result in us being unable to fully utilize the facilities, may have a material adverse effect on our ability to operate our business, particularly on a daily basis, and have significant negative consequences on our financial and operating conditions. Loss of access to these facilities may result in increased costs, delays in the development of our product candidates or interruption of our business operations. As part of our risk management policy, we maintain insurance coverage at levels that we believe are appropriate for our business. However, in the event of an accident or incident at these facilities, we cannot assure you that the amounts of insurance will be sufficient to satisfy any damages and losses. If our facilities are unable to operate because of an accident or incident or for any other reason, even for a short period of time, any or all of our research and development programs may be harmed. Any business interruption may have a material adverse effect on our business, financial position, results of operations and prospects.

***Our ability to utilize our net operating loss carryforwards and certain other tax attributes may be limited.***

We have incurred substantial losses during our history, do not expect to become profitable for the foreseeable future and may never achieve profitability. To the extent that we continue to generate taxable losses, unused losses will carry forward to offset future taxable income, if any, until such unused losses expire. We may be unable to use these losses to offset income before such unused losses expire. Under

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Section 382 of the Internal Revenue Code of 1986, as amended, if a corporation undergoes an ownership change, which is generally defined as a greater than 50 percentage point 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 may be further limited. We have not performed an analysis on whether we have experienced any ownership changes in the past. It is possible that we have experienced an ownership change, including pursuant to the initial public offering of our common stock, which closed on February 4, 2014, and our net operating losses are subject to such limitation. As of December 31, 2013, we had U.S. federal and Massachusetts net operating loss carryforwards of \$65.1 million and \$64.6 million, respectively. Any limit on these loss carryforwards if we have or do experience an ownership change could have an adverse effect on our business, financial position, results of operations and prospects.

***The investment of our cash and cash equivalents and held-to-maturity investments is subject to risks which may cause losses and affect the liquidity of these investments.***

As of June 30, 2014, we had \$120.3 million in cash and cash equivalents and held-to-maturity investments. We historically have invested substantially all of our available cash and cash equivalents in corporate bonds, commercial paper, securities issued by the U.S. government, certificates of deposit and money market funds meeting the criteria of our investment policy, which is focused on the preservation of our capital. These investments are subject to general credit, liquidity, market and interest rate risks, including the impact of U.S. sub-prime mortgage defaults that have affected various sectors of the financial markets and caused credit and liquidity issues. We may realize losses in the fair value of these investments or a complete loss of these investments, which would have a negative effect on our condensed consolidated financial statements.

In addition, should our investments cease paying or reduce the amount of interest paid to us, our interest income would suffer. The market risks associated with our investment portfolio may have an adverse effect on our results of operations, liquidity and financial condition.

***Changes in accounting rules and regulations, or interpretations thereof, could result in unfavorable accounting charges or require us to change our compensation policies.***

Accounting methods and policies for biopharmaceutical companies, including policies governing revenue recognition, research and development and related expenses and accounting for stock-based compensation, are subject to review, interpretation and guidance from our auditors and relevant accounting authorities, including the Securities and Exchange Commission and the Public Company Accounting Oversight Board. Changes to accounting methods or policies, or interpretations thereof, may require us to reclassify, restate or otherwise change or revise our financial statements, including those contained in this Quarterly Report on Form 10-Q.

**Risks Related to Intellectual Property**

***If we are not able to obtain and enforce patent protection for our technologies or product candidates, development and commercialization of our product candidates may be adversely affected.***

Our success depends in part on our ability to obtain and maintain patents and other forms of intellectual property rights, including in-licenses of intellectual property rights of others, for our product candidates, methods used to manufacture our product candidates and methods for treating patients using our product candidates, as well as our ability to preserve our trade secrets, to prevent third parties from infringing upon our proprietary rights and to operate without infringing upon the proprietary rights of others. As of August 1, 2014, our patent estate, including the patents and patent applications that we have licensed from COH included approximately 19 issued patents and approximately

70 pending patent applications for research and development of our DsiRNA molecules and delivery technologies. We may not be able to apply for patents on certain aspects of our product candidates or delivery technologies in a timely fashion or at all. Our existing issued and granted patents and any future patents we obtain may not be sufficiently broad to prevent others from using our technology or from developing competing products and technology. There is no guarantee that any of our pending patent applications will result in issued or granted patents, that any of our issued or granted patents will not later be found to be invalid or unenforceable or that any issued or granted patents will include claims that are sufficiently broad to cover our product candidates or delivery technologies or to provide meaningful protection from our competitors. Moreover, the patent position of biotechnology and pharmaceutical companies can be highly uncertain because it involves complex legal and factual questions. We will be able to protect our proprietary rights from unauthorized use by third parties only to the extent that our current and future proprietary technology and product candidates are covered by valid and enforceable patents or are effectively maintained as trade secrets. If third parties disclose or misappropriate our proprietary rights, it may materially and adversely impact our position in the market.

The U.S. Patent and Trademark Office (USPTO) and various foreign governmental patent agencies require compliance with a number of procedural, documentary, fee payment and other provisions during the patent process. There are situations in which noncompliance can result in abandonment or lapse of a patent or patent application, resulting in partial or complete loss of patent rights in the relevant jurisdiction. In such an event, competitors might be able to enter the market earlier than would otherwise have been the case. The standards applied by the

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USPTO and foreign patent offices in granting patents are not always applied uniformly or predictably. For example, there is no uniform worldwide policy regarding patentable subject matter or the scope of claims allowable in biotechnology and pharmaceutical patents. As such, we do not know the degree of future protection that we will have on our proprietary products and technology. While we will endeavor to try to protect our product candidates with intellectual property rights such as patents, as appropriate, the process of obtaining patents is time-consuming, expensive and sometimes unpredictable.

In addition, there are numerous recent changes to the patent laws and proposed changes to the rules of the USPTO which may have a significant impact on our ability to protect our technology and enforce our intellectual property rights. For example, the Leahy-Smith America Invents Act (AIA) enacted in 2011 involves significant changes in patent legislation. The Supreme Court has ruled on several patent cases in recent years, some of which cases either narrow the scope of patent protection available in certain circumstances or weaken the rights of patent owners in certain situations. The recent decision by the Supreme Court in *Association for Molecular Pathology v. Myriad Genetics, Inc.* precludes a claim to a nucleic acid having a stated nucleotide sequence which is identical to a sequence found in nature and unmodified. We currently are not aware of an immediate impact of this decision on our patents or patent applications because we are developing nucleic acid products which contain modifications that we believe are not found in nature. However, this decision has yet to be clearly interpreted by courts and by the USPTO. We cannot assure you that the interpretations of this decision or subsequent rulings will not adversely impact our patents or patent applications. In addition to increasing uncertainty with regard to our ability to obtain patents in the future, this combination of events has created uncertainty with respect to the value of patents, once obtained. Depending on decisions by the U.S. Congress, the federal courts and the USPTO, 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.

Once granted, patents may remain open to opposition, interference, re-examination, post-grant review, inter partes review, nullification or derivation action in court or before patent offices or similar proceedings for a given period after allowance or grant, during which time third parties can raise objections against such initial grant. In the course of such proceedings, which may continue for a protracted period of time, the patent owner may be compelled to limit the scope of the allowed or granted claims thus attacked, or may lose the allowed or granted claims altogether. In addition, there can be no assurance that:

Others will not or may not be able to make, use or sell compounds that are the same as or similar to our product candidates but that are not covered by the claims of the patents that we own or license.

We or our licensors, collaborators or any future collaborators are the first to make the inventions covered by each of our issued patents and pending patent applications that we own or license.

We or our licensors, collaborators or any future collaborators are the first to file patent applications covering certain aspects of our inventions.

Others will not independently develop similar or alternative technologies or duplicate any of our technologies without infringing our intellectual property rights.



A third party may not challenge our patents and, if challenged, a court may not hold that our patents are valid, enforceable and infringed.

Any issued patents that we own or have licensed will provide us with any competitive advantages, or will not be challenged by third parties.

