FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

| OMB APPROVAL | | | | | | | | | | |
|---------------------|-----------|--|--|--|--|--|--|--|--|--|
| OMB Number: | 3235-0287 | | | | | | | | | |
| Estimated average I | ourden | | | | | | | | | |
| 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)

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

| | | | | | | | | (, | | | | | | | | | | | |
|--|---|------------|--|---------|---|---|-------|--------------|--|---|----------------------|--|---------------------------------------|--|---|--|---|---|---------------------------------------|
| Name and Address of Reporting Person* Kalowski Lee | | | | | 2. Issuer Name and Ticker or Trading Symbol BICYCLE THERAPEUTICS plc [BCYC] | | | | | | | | | k all applic Directo | able) | g Person(s) to Iss 10% Ov Other (s | | wner | |
| | (Last) (First) (Middle) C/O BICYCLE THERAPEUTICS PLC B900, BABRAHAM RESEARCH CAMPUS | | | | | 3. Date of Earliest Transaction (Month/Day/Year) 01/14/2021 | | | | | | | | | below) President and | | | below) | |
| (Street) CAMBRIDGE X0 CB22 3AT | | | | Γ | 4. If Amendment, Date of Original Filed (Month/Day/Year) | | | | | | | | | Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person Form filed by More than One Reporting Person | | | | | |
| (City) | (S | tate) | (Zip) | | | | | | | | | | | | | | | | |
| | | Tab | le I - Noi | n-Deriv | ative | e Se | curit | ies Ac | quired, | Dis | posed o | f, or Be | nefici | ally | Owned | | | | |
| | | | 2. Transaction Date (Month/Day/Year) | | ear) | 2A. Deemed Execution Date, if any (Month/Day/Year | | Code (Instr. | | 4. Securities Acquired (A Disposed Of (D) (Instr. 3, 5) | | | | Securitie Benefici | 5. Amount of Securities Beneficially Owned Following | | : Direct | 7. Nature of Indirect Beneficial Ownership (Instr. 4) | |
| | | | | | | | | | Code | v | Amount | (A) (D) | Pric | Э | Transaction(s) (Instr. 3 and 4) | | | | |
| Ordinary Shares | | | | 01/14 | 4/2021 | | | | M ⁽¹⁾ | | 10,000 | 0 A | A \$1.6 ⁽² | | 10,000 | | D | | |
| Ordinary Shares 01 | | | | 01/14 | 4/2021 | | | | S ⁽¹⁾ | | 10,000 | 0 D | | 25 | 0 | | | D | |
| | | ٦ | Table II - | | | | | | | | osed of, onvertil | | | | wned | | | | |
| 1. Title of Derivative Security (Instr. 3) | erivative Conversion Date ecurity or Exercise (Month/Day/Year) if any | | | Date, | Code (Insti | | | | 6. Date Exercisable and Expiration Date (Month/Day/Year) | | | 7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4) | | | Price of Perivative Security Instr. 5) | 9. Numbe derivative Securities Beneficia Owned Following Reported Transacti (Instr. 4) | Ownersh Form: Direct (D or Indirect (I) (Instr. | Ownership | Beneficial Ownership (Instr. 4) |
| | | | | c | Code | v | (A) | (D) | Date Exercisa | | Expiration Date | Title | Amous or Number of Shares | er | | | | | |
| Stock Option (Right to | \$1.6 ⁽²⁾ | 01/14/2021 | | | M ⁽¹⁾ | | | 10,000 | (3) | | 07/23/2027 | Ordinary Shares | 10,00 | 0 | \$0.00 | 94,77 | 4 | D | |

Explanation of Responses:

- 1. Shares exercised and sold pursuant to a Rule 10b5-1 trading plan.
- 2. This exercise price is in US Dollars and reflects the conversion of GBP to USD at an exchange rate of \$1.36521 to GBP 1.00 as of January 14, 2021.
- 3. 20% of the shares subject to the option vested on July 24, 2018, 60% of the shares subject to the option will vest each month thereafter in 36 equal monthly installments, and the remaining 20% of the shares subject to the option will vest on the earlier of (i) the fourth anniversary of the grant date and (ii) the date on which the Issuer's board of directors determines that the Issuer has satisfied certain performance vesting milestones, in all cases provided that the Reporting Person remains continuously employed with us through each applicable vesting date.

Remarks:

/s/ Lee Kalowski

01/15/2021

** 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.