SECURITIES AND EXCHANGE COMMISSION Washington, D.C. 20549

PRE-EFFECTIVE AMENDMENT NO. 1 TO FORM F-6 REGISTRATION STATEMENT UNDER THE SECURITIES ACT OF 1933 FOR AMERICAN DEPOSITARY SHARES EVIDENCED BY

THE SECURITIES ACT OF 1933 FOR AMERICAN DEPOSITARY SHARES EVIDENCED BY AMERICAN DEPOSITARY RECEIPTS

ORCHARD THERAPEUTICS PLC

(Exact name of issuer of deposited securities as specified in its charter)

N/A

(Translation of issuer's name into English)

England and Wales

(Jurisdiction of incorporation or organization of issuer)

CITIBANK, N.A.

(Exact name of depositary as specified in its charter)

388 Greenwich Street New York, New York 10013 +1 877 248 4237

(Address, including zip code, and telephone number, including area code, of depositary's principal executive offices)

Cogency Global Inc. 10 East 40th Street 10th Floor New York, New York 10016 +1 212 947 7200

(Address, including zip code, and telephone number, including area code, of agent for service)

Copies to:

Mitchell S. Bloom Michael H. Bison Goodwin Procter LLP 100 Northern Avenue Boston, MA 02210 (617) 570-1000

It is proposed that this filing become effective under Rule 466:

immediately upon filing.

Herman H. Raspé, Esq.

Patterson Belknap Webb & Tyler LLP

1133 Avenue of the Americas

New York, New York 10036

212-336-2000

 \Box on (Date) at (Time).

If a separate registration statement has been filed to register the deposited shares, check the following box: 🗵

CALCULATION OF REGISTRATION FEE					
Title of Each Class of	Amount to be	Proposed Maximum	Proposed Maximum	Amount of	
Securities to be Registered	Registered	Aggregate Price Per Unit*	Aggregate Offering	Registration Fee	
			Price**		
American Depositary Shares (<u>ADS(s)</u>), each ADS					
representing the right to receive one (1) Ordinary	N/A	N/A	N/A	N/A	
Share of Orchard Therapeutics plc					

* Each unit represents 100 ADSs.

** Estimated solely for the purpose of calculating the registration fee. Pursuant to Rule 457(k), such estimate is computed on the basis of the maximum aggregate fees or charges to be imposed in connection with the issuance of ADSs.

The Registrant hereby amends this Registration Statement on such date or dates as may be necessary to delay its effective date until the Registrant shall file a further amendment which specifically states that this Registration Statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act of 1933, or until this Registration Statement shall become effective on such date as the Commission, acting pursuant to said Section 8(a), may determine.

This Pre-Effective Amendment No. 1 to Registration Statement may be executed in any number of counterparts, each of which shall be deemed an original, and all of such counterparts together shall constitute one and the same instrument.

PART I

INFORMATION REQUIRED IN PROSPECTUS

Cross Reference Sheet

Location in Form of American

Item 1. DESCRIPTION OF SECURITIES TO BE REGISTERED

Item Number and Caption			Depositary Receipt (" <u>Receipt</u> ") <u>Filed Herewith as Prospectus</u>
1.	Name of Depositary and address of its principal executive office		Face of Receipt - Introductory Article.
2.	Title of Receipts and identity of deposited securities		Face of Receipt - Top Center.
	Terms of Deposit:		
	(i)	The amount of deposited securities represented by one American Depositary Share ("ADSs")	Face of Receipt - Upper right corner.
	(ii)	The procedure for voting, if any, the deposited securities	<u>Reverse of Receipt</u> - Paragraphs (17) and (18).
	(iii)	The collection and distribution of dividends	<u>Reverse of Receipt</u> - Paragraph (15).
	(iv)	The transmission of notices, reports and proxy soliciting material	<u>Face of Receipt</u> - Paragraph (14); <u>Reverse of Receipt</u> - Paragraph (18).
	(v)	The sale or exercise of rights	<u>Reverse of Receipt</u> – Paragraphs (15) and (17).
	(vi)	The deposit or sale of securities resulting from dividends, splits or plans of reorganization	<u>Face of Receipt</u> - Paragraphs (3) and (6); <u>Reverse of Receipt</u> - Paragraphs (15) and (17).
	(vii)	Amendment, extension or termination of the deposit agreement	<u>Reverse of Receipt</u> - Paragraphs (23) and (24) (no provision for extensions).
	(viii)	Rights of holders of Receipts to inspect the transfer books of the Depositary and the list of holders of ADSs	<u>Face of Receipt</u> - Paragraph (14).

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Item Number and Caption			Filed Herewith as Prospectus	
	(ix)	Restrictions upon the right to deposit or withdraw the underlying securities	<u>Face of Receipt</u> – Paragraphs (2), (3), (4), (6), (7), (9) and (10).	
	(x)	Limitation upon the liability of the Depositary	<u>Face of Receipt</u> - Paragraph (8); <u>Reverse of Receipt</u> - Paragraphs (20) and (21).	
3.	Fees a ADSs	nd charges which may be imposed directly or indirectly on holders of	Face of Receipt - Paragraph (11).	
Item	2.	AVAILABLE INFORMATION	Face of Receipt - Paragraph (14).	

