FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

OMB APPROVAL

| OMB Number: | 3235-028 | | | | | | | |
|--------------------------|----------|--|--|--|--|--|--|--|
| Estimated average burden | | | | | | | | |

37 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 2. Issuer Name and Ticker or Trading Symbol 5. Relationship of Reporting Person(s) to Issuer 1. Name and Address of Reporting Person* (Check all applicable) BICYCLE THERAPEUTICS plc [BCYC] Leone Peter Brian Director 10% Owner Officer (give title Other (specify X below) below) 3. Date of Earliest Transaction (Month/Day/Year) (First) (Middle) (Last) Chief Business Officer 05/28/2019 C/O BICYCLE THERAPEUTICS PLC B900, BABRAHAM RESEARCH CAMPUS 4. If Amendment, Date of Original Filed (Month/Day/Year) 6. Individual or Joint/Group Filing (Check Applicable Line) (Street) X Form filed by One Reporting Person **CAMBRIDGE CB22 3AT** X0Form filed by More than One Reporting Person (City) (State) (Zip) Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned 4. Securities Acquired (A) or Disposed Of (D) (Instr. 3, 4 and 5) 1. Title of Security (Instr. 3) 2A. Deemed 6. Ownership 7. Nature 2. Transaction 5. Amount of Securities Beneficially Form: Direct (D) or Indirect of Indirect Beneficial **Execution Date** Transaction (Month/Day/Year) Code (Instr. if any (Month/Day/Year) 8) Owned Following (I) (Instr. 4) Ownership (Instr. 4) Reported (A) or Price Transaction(s)
(Instr. 3 and 4) Amount Code

| | | | | | | | | | | (6) | | (ilisti. 3 d | anu 4) | | |
|--|---|--|---|------------------------------|---|--------|-----|--|--------------------|--|--|---|--|--|--|
| 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. Transa Code (8) | | | | 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 | Title | Amount or Number of Shares | | | | |
| Stock Option (Right to Buy) | \$14 | 05/28/2019 | | A | | 43,269 | | (1) | 05/28/2029 | Ordinary Shares | 43,269 | \$0 | 43,269 | D | |

Explanation of Responses:

1. This option shall vest in 36 equal monthly installments at the end of each calendar month following the date of the grant.

/s/ Lee Kalowski, as Attorney-

05/30/2019

** Signature of Reporting Person

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.