

---

---

**SECURITIES AND EXCHANGE COMMISSION**

Washington, D.C. 20549

---

**SCHEDULE 13G**

(Rule 13d-102)

**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**

(Amendment No. 6)\*

---

**KARYOPHARM THERAPEUTICS INC.**

(Name of Issuer)

**Common Stock par value \$0.0001 per share**  
(Title of Class of Securities)

**48576U106**  
(CUSIP Number)

**December 31, 2019**  
(Date of Event Which Requires Filing of this Statement)

---

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).

---

---

1.	Names of Reporting Persons <b>Michael G. Kauffman</b>	
2.	Check the Appropriate Box if a Member of a Group (See Instructions) (a) <input type="checkbox"/> (b) <input type="checkbox"/>	
3.	SEC Use Only	
4.	Citizenship or Place of Organization <b>United States of America</b>	
Number of Shares Beneficially Owned by Each Reporting Person With	5.	Sole Voting Power <b>1,814,674</b>
	6.	Shared Voting Power <b>2,106,760</b>
	7.	Sole Dispositive Power <b>1,814,674</b>
	8.	Shared Dispositive Power <b>2,106,760</b>
9.	Aggregate Amount Beneficially Owned by Each Reporting Person <b>3,921,434</b>	
10.	Check if the Aggregate Amount in Row (9) Excludes Certain Shares (See Instructions) <input type="checkbox"/>	
11.	Percent of Class Represented by Amount in Row (9) <b>5.77%(1)</b>	
12.	Type of Reporting Person (See Instructions) <b>IN</b>	

(1) Based upon 65,370,448 shares of Karyopharm Therapeutics Inc. Common Stock outstanding on December 31, 2019.

1.	Names of Reporting Persons	
	Sharon Shacham	
2.	Check the Appropriate Box if a Member of a Group (See Instructions) (a) <input type="checkbox"/> (b) <input type="checkbox"/>	
3.	SEC Use Only	
4.	Citizenship or Place of Organization	
	United States of America	
Number of Shares Beneficially Owned by Each Reporting Person With	5.	Sole Voting Power
		2,106,760
	6.	Shared Voting Power
		1,814,674
	7.	Sole Dispositive Power
		2,106,760
	8.	Shared Dispositive Power
		1,814,674
9.	Aggregate Amount Beneficially Owned by Each Reporting Person	
	3,921,434	
10.	Check if the Aggregate Amount in Row (9) Excludes Certain Shares (See Instructions)	
	<input type="checkbox"/>	
11.	Percent of Class Represented by Amount in Row (9)	
	5.77%(1)	
12.	Type of Reporting Person (See Instructions)	
	IN	

(1) Based upon 65,370,448 shares of Karyopharm Therapeutics Inc. Common Stock outstanding on December 31, 2019.

## Item 1.

- (a) Name of Issuer:  
Karyopharm Therapeutics Inc.
- (b) Address of Issuer's Principal Executive Offices:  
85 Wells Avenue, 2<sup>nd</sup> Floor, Newton, MA 02459.

## Item 2.

- (a) Name of Person Filing:  
This Schedule is filed by Michael G. Kauffman and Sharon Shacham (each a "Reporting Person" and collectively, "Reporting Persons").
- (b) Address of Principal Business Office or, if none, Residence:  
c/o Karyopharm Therapeutics Inc., 85 Wells Avenue, 2<sup>nd</sup> Floor, Newton, MA 02459.
- (c) Citizenship:  
See Row 4 of cover page for each Reporting Person.
- (d) Title of Class of Securities:  
Common stock par value \$0.0001 per share.
- (e) CUSIP Number:  
48576U106

## Item 3. If this statement is filed pursuant to §§240.13d-1(b) or 240.13d-2(b) or (c), check whether the person filing is a:

- (a)  Broker or dealer registered under section 15 of the Act (15 U.S.C. 78o).
- (b)  Bank as defined in section 3(a)(6) of the Act (15 U.S.C. 78c).
- (c)  Insurance company as defined in section 3(a)(19) of the Act (15 U.S.C. 78c).
- (d)  Investment company registered under section 8 of the Investment Company Act of 1940 (15 U.S.C. 80a-8).
- (e)  An investment adviser in accordance with §240.13d-1(b)(1)(ii)(E);
- (f)  An employee benefit plan or endowment fund in accordance with §240.13d-1(b)(1)(ii)(F);
- (g)  A parent holding company or control person in accordance with §240.13d-1(b)(1)(ii)(G);
- (h)  A savings associations as defined in Section 3(b) of the Federal Deposit Insurance Act (12 U.S.C. 1813);
- (i)  A church plan that is excluded from the definition of an investment company under section 3(c)(14) of the Investment Company Act of 1940 (15 U.S.C. 80a-3);
- (j)  A non-U.S. institution in accordance with §240.13d-1(b)(1)(ii)(J).
- (k)  Group, in accordance with §240.13d-1(b)(1)(ii)(K).

If filing as a non-U.S. institution in accordance with §240.13d-1(b)(1)(ii)(J), please specify the type of institution:

**Item 4. Ownership**

Provide the following information regarding the aggregate number and percentage of the class of securities of the issuer identified in Item 1.

(a) Amount beneficially owned:

For each Reporting Person, 3,921,434 shares of Common Stock, consisting of (a) 542,139 shares of Common Stock held by Dr. Kauffman, (b) 21,800 restricted stock units held by Dr. Kauffman that will vest within 60 days after December 31, 2019, (c) 1,250,735 shares of Common Stock underlying options held by Dr. Kauffman that are exercisable as of December 31, 2019 or will become exercisable within 60 days after such date, (d) 743,506 shares of Common Stock held by Dr. Shacham, (e) 21,800 restricted stock units held by Dr. Shacham that will vest within 60 days after December 31, 2019 and (f) 1,341,454 shares of Common Stock underlying options held by Dr. Shacham that are exercisable as of December 31, 2019 or will become exercisable within 60 days after such date.

(b) Percent of class:

See Row 11 of cover page for each Reporting Person.

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

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

See Row 5 of cover page for each Reporting Person.

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

See Row 6 of cover page for each Reporting Person.

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

See Row 7 of cover page for each Reporting Person.

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

See Row 8 of cover page for each Reporting Person.

**Item 5. Ownership of Five 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 five percent of the class of securities, check the following .

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

Not applicable.

**Item 7. Identification and Classification of the Subsidiary Which Acquired the Security Being Reported 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. Certification

Not applicable.

**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.

February 13, 2020

\_\_\_\_\_  
Date

/s/ Michael G. Kauffman

\_\_\_\_\_  
Signature

Michael G. Kauffman, M.D., Ph.D.

\_\_\_\_\_  
Name

February 13, 2020

\_\_\_\_\_  
Date

/s/ Sharon Shacham

\_\_\_\_\_  
Signature

Sharon Shacham, Ph.D., M.B.A.

\_\_\_\_\_  
Name

Exhibit Index

- Exhibit 1. Joint Filing Agreement, dated as of February 14, 2014, by and between the Reporting Persons (incorporated by reference to Exhibit 1 to the Schedule 13G filed by the Reporting Persons with the Securities and Exchange Commission on February 14, 2014).