The Company is subject to the periodic reporting requirements of the United States Securities Exchange Act of 1934, as amended, and, accordingly, files certain reports with, and submits certain reports to, the United States Securities and Exchange Commission (the "Commission"). These reports can be retrieved from the Commission's internet website (www.sec.gov), and can be inspected and copied at the public reference facilities maintained by the Commission at 100 F Street, N.E., Washington D.C. 20549.

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Location in Form of American ositary Receipt ("Receipt")

PROSPECTUS

The Prospectus consists of the proposed form of American Depositary Receipt included as Exhibit A to the Form of Deposit Agreement filed as Exhibit (a) to this Registration Statement on Form F-6 and is incorporated herein by reference.

PART II

INFORMATION NOT REQUIRED IN PROSPECTUS

Item 3. EXHIBITS

- (a) Form of Deposit Agreement, by and among Orchard Therapeutics plc (the "<u>Company</u>"), Citibank, N.A., as depositary (the "<u>Depositary</u>"), and all Holders and Beneficial Owners of American Depositary Shares issued thereunder ("<u>Deposit Agreement</u>"). — Previously filed as exhibit (a) to Registration Statement on Form F-6 (Reg. No. 333-227905) with the Commission on October 19, 2018, and incorporated herein by reference.
- (b) Any other agreement to which the Depositary is a party relating to the issuance of the American Depositary Shares registered hereunder or the custody of the deposited securities represented thereby. None.
- (c) Every material contract relating to the deposited securities between the Depositary and the issuer of the deposited securities in effect at any time within the last three years. None.
- (d) Opinion of counsel for the Depositary as to the legality of the securities to be registered. Previously filed as Exhibit (d) to Registration Statement on Form F-6 (Reg. No. 333-227905) with the Commission on October 19, 2018, and incorporated herein by reference.
- (e) Certificate under Rule 466. None.
- (f) Powers of Attorney for certain officers and directors and the authorized representative of the Company. Previously filed.

Item 4. UNDERTAKINGS

- (a) The Depositary undertakes to make available at the principal office of the Depositary in the United States, for inspection by holders of ADSs, any reports and communications received from the issuer of the deposited securities which are both (1) received by the Depositary as the holder of the deposited securities, and (2) made generally available to the holders of the underlying securities by the issuer.
- (b) If the amount of fees charged is not disclosed in the prospectus, the Depositary undertakes to prepare a separate document stating the amount of any fee charged and describing the service for which it is charged and to deliver promptly a copy of such fee schedule without charge to anyone upon request. The Depositary undertakes to notify each registered holder of an ADS thirty (30) days before any change in the fee schedule.

SIGNATURES

Pursuant to the requirements of the Securities Act of 1933, as amended, Citibank, N.A., acting solely on behalf of the legal entity to be created by the Deposit Agreement, by and among Orchard Therapeutics plc, Citibank, N.A., as depositary, and all Holders and Beneficial Owners of American Depositary Shares to be issued thereunder, certifies that it has reasonable grounds to believe that all the requirements for filing on Form F-6 are met and has duly caused this Pre-Effective Amendment No. 1 to Registration Statement on Form F-6 to be signed on its behalf by the undersigned, thereunto duly authorized, in the City of New York, State of New York, on the 29th day of October, 2018.

Legal entity created by the Deposit Agreement under which the American Depositary Shares registered hereunder are to be issued, each American Depositary Share representing the right to receive one (1) Ordinary Share of Orchard Therapeutics plc.

CITIBANK, N.A., solely in its capacity as Depositary

By: /s/ Leslie Deluca Name: Leslie Deluca Title: Attorney-in-fact

SIGNATURES

Pursuant to the requirements of the Securities Act of 1933, as amended, Orchard Therapeutics plc certifies that it has reasonable grounds to believe that all the requirements for filing on Form F-6 are met and has duly caused this Pre-Effective Amendment No. 1 to Registration Statement on Form F-6 to be signed on its behalf by the undersigned thereunto duly authorized, in the City of London, United Kingdom, on October 29, 2018.

ORCHARD THERAPEUTICS PLC

By: /s/ Mark Rothera

Name: Mark Rothera Title: President and Chief Executive Officer Pursuant to the requirements of the Securities Act of 1933, as amended, this Pre-Effective Amendment No. 1 to Registration Statement on Form F-6 has been signed by the following persons in the following capacities on October 29, 2018.

Signature	Title
/s/ Mark Rothera Mark Rothera	President, Chief Executive Officer and Director (Principal Executive Officer)
/s/ Frank E. Thomas Frank E. Thomas	Chief Financial Officer and Chief Business Officer (Principal Financial Officer and Principal Accounting Officer)
* James A. Geraghty	Chairman of the Board of Directors
* Joanne T. Beck, Ph.D.	Director
* Marc Dunoyer	Director
* Jon Ellis, Ph.D.	Director
* Bobby Gaspar, M.D., Ph.D.	Director
* Alex Pasteur, Ph.D.	Director
*	Director
Charles A. Rowland, Jr.	Director
Hong Fang Song * Elise Wang	Director
LITSC Mank	II-5

*By <u>/s/ Frank E. Thomas</u> Name