

**240.13d-102 Schedule 13G - Information to be included in statements filed pursuant to 240.13d-1(b), (c), and (d) and amendments thereto filed pursuant to 240.13d-2.**

Securities and Exchange Commission, Washington, D.C. 20549

Schedule 13G

Under the Securities Exchange Act of 1934

(Amendment No.)\*

(Name of Issuer)

Karyopharm Therapeutics Inc.

(Title of Class of Securities)

Common Stock, Par Value \$0.0001 Per Share

(CUSIP Number)

48576U106

(Date of Event Which Requires Filing of this Statement)

February 22, 2019

Check the appropriate box to designate the rule pursuant to which this Schedule is filed:

Rule 13d-1(b)

Rule 13d-1(c)

Rule 13d-1(d)

\*The remainder of this cover page shall be filled out for a reporting person's initial filing on this form with respect to the subject class of securities, and for any subsequent amendment containing information which would alter the disclosures provided in a prior cover page.

The information required in the remainder of this cover page shall not be deemed to be "filed" for the purpose of Section 18 of the Securities Exchange Act of 1934 ("Act") or otherwise subject to the liabilities of that section of the Act but shall be subject to all other provisions of the Act (however, see the Notes).

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CUSIP No. 48576U106	
(1) Names of reporting persons Point72 Asset Management, L.P.	
(2) Check the appropriate box if a member of a group	(a)
(see instructions)	(b)X
(3) SEC use only	
(4) Citizenship or place of organization Delaware	
Number of shares beneficially owned by each reporting person with:	
(5) Sole voting power 0	
(6) Shared voting power 3,408,141 (see Item 4)	
(7) Sole dispositive power 0	
(8) Shared dispositive power 3,408,141 (see Item 4)	
(9) Aggregate amount beneficially owned by each reporting person 3,408,141 (see Item 4)	
(10) Check if the aggregate amount in Row (9) excludes certain shares (see instructions)	
(11) Percent of class represented by amount in Row (9) 5.6% (see Item 4)	
(12) Type of reporting person (see instructions) PN	

CUSIP No. 48576U106	
(1) Names of reporting persons Point72 Capital Advisors, Inc.	
(2) Check the appropriate box if a member of a group	(a)
(see instructions)	(b)X
(3) SEC use only	
(4) Citizenship or place of organization Delaware	
Number of shares beneficially owned by each reporting person with:	
(5) Sole voting power 0	
(6) Shared voting power 3,408,141 (see Item 4)	
(7) Sole dispositive power 0	
(8) Shared dispositive power 3,408,141 (see Item 4)	
(9) Aggregate amount beneficially owned by each reporting person 3,408,141 (see Item 4)	
(10) Check if the aggregate amount in Row (9) excludes certain shares (see instructions)	
(11) Percent of class represented by amount in Row (9) 5.6% (see Item 4)	
(12) Type of reporting person (see instructions) CO	

CUSIP No. 48576U106	
(1) Names of reporting persons	Cubist Systematic Strategies, LLC
(2) Check the appropriate box if a member of a group	(a)
(see instructions)	(b)X
(3) SEC use only	
(4) Citizenship or place of organization	Delaware
Number of shares beneficially owned by each reporting person with:	
(5) Sole voting power	0
(6) Shared voting power	97,396 (see Item 4)
(7) Sole dispositive power	0
(8) Shared dispositive power	97,396 (see Item 4)
(9) Aggregate amount beneficially owned by each reporting person	97,396 (see Item 4)
(10) Check if the aggregate amount in Row (9) excludes certain shares (see instructions)	
(11) Percent of class represented by amount in Row (9)	0.2% (see Item 4)
(12) Type of reporting person (see instructions)	OO

CUSIP No. 48576U106	
(1) Names of reporting persons Point72 Hong Kong Limited	
(2) Check the appropriate box if a member of a group	(a)
(see instructions)	(b)X
(3) SEC use only	
(4) Citizenship or place of organization Hong Kong	
Number of shares beneficially owned by each reporting person with:	
(5) Sole voting power 0	
(6) Shared voting power 119 (see Item 4)	
(7) Sole dispositive power 0	
(8) Shared dispositive power 119 (see Item 4)	
(9) Aggregate amount beneficially owned by each reporting person 119 (see Item 4)	
(10) Check if the aggregate amount in Row (9) excludes certain shares (see instructions)	
(11) Percent of class represented by amount in Row (9) <0.1% (see Item 4)	
(12) Type of reporting person (see instructions) OO	

