

| OMB APPROVAL | |
|---|-----------|
| OMB Number: | 3235-0287 |
| Estimated average burden hours per response: | 0.5 |

Check this box if no longer subject to Section 16. Form 4 or Form 5 obligations may continue. See Instruction 1(b).

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

Filed pursuant to Section 16(a) of the Securities Exchange Act of 1934 or Section 30(h) of the Investment Company Act of 1940

| | | |
|--|---|--|
| 1. Name and Address of Reporting Person* <u>Chione Ltd</u> (Last) (First) (Middle) SIMOU MENARDOU 5, KIFISIA COURT, OFFICE 225 (Street) 6015 LARNACA G4 CY (City) (State) (Zip) | 2. Issuer Name and Ticker or Trading Symbol <u>Karyopharm Therapeutics Inc. [KPTI]</u> | 5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director <input checked="" type="checkbox"/> 10% Owner Officer (give title below) Other (specify below) |
| | 3. Date of Earliest Transaction (Month/Day/Year) <u>02/25/2019</u> | |
| 4. If Amendment, Date of Original Filed (Month/Day/Year) | | |

Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned

| 1. Title of Security (Instr. 3) | 2. Transaction Date (Month/Day/Year) | 2A. Deemed Execution Date, if any (Month/Day/Year) | 3. Transaction Code (Instr. 8) | | 4. Securities Acquired (A) or Disposed Of (D) (Instr. 3, 4 and 5) | | | 5. Amount of Securities Beneficially Owned Following Reported Transaction(s) (Instr. 3 and 4) | 6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4) | 7. Nature of Indirect Beneficial Ownership (Instr. 4) |
|---------------------------------|--------------------------------------|--|--------------------------------|---|---|------------|----------------------------|---|--|---|
| | | | Code | V | Amount | (A) or (D) | Price | | | |
| COMMON STOCK | 02/25/2019 | | S | | 800,000 | D | \$4.9936 ⁽¹⁾⁽²⁾ | 6,831,267 ⁽³⁾⁽⁴⁾⁽⁵⁾ | D | |
| COMMON STOCK | 02/26/2019 | | S | | 123,400 | D | \$5.0017 ⁽¹⁾⁽²⁾ | 6,707,867 ⁽³⁾⁽⁴⁾⁽⁵⁾ | D | |
| COMMON STOCK | 02/27/2019 | | S | | 558,243 | D | \$4.5067 ⁽¹⁾⁽²⁾ | 6,149,624 ⁽³⁾⁽⁴⁾⁽⁵⁾ | D | |

Table II - Derivative Securities Acquired, Disposed of, or Beneficially Owned (e.g., puts, calls, warrants, options, convertible securities)

| 1. Title of Derivative Security (Instr. 3) | 2. Conversion or Exercise Price of Derivative Security | 3. Transaction Date (Month/Day/Year) | 3A. Deemed Execution Date, if any (Month/Day/Year) | 4. Transaction Code (Instr. 8) | | 5. Number of Derivative Securities Acquired (A) or Disposed of (D) (Instr. 3, 4 and 5) | | 6. Date Exercisable and Expiration Date (Month/Day/Year) | | 7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4) | 8. Price of Derivative Security (Instr. 5) | 9. Number of derivative Securities Beneficially Owned Following Reported Transaction(s) (Instr. 4) | 10. Ownership Form: Direct (D) or Indirect (I) (Instr. 4) | 11. Nature of Indirect Beneficial Ownership (Instr. 4) |
|--|--|--------------------------------------|--|--------------------------------|---|--|-----|--|-----------------|---|--|--|---|--|
| | | | | Code | V | (A) | (D) | Date Exercisable | Expiration Date | | | | | |

| | | |
|--|--|--|
| 1. Name and Address of Reporting Person* <u>Chione Ltd</u> (Last) (First) (Middle) SIMOU MENARDOU 5, KIFISIA COURT, OFFICE 225 (Street) 6015 LARNACA G4 CY (City) (State) (Zip) | | |
|--|--|--|

1. Name and Address of Reporting Person*

[Czernik Marcin](#)

(Last) (First) (Middle)

SIMOU MENARDOU 5,
KIFISIA COURT, OFFICE 225

(Street)

6015 LARNACA G4 00000

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[Hadjimichael Andreas](#)

(Last) (First) (Middle)

