FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

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

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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* Keen Nicholas						2. Issuer Name and Ticker or Trading Symbol BICYCLE THERAPEUTICS plc [BCYC]							5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director 10% Owner Officer (give title Other (specify						
(Last) (First) (Middle) C/O BICYCLE THERAPEUTICS PLC B900, BABRAHAM RESEARCH CAMPUS					3. Date of Earliest Transaction (Month/Day/Year) 03/12/2021							X Officer (give title Officer Scientific Officer							
(Street) CAMBRIDGE X0 CB22 3A					4. If <i>i</i>	4. If Amendment, Date of Original Filed (Month/Day/Year)								G. Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person Form filed by More than One Reporting Person					
(City)	(Sta	ate) (Z	Zip)																
Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned																			
Date				2. Transacti Date (Month/Day	n/Day/Year) if		2A. Deemed Execution Date, if any (Month/Day/Year)		action (Instr.	4. Securities Acquired (A) Disposed Of (D) (Instr. 3, 4			and 5) Securi Benefi		ties cially Following	Form	: Direct Indirect str. 4)	7. Nature of Indirect Beneficial Ownership (Instr. 4)	
								Code	v	Amount	(A) or (D)	Price	Tran		action(s) 3 and 4)			(111501. 4)	
Ordinary Shares 03/12/2					021	21		S ⁽¹⁾		9,500	D	\$28.4	48 ⁽²⁾ 38		3,821		D		
Ordinary Shares 03/12/2				021)21		S ⁽¹⁾		283	D	\$29	\$29		38,538		D			
Ordinary Shares 03/15/20					021	21		S ⁽¹⁾		217	D	\$28	\$28		38,321		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)	ve Conversion Date Execution I or Exercise (Month/Day/Year) if any		ion Date, Transa Code		action (Instr. Securities Acquired (A) or Disposed of (D) (Instr. 3, 4 and 5)		6. Date Exer Expiration D (Month/Day/		ate	7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4)		8. Price of Derivative Security (Instr. 5)		9. Number derivative Securities Beneficially Owned Following Reported Transaction (Instr. 4)	y G	LO. Dwnership Form: Direct (D) or Indirect (I) (Instr. 4)	11. Nature of Indirect Beneficial Ownership (Instr. 4)		
					Code	v	(A) (D)	Date Exerc	isable	Expiration Date		Amount or Number of Shares							

Explanation of Responses:

2. The price reported in Column 4 is a weighted average price. These shares were sold in multiple transactions at prices ranging from \$28.00 to \$28.925 inclusive. The Reporting Person undertakes 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 the ranges set forth in this footnote.

Remarks:

/s/ Lee Kalowski, Attorney-in-

03/16/2021

Fact

** 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.

^{1.} Shares were sold pursuant to a Rule 10b5-1 trading plan.