CUSIP No. 48576U106	
(1) Names of reporting persons	Steven A. Cohen
(2) Check the appropriate box if a member of a group	(a)
(see instructions)	(b)X
(3) SEC use only	
(4) Citizenship or place of organization	United States
Number of shares beneficially owned by each reporting person with:	
(5) Sole voting power	0
(6) Shared voting power	3,505,656 (see Item 4)
(7) Sole dispositive power	0
(8) Shared dispositive power	3,505,656 (see Item 4)
(9) Aggregate amount beneficially owned by each reporting person	3,505,656 (see Item 4)
(10) Check if the aggregate amount in Row (9) excludes certain shares (see instructions)	
(11) Percent of class represented by amount in Row (9)	5.8% (see Item 4)
(12) Type of reporting person (see instructions)	IN

Item 1(a) Name of issuer:

Karyopharm Therapeutics Inc.

Item 1(b) Address of issuer's principal executive offices:

85 Wells Avenue, 2nd Floor, Newton, MA 02459, United States

2(a) Name of person filing:

This statement is filed by: (i) Point72 Asset Management, L.P. ("Point72 Asset Management") with respect to shares of common stock, par value \$0.0001 per share ("Shares"), of the Issuer held by certain investment funds it manages; (ii) Point72 Capital Advisors, Inc. ("Point72 Capital Advisors Inc.") with respect to Shares held by certain investment funds managed by Point72 Asset Management; (iii) Cubist Systematic Strategies, LLC ("Cubist Systematic Strategies") with respect to Shares held by certain investment funds it manages; (iv) Point72 Hong Kong Limited ("Point72 Hong Kong") with respect to Shares held by certain investment funds it manages; and (v) Steven A. Cohen ("Mr. Cohen") with respect to Shares beneficially owned by Point72 Asset Management, Point72 Capital Advisors Inc., Cubist Systematic Strategies, and Point72 Hong Kong.

Point72 Asset Management, Point72 Capital Advisors Inc., Cubist Systematic Strategies, Point72 Hong Kong, and Mr. Cohen have entered into a Joint Filing Agreement, a copy of which is filed with this Schedule 13G as Exhibit 99.1, pursuant to which they have agreed to file this Schedule 13G jointly in accordance with the provisions of Rule 13d-1(k) of the Act.

2(b) Address or principal business office or, if none, residence:

The address of the principal business office of (i) Point72 Asset Management, Point72 Capital Advisors Inc., and Mr. Cohen is 72 Cummings Point Road, Stamford, CT 06902; (ii) Cubist Systematic Strategies is 330 Madison Avenue, New York, NY 10017; and (iii) Point72 Hong Kong is 12th Floor, Chater House, 8 Connaught Central, Hong Kong.

2(c) Citizenship:

Point72 Asset Management is a Delaware limited partnership. Point72 Capital Advisors Inc. is a Delaware corporation. Cubist Systematic Strategies is a Delaware limited liability company. Point72 Hong Kong is a Hong Kong limited liability company. Mr. Cohen is a United States citizen.

2(d) Title of class of securities:

Common Stock, Par Value \$0.0001 Per Share

2(e) CUSIP Number:

48576U106

*Item 3.*

Not applicable

*Item 4. Ownership*

As of the close of business on February 26, 2019:

1. Point72 Asset Management, L.P.

(a) Amount beneficially owned: 3,408,141

(b) Percent of class: 5.6%

(c) Number of shares as to which the person has:

(i) Sole power to vote or to direct the vote: 0

(ii) Shared power to vote or to direct the vote: 3,408,141

(iii) Sole power to dispose or to direct the disposition of: 0

(iv) Shared power to dispose or to direct the disposition of: 3,408,141

2. Point72 Capital Advisors, Inc.

(a) Amount beneficially owned: 3,408,141

(b) Percent of class: 5.6%

(c) Number of shares as to which the person has:

(i) Sole power to vote or to direct the vote: 0

(ii) Shared power to vote or to direct the vote: 3,408,141

(iii) Sole power to dispose or to direct the disposition of: 0

(iv) Shared power to dispose or to direct the disposition of: 3,408,141

3. Cubist Systematic Strategies, LLC

(a) Amount beneficially owned: 97,396

(b) Percent of class: 0.2%

(c) Number of shares as to which the person has:

(i) Sole power to vote or to direct the vote: 0

(ii) Shared power to vote or to direct the vote: 97,396

(iii) Sole power to dispose or to direct the disposition of: 0

(iv) Shared power to dispose or to direct the disposition of: 97,396

4. Point72 Hong Kong Limited

(a) Amount beneficially owned: 119

(b) Percent of class: <0.1%

(c) Number of shares as to which the person has:

