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

FORM S-8
REGISTRATION STATEMENT
UNDER
THE SECURITIES ACT OF 1933

Karyopharm Therapeutics Inc.
(Exact name of registrant as specified in its charter)

Delaware
(State or other jurisdiction of
incorporation or organization)

26-3931704
(I.R.S. Employer
Identification No.)

85 Wells Avenue, 2nd Floor
Newton, MA
(Address of Principal Executive Offices)

02459
(Zip Code)

Inducement Stock Option Awards (August 2018 – July 2019)
(Full title of the plan)

Michael G. Kauffman
Chief Executive Officer
Karyopharm Therapeutics Inc.
85 Wells Avenue, 2nd Floor
Newton, MA 02459
(Name and address of agent for service)

(617) 658-0600
(Telephone number, including area code, of agent for service)

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

Large accelerated filer Accelerated filer
Non-accelerated filer (Do not check if a smaller reporting company) Smaller reporting company
Emerging growth company

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

CALCULATION OF REGISTRATION FEE

Title of Securities to be Registered	Amount to be Registered (1)	Proposed Maximum Offering Price Per Share	Proposed Maximum Aggregate Offering Price	Amount of Registration Fee
Common Stock, \$0.0001 par value per share	150,000 shares(2)	\$5.06(4)	\$759,000(4)	\$91.99
Common Stock, \$0.0001 par value per share	45,000 shares(3)	\$21.05(4)	\$947,250(4)	\$114.81
Common Stock, \$0.0001 par value per share	74,000 shares(3)	\$17.03(4)	\$1,260,220(4)	\$152.74
Common Stock, \$0.0001 par value per share	113,000 shares(3)	\$10.54(4)	\$1,191,020(4)	\$144.35
Common Stock, \$0.0001 par value per share	140,500 shares(3)	\$10.40(4)	\$1,461,200(4)	\$177.10
Common Stock, \$0.0001 par value per share	72,500 shares(3)	\$9.37(4)	\$679,325(4)	\$82.33
Common Stock, \$0.0001 par value per share	413,000 shares(3)	\$8.47(4)	\$3,498,110(4)	\$423.97
Common Stock, \$0.0001 par value per share	75,000 shares(3)	\$4.13(4)	\$309,750(4)	\$37.54
Common Stock, \$0.0001 par value per share	38,000 shares(3)	\$5.84(4)	\$221,920(4)	\$26.90
Common Stock, \$0.0001 par value per share	75,500 shares(3)	\$4.67(4)	\$352,585(4)	\$42.73
Common Stock, \$0.0001 par value per share	95,500 shares(3)	\$5.64(4)	\$538,620(4)	\$65.28
Common Stock, \$0.0001 par value per share	55,500 shares(3)	\$5.99(4)	\$332,445(4)	\$40.29
Common Stock, \$0.0001 par value per share	67,500 shares(3)	\$8.81(4)	\$594,675(4)	\$72.07

- (1) In accordance with Rule 416 under the Securities Act of 1933, as amended, this registration statement shall be deemed to cover any additional securities that may from time to time be offered or issued to prevent dilution resulting from stock splits, stock dividends or similar transactions.
- (2) Consists of shares issuable under a new hire inducement stock option award granted on February 25, 2019 to Michael Mason in accordance with Nasdaq Listing Rule 5635(c)(4), as an inducement material to his entering into employment with the registrant as Senior Vice President, Chief Financial Officer and Treasurer.
- (3) Consists of shares issuable under new hire inducement stock option awards granted between August 31, 2018 and July 31, 2019 to certain employees in accordance with Nasdaq Listing Rule 5635(c)(4), as an inducement material to their entering into employment with the registrant.
- (4) Estimated solely for the purpose of calculating the registration fee pursuant to Rule 457(h) of the Securities Act of 1933, as amended. The price per share and aggregate offering price are calculated on the basis of the exercise price of the applicable inducement stock option award.

PART I
INFORMATION REQUIRED IN THE SECTION 10(a) PROSPECTUS

Item 1. Plan Information.

The information required by Item 1 is included in documents sent or given to recipients of the new hire inducement stock option awards covered by this registration statement pursuant to Rule 428(b)(1) of the Securities Act of 1933, as amended (the "Securities Act").

Item 2. Registrant Information and Employee Plan Annual Information.

