

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

Schedule 13G/A

(Amendment No. 1)

Under the Securities Exchange Act of 1934*

ZIOPHARM Oncology Inc

(Name of Issuer)

Common Stock

(Title of Class of Securities)

98973P101

(CUSIP Number)

December 31, 2010

(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).

Persons who respond to the collection of information contained in this form are not required to respond unless the form displays a currently valid OMB control number.

1. NAMES OF REPORTING PERSONS

Great Point Partners, LLC

I.R.S. IDENTIFICATION NO. OF ABOVE PERSON (ENTITIES ONLY):

37-1475292

2. CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP(a) (b) **3. SEC USE ONLY****4. CITIZENSHIP OR PLACE OF ORGANIZATION**

USA

NUMBER OF SHARES	5. SOLE VOTING POWER	0
BENEFICIALLY	6. SHARED VOTING POWER	806,500 ¹
OWNED BY EACH	7. SOLE DISPOSITIVE POWER	0
REPORTING PERSON	8. SHARED DISPOSITIVE POWER	806,500 ¹
WITH		

9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON806,500¹**10. CHECK BOX IF THE AGGREGATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHARES (See Instructions)** **11. PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW (9)**1.63%²**12. TYPE OF REPORTING PERSON (See Instructions)**

OO

¹ Consists of 421,534 shares of Common Stock underlying a warrant held by Biomedical Value Fund, LP ("BVF"), 274,210 shares of Common Stock underlying a warrant held by Biomedical Offshore Value Fund, Ltd. ("BOVF") and 110,756 shares of Common Stock underlying a warrant held by Biomedical Institutional Value Fund, LP. ("BIVF").

² Based on a total of 48,557,678 shares of Common Stock outstanding as reported in the Issuer's Form 10-Q filed on November 4, 2010.

1. NAMES OF REPORTING PERSONS

Dr. Jeffrey R. Jay, M.D.

I.R.S. IDENTIFICATION NO. OF ABOVE PERSON (ENTITIES ONLY):

2. CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP(a) (b) **3. SEC USE ONLY****4. CITIZENSHIP OR PLACE OF ORGANIZATION**

USA

NUMBER OF SHARES	5. SOLE VOTING POWER	0
BENEFICIALLY	6. SHARED VOTING POWER	806,500 ¹
OWNED BY EACH	7. SOLE DISPOSITIVE POWER	0
REPORTING PERSON	8. SHARED DISPOSITIVE POWER	806,500 ¹
WITH		

9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON806,500¹**10. CHECK BOX IF THE AGGREGATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHARES (See Instructions)** **11. PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW (9)**1.63%²**12. TYPE OF REPORTING PERSON (See Instructions)**

IN

1. NAMES OF REPORTING PERSONS

Mr. David Kroin

I.R.S. IDENTIFICATION NO. OF ABOVE PERSON (ENTITIES ONLY):

2. CHECK THE APPROPRIATE BOX IF A MEMBER OF A GROUP(a) (b) **3. SEC USE ONLY****4. CITIZENSHIP OR PLACE OF ORGANIZATION**

USA

NUMBER OF SHARES	5. SOLE VOTING POWER	0
BENEFICIALLY	6. SHARED VOTING POWER	806,500 ¹
OWNED BY EACH	7. SOLE DISPOSITIVE POWER	0
REPORTING PERSON	8. SHARED DISPOSITIVE POWER	806,500 ¹
WITH		

9. AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON806,500¹**10. CHECK BOX IF THE AGGREGATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHARES (See Instructions)** **11. PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW (9)**1.63%²**12. TYPE OF REPORTING PERSON (See Instructions)**

IN

Item 1.

- (a) Name of Issuer
ZIOPHARM Oncology Inc.
- (b) Address of Issuer's Principal Executive Offices
1180 Avenue of the Americas, 19th Floor, New York, NY 10036

Item 2.

- (a) Name of Person Filing
Great Point Partners, LLC
Dr. Jeffrey R. Jay, M.D.
Mr. David Kroin

The Reporting Persons have entered into a Joint Filing Agreement, dated February 14, 2011, a copy of which is filed with this Schedule 13G/A as Exhibit A, pursuant to which the Reporting Persons have agreed to file this statement jointly in accordance with the provisions of Rule 13d-1(k)(1) under the Act.

- (b) Address of Principal Business Office, or if none, Residence
The address of the principal business office of each of the Reporting Persons is
165 Mason Street, 3rd Floor
Greenwich, CT 06830

- (c) Citizenship

Great Point Partners, LLC is a limited liability company organized under the laws of the State of Delaware. Dr. Jeffrey R. Jay, M.D. is a citizen of the United States. Mr. David Kroin is a citizen of the United States.

- (d) Title of Class of Securities

Common Stock

- (e) CUSIP Number

98973P101

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:

Not Applicable.

- (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) o An employee benefit plan or endowment fund in accordance with §240.13d-1(b)(1)(ii)(F).
- (g) o A parent holding company or control person in accordance with §240.13d-1(b)(1)(ii)(G).
- (h) o A savings associations as defined in Section 3(b) of the Federal Deposit Insurance Act (12 U.S.C. 1813).
- (i) o 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) o Group, in accordance with §240.13d-1(b)(1)(ii)(J).

