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

(Print or Type Perponses)

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

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SEC 1474 (9-02)

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 F FROST PHILLIP MD ET AL (Last) (First) OPKO HEALTH, INC., 4400 BLVD. | (Middle) | Issuer Name and Ticker or Trading Symbol OPKO HEALTH, INC. [OPK] Date of Earliest Transaction (Month/Day/Year) 05/10/2022 | | | | | | 5. Relationship of Reporting Person(s) to Issuer (Check all applicable) _X_Director _X_10% Owner X_Officer (give title below)Other (specify below) CEO & Chairman | | | |
|--|--|--|-------------|--|-------------|--------|--|---|---|------------------------|--|
| (Street) MIAMI, FL 33137 | 4. If Amendment, Date Original Filed(Month/Day/Year) | | | | | | 6. Individual or Joint/Group Filing(Check Applicable Line) Form filed by One Reporting Person X_Form filed by More than One Reporting Person | | | | |
| (City) (State) | (Zip) | Т | able I - No | on-D | erivative S | ecurit | ies Acquir | ired, Disposed of, or Beneficially Owned | | | |
| 1.Title of Security (Instr. 3) | | 2A. Deemed Execution Date, if any (Month/Day/Year) | | nstr. 8) (Instr. 3, 4 and 5) (A) (A) or | | Ď) | 5. Amount of Securities Beneficially Owned Following Reported Transaction(s) (Instr. 3 and 4) | Form: | 7. Nature of Indirect Beneficial Ownership (Instr. 4) | | |
| Common Stock | | | | | | | | 3,068,951 | D | | |
| Common Stock | 05/10/2022 | | Р | | 400,000 | A | \$ 2.7839 (1) | 195,156,694 | I | See Footnote (2) | |
| Common Stock | | | | | | | | 30,127,177 | | See Footnote (3) | |

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

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.

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

| (e.g., puts, calls, warrants, options, convertible securities) | | | | | | | | | | | | | | | |
|--|-------------|------------------|--------------------|------------|----|--------|--------|--------------|------------|--------|---------|------------|----------------|-------------|-------------|
| 1. Title of | | | 3A. Deemed | 4. | | 5. | | 6. Date Exer | | | | | 9. Number of | | 11. Nature |
| Derivative | Conversion | Date | Execution Date, if | Transacti | on | Numl | ber | and Expirati | on Date | Amou | unt of | Derivative | Derivative | Ownership | of Indirect |
| Security | or Exercise | (Month/Day/Year) | any | Code | | of | | (Month/Day | /Year) | Unde | rlying | Security | Securities | Form of | Beneficial |
| (Instr. 3) | Price of | | (Month/Day/Year) | (Instr. 8) | | Deriv | ative | | | Secur | rities | (Instr. 5) | Beneficially | Derivative | Ownership |
| | Derivative | | | | | Secur | rities | | | (Instr | . 3 and | | Owned | Security: | (Instr. 4) |
| | Security | | | | | Acqu | ired | | | 4) | | | Following | Direct (D) | |
| | | | | | | (A) o | r | | | | | | Reported | or Indirect | |
| | | | | | | Dispo | osed | | | | | | Transaction(s) | (I) | |
| | | | | | | of (D |) | | | | | | (Instr. 4) | (Instr. 4) | |
| | | | | | | (Instr | . 3, | | | | | | | | |
| | | | | | | 4, and | 15) | | | | | | | | |
| | | | | | | | | | | | Amount | 1 | | | |
| | | | | | | | | Dete | F | | or | | | | |
| | | | | | | | | * | Expiration | Title | Number | | | | |
| | | | | | | | | Exercisable | Date | | of | | | | |
| | | | | Code | V | (A) | (D) | | | | Shares | | | | |

Reporting Owners

| | Relationships | | | | | | | |
|---|---------------|--------------|----------------|-------|--|--|--|--|
| Reporting Owner Name / Address | Director | 10% Owner | Officer | Other | | | | |
| FROST PHILLIP MD ET AL OPKO HEALTH, INC. 4400 BISCAYNE BLVD. MIAMI, FL 33137 | Х | Х | CEO & Chairman | | | | | |
| Frost Gamma Investments Trust 4400 BISCAYNE BLVD. MIAMI, FL 33137 | | Х | | | | | | |

Signatures

| Phillip Frost, M.D., Individually and as Trustee | 05/11/2022 |
|--|------------|
| **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 price reported is a weighted average price. These shares were purchased in multiple transactions at prices ranging from \$2.605 to \$2.965, inclusive. The reporting person (1) undertakes to provide to OPKO Health, Inc. (the "Company"), any security holder of the Company or the staff of the Securities and Exchange Commission, upon request, full information regarding the number of shares purchased at each separate price within the range set forth in footnote (1) to this Form 4.

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

(2) 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 held by Frost Nevada Investments Trust, of which the Reporting Person is the trustee and Frost-Nevada, L.P. is the sole and exclusive beneficiary. The

(3) Reporting Person is one of seven limited partners of Frost-Nevada, L.P. and the sole shareholder of Frost-Nevada Corporation, the sole general partner of Frost-Nevada, L.P. 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.

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: May 10, 2022

Relationship to Issuer: 10% Owner

FROST GAMMA INVESTMENTS TRUST

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