We may develop additional proprietary technologies that are patentable.

The patents of others will not have an adverse effect on our business.

Our competitors do not conduct research and development activities in countries where we do not have enforceable patent rights and then use the information learned from such activities to develop competitive products for sale in our major commercial markets.

***We license patent rights from third-party owners or licensees. If such owners or licensees do not properly or successfully obtain, maintain or enforce the patents underlying such licenses, or if they retain or license to others any competing rights, our competitive position and business prospects may be adversely affected.***

We do, and will continue to, rely on intellectual property rights licensed from third parties to protect our technology. We are a party to a number of licenses that give us rights to third-party intellectual property that is necessary or useful for our business. In particular, we have a license from COH (on behalf of itself and IDT) to certain patent rights, which provide platform intellectual property for research and development of our DsiRNA molecules. Pursuant to this agreement, we have a worldwide license from COH (subject to the pre-existing non-exclusive license) for the exploitation of key intellectual property rights in this respect, and COH and IDT retain ownership of the patents and patent applications to which we are licensed under the agreement. We also intend to license additional third-party intellectual property in the

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future. Our success will depend in part on the ability of our licensors to obtain, maintain and enforce patent protection for our licensed intellectual property, in particular, those patents to which we have secured exclusive rights. Our licensors may not successfully prosecute the patent applications licensed to us. Even if patents issue or are granted, our licensors may fail to maintain these patents, may determine not to pursue litigation against other companies that are infringing these patents, or may pursue litigation less aggressively than we would. Further, we may not obtain exclusive rights, which would allow for third parties to develop competing products. Without protection for, or exclusive right to, the intellectual property we license, other companies might be able to offer substantially identical products for sale, which could adversely affect our competitive business position and harm our business prospects. In addition, we sublicense our rights under our third-party licenses to KHK and may sublicense such rights to current or future collaborators or any future strategic partners. Any impairment of these sublicensed rights could result in reduced revenue under our collaboration agreement with KHK or result in termination of an agreement by one or more of our collaborators or any future strategic partners.

***Certain third parties may also have rights in the patents related to DsiRNA included in the license granted to us by COH, including the core DsiRNA patent (U.S. 8,084,599), which could allow them to develop, market and sell product candidates in competition with ours.***

To the extent that we do not have exclusive rights in the patents covered by the license granted to us by COH, we cannot prevent third parties from developing DsiRNA based product candidates in competition with ours. Prior to entering into the license with us, COH had entered into a non-exclusive license with a third party with respect to such patent rights to manufacture, use, import, offer for sale and sell products covered by the licensed patent rights for the treatment or prevention of disease in humans (excluding viruses and delivery of products into the eye or ear). While we believe that such non-exclusive license has been terminated, COH has informed us that a sublicensee to that non-exclusive license was permitted to enter into an equivalent non-exclusive license which, to our knowledge, is subsisting with Arrowhead Research Corporation (Arrowhead), as successor to the non-exclusive license holder. As successor to the non-exclusive license holder, we believe that Arrowhead has substantially similar access to the same patent rights related to DsiRNA granted to us under our license with COH. Arrowhead is developing RNA-based therapeutics for the treatment of diseases of the liver, which may directly compete with our product candidates. In addition, the U.S. government has certain rights to the inventions covered by the patent rights and COH, as an academic research and medical center, has the right to practice the licensed patent rights for educational, research and clinical uses. If Arrowhead or another party develops, manufactures, markets and sells any product covered by the same patent rights and technologies that compete with ours, it could significantly undercut the value of any of our product candidates, which would materially adversely affect our revenue, financial condition and results of operations.

***Other companies or organizations may challenge our or our licensors' patent rights or may assert patent rights that prevent us from developing and commercializing our products.***

RNAi therapeutics is a relatively new scientific field, the commercial exploitation of which has resulted in many different patents and patent applications from organizations and individuals seeking to obtain patent protection in the field. We have obtained grants and issuances of RNAi, RNAi therapeutic and DsiRNA patents and have licensed many of these patents from third parties on an exclusive or non-exclusive basis. The issued patents and pending patent applications in the U.S. and in key markets around the world that we own or license claim many different methods, compositions and processes relating to the discovery, development, manufacture and commercialization of RNAi therapeutics and DsiRNA therapeutics. Specifically, we own and have licensed a portfolio of patents, patent applications and other intellectual property covering: (1) certain aspects of the structure and uses of DsiRNAs, including their manufacture and use as therapeutics, and DsiRNA-related mechanisms, (2) chemical modifications to DsiRNAs that improve their suitability for therapeutic uses, (3) DsiRNAs directed to specific gene sequences and drug targets as treatments for particular diseases and (4) delivery technologies, such as in the field of lipid chemistry, lipid

nanoparticles and lipid nanoparticle formulation.

As the field of RNAi therapeutics matures, patent applications are being processed by national patent offices around the world. There is uncertainty about which patents will issue, and, if they do, as to when, to whom, and with what claims. It is likely that there will be significant litigation in the courts and other proceedings, such as interference, reexamination and opposition proceedings, in various patent offices relating to patent rights in the RNAi therapeutics field. In many cases, the possibility of appeal or opposition exists for either us or our opponents, and it may be years before final, unappealable rulings are made with respect to these patents in certain jurisdictions. The timing and outcome of these and other proceedings is uncertain and may adversely affect our business if we are not successful in defending the patentability and scope of our pending and issued patent claims or if third parties are successful in obtaining claims that cover our DsiRNA technology or any of our product candidates. In addition, third parties may attempt to invalidate our intellectual property rights. Even if our rights are not directly challenged, disputes could lead to the weakening of our intellectual property rights. Our defense against any attempt by third parties to circumvent or invalidate our intellectual property rights could be costly to us, could require significant time and attention of our management and could have a material adverse effect on our business and our ability to successfully compete in the field of RNAi therapeutics.

There are many issued and pending patents that claim aspects of oligonucleotide chemistry and modifications that we may need to apply to our DsiRNA therapeutic candidates. There are also many issued patents that claim targeting genes or portions of genes that may be relevant for DsiRNA drugs we wish to develop. Thus, it is possible that one or more organizations will hold patent rights to which we will need a license. If those organizations refuse to grant us a license to such patent rights on reasonable terms, we may be unable to market products or perform research and development or other activities covered by these patents.

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Obtaining a valid and enforceable issued or granted patent covering our technology in the U.S. and worldwide can be extremely costly. In jurisdictions where we have not obtained patent protection, competitors may use our technology to develop their own products and further, may export otherwise infringing products to territories where we have patent protection, but where it is more difficult to enforce a patent as compared to the U.S. Competitor products may compete with our future products in jurisdictions where we do not have issued or granted patents or where our issued or granted patent claims or other intellectual property rights are not sufficient to prevent competitor activities in these jurisdictions. The legal systems of certain countries, particularly certain developing countries, make it difficult to enforce patents and such countries may not recognize other types of intellectual property protection, particularly that relating to biopharmaceuticals. This could make it difficult for us to prevent the infringement of our patents or marketing of competing products in violation of our proprietary rights generally in certain jurisdictions. Proceedings to enforce our patent rights in foreign jurisdictions could result in substantial cost and divert our efforts and attention from other aspects of our business.

We generally file a provisional patent application first (a priority filing) at the USPTO. A U.S. utility application and international application under the Patent Cooperation Treaty (PCT) are usually filed within twelve months after the priority filing. Based on the PCT filing, national and regional patent applications may be filed in the European Union, Japan, Australia and Canada and, depending on the individual case, also in any or all of, inter alia, China, India, South Korea, Singapore, Taiwan and South Africa. We have so far not filed for patent protection in all national and regional jurisdictions where such protection may be available. In addition, we may decide to abandon national and regional patent applications before grant. Finally, the grant proceeding of each national or regional patent is an independent proceeding which may lead to situations in which applications might in some jurisdictions be refused by the relevant registration authorities, while granted by others. It is also quite common that depending on the country, various scopes of patent protection may be granted on the same product candidate or technology.