The written statement required by Item 2 is included in documents sent or given to recipients of the new hire inducement stock option awards covered by this registration statement pursuant to Rule 428(b)(1) of the Securities Act.

PART II
INFORMATION REQUIRED IN THE REGISTRATION STATEMENT

Item 3. Incorporation of Documents by Reference.

The registrant is subject to the informational and reporting requirements of Sections 13(a), 14, and 15(d) of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), and in accordance therewith files reports, proxy statements and other information with the Securities and Exchange Commission (the "Commission"). The following documents, which are on file with the Commission, are incorporated in this registration statement by reference:

- (a) The registrant's latest annual report filed pursuant to Section 13(a) or 15(d) of the Exchange Act or the latest prospectus filed pursuant to Rule 424(b) under the Securities Act that contains audited financial statements for the registrant's latest fiscal year for which such statements have been filed.
- (b) All other reports filed pursuant to Section 13(a) or 15(d) of the Exchange Act since the end of the fiscal year covered by the document referred to in (a) above.
- (c) The description of the securities contained in the registrant's registration statement on [Form 8-A](#) filed under the Exchange Act, including any amendment or report filed for the purpose of updating such description.

All documents subsequently filed by the registrant pursuant to Sections 13(a), 13(c), 14 and 15(d) of the Exchange Act, prior to the filing of a post-effective amendment which indicates that all securities offered hereby have been sold or which deregisters all securities then remaining unsold, shall be deemed to be incorporated by reference in this registration statement and to be part hereof from the date of the filing of such documents. Any statement contained in a document incorporated or deemed to be incorporated by reference herein shall be deemed to be modified or superseded for the purposes of this registration statement to the extent that a statement contained herein or in any other subsequently filed document which also is or is deemed to be incorporated by reference herein modifies or supersedes such statement. Any statement so modified or superseded shall not be deemed, except as so modified or superseded, to constitute a part of this registration statement.

Item 4. Description of Securities.

Not applicable.

Item 5. Interests of Named Experts and Counsel.

Not applicable.

Item 6. Indemnification of Directors and Officers.

Section 102 of the General Corporation Law of the State of Delaware permits a corporation to eliminate the personal liability of directors of a corporation to the corporation or its stockholders for monetary damages for a breach of fiduciary duty as a director, except where the director breached his duty of loyalty, failed to act in good faith, engaged in intentional misconduct or knowingly violated a law, authorized the payment of a dividend or approved a stock repurchase in violation of Delaware corporate law or obtained an improper personal benefit. The registrant's certificate of incorporation provides that no director of the registrant shall be personally liable to it or its stockholders for monetary damages for any breach of fiduciary duty as a director, notwithstanding any provision of law imposing such liability, except to the extent that the General Corporation Law of the State of Delaware prohibits the elimination or limitation of liability of directors for breaches of fiduciary duty.

Section 145 of the General Corporation Law of the State of Delaware provides that a corporation has the power to indemnify a director, officer, employee, or agent of the corporation, or a person serving at the request of the corporation for another corporation, partnership, joint venture, trust or other enterprise in related capacities against expenses (including attorneys' fees), judgments, fines and amounts paid in settlement actually and reasonably incurred by the person in connection with an action, suit or proceeding to which he was or is or is threatened to be made a party to any threatened, pending, or completed action, suit or proceeding by reason of such position, if such person acted in good faith and in a manner he reasonably believed to be in or not opposed to the best interests of the corporation, and, in any criminal action or proceeding, had no reasonable cause to believe his conduct was unlawful, except that, in the case of actions brought by or in the right of the corporation, no indemnification shall be made with respect to any claim, issue or matter as to which such person shall have been adjudged to be liable to the corporation unless and only to the extent that the Court of Chancery or other adjudicating court determines that, despite the adjudication of liability but in view of all of the circumstances of the case, such person is fairly and reasonably entitled to indemnity for such expenses which the Court of Chancery or such other court shall deem proper.