(i) Sole power to vote or to direct the vote: 0

(ii) Shared power to vote or to direct the vote: 119

(iii) Sole power to dispose or to direct the disposition of: 0

(iv) Shared power to dispose or to direct the disposition of: 119

5. Steven A. Cohen

(a) Amount beneficially owned: 3,505,656

(b) Percent of class: 5.8%

(c) Number of shares as to which the person has:

(i) Sole power to vote or to direct the vote: 0

(ii) Shared power to vote or to direct the vote: 3,505,656

(iii) Sole power to dispose or to direct the disposition of: 0

(iv) Shared power to dispose or to direct the disposition of: 3,505,656

Point72 Asset Management, Point72 Capital Advisors Inc., Cubist Systematic Strategies, Point72 Hong Kong, and Mr. Cohen own directly no Shares. Pursuant to an investment management agreement, Point72 Asset Management maintains investment and voting power with respect to the securities held by certain investment funds it manages. Point72 Capital Advisors Inc. is the general partner of Point72 Asset Management. Pursuant to an investment management agreement, Cubist Systematic Strategies maintains investment and voting power with respect to the securities held by certain investment funds it manages. Pursuant to an investment management agreement, Point72 Hong Kong maintains investment and voting power with respect to the securities held by certain investment funds it manages. Mr. Cohen controls each of Point72 Asset Management, Point72 Capital Advisors Inc., Cubist Systematic Strategies, and Point72 Hong Kong. By reason of the provisions of Rule 13d-3 of the Securities Exchange Act of 1934, as amended, each of (i) Point72 Asset Management, Point72 Capital Advisors Inc., and Mr. Cohen may be deemed to beneficially own 3,408,141 Shares (constituting approximately 5.6% of the Shares outstanding); (ii) Cubist Systematic Strategies and Mr. Cohen may be deemed to beneficially own 97,396 Shares (constituting approximately 0.2% of the Shares outstanding); and (iii) Point72 Hong Kong and Mr. Cohen may be deemed to beneficially own 119 Shares (constituting <0.1% of the Shares outstanding). Each of Point72 Asset Management, Point72 Capital Advisors Inc., Cubist Systematic Strategies, Point72 Hong Kong, and Mr. Cohen disclaims beneficial ownership of any of the securities covered by this statement.

*Item 5. Ownership of 5 Percent or Less of a Class.*

If this statement is being filed to report the fact that as of the date hereof the reporting person has ceased to be the beneficial owner of more than 5 percent of the class of securities, check the following .

*Item 6. Ownership of More than 5 Percent on Behalf of Another Person.*

Not applicable

*Item 7. Identification and Classification of the Subsidiary Which Acquired the Security Being Reporting on by the Parent Holding Company or Control Person.*

Not applicable

*Item 8. Identification and Classification of Members of the Group*

Not applicable

*Item 9. Notice of Dissolution of Group.*

Not applicable

*Item 10. Certifications*

By signing below I certify that, to the best of his/her knowledge and belief, the securities referred to above were not acquired and are not held for the purpose of or with the effect of changing or influencing the control of the issuer of the securities and were not acquired and are not held in connection with or as a participant in any transaction having that purpose or effect.

*Signature.* After reasonable inquiry and to the best of my knowledge and belief, I certify that the information set forth in this statement is true, complete and correct.

Dated: February 27, 2019

POINT72 ASSET MANAGEMENT, L.P.

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

POINT72 CAPITAL ADVISORS, INC.

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

CUBIST SYSTEMATIC STRATEGIES, LLC

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

POINT72 HONG KONG LIMITED

By: /s/ Jason M. Colombo

Name: Jason M. Colombo

Title: Authorized Person

STEVEN A. COHEN

By: /s/ Jason M. Colombo

Name: Jason M. Colombo

Title: Authorized Person

JOINT FILING AGREEMENT

The undersigned hereby agree that the foregoing statement on Schedule 13G is filed on behalf of each of the undersigned in accordance with the provisions of Rule 13d-1(k) under the Securities Exchange Act of 1934, as amended, and that all subsequent amendments to this statement on Schedule 13G may be filed on behalf of each of the undersigned without the necessity of filing additional joint filing agreements.

Dated: February 27, 2019

POINT72 ASSET MANAGEMENT, L.P.

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

POINT72 CAPITAL ADVISORS, INC.

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

CUBIST SYSTEMATIC STRATEGIES, LLC

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

POINT72 HONG KONG LIMITED

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person

STEVEN A. COHEN

By: /s/ Jason M. Colombo

Name: Jason M. Colombo  
Title: Authorized Person