The laws of some jurisdictions do not protect intellectual property rights to the same extent as the laws in the U.S., and many companies have encountered significant difficulties in protecting and defending such rights in such jurisdictions. 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 from others in those jurisdictions. Many countries have compulsory licensing laws under which a patent owner may be compelled to grant licenses to third parties. In addition, many 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 with respect to any patents relevant to our business, our competitive position in the relevant jurisdiction may be impaired and our business and results of operations may be adversely affected.

***We or our licensors, collaborators or any future strategic partners may become subject to third party claims or litigation alleging infringement of patents or other proprietary rights or seeking to invalidate patents or other proprietary rights, and we may need to resort to litigation to protect or enforce our patents or other proprietary rights, all of which could be costly, time consuming, delay or prevent the development and commercialization of our product candidates, or put our patents and other proprietary rights at risk.***

We or our licensors, collaborators or any future strategic partners may be subject to third-party claims for infringement or misappropriation of patent or other proprietary rights. We are generally obligated under our license or collaboration agreements to indemnify and hold harmless our licensors or collaborator for damages arising from intellectual property infringement by us. If we or our licensors, collaborators or any future strategic partners are found

to infringe a third party patent or other intellectual property rights, we could be required to pay damages, potentially including treble damages, if we are found to have willfully infringed. In addition, we or our licensors, collaborators or any future strategic partners may choose to seek, or be required to seek, a license from a third party, which may not be available on acceptable terms, if at all. Even if a license can be obtained on acceptable terms, the rights may be non-exclusive, which could give our competitors access to the same technology or intellectual property rights licensed to us. If we fail to obtain a required license, we or our collaborator, or any future collaborator, may be unable to effectively market product candidates based on our technology, which could limit our ability to generate revenue or achieve profitability and possibly prevent us from generating revenue sufficient to sustain our operations. In addition, we may find it necessary to pursue claims or initiate lawsuits to protect or enforce our patent or other intellectual property rights. The cost to us in defending or initiating any litigation or other proceeding relating to patent or other proprietary rights, even if resolved in our favor, could be substantial, and litigation would divert our management's attention. Some of our competitors may be able to sustain the costs of complex patent litigation more effectively than we can because they have substantially greater resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could delay our research and development efforts and limit our ability to continue our operations.

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If we were to initiate legal proceedings against a third party to enforce a patent covering one of our products or our technology, the defendant could counterclaim that our patent is invalid or unenforceable. In patent litigation in the U.S., defendant counterclaims alleging invalidity or unenforceability are commonplace. Grounds for a validity challenge could be an alleged failure to meet any of several statutory requirements, for example, lack of novelty, obviousness or non-enablement. Grounds for an unenforceability assertion could be an allegation that someone connected with prosecution of the patent withheld relevant information from the USPTO, or made a misleading statement, during prosecution. The outcome following legal assertions of invalidity and unenforceability during patent litigation is unpredictable. With respect to the validity question, for example, we cannot be certain that there is no invalidating prior art, of which we and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity or unenforceability, we would lose at least part, and perhaps all, of the patent protection on one or more of our products or certain aspects of our platform technology. Such a loss of patent protection could have a material adverse impact on our business. Patents and other intellectual property rights also will not protect our technology if competitors design around our protected technology without legally infringing our patents or other intellectual property rights.

***Intellectual property rights of third parties could adversely affect our ability to commercialize our product candidates, and we might be required to litigate or obtain licenses from third parties in order to develop or market our product candidates. Such litigation or licenses could be costly or not available on commercially reasonable terms.***

Because the RNAi intellectual property landscape is still evolving, it is difficult to conclusively assess our freedom to operate without infringing on third party rights. There are numerous companies that have pending patent applications and issued patents broadly directed to RNAi generally and to RNAi delivery technologies. Our competitive position may suffer if patents issued to third parties or other third party intellectual property rights cover our products or elements thereof, or our manufacture or uses relevant to our development plans. In such cases, we may not be in a position to develop or commercialize products or product candidates unless we successfully pursue litigation to nullify or invalidate the third party intellectual property right concerned, or enter into a license agreement with the intellectual property right holder, if available on commercially reasonable terms. For instance, we received a letter from Alnylam in March 2010, claiming that we require access to certain patent and patent applications owned or controlled by Alnylam and demanding that we cease and desist from alleged infringing activities unless and until we obtain a license from Alnylam for the necessary intellectual property. We have disputed Alnylam's claims and engaged in several discussions with Alnylam. We have not received any further correspondence from Alnylam since 2010 regarding this claim. However, there can be no assurance that Alnylam, which in March 2014 acquired the RNAi-related intellectual property of Sirna Therapeutics, Inc. from Merck & Co., will not continue to pursue this or other claims against us. We are aware of issued patents, and there may be others of which we are not aware, held by third parties that, if found to be valid and enforceable, could be alleged to be infringed by our DsiRNA molecules. We are also aware of pending patent applications, and there may be others of which we are not aware, that if they result in issued patents, could be alleged to be infringed by our DsiRNA molecules. If such an infringement claim should be brought and be successful, we may be required to pay substantial damages, be forced to abandon our product candidates or seek a license from any patent holders. No assurances can be given that a license will be available on commercially reasonable terms, if at all.

For example, we are aware of a European patent, granted in 2006, and assigned to Alnylam (EP 1 352 061 B1) (EP 061 patent), that broadly covers various RNAi constructs, including potentially our DsiRNA molecules. The EP 061 patent has been validated and is currently in force in Austria, Germany, Ireland, Liechtenstein, Switzerland, and the United Kingdom. It has not yet been validated in any other European countries. Another third party, Sirna Therapeutics, Inc. (Sirna Therapeutics), formerly a wholly-owned subsidiary of Merck & Co., Inc., filed an opposition seeking to revoke the EP 061 patent as invalid. In August 2009, the Opposition Division of the European Patent Office

(EPO) rejected the opposition and upheld all of the claims of the EP 061 patent as originally granted. That decision was appealed by Sirna Therapeutics to the Technical Board of Appeal of the EPO. In March 2014, Alnylam acquired Sirna Therapeutics from Merck & Co., Inc. and subsequently withdrew the appeal. It is therefore no longer possible to challenge this patent centrally in the EPO, though we could challenge this patent in the individual European countries where it has been validated. If the EP 061 patent remains in force in each validated European country, we could be prevented from commercializing our DsiRNA products in each of those countries and we could be sued for patent infringement in such countries. We are aware that others are pursuing patent applications directed to similar subject matter in the U.S. and other jurisdictions and reinstatement of a revoked European patent broadly covering various RNA constructs. If any one of these applications were ultimately to issue as patents or the revoked patent were reinstated with claims that cover our DsiRNA molecules, their methods of use or methods of delivery, we could be sued for patent infringement in each of those countries as well. If we were unsuccessful in defending ourselves in any of these actions, we may be required to pay substantial damages, be forced to abandon our product candidates or seek a license from any patent holders, in each case, in such countries. We believe that the expected expiration date of the EP 061 patent and any foreign counterparts that might issue is early 2022.

It is also possible that we have failed to identify relevant third party patents or applications. For example, U.S. applications filed before November 29, 2000 and certain U.S. applications filed after that date that will not be filed outside the U.S. remain confidential until patents issue. Patent applications in the U.S. and elsewhere are published approximately 18 months after the earliest filing for which priority is claimed, with such earliest filing date being commonly referred to as the priority date. Therefore, patent applications covering our products or platform technology could have been filed by others without our knowledge. Additionally, pending patent applications which have been published can, subject to certain limitations, be later amended in a manner that could cover our platform technologies, our products or the use of our products. Third party intellectual property right holders may also actively bring infringement claims against us. We cannot guarantee that

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we will be able to successfully settle or otherwise resolve such infringement claims. If we are unable to successfully settle future claims on terms acceptable to us, we may be required to engage in or continue costly, unpredictable and time-consuming litigation and may be prevented from or experience substantial delays in marketing our products. If we fail in any such dispute, in addition to being forced to pay damages, we may be temporarily or permanently prohibited from commercializing any of our product candidates that are held to be infringing. We might, if possible, also be forced to redesign product candidates so that we no longer infringe the third party intellectual property rights. Any of these events, even if we were ultimately to prevail, could require us to divert substantial financial and management resources that we would otherwise be able to devote to our business.