The registrant's certificate of incorporation provides that it will indemnify each person who was or is a party or threatened to be made a party to any threatened, pending or completed action, suit or proceeding, whether civil, criminal, administrative or investigative (other than an action by or in the right of the registrant) by reason of the fact that he or she is or was, or has agreed to become, a director or officer of the registrant, or is or was serving, or has agreed to serve, at the registrant's request, as a director, officer, partner, employee or trustee of, or in a similar capacity with, another corporation, partnership, joint venture, trust or other enterprise (including any employee benefit plan) (all such persons being referred to as an "Indemnitee"), or by reason of any action alleged to have been taken or omitted in such capacity, against all expenses (including attorneys' fees), liabilities, losses, judgments, fines (including excise taxes and penalties arising under the Employee Retirement Income Security Act of 1974), and amounts paid in settlement actually and reasonably incurred in connection with such action, suit or proceeding and any appeal therefrom, if such Indemnitee acted in good faith and in a manner he or she reasonably believed to be in, or not opposed to, the best interests of the registrant, and, with respect to any criminal action or proceeding, he or she had no reasonable cause to believe his or her conduct was unlawful.

The registrant's certificate of incorporation also provides that it will indemnify any Indemnitee who was or is a party to any threatened, pending, or completed action or suit by or in the right of the registrant to procure a judgment in its favor by reason of the fact that the Indemnitee is or was, or has agreed to become, a director or officer of the registrant, or is or was serving, or has agreed to serve, at the registrant's request, as a director, officer, partner, employee or trustee of, or in a similar capacity with, another corporation, partnership, joint venture, trust or other enterprise (including any employee benefit plan), or by reason of any action alleged to have been taken or omitted in such capacity, against all expenses (including attorneys' fees) and, to the extent permitted by law, amounts paid in settlement actually and reasonably incurred in connection with such action, suit or proceeding and any appeal therefrom, if the Indemnitee acted in good faith and in a manner he or she reasonably believed to be in, or not opposed to, the best interests of the registrant, except that no indemnification shall be made with respect to any claim, issue or matter as to which such person shall have been adjudged to be liable to the registrant, unless a court determines that, despite such adjudication but in view of all of the circumstances, he or she is entitled to indemnification of such expenses. Notwithstanding the foregoing, to the extent that any Indemnitee has been successful, on the merits or otherwise, he or she will be indemnified by the registrant against all expenses (including attorneys' fees) actually and reasonably incurred in connection therewith. Expenses must be advanced to an Indemnitee under certain circumstances.

The registrant has entered into indemnification agreements with each of its directors. These indemnification agreements may require the registrant, among other things, to indemnify directors for some expenses, including attorneys' fees, judgments, fines and settlement amounts incurred by a director in any action or proceeding arising out of his or her service as one of the registrant's directors, or any of the registrant's subsidiaries or any other company or enterprise to which the person provides services at the registrant's request.

The registrant maintains a general liability insurance policy which covers certain liabilities of the directors and officers of the registrant arising out of claims based on acts or omissions in their capacities as directors or officers.

Item 7. Exemption from Registration Claimed.

Not applicable.

Item 8. Exhibits.

<u>Number</u>	<u>Description</u>
4.1(1)	Restated Certificate of Incorporation of the Registrant, as amended
4.2(2)	Amended and Restated By-Laws of the Registrant
5.1	Opinion of Wilmer Cutler Pickering Hale and Dorr LLP, counsel to the Registrant
23.1	Consent of Wilmer Cutler Pickering Hale and Dorr LLP (included in Exhibit 5.1)
23.2	Consent of Ernst & Young LLP
24.1	Power of attorney (included on the signature pages of this registration statement)
99.1(3)	Nonstatutory Stock Option Agreement, dated February 25, 2019, between the Registrant and Michael Mason
99.2(4)	Form of Nonstatutory Stock Option Agreement for Inducement Grants

(1)	Previously filed with the Securities and Exchange Commission as Exhibit 3.1 to the Registrant's Quarterly Report on Form 10-Q (File No. 001-36167) filed with the Securities and Exchange Commission on August 7, 2019 and incorporated herein by reference.
(2)	Previously filed with the Securities and Exchange Commission as Exhibit 3.2 to the Registrant's Current Report on Form 8-K (File No. 001-36167) filed with the Securities and Exchange Commission on November 18, 2013 and incorporated herein by reference.
(3)	Previously filed with the Securities and Exchange Commission as Exhibit 10.2 to the Registrant's Current Report on Form 8-K (File No. 001-36167) filed with the Securities and Exchange Commission on February 25, 2019 and incorporated herein by reference.
(4)	Previously filed with the Securities and Exchange Commission as Exhibit 10.3 to the Registrant's Quarterly Report on Form 10-Q (File No. 001-36167) filed with the Securities and Exchange Commission on May 10, 2018 and incorporated herein by reference.