Item 4. Ownership

Biomedical Value Fund, L.P. (“BVF”) is the owner of a warrant to purchase 421,534 shares of Common Stock of the Issuer (the “BVF Shares”). Great Point Partners, LLC (“Great Point”) is the investment manager of BVF, and by virtue of such status may be deemed to be the beneficial owner of the BVF Shares. Each of Dr. Jeffrey R. Jay, M.D. (“Dr. Jay”), as senior managing member of Great Point, and Mr. David Kroin (“Mr. Kroin”), as special managing member of Great Point, has voting and investment power with respect to the BVF Shares, and therefore may be deemed to be the beneficial owner of the BVF Shares.

Biomedical Offshore Value Fund, Ltd. (“BOVF”) is the owner of a warrant to purchase 274,210 shares of Common Stock of the Issuer (the “BOVF Shares”). Great Point is the investment manager of BOVF, and by virtue of such status may be deemed to be the beneficial owner of the BOVF Shares. Each of Dr. Jay, as senior managing member of Great Point, and Mr. Kroin, as special managing member of Great Point, has voting and investment power with respect to the BOVF Shares, and therefore may be deemed to be the beneficial owner of the BOVF Shares.

Biomedical Institutional Value Fund, LP (“BIVF”) is the owner of a warrant to purchase 110,756 shares of Common Stock of the Issuer (the “BIVF Shares”). Great Point is the investment manager of BIVF, and by virtue of such status may be deemed to be the beneficial owner of the BIVF Shares. Each of Dr. Jay, as senior managing member of Great Point, and Mr. Kroin, as special managing member of Great Point, has voting and investment power with respect to the BIVF Shares, and therefore may be deemed to be the beneficial owner of the BIVF Shares.

Notwithstanding the above, Great Point, Dr. Jay and Mr. Kroin disclaim beneficial ownership of the BVF Shares, BOVF Shares, and BIVF Shares, except to the extent of their respective pecuniary interests.

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

1. Great Point Partners, LLC

- (a) Amount beneficially owned: 806,500³

³

Consists of 421,534 shares of Common Stock underlying a warrant held by Biomedical Value Fund, LP (“BVF”), 274,210 shares of Common Stock underlying a warrant held by Biomedical Offshore Value Fund, Ltd. (“BOVF”) and 110,756 shares of Common Stock underlying a warrant held by Biomedical Institutional Value Fund, LP. (“BIVF”).

- (b) Percent of class: 1.63%⁴
- (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: 806,500³
 - (iii) Sole power to dispose or to direct the disposition of: - 0 -.
 - (iv) Shared power to dispose or to direct the disposition of: 806,500³

2. **Dr. Jeffrey R. Jay, M.D.**

- (a) Amount beneficially owned: 806,500³
- (b) Percent of class: 1.63%⁴
- (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: 806,500³
 - (iii) Sole power to dispose or to direct the disposition of: 0.
 - (iv) Shared power to dispose or to direct the disposition of: 806,500³

3. **Mr. David Kroin**

- (a) Amount beneficially owned: 806,500³
- (b) Percent of class: 1.63%⁴
- (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: 806,500³
 - (iii) Sole power to dispose or to direct the disposition of: 0.
 - (iv) Shared power to dispose or to direct the disposition of: 806,500³

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 each of the Reporting Persons has ceased to be the beneficial owner of more than five percent of the class of securities, check the following [X].

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

See Item 4.

⁴ Based on a total of 48,557,678 shares of Common Stock outstanding as reported in the Issuer's Form 10-Q filed on November 4, 2010.

Item 7. Identification and Classification of the Subsidiary Which Acquired the Security Being Reported on by the Parent Holding Company

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

By signing below I certify that, to the best of my 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 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.

Date: February 14, 2011

GREAT POINT PARTNERS, LLC

By: /s/ Dr. Jeffrey R. Jay, M.D.

Dr. Jeffrey R. Jay, M.D.,

as senior managing member

/s/ Dr. Jeffrey R. Jay, M.D.

DR. JEFFREY R. JAY, M.D.

/s/ Mr. David Kroin

Mr. David Kroin

AGREEMENT REGARDING THE JOINT FILING OF SCHEDULE 13G/A

The undersigned hereby agree as follows:

(i) Each of them is individually eligible to use the Schedule 13G/A to which this Exhibit is attached, and such Schedule 13G/A is filed on behalf of each of them; and

(ii) Each of them is responsible for the timely filing of such Schedule 13G/A and any amendments thereto, and for the completeness and accuracy of the information concerning such person contained therein; but none of them is responsible for the completeness or accuracy of the information concerning the other persons making the filing, unless such person knows or has reason to believe that such information is inaccurate.

Date: February 14, 2011

GREAT POINT PARTNERS, LLC

By: /s/ Dr. Jeffrey R. Jay, M.D.
Dr. Jeffrey R. Jay, M.D.,
as senior managing member

/s/ Dr. Jeffrey R. Jay, M.D.
DR. JEFFREY R. JAY, M.D.

/s/ Mr. David Kroin
Mr. David Kroin