***If we fail to comply with our obligations under any license, collaboration or other agreements, we may be required to pay damages and could lose intellectual property rights that are necessary for developing and protecting our product candidates and delivery technologies or we could lose certain rights to grant sublicenses.***

Our current licenses impose, and any future licenses we enter into are likely to impose, various development, commercialization, funding, milestone, royalty, diligence, sublicensing, insurance, patent prosecution and enforcement, and other obligations on us. If we breach any of these obligations, or use the intellectual property licensed to us in an unauthorized manner, we may be required to pay damages and the licensor may have the right to terminate the license, which could result in us being unable to develop, manufacture and sell products that are covered by the licensed technology or enable a competitor to gain access to the licensed technology. Moreover, our licensors may own or control intellectual property that has not been licensed to us and, as a result, we may be subject to claims, regardless of their merit, that we are infringing or otherwise violating the licensor's rights. In addition, while we cannot currently determine the amount of the royalty obligations we would be required to pay on sales of future products, if any, the amounts may be significant. The amount of our future royalty obligations will depend on the technology and intellectual property we use in products that we successfully develop and commercialize, if any. Therefore, even if we successfully develop and commercialize products, we may be unable to achieve or maintain profitability.

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

In addition to seeking patent protection for certain aspects of our product candidates and delivery technologies, we also consider trade secrets, including confidential and unpatented know-how important to the maintenance of our competitive position. We protect trade secrets and confidential and unpatented know-how, in part, by entering into non-disclosure and confidentiality agreements with parties who have access to such knowledge, such as our employees, corporate collaborators, outside scientific collaborators, CROs, contract manufacturers, consultants, advisors and other third parties. We also enter into confidentiality and invention or patent assignment agreements with our employees and consultants that obligate them to maintain confidentiality and assign their inventions to us. 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 is unpredictable. In addition, some courts in the U.S. and certain foreign jurisdictions are less willing or unwilling to protect trade secrets. If any of our trade secrets were to be lawfully obtained or independently developed by a competitor, we would have no right to prevent them from using that technology or information to compete with us. If any of our trade secrets were to be disclosed to or independently developed by a competitor, our competitive position would be harmed.

***We may be subject to claims that we or our employees or consultants have wrongfully used or disclosed alleged trade secrets of our employees or consultants former employers or their clients. These claims may be costly to defend and if we do not successfully do so, we may be required to pay monetary damages and may lose valuable***



***intellectual property rights or personnel.***

Many of our employees were previously employed at universities or biotechnology or pharmaceutical companies, including our competitors or potential competitors. Although no claims against us are currently pending, we may be subject to claims that these employees or we have inadvertently or otherwise used or disclosed trade secrets or other proprietary information of their former employers. Litigation may be necessary to defend against these claims. If we fail in defending such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights or personnel. A loss of key research personnel or their work product could hamper our ability to commercialize, or prevent us from commercializing, our product candidates, which could severely harm our business. Even if we are successful in defending against these claims, litigation could result in substantial costs and be a distraction to management.

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

Our trademarks or trade names may be challenged, infringed, circumvented or declared generic or determined to be infringing on other marks. We may not be able to protect our rights to these trademarks and trade names or may be forced to stop using these names, which we need for name recognition by potential partners or customers in our markets of interest. If we are unable to establish name recognition based on our trademarks and trade names, we may not be able to compete effectively and our business may be adversely affected.

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

Our product candidates are subject to extensive governmental regulations relating to, among other things, research, testing, development, manufacturing, safety, efficacy, approval, recordkeeping, reporting, labeling, storage, packaging, advertising and promotion, pricing, marketing and distribution of drugs. Rigorous preclinical testing and clinical trials and an extensive regulatory approval process are required to be successfully completed in the U.S. and in many foreign jurisdictions before a new drug can be marketed. Satisfaction of these and other regulatory requirements is costly, time consuming, uncertain and subject to unanticipated delays. It is possible that none of the product candidates we may develop will obtain the regulatory approvals necessary for us or our collaborators to begin selling them.

We have very limited experience in conducting and managing the clinical trials necessary to obtain regulatory approvals, including approval by the FDA. The time required to obtain FDA and other approvals is unpredictable but typically takes many years following the commencement of clinical trials, depending upon the type, complexity and novelty of the product candidate. The standards that the FDA and its foreign counterparts use when regulating us are not always applied predictably or uniformly and can change. Any analysis we perform of data from preclinical and clinical activities is subject to confirmation and interpretation by regulatory authorities, which could delay, limit or prevent regulatory approval. We may also encounter unexpected delays or increased costs due to new government regulations, for example, from future legislation or administrative action, or from changes in FDA policy during the period of product development, clinical trials and FDA regulatory review. It is impossible to predict whether legislative changes will be enacted, or whether FDA or foreign regulations, guidance or interpretations will be changed, or what the impact of such changes, if any, may be.

Because the drugs we are developing may represent a new class of drug, the FDA and its foreign counterparts have not yet established any definitive policies, practices or guidelines in relation to these drugs. While we believe the product candidates that we are currently developing are regulated as new drugs under the Federal Food, Drug, and Cosmetic Act, the FDA could decide to regulate them or other products we may develop as biologics under the Public Health Service Act. The lack of policies, practices or guidelines may hinder or slow review by the FDA of any regulatory filings that we may submit. Moreover, the FDA may respond to these submissions by defining requirements we may not have anticipated. Such responses could lead to significant delays in the clinical development of our product candidates. In addition, because there may be approved treatments for some of the diseases for which we may seek approval, in order to receive regulatory approval, we may need to demonstrate through clinical trials that the product candidates we develop to treat these diseases, if any, are not only safe and effective, but safer or more effective than existing products. Furthermore, in recent years, there has been increased public and political pressure on the FDA with respect to the approval process for new drugs, and the FDA's standards, especially regarding drug safety, appear to have become more stringent.

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

restrictions may limit the size of the market for the product and affect reimbursement by third-party payors.

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

***If we or our collaborators, manufacturers or service providers fail to comply with healthcare laws and regulations, we or they could be subject to enforcement actions, which could affect our ability to develop, market and sell our products and may harm our reputation.***

We and our collaborators are subject to federal, state, and foreign healthcare laws and regulations pertaining to fraud and abuse and patients' rights. These laws and regulations include:

the U.S. federal healthcare program anti-kickback law, which prohibits, among other things, persons from soliciting, receiving or providing remuneration, directly or indirectly, to induce either the referral of an individual for a healthcare item or service, or the purchasing or ordering of an item or service, for which payment may be made under a federal healthcare program such as Medicare or Medicaid;

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the U.S. federal false claims law, which prohibits, among other things, individuals or entities from knowingly presenting or causing to be presented, claims for payment by government funded programs such as Medicare or Medicaid that are false or fraudulent, and which may apply to us by virtue of statements and representations made to customers or third parties;

the U.S. federal Health Insurance Portability and Accountability Act (HIPAA) and Health Information Technology for Economic and Clinical Health (HITECH) Act, which prohibit executing a scheme to defraud healthcare programs, impose requirements relating to the privacy, security, and transmission of individually identifiable health information, and require notification to affected individuals and regulatory authorities of certain breaches of security of individually identifiable health information;

the federal Open Payments regulations under the National Physician Payment Transparency Program have been issued under the Patient Protection and Affordable Care Act, as amended by the Health Care and Education Affordability Reconciliation Act, and will require that manufacturers of pharmaceutical and biological drugs covered by Medicare, Medicaid, and Children's Health Insurance Programs report all consulting fees, travel reimbursements, research grants, and other payments or gifts with values over \$10 made to physicians and teaching hospitals; and

state laws comparable to each of the above federal laws, such as, for example, anti-kickback and false claims laws applicable to commercial insurers and other non-federal payors, requirements for mandatory corporate regulatory compliance programs, and laws relating to patient data privacy and security.