Item 9. Undertakings.

1. The undersigned registrant hereby undertakes:

(1) To file, during any period in which offers or sales are being made, a post-effective amendment to this registration statement:

(i) To include any prospectus required by Section 10(a)(3) of the Securities Act;

(ii) To reflect in the prospectus any facts or events arising after the effective date of the registration statement (or the most recent post-effective amendment thereof) which, individually or in the aggregate, represent a fundamental change in the information set forth in the registration statement; and

(iii) To include any material information with respect to the plan of distribution not previously disclosed in the registration statement or any material change to such information in the registration statement;

provided, however, that paragraphs (i) and (ii) do not apply if the information required to be included in a post-effective amendment by those paragraphs is contained in reports filed with or furnished to the Commission by the registrant pursuant to Section 13 or Section 15(d) of the Exchange Act that are incorporated by reference in the registration statement.

(2) That, for the purpose of determining any liability under the Securities Act, each such post-effective amendment shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial *bona fide* offering thereof.

(3) To remove from registration by means of a post-effective amendment any of the securities being registered which remain unsold at the termination of the offering.

2. The undersigned registrant hereby undertakes that, for purposes of determining any liability under the Securities Act, each filing of the registrant's annual report pursuant to Section 13(a) or Section 15(d) of the Exchange Act that is incorporated by reference in the registration statement shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial *bona fide* offering thereof.

3. Insofar as indemnification for liabilities arising under the Securities Act may be permitted to directors, officers and controlling persons of the registrant pursuant to the foregoing provisions, or otherwise, the registrant has been advised that in the opinion of the Securities and Exchange Commission such indemnification is against public policy as expressed in the Securities Act and is, therefore, unenforceable. In the event that a claim for indemnification against such liabilities (other than the payment by the registrant of expenses incurred or paid by a director, officer or controlling person of the registrant in the successful defense of any action, suit or proceeding) is asserted by such director, officer or controlling person in connection with the securities being registered, the registrant will, unless in the opinion of its counsel the matter has been settled by controlling precedent, submit to a court of appropriate jurisdiction the question whether such indemnification by it is against public policy as expressed in the Securities Act and will be governed by the final adjudication of such issue.

SIGNATURES

Pursuant to the requirements of the Securities Act of 1933, the registrant certifies that it has reasonable grounds to believe that it meets all of the requirements for filing on Form S-8 and has duly caused this registration statement to be signed on its behalf by the undersigned, thereunto duly authorized, in the City of Newton, Commonwealth of Massachusetts, on this 7th day of August, 2019.

KARYOPHARM THERAPEUTICS INC.

By: /s/ Michael G. Kauffman
Name: Michael G. Kauffman, M.D., Ph.D.
Title: Chief Executive Officer

POWER OF ATTORNEY AND SIGNATURES

We, the undersigned officers and directors of Karyopharm Therapeutics Inc., hereby severally constitute and appoint Michael Kauffman, M.D., Ph.D., Michael Mason and Christopher B. Primiano, and each of them singly, our true and lawful attorneys with full power to them, and each of them singly, to sign for us and in our names in the capacities indicated below, the registration statement on Form S-8 filed herewith and any and all subsequent amendments to said registration statement, and generally to do all such things in our names and on our behalf in our capacities as officers and directors to enable Karyopharm Therapeutics Inc. to comply with the provisions of the Securities Act of 1933, as amended, and all requirements of the Securities and Exchange Commission, hereby ratifying and confirming our signatures as they may be signed by our said attorneys, or any of them, to said registration statement and any and all amendments thereto.

Pursuant to the requirements of the Securities Act of 1933, this registration statement has been signed by the following persons in the capacities and on the dates indicated.