SIMOU MENARDOU 5,
KIFISIA COURT, OFFICE 225

(Street)

6015 LARNACA G4 00000

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[Hadjimichael George](#)

(Last) (First) (Middle)

SIMOU MENARDOU 5,
KIFISIA COURT, OFFICE 225

(Street)

6015 LARNACA G4 00000

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[Smolokowski Wiaczeslaw](#)

(Last) (First) (Middle)

CHALET LENOTCHKA CH.DE BARNOUD
1885 CHESIERES

(Street)

SWITZERLAND G4 00000

(City) (State) (Zip)

Explanation of Responses:

- The prices reported in Column 4 are weighted average prices. The shares were sold on 02/25/2019 at prices ranging from \$4.80 to \$5.31, inclusive, on 02/26/2019 at prices ranging from \$5.00 to \$5.08, inclusive, and on 02/27/2019 at prices ranging from \$4.40 to \$4.675, inclusive.
- The reporting persons undertake to provide to the issuer, any security holder of the issuer, or the staff of the Securities and Exchange Commission, upon request, full information regarding the number of shares sold at each separate price within each of the ranges set forth in footnote 1 above.
- Shares of Common Stock are owned directly by Chione Limited ("Chione"). Chione's directors, Marcin Czernik, Andreas Hadjimichael and George Hadjimichael, may be deemed to share voting and investment power and beneficial ownership of the shares of Common Stock directly owned by Chione. Wiaczeslaw Smolokowski, the sole shareholder of Chione, may also be deemed to share voting and investment power and beneficial ownership of the shares of Common Stock directly owned by Chione.
- Each reporting person states that neither the filing of this Form 4 nor anything herein shall be deemed an admission that such person or any other person is, for purposes of Section 16 of the Securities Exchange Act of 1934, as amended (the "Act"), or otherwise, the beneficial owner of any securities covered by this Form 4.
- Each reporting person may be deemed to be a member of a group with respect to the issuer or securities of the issuer for purposes of Section 13(d) or 13(g) of the Act. Each reporting person declares that neither the filing of this Form 4 nor anything herein shall be construed as an admission that such person or any other person is, for the purposes of Section 13(d) or 13(g) of the Act or any other purpose, a member of a group with respect to the issuer or securities of the issuer.

Remarks:

Exhibit Index Exhibit 24.1 - Power of Attorney, dated February 8, 2019, made by Marcin Czernik and Chione Limited in favor of Simon Prisk (previously filed) Exhibit 24.2 - Power of Attorney, dated February 8, 2019, made by Andreas Hadjimichael and Chione Limited in favor of Simon Prisk (previously filed) Exhibit 24.3 - Power of Attorney, dated February 8, 2019, made by George Hadjimichael and Chione Limited in favor of Simon Prisk (previously filed) Exhibit 24.4 - Power of Attorney, dated February 8, 2019, made by Wiaczeslaw Smolokowski in favor of Simon Prisk Exhibit (previously filed) 99.1 - Joint Filer Information (filed herewith) Exhibit 99.2 - Joint Filing Agreement (filed herewith). All previously filed exhibits were filed with the Schedule 13G/A of the reporting persons, filed on February 13, 2019.

/s/ Chione Limited, by /s/
Simon Prisk, as attorney-in
fact by power of attorney 02/27/2018
/s/ Marcin Czernik, by /s/
Simon Prisk, as attorney-in
fact by power of attorney 02/27/2018
/s/ Andreas Hadjimichael, by
/s/ Simon Prisk, as attorney-in
fact by power of attorney 02/27/2018
/s/ George Hadjimichael, by
/s/ Simon Prisk, as attorney-in
fact by power of attorney 02/27/2018
/s/ Wlaczslaw Smolokowski,
by /s/ Simon Prisk, as
attorney-in fact by power of
attorney 02/27/2018

** Signature of Reporting Person Date

Reminder: Report on a separate line for each class of securities beneficially owned directly or indirectly.

* If the form is filed by more than one reporting person, see Instruction 4 (b)(v).

** Intentional misstatements or omissions of facts constitute Federal Criminal Violations See 18 U.S.C. 1001 and 15 U.S.C. 78ff(a).

Note: File three copies of this Form, one of which must be manually signed. If space is insufficient, see Instruction 6 for procedure.

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 Number.

