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

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

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

#### STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP OF SECURITIES

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<br>FROST PHILLIP MD ET AI | 2. Issuer Name <b>a</b><br>Opko Health, In |   |                                  | ading Sy | mbol                               | 5. Relationship of Reporting Person(s) to Issuer<br>(Check all applicable)<br>-X_Director<br>X_Officer (give title below)<br>CEO & Chairman |   |  |  |                                |
|--|--|---|----------------------------------|----------|------------------------------------|---|---|--|--|--------------------------------|
| (Last) (First)<br>OPKO HEALTH, INC., 4400<br>BLVD.         | 3. Date of Earliest 09/13/2016             | Transactic  | on (M                            | onth/Day | /Year)                             |   |   |  |  |                                |
| (Street)<br>MIAMI, FL 33137                                |  | 4. If Amendment,  | Date Origi                       | nal F    | iled(Montl                         | h/Day/Ye  | 6. Individual or Joint/Group Filir<br>Form filed by One Reporting Person<br>_X_Form filed by More than One Reportin |  | able Line)   |                                |
| (City) (State)   | (Zip)                                      | Tab   | le I - Non                       | -Deri    | vative So                          | ecuritie  | es Acqui  | ired, Disposed of, or Beneficially   | y Owned  |                                |
| 1.Title of Security<br>(Instr. 3)                          | Date<br>(Month/Day/Year)                   | 2A. Deemed<br>Execution Date, if<br>any<br>(Month/Day/Year) | 3. Transac<br>Code<br>(Instr. 8) | ction    | 4. Secur<br>(A) or D<br>(Instr. 3) | oisposed  | 1 of (D)<br>5)  | 5. Amount of Securities<br>Beneficially Owned Following<br>Reported Transaction(s)<br>(Instr. 3 and 4) | 6.<br>Ownership<br>Form:<br>Direct (D)<br>or Indirect<br>(I) | Beneficia<br>Ownershi          |
| Common Stock   |  |   | Code                             | V        | Amount                             | · ·   | Price   | 3,068,951  | (I)<br>(Instr. 4)  |                                |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 1,000                              | А   | \$<br>9.78  | 160,136,143  | I  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 1,000                              | А   | \$ 9.8  | 160,137,143  | I  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 500                                | А   | \$<br>9.82  | 160,137,643  | Ι  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 500                                | А   | \$<br>9.83  | 160,138,143  | I  | See<br>Footnote<br>( <u>1)</u> |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 500                                | А   | \$<br>9.93  | 160,138,643  | I  | See<br>Footnote<br>( <u>1)</u> |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 500                                | А   | \$<br>9.95  | 160,139,143  | I  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 500                                | А   | \$<br>9.97  | 160,139,643  | I  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 1,000                              | А   | \$ 10   | 160,140,643  | I  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 500                                | А   | \$<br>10.02   | 160,141,143  | I  | See<br>Footnote<br>( <u>1)</u> |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 2,000                              | А   | \$<br>10.04   | 160,143,143  | Ι  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 1,000                              | А   | \$<br>10.05   | 160,144,143  | I  | See<br>Footnote<br>(1)         |
| Common Stock   | 09/13/2016                                 |   | Р                                |          | 1,000                              | А   | \$<br>10.06   | 160,145,143  | Ι  | See<br>Footnote<br>(1)         |
| Common Stock   |  |   |                                  |          |                                    |   |   | 20,091,062   | I  | See<br>Footnote<br>(2)         |

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

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SEC 1474 (9-02)

| Table II - Derivative Securities Acquired, | Disposed of, or Beneficially Owned |
|--|------------------------------------|
|--|------------------------------------|