<u>Signature</u>	<u>Title</u>	<u>Date</u>
<u>/s/ Michael G. Kauffman</u> Michael G. Kauffman, M.D., Ph.D.	Chief Executive Officer and Director (principal executive officer)	August 7, 2019
<u>/s/ Michael Mason</u> Michael Mason	Senior Vice President, Chief Financial Officer and Treasurer (principal financial and accounting officer)	August 7, 2019
<u>/s/ Garen G. Bohlin</u> Garen G. Bohlin	Director	August 7, 2019
<u>/s/ Mikael Dolsten</u> Mikael Dolsten, M.D., Ph.D.	Director	August 7, 2019
<u>/s/ J. Scott Garland</u> J. Scott Garland	Director	August 7, 2019
<u>/s/ Barry E. Greene</u> Barry E. Greene	Director	August 7, 2019
<u>/s/ Deepika R. Pakianathan</u> Deepika R. Pakianathan, Ph.D.	Director	August 7, 2019
<u>/s/ Mansoor Raza Mirza</u> Mansoor Raza Mirza, M.D.	Director	August 7, 2019

August 7, 2019

Karyopharm Therapeutics Inc.
85 Wells Avenue, 2nd Floor
Newton, MA 02459

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wilmerhale.com

Re: Inducement Stock Option Awards for the Period of August 2018 through July 2019

Ladies and Gentlemen:

We have assisted in the preparation of a Registration Statement on Form S-8 (the "Registration Statement") to be filed with the Securities and Exchange Commission (the "Commission") under the Securities Act of 1933, as amended (the "Securities Act"), relating to an aggregate of 1,415,000 shares of common stock, \$0.0001 par value per share (the "Shares"), of Karyopharm Therapeutics Inc., a Delaware corporation (the "Company"), issuable pursuant to nonstatutory stock option agreements providing for employee inducement grants between the Company and various employees, which were entered into in connection with the commencement of such employees' employment with the Company pursuant to Nasdaq Stock Market Rule 5635(c)(4) (the "Inducement Award Agreements").

We have examined the Certificate of Incorporation and By-Laws of the Company, each as amended and restated to date, and originals, or copies certified to our satisfaction, of all pertinent records of the meetings of the directors and stockholders of the Company, the Registration Statement and such other documents relating to the Company as we have deemed material for the purposes of this opinion.

In our examination of the foregoing documents, we have assumed the genuineness of all signatures, the authenticity of all documents submitted to us as originals, the conformity to original documents of all documents submitted to us as certified, photostatic or other copies, the authenticity of the originals of any such documents and the legal competence of all signatories to such documents.

We assume that the appropriate action will be taken, prior to the offer and sale of the Shares in accordance with the Inducement Award Agreements, to register and qualify the Shares for sale under all applicable state securities or "blue sky" laws.

We express no opinion herein as to the laws of any state or jurisdiction other than the state laws of The Commonwealth of Massachusetts and the General Corporation Law of the State of Delaware.

It is understood that this opinion is to be used only in connection with the offer and sale of the Shares while the Registration Statement is in effect.

Please note that we are opining only as to the matters expressly set forth herein, and no opinion should be inferred as to any other matters.

Wilmer Cutler Pickering Hale and Dorr LLP, 60 State Street, Boston, Massachusetts 02109

Beijing Berlin Boston Brussels Denver Frankfurt London Los Angeles New York Palo Alto San Francisco Washington

Karyopharm Therapeutics Inc.
August 7, 2019
Page 2

Based on the foregoing, we are of the opinion that the Shares have been duly authorized for issuance and, when the Shares are issued and paid for in accordance with the terms and conditions of the Inducement Award Agreements, the Shares will be validly issued, fully paid and nonassessable.

We hereby consent to the filing of this opinion with the Commission in connection with the Registration Statement in accordance with the requirements of Item 601(b)(5) of Regulation S-K under the Securities Act. In giving such consent, we do not hereby admit that we are in the category of persons whose consent is required under Section 7 of the Securities Act or the rules and regulations of the Commission.

Very truly yours,

WILMER CUTLER PICKERING
HALE AND DORR LLP

By: /s/ Jason L. Kropp
Jason L. Kropp, a Partner

Consent of Independent Registered Public Accounting Firm

We consent to the incorporation by reference in the Registration Statement (Form S-8) pertaining to Inducement Stock Option Awards (August 2018 – July 2019) of Karyopharm Therapeutics Inc. of our reports dated February 28, 2019, with respect to the consolidated financial statements of Karyopharm Therapeutics Inc., and the effectiveness of internal control over financial reporting of Karyopharm Therapeutics Inc., included in its Annual Report (Form 10-K) for the year ended December 31, 2018, filed with the Securities and Exchange Commission.

/s/ Ernst & Young LLP

Boston, Massachusetts

August 7, 2019