Joint Filer Information

(Other Reporting Persons)

| | |
|---|--|
| Name: | Marcin Czemik |
| Address: | Simou Menardou 5 Kifisia Court, Office 225 6015 Lamaca, Cyprus |
| Designated Filer: | Chione Limited |
| Issuer & Ticker Symbol: | Karyopharm Therapeutics Inc. [KPTI] |
| Date of Earliest Transaction Required to Be Reported: | 02/25/2019 |
| Relationship of Reporting Persons to Issuer: | 10% Owner |
| If Amendment, Date Original Filed (Month/Day/Year): | Not Applicable |
| Individual or Joint/Group Filing: | Form filed by More than One Reporting Person |
| Signature and Date: | Marcin Czemik, by /s/ Simon Prisk, as attorney-in fact by power of attorney 02/27/2019 |
| Name: | Andreas Hadjimichael |
| Address: | Simou Menardou 5 Kifisia Court, Office 225 6015 Lamaca, Cyprus |
| Designated Filer: | Chione Limited |
| Issuer & Ticker Symbol: | Karyopharm Therapeutics Inc. [KPTI] |
| Date of Earliest Transaction Required to Be Reported: | 02/25/2019 |
| Relationship of Reporting Persons to Issuer: | 10% Owner |
| If Amendment, Date Original Filed (Month/Day/Year): | Not Applicable |
| Individual or Joint/Group Filing: | Form filed by More than One Reporting Person |
| Signature and Date: | Andreas Hadjimichael, by /s/ Simon Prisk, as attorney-in fact by power of attorney 02/27/2019 |

Name: George Hadjimichael
Address: Simou Menardou 5
Kifisia Court, Office 225
6015 Lamaca, Cyprus

Designated Filer: Chione Limited
Issuer & Ticker Symbol: Karyopharm Therapeutics Inc. [KPTI]
Date of Earliest Transaction Required to Be Reported: 02/25/2019
Relationship of Reporting Persons to Issuer: 10% Owner
If Amendment, Date Original Filed (Month/Day/Year): Not Applicable
Individual or Joint/Group Filing: Form filed by More than One Reporting Person
Signature and Date: George Hadjimichael, by /s/ Simon Prisk, as attorney-in fact by power of attorney 02/27/2019

Name: Wiaczeslaw Smolokowski
Address: Chalet Lenotchka
Ch.de Bamoud
1885 Chesieres
Switzerland

Designated Filer: Chione Limited
Issuer & Ticker Symbol: Karyopharm Therapeutics Inc. [KPTI]
Date of Earliest Transaction Required to Be Reported: 02/25/2019
Relationship of Reporting Persons to Issuer: 10% Owner
If Amendment, Date Original Filed (Month/Day/Year): Not Applicable
Individual or Joint/Group Filing: Form filed by More than One Reporting Person
Signature and Date: Wiaczeslaw Smolokowski, by /s/ Simon Prisk, as attorney-in fact by power of attorney 02/27/2019

JOINT FILING AGREEMENT

Joint Filing Agreement, dated February 27, 2019, by and among the undersigned, pursuant to and in accordance with the Securities Exchange Act of 1934, as amended (the "Exchange Act"), and the rules and regulations thereunder, each party hereto hereby agrees to the joint filing, on behalf of each of them, of any filing required by such party under Section 16 of the Exchange Act or any rule or regulation thereunder (including any amendment, restatement, supplement, and/or exhibit thereto) with the Securities and Exchange Commission (and, if such security is registered on a national securities exchange, also with the exchange), and further agrees to the filing, furnishing, and/or incorporation by reference of this agreement as an exhibit thereto. This agreement shall remain in full force and effect until revoked by any party hereto in a signed writing provided to each other party hereto, and then only with respect to such revoking party.

IN WITNESS WHEREOF, each party hereto, being duly authorized, has caused this Agreement to be executed and effective as of the date first written above.

Chione Limited, by /s/ Simon Prisk, as attorney-in fact by power of attorney dated 02/08/2019

Marcin Czernik, by /s/ Simon Prisk, as attorney-in fact by power of attorney dated 02/08/2019

Andreas Hadjimichael, by /s/ Simon Prisk, as attorney-in fact, by power of attorney dated 02/08/2019

George Hadjimichael, by /s/ Simon Prisk, as attorney-in fact, by power of attorney dated 02/08/2019

Wiaczeslaw Smolokowski, by /s/ Simon Prisk, as attorney-in fact, by power of attorney dated 02/08/2019