| _ | (e.g., puts, calls, warrants, options, convertible securities) |             |                  |                    |            |      |         |       |                     |                    |        |          |             |                |             |             |
|---|--|-------------|------------------|--------------------|------------|------|---------|-------|---------------------|--------------------|--------|----------|-------------|----------------|-------------|-------------|
| 1 | . Title of   | 2.          | 3. Transaction   | 3A. Deemed         | 4.         | 4    | 5. Nu   | mber  | 6. Date Exer        | cisable            | 7. Tit | tle and  | 8. Price of | 9. Number of   | 10.         | 11. Nature  |
| I | Derivative   | Conversion  | Date             | Execution Date, if | Transactio | on ( | of      |       | and Expirati        | on Date            | Amo    | unt of   | Derivative  | Derivative     | Ownership   | of Indirect |
| 5 | Security   | or Exercise | (Month/Day/Year) | any                | Code       | ]    | Deriv   | ative | re (Month/Day/Year) |                    | Unde   | erlying  | Security    | Securities     | Form of     | Beneficial  |
| ( | Instr. 3)  | Price of    |                  | (Month/Day/Year)   | (Instr. 8) |      | Secur   | ities |                     |                    | Secu   | rities   | (Instr. 5)  | Beneficially   | Derivative  | Ownership   |
|   |  | Derivative  |                  |                    |            | 4    | Acqui   | ired  |                     |                    | (Instr | r. 3 and |             | Owned          | Security:   | (Instr. 4)  |
|   |  | Security    |                  |                    |            | (    | (A) oi  | r     |                     |                    | 4)     |          |             | Following      | Direct (D)  |             |
|   |  |             |                  |                    |            | ]    | Dispo   | sed   |                     |                    |        |          |             | Reported       | or Indirect |             |
|   |  |             |                  |                    |            | 0    | of (D)  | )     |                     |                    |        |          |             | Transaction(s) | (I)         |             |
|   |  |             |                  |                    |            |      | (Instr. | · · · |                     |                    |        |          |             | (Instr. 4)     | (Instr. 4)  |             |
|   |  |             |                  |                    |            | 4    | 4, and  | 15)   |                     |                    |        |          |             |                |             |             |
|   |  |             |                  |                    |            |      |         |       |                     |                    |        | Amount   |             |                |             |             |
|   |  |             |                  |                    |            |      |         |       | Dit                 | E intim            |        | or       |             |                |             |             |
|   |  |             |                  |                    |            |      |         |       | Date<br>Exercisable | Expiration<br>Date | Title  | Number   |             |                |             |             |
|   |  |             |                  |                    |            |      |         |       | Exercisable         | Date               |        | of       |             |                |             |             |
|   |  |             |                  |                    | Code V     | V    | (A)     | (D)   |                     |                    |        | Shares   |             |                |             |             |

## **Reporting Owners**

| Dama         | uting Original Names / Addison  | Relationships |           |                |       |  |  |  |  |
|--------------|---|---------------|-----------|----------------|-------|--|--|--|--|
| керо         | Reporting Owner Name / Address  |               | 10% Owner | Officer        | Other |  |  |  |  |
| OPK0<br>4400 | ST PHILLIP MD ET AL<br>D HEALTH, INC.<br>BISCAYNE BLVD.<br>MI, FL 33137 | х             | Х         | CEO & Chairman |       |  |  |  |  |
| 4400         | Gamma Investments Trust<br>BISCAYNE BLVD.<br>MI, FL 33137               |               | Х         |                |       |  |  |  |  |

# Signatures

| Phillip Frost, M.D., Individually and as Trustee | 09/14/2016 |
|--|------------|
| **Signature of Reporting Person                  | Date       |

# **Explanation of Responses:**

- \* 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).
- The securities are held by Frost Gamma Investments Trust, of which Phillip Frost M.D., is the trustee. Frost Gamma L.P. is the sole and exclusive beneficiary of Frost Gamma Investments Trust. Dr. Frost is one of two limited partners of Frost Gamma L.P. The general partner of Frost Gamma L.P. is Frost Gamma, Inc., and the sole (1) shareholder of Frost Gamma, Inc. is Frost-Nevada Corporation. Dr. Frost is also the sole shareholder of Frost-Nevada Corporation. The reporting person disclaims
- beneficial ownership of these securities, except to the extent of any pecuniary interest therein and this report shall not be deemed an admission that the reporting person is the beneficial owner of these securities for purposes of Section 16 or for any other purpose.
- These securities are owned directly by The Frost Group, LLC. Frost Gamma Investments Trust is a principal member of The Frost Group, LLC. The reporting person (2) disclaims beneficial ownership of these securities, except to the extent of any pecuniary interest therein and this report shall not be deemed an admission that the reporting person is the beneficial owner of these securities for purposes of Section 16 or for any other purpose.

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

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### JOINT FILER INFORMATION

Name: Frost Gamma Investments Trust

Address: 4400 Biscayne Blvd. Miami, FL 33137

Designated Filer: Phillip Frost, M.D.

Issuer Name and Ticker Symbol: OPKO Health, Inc. (OPK)

Date of Earliest Transaction: September 13, 2016

Relationship to Issuer: 10% Owner

FROST GAMMA INVESTMENTS TRUST

By: /s/ Phillip Frost, M.D., as Trustee Phillip Frost, M.D., Trustee