If our operations are found to be in violation of any such requirements, we may be subject to penalties, including civil or criminal penalties, monetary damages, the curtailment or restructuring of our operations, loss of eligibility to obtain approvals from the FDA, or exclusion from participation in government contracting, healthcare reimbursement or other government programs, including Medicare and Medicaid, any of which could adversely affect our financial results. Although effective compliance programs can mitigate the risk of investigation and prosecution for violations of these laws, these risks cannot be entirely eliminated. Any action against us for an alleged or suspected violation could cause us to incur significant legal expenses and could divert our management's attention from the operation of our business, even if our defense is successful. In addition, achieving and sustaining compliance with applicable laws and regulations may be costly to us in terms of money, time and resources.

If we or our collaborators, manufacturers or service providers fail to comply with applicable federal, state or foreign laws or regulations, we could be subject to enforcement actions, which could affect our ability to develop, market and sell our products successfully and could harm our reputation and lead to reduced acceptance of our products by the market. These enforcement actions include, among others:

adverse regulatory inspection findings;

warning letters;

voluntary or mandatory product recalls or public notification or medical product safety alerts to healthcare professionals;

restrictions on, or prohibitions against, marketing our products;

restrictions on, or prohibitions against, importation or exportation of our products;

suspension of review or refusal to approve pending applications or supplements to approved applications;

exclusion from participation in government-funded healthcare programs;

exclusion from eligibility for the award of government contracts for our products;

suspension or withdrawal of product approvals;

product seizures;

injunctions; and

civil and criminal penalties and fines.

***Any drugs we develop may become subject to unfavorable pricing regulations, third-party reimbursement practices or healthcare reform initiatives, thereby harming our business.***

The regulations that govern marketing approvals, pricing and reimbursement for new drugs vary widely from country to country. 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. In some foreign markets, prescription pharmaceutical pricing remains subject to continuing

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governmental control even after initial approval is granted. Although we intend to monitor these regulations, our programs are currently in the early stages of development and we will not be able to assess the impact of price regulations for a number of years. As a result, we might obtain regulatory approval for a product in a particular country, but then be subject to price regulations that delay our commercial launch of the product and negatively impact the revenues we are able to generate from the sale of the product in that country.

Our ability to commercialize any products successfully also will depend in part on the extent to which reimbursement for these products and related treatments will be available from government health administration authorities, private health insurers and other organizations. Even if we succeed in bringing one or more products to the market, these products may not be considered cost-effective, and the amount reimbursed for any products may be insufficient to allow us to sell our products on a competitive basis. Because our programs are in the early stages of development, we are unable at this time to determine their cost effectiveness or the likely level or method of reimbursement. Increasingly, the third-party payors who reimburse patients or healthcare providers, such as government and private insurance plans, are requiring that drug companies provide them with predetermined discounts from list prices, and are seeking to reduce the prices charged or the amounts reimbursed for pharmaceutical products. If the price we are able to charge for any products we develop, or the reimbursement provided for such products, is inadequate in light of our development and other costs, our return on investment could be adversely affected.

We currently expect that certain/some drugs we develop may need to be administered under the supervision of a physician on an outpatient basis. Under currently applicable U.S. law, certain drugs that are not usually self-administered (including injectable drugs) may be eligible for coverage under the Medicare Part B program if:

they are incident to a physician's services;

they are reasonable and necessary for the diagnosis or treatment of the illness or injury for which they are administered according to accepted standards of medical practice; and

they have been approved by the FDA and meet other requirements of the statute.

There may be significant delays in obtaining coverage for newly-approved drugs, and coverage may be more limited than the purposes for which the drug is approved by the FDA. Moreover, eligibility for coverage does not imply that any drug will be reimbursed in all cases or at a rate that covers our costs, including research, development, manufacture, sale and distribution. Interim payments for new drugs, if applicable, may also not be sufficient to cover our costs and may not be made permanent. Reimbursement may be based on payments allowed for lower-cost drugs that are already reimbursed, may be incorporated into existing payments for other services and may reflect budgetary constraints or imperfections in Medicare data. 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 U.S. Third-party payors often rely upon Medicare coverage policy and payment limitations in setting their own reimbursement rates. Our inability to promptly obtain coverage and adequate reimbursement rates from both government-funded and private payors for new drugs that we develop and for which we obtain regulatory approval could have a material adverse effect on our operating results, our ability to raise capital needed to commercialize products and our financial condition.

We believe that the efforts of governments and third-party payors to contain or reduce the cost of healthcare and legislative and regulatory proposals to broaden the availability of healthcare will continue to affect the business and financial condition of pharmaceutical and biopharmaceutical companies. A number of legislative and regulatory changes in the healthcare system in the U.S. and other major healthcare markets have been proposed in recent years, and such efforts have expanded substantially in recent years. These developments have included prescription drug benefit legislation that was enacted and took effect in January 2006, healthcare reform legislation enacted by certain states, and major healthcare reform legislation that was passed by Congress and enacted into law in the U.S. in 2010. These developments could, directly or indirectly, affect our ability to sell our products, if approved, at a favorable price.

In March 2010, President Obama signed into law the Patient Protection and Affordable Care Act, as amended by the Health Care and Education Affordability Reconciliation Act (ACA), a sweeping law intended to broaden access to health insurance, reduce or constrain the growth of healthcare spending and enhance remedies against fraud and abuse. The ACA also contains provisions that will affect companies in the pharmaceutical industry and other healthcare related industries by imposing additional costs and changes to business practices. Provisions affecting pharmaceutical companies include the following.

Mandatory rebates for drugs sold into the Medicaid program have been increased, and the rebate requirement has been extended to drugs used in risk-based Medicaid managed care plans.

The 340B Drug Pricing Program under the Public Health Services Act has been extended to require mandatory discounts for drug products sold to certain critical access hospitals, cancer hospitals and other covered entities.

Pharmaceutical companies are required to offer discounts on brand-name drugs to patients who fall within the Medicare Part D coverage gap, commonly referred to as the Donut Hole.

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Pharmaceutical companies are required to pay an annual non-tax deductible fee to the federal government based on each company's market share of prior year total sales of branded products to certain federal healthcare programs, such as Medicare, Medicaid, Department of Veterans Affairs and Department of Defense. Since we expect our branded pharmaceutical sales to constitute a small portion of the total federal health program pharmaceutical market, we do not expect this annual assessment to have a material impact on our financial condition.

For product candidates classified as biologics, approval of an application for a follow-on biologic product may not become effective until 12 years after the date on which the reference innovator biologic product was first licensed by the FDA, with a possible six-month extension for pediatric products. After this exclusivity ends, it will be easier for biosimilar manufacturers to enter the market, which is likely to reduce the pricing for such products and could affect our profitability.

The full effects of the U.S. healthcare reform legislation cannot be known until the new law is fully implemented through regulations or guidance issued by the Centers for Medicare & Medicaid Services and other federal and state healthcare agencies. The financial impact of the U.S. healthcare reform legislation over the next few years will depend on a number of factors, including but not limited to, the policies reflected in implementing regulations and guidance and changes in sales volumes for products affected by the new system of rebates, discounts and fees. The new legislation may also have a positive impact on our future net sales, if any, by increasing the aggregate number of persons with healthcare coverage in the U.S., but such increases are unlikely to be realized until approximately 2014 at the earliest.

Moreover, we cannot predict what healthcare reform initiatives may be adopted in the future. Further federal and state legislative and regulatory developments are likely, and we expect ongoing initiatives in the U.S. to increase pressure on drug pricing. Such reforms could have an adverse effect on anticipated revenues from product candidates that we may successfully develop and for which we may obtain regulatory approval and may affect our overall financial condition and ability to develop product candidates.

***Our ability to obtain services, reimbursement or funding from the federal government may be impacted by possible reductions in federal spending.***

U.S. federal government agencies currently face potentially significant spending reductions. Under the Budget Control Act of 2011, the failure of Congress to enact deficit reduction measures of at least \$1.2 trillion for the years 2013 through 2021 triggered automatic cuts to most federal programs. These cuts would include aggregate reductions to Medicare payments to providers of up to two percent per fiscal year, starting in 2013. Under the American Taxpayer Relief Act of 2012, which was enacted on January 1, 2013, the imposition of these automatic cuts was delayed until March 1, 2013. Certain of these automatic cuts have been implemented. The full impact on our business of these automatic cuts is uncertain. If federal spending is reduced, anticipated budgetary shortfalls may also impact the ability of relevant agencies, such as the FDA or the National Institutes of Health to continue to function at current levels. Amounts allocated to federal grants and contracts may be reduced or eliminated. These reductions may also impact the ability of relevant agencies to timely review and approve drug research and development, manufacturing, and marketing activities, which may delay our ability to develop, market and sell any products we may develop.

***If any of our product candidates receives marketing approval and we or others later identify undesirable side effects caused by the product candidate, our ability to market and derive revenue from the product candidates could be compromised.***



In the event that any of our product candidates receive regulatory approval and we or others identify undesirable side effects caused by one of our products, any of the following adverse events could occur, which could result in the loss of significant revenue to us and materially and adversely affect our results of operations and business:

regulatory authorities may withdraw their approval of the product or seize the product;

we may be required to recall the product or change the way the product is administered to patients;

additional restrictions may be imposed on the marketing of the particular product or the manufacturing processes for the product or any component thereof;

we may be subject to fines, injunctions or the imposition of civil or criminal penalties;

regulatory authorities may require the addition of labeling statements, such as a black box warning or a contraindication;

we may be required to create a Medication Guide outlining the risks of such side effects for distribution to patients;

we could be sued and held liable for harm caused to patients;

the product may become less competitive; and

our reputation may suffer.

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**Risks Related to Our Common Stock**

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

We are an emerging growth company as defined in the Jumpstart Our Business Act (JOBS Act). For as long as we continue to be an emerging growth company, we may take advantage of exemptions from various reporting requirements that are applicable to other public companies that are not emerging growth companies, including (1) not being required to comply with the auditor attestation requirements of Section 404(b) of the Sarbanes-Oxley Act of 2002 (Sarbanes-Oxley Act), (2) reduced disclosure obligations regarding executive compensation in our periodic reports and proxy statements and (3) exemptions from the requirements of holding a nonbinding advisory vote on executive compensation and stockholder approval of any golden parachute payments not previously approved. We could be an emerging growth company for up to five years, although circumstances could cause us to lose that status earlier, including if the market value of our common stock held by non-affiliates exceeds \$700.0 million as of any June 30 before that time or if we have total annual gross revenue of \$1.0 billion or more during any fiscal year before that time, in which cases we would no longer be an emerging growth company as of the following December 31, or if we issue more than \$1.0 billion in non-convertible debt during any three-year period before that time, in which case we would no longer be an emerging growth company immediately. Even after we no longer qualify as an emerging growth company, we may still qualify as a smaller reporting company which would allow us to take advantage of many of the same exemptions from disclosure requirements including not being required to comply with the auditor attestation requirements of Section 404(b) of the Sarbanes-Oxley Act and reduced disclosure obligations regarding executive compensation in our periodic reports and proxy statements. We cannot predict if investors will find our common stock less attractive because we may rely on these exemptions. If some investors find our common stock less attractive as a result, there may be a less active trading market for our common stock and our share price may be more volatile.

Under the JOBS Act, emerging growth companies can also delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have irrevocably elected not to avail ourselves of this exemption from new or revised accounting standards and, therefore, will be subject to the same new or revised accounting standards as other public companies that are not emerging growth companies.

*Our stock price may be volatile and purchasers of our common stock could incur substantial losses.*

Our stock price is volatile. From January 30, 2014, the first day of trading of our common stock, through August 6, 2014, our stock had high and low closing sale prices in the range of \$46.00 and \$14.89 per share. The market price for our common stock may be influenced by many factors, including the other risks described in this section of this Quarterly Report on Form 10-Q titled "Risk Factors" and the following:

the success of competitive products or technologies;

results of preclinical and clinical studies of our product candidates, or those of our competitors, our existing collaborator or any future collaborators;

regulatory or legal developments in the U.S. and other countries, especially changes in laws or regulations applicable to our products;

introductions and announcements of new products by us, our commercialization partners, or our competitors, and the timing of these introductions or announcements;

actions taken by regulatory agencies with respect to our products, clinical studies, manufacturing process or sales and marketing terms;

actual or anticipated variations in our financial results or those of companies that are perceived to be similar to us;

the success of our efforts to acquire or in-license additional technologies, products or product candidates;

developments concerning our collaborations, including but not limited to those with our sources of manufacturing supply and our commercialization partners;

announcements by us or our competitors of significant acquisitions, strategic partnerships, joint ventures or capital commitments;

developments or disputes concerning patents or other proprietary rights, including patents, litigation matters and our ability to obtain patent protection for our products;

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our ability or inability to raise additional capital and the terms on which we raise it;

the recruitment or departure of key personnel;

changes in the structure of healthcare payment systems;

market conditions in the pharmaceutical and biotechnology sectors;

actual or anticipated changes in earnings estimates or changes in stock market analyst recommendations regarding our common stock, other comparable companies or our industry generally;

our failure or the failure of our competitors to meet analysts' projections or guidance that we or our competitors may give to the market;

fluctuations in the valuation of companies perceived by investors to be comparable to us;

announcement and expectation of additional financing efforts;

speculation in the press or investment community;

trading volume of our common stock;

sales of our common stock by us or our stockholders;

the absence of lock-up agreements in connection with the initial public offering of our common stock with the holders of substantially all of our outstanding shares;

the concentrated ownership of our common stock;

changes in accounting principles;

terrorist acts, acts of war or periods of widespread civil unrest;

natural disasters and other calamities; and

general economic, industry and market conditions.

In addition, the stock markets in general, and the markets for pharmaceutical, biopharmaceutical and biotechnology stocks in particular, have experienced extreme volatility that has been often unrelated to the operating performance of the issuer. These broad market and industry factors may seriously harm the market price of our common stock, regardless of our operating performance.

***The future issuance of equity or of debt securities that are convertible into equity will dilute our share capital.***

We may choose to raise additional capital in the future, depending on market conditions, strategic considerations and operational requirements. To the extent that additional capital is raised through the issuance of shares or other securities convertible into shares, our stockholders will be diluted. Future issuances of our common stock or other equity securities, or the perception that such sales 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. We cannot predict 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.

***None of our outstanding shares are subject to lock-up agreements in connection with the initial public offering of our common stock. The sale of a significant number of our shares may cause the market price of our common stock to drop significantly.***

Our executive officers as of the initial public offering of our common stock have entered into lock-up agreements with the underwriters for the initial public offering under which they have agreed, subject to specific exceptions, not to sell shares of our common stock or securities exchangeable or exercisable therefor owned by them for a period of 180 days after the date of the final prospectus for the offering. These restrictions expired on July 28, 2014. No other stockholders entered into lock-up agreements in connection with the initial public offering. As a result, none of our stockholders or securities holders remain subject to any lock-up agreements in connection with the initial public offering. As a result, all shares of our common stock, including shares of common stock issuable upon exercise of outstanding options and warrants, are available for sale in the public market except as limited by vesting and exercise requirements, our insider trading policy and volume limitations applicable to affiliates under Rule 144. The sale of a significant number of shares of our common stock in the public market or the perception that such sale may occur could significantly reduce the market price of our common stock.

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In addition, we filed a registration statement on Form S-8 on February 6, 2014 registering the issuance of a total of 4,369,672 shares of common stock subject to options or other equity awards issued or reserved for future issuance under our equity incentive plans. The shares registered under such registration statement are available for sale in the public market subject to option vesting and exercise requirements, our insider trading policy and volume limitations applicable to affiliates under Rule 144.

Moreover, the holders of a substantial number of shares of our common stock, including shares of common stock issuable upon exercise of outstanding warrants, have rights, subject to some conditions, to require us to file registration statements covering their shares or to include their shares in registration statements that we may file for ourselves or other stockholders. If such holders, by exercising their registration rights, cause a large number of securities to be registered and sold into the public market, these sales could reduce the market price for our common stock.

***The employment agreements with our executive officers may require us to pay severance benefits to officers who are terminated in connection with a change of control of us, which could harm our financial condition.***

Our executive officers are parties to employment agreements providing, in the event of a termination of employment in connection with a change of control of us, for cash payments of up to approximately \$2.1 million for severance and other benefits and acceleration of vesting of up to all outstanding stock options. The accelerated vesting of options could result in dilution to our existing stockholders and reduce the market price of our common stock. The payment of these severance benefits could harm our financial condition. In addition, these potential severance payments may discourage or prevent third parties from seeking a business combination with us.

***If securities or industry analysts do not publish 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 will be influenced by the research and reports that industry or securities analysts publish about us or our business. If any of the analysts who cover us issue an adverse or misleading opinion regarding us, our business model, our intellectual property or our stock performance, or if our target studies and operating results fail to meet the expectations of analysts, our stock price would likely decline. If one or more of these analysts 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 our stock price or trading volume to decline.

***Our principal stockholders and management own a significant percentage of our stock and will be able to exert significant control over matters subject to stockholder approval.***

As of August 6, 2014, our executive officers and directors, together with holders of five percent or more of our outstanding common stock and their respective affiliates, beneficially own, in the aggregate, approximately 81.6 percent of our outstanding common stock, including shares subject to outstanding options and warrants that are exercisable within 60 days after such date, based on the Forms 3 and 4 and Schedules 13D and 13G filed by them with the SEC. As a result, these stockholders, if acting together, will continue to have significant influence over the outcome of corporate actions requiring stockholder approval, including the election of directors, any merger, consolidation or sale of all or substantially all of our assets and any other significant corporate transaction. The interests of these stockholders may not be the same as or may even conflict with the interests of our other stockholders. For example, these stockholders could delay or prevent a change of control of our company, even if such a change of control would benefit our other stockholders, which could deprive our stockholders of an opportunity to receive a premium for their common stock as part of a sale of our company or our assets and might affect the prevailing market price of our common stock. The significant concentration of stock ownership may adversely affect

the trading price of our common stock due to investors' perception that conflicts of interest may exist or arise.

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

Provisions in our certificate of incorporation and our bylaws may delay or prevent an acquisition of us or a change in our management. In addition, these provisions may frustrate or prevent any attempts by our stockholders to replace or remove our current management by making it more difficult for stockholders to replace members of our board of directors. Because our board of directors is responsible for appointing the members of our management team, these provisions could in turn affect any attempt by our stockholders to replace current members of our management team. These provisions include:

a prohibition on actions by our stockholders by written consent;

a requirement that special meetings of stockholders, which the Company is not obligated to call more than once per calendar year, be called only by the chairman of our board of directors, our chief executive officer, our board of directors pursuant to a resolution adopted by a majority of the total number of authorized directors, or, subject to certain conditions, by our secretary at the request of the stockholders holding of record, in the aggregate, shares entitled to cast not less than ten percent of the votes at a meeting of the stockholders (assuming all shares entitled to vote at such meeting were present and voted);

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advance notice requirements for election to our board of directors and for proposing matters that can be acted upon at stockholder meetings; and

the authority of the board of directors to issue preferred stock with such terms as the board of directors may determine.

Moreover, because we are incorporated in Delaware, we are governed by the provisions of Section 203 of the Delaware General Corporation Law, as amended, which prohibits a person who owns in excess of 15 percent of our outstanding voting stock from merging or combining with us for a period of three years after the date of the transaction in which the person acquired in excess of 15 percent of our outstanding voting stock, unless the merger or combination is approved in a prescribed manner. These provisions would apply even if the proposed merger or acquisition could be considered beneficial by some stockholders.

***We will incur increased 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, and particularly after we are no longer an emerging growth company, we will incur significant legal, accounting and other expenses that we did not incur as a private company. The Sarbanes-Oxley Act, the Dodd-Frank Wall Street Reform and Consumer Protection Act, the listing requirements of the NASDAQ 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 will need to devote a substantial amount of time to these compliance initiatives. Moreover, these rules and regulations will increase our legal and financial compliance costs and will make some activities more time-consuming and costly. For example, we expect that these rules and regulations may 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. However, these rules and regulations are 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.

We are not currently required to comply with the SEC's rules that implement Section 404 of the Sarbanes-Oxley Act (Section 404), and are therefore not required to make a formal assessment of the effectiveness of our internal control over financial reporting for that purpose. Pursuant to Section 404, we will be required to furnish a report by our management on our internal control over financial reporting. However, while we remain an emerging growth company, we will not be required to include an attestation report on internal control over financial reporting issued by our independent registered public accounting firm. To achieve compliance with Section 404 within the prescribed period, we will be engaged in a process to document and evaluate our internal control over financial reporting, which is both costly and challenging. In this regard, 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 we will not 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. If we identify one or more material weaknesses, it could result in an adverse reaction in the financial markets due to a loss of confidence in the reliability of our financial statements. In addition, if we are not able to continue to meet these requirements, we may not be able to remain listed on The NASDAQ Market.



***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. As a result, capital appreciation, if any, of our common stock will be sole source of gain of our stockholders for the foreseeable future.

***We may incur significant costs from class action litigation due to our expected stock volatility.***

Our stock price may fluctuate for many reasons, including as a result of public announcements regarding the progress of our development efforts or the development efforts of our collaborators or competitors, the addition or departure of our key personnel, variations in our quarterly operating results and changes in market valuations of pharmaceutical and biotechnology companies. This risk is especially relevant to us because pharmaceutical and biotechnology companies have experienced significant stock price volatility in recent years. When the market price of a stock has been volatile as our stock price may be, holders of that stock have occasionally brought securities class action litigation against the company that issued the stock. If any of our stockholders were to bring a lawsuit of this type against us, even if the lawsuit is without merit, we could incur substantial costs defending the lawsuit. The lawsuit could also divert the time and attention of our management.

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*Our amended and restated bylaws designates the Court of Chancery of the State of Delaware as the sole and exclusive forum for certain types of actions and proceedings that may be initiated by our stockholders, which could limit our stockholders' ability to obtain a favorable judicial forum for disputes with us or our directors, officers or employees.*

Our amended and restated bylaws provide that, subject to limited exceptions, the Court of Chancery of the State of Delaware will be the sole and exclusive forum for any derivative action or proceeding brought on our behalf, 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, any action asserting a claim against us arising pursuant to any provision of the Delaware General Corporation Law, as amended, our amended and restated certificate of incorporation or our amended and restated bylaws, any action to interpret, apply, enforce or determine the validity of our amended and restated certificate of incorporation or our amended and restated bylaws or any other action asserting a claim against us that is governed by the internal affairs doctrine. Any person or entity purchasing or otherwise acquiring any interest in shares of our capital stock shall be deemed to have notice of and to have consented to the provisions of our amended and restated certificate of incorporation described above. This choice of forum provision may limit a stockholder's ability to bring a claim in a judicial forum that it finds favorable for disputes with us or our directors, officers or other employees, which may discourage such lawsuits against us and our directors, officers and employees. Alternatively, if a court were to find these provisions of our amended and restated certificate of incorporation inapplicable to, or unenforceable in respect of, one or more of the specified types of actions or proceedings, we may incur additional costs associated with resolving such matters in other jurisdictions, which could adversely affect our business and financial condition.

**Item 2. UNREGISTERED SALES OF EQUITY SECURITIES AND USE OF PROCEEDS*****(a) Unregistered Sales of Equity Securities***

During the six months ended June 30, 2014, we issued a total of 12,702 shares of common stock to Hercules Technology II, L.P. (Hercules) upon the cashless net exercise of certain warrants held by Hercules to purchase a total of 47,400 shares of our common stock at an exercise price of \$25.00 per share. We deemed such issuance to be exempt from registration under the Securities Act of 1933, as amended (Securities Act), in reliance on Section 4(2) of the Securities Act, including Regulation D and Rule 506 promulgated thereunder, relative to transactions by an issuer not involving a public offering.

During the six months ended June 30, 2014, a total of 153,953 shares of our common stock were issued in unregistered sales of securities upon exercise of stock options granted pursuant to our 2007 Employee, Director and Consultant Stock Plan, as amended, and 2010 Employee, Director and Consultant Equity Incentive Plan, as amended, for aggregate cash consideration of \$526,519. The issuances of these shares of common stock were exempt from registration under the Securities Act in reliance on Rule 701 of the Securities Act as offers and sales of securities under compensatory benefit plans and contracts relating to compensation in compliance with Rule 701. Each of the recipients of shares of common stock in a transaction exempt from registration either received or had adequate access, through employment, business or other relationships, to information about us.

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

On February 4, 2014, we completed the initial public offering of our common stock and sold and issued a total of 6,900,000 shares of our common stock, including 900,000 shares sold pursuant to the exercise in full by the underwriters of the option to purchase additional shares, at a public offering price of \$15.00 per share for aggregate gross proceeds of \$103.5 million before deducting underwriting commissions and discounts and offering expenses

payable by us.

Jefferies LLC, Leerink Partners LLC, and Stifel, Nicolaus & Company, Incorporated acted as joint book-running managers for our initial public offering. Robert W. Baird & Co. Incorporated acted as co-lead manager for our initial public offering. We received net proceeds of approximately \$92.7 million in our initial public offering after deducting underwriting commissions and discounts of \$7.2 million and offering expenses of \$3.5 million. None of the offering expenses were paid to our directors, officers or persons owning ten percent or more of any class of our equity securities, or to any of their associates or our affiliates.

The shares of common stock were registered under the Securities Act on a registration statement on Form S-1 (File No. 333-193150). The SEC declared the registration statement effective on January 29, 2014. Shares of our common stock began trading on The NASDAQ Global Select Market on January 30, 2014. On February 4, 2014, following the sale of 6,900,000 shares of our common stock, our initial public offering ended.

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There has been no material change in the planned or actual use of proceeds from our initial public offering as described in the final prospectus dated as of January 29, 2014 for the initial public offering and filed with the SEC pursuant to Rule 424(b) under the Securities Act on January 30, 2014.

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

We did not repurchase any of our equity securities during the six month period ended June 30, 2014.

**Item 3. DEFAULTS UPON SENIOR SECURITIES**

Not applicable.

**Item 4. MINE SAFETY DISCLOSURES**

Not applicable.

**Item 5. OTHER INFORMATION**

None.

**Item 6. EXHIBITS****Exhibit**

| <b>Number</b> | <b>Description of Documents</b>  |
|---------------|--|
| 3.1(1)        | Amended and Restated Certificate of Incorporation of the Company.  |
| 3.2(1)        | Amended and Restated Bylaws of the Company.  |
| 4.1(2)        | Specimen Common Stock Certificate.   |
| 10.1(2)       | 2014 Performance Incentive Plan.++   |
| 10.2(2)       | 2014 Employee Stock Purchase Plan.++   |
| 10.3(2)       | Form of Indemnification Agreement by and between the Company and each of its directors.++                |
| 10.4(2)       | Letter agreement dated as of January 24, 2014, by and between the Company and James E. Dentzer.++        |
| 31.1(3)       | Certification of the Company's principal executive officer required by Rule 13a-14(a) or Rule 15d-14(a). |
| 31.2(3)       | Certification of the Company's principal financial officer required by Rule 13a-14(a) or Rule 15d-14(a). |
| 32.1*         | Section 1350 Certifications.   |

101\*\* The following materials from the Company's Quarterly Report on Form 10-Q for the quarter ended June 30, 2014, formatted in XBRL (eXtensible Business Reporting Language), include: (i) the Condensed Consolidated Balance Sheets as of June 30, 2014 (unaudited) and December 31, 2013, (ii) the Condensed Consolidated Statements of Operations (unaudited) for the three and six months ended June 30, 2014 and 2013, (iii) the Condensed Consolidated Statements of Cash Flows (unaudited) for the six months ended June 30, 2014 and 2013, and (iv) the Notes to Condensed Consolidated Financial Statements (unaudited).

++ Management contract or compensatory plan or arrangement.

- (1) Incorporated by reference to the indicated exhibit in the Company's Current Report on Form 8-K filed on February 5, 2014.
- (2) Incorporated by reference to the indicated exhibit in the Company's Amendment No. 3 to Registration Statement on Form S-1 (No. 333-193150) filed on January 28, 2014.
- (3) Filed herewith.

\* Exhibit 32.1 is being furnished and shall not be deemed to be filed for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (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 of 1933, as amended (Securities Act), or the Exchange Act, except as otherwise stated in such filing.

\*\* Exhibit 101 is being furnished and, in accordance with Rule 406T of Regulation S-T, shall not be deemed to be filed for purposes of Section 18 of the Exchange Act or otherwise subject to 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.

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**SIGNATURES**

Pursuant to 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.

DICERNA PHARMACEUTICALS, INC.

Date: August 7, 2014

By: /s/ James E. Dentzer

**James E. Dentzer**  
**Chief Financial Officer (Principal Financial**  
**Officer and Principal Accounting Officer)**

**Table of Contents****EXHIBIT INDEX****Exhibit**

| <b>Number</b> | <b>Description of Documents</b>   |
|---------------|---|
| 3.1(1)        | Amended and Restated Certificate of Incorporation of the Company.   |
| 3.2(1)        | Amended and Restated Bylaws of the Company.   |
| 4.1(2)        | Specimen Common Stock Certificate.  |
| 10.1(2)       | 2014 Performance Incentive Plan.++  |
| 10.2(2)       | 2014 Employee Stock Purchase Plan.++  |
| 10.3(2)       | Form of Indemnification Agreement by and between the Company and each of its directors.++   |
| 10.4(2)       | Letter agreement dated as of January 24, 2014, by and between the Company and James E. Dentzer.++   |
| 31.1(3)       | Certification of the Company's principal executive officer required by Rule 13a-14(a) or Rule 15d-14(a).  |
| 31.2(3)       | Certification of the Company's principal financial officer required by Rule 13a-14(a) or Rule 15d-14(a).  |
| 32.1*         | Section 1350 Certifications.  |
| 101**         | The following materials from the Company's Quarterly Report on Form 10-Q for the quarter ended June 30, 2014, formatted in XBRL (eXtensible Business Reporting Language), include: (i) the Condensed Consolidated Balance Sheets as of June 30, 2014 (unaudited) and December 31, 2013, (ii) the Condensed Consolidated Statements of Operations (unaudited) for the three and six months ended June 30, 2014 and 2013, (iii) the Condensed Consolidated Statements of Cash Flows (unaudited) for the six months ended June 30, 2014 and 2013, and (iv) the Notes to Condensed Consolidated Financial Statements (unaudited). |

++ Management contract or compensatory plan or arrangement.

- (1) Incorporated by reference to the indicated exhibit in the Company's Current Report on Form 8-K filed on February 5, 2014.
- (2) Incorporated by reference to the indicated exhibit in the Company's Amendment No. 3 to Registration Statement on Form S-1 (No. 333-193150) filed on January 28, 2014.
- (3) Filed herewith.

\* Exhibit 32.1 is being furnished and shall not be deemed to be filed for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (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 of 1933, as amended (Securities Act), or the Exchange Act, except as otherwise stated in such filing.

\*\* Exhibit 101 is being furnished and, in accordance with Rule 406T of Regulation S-T, shall not be deemed to be filed for purposes of Section 18 of the Exchange Act or otherwise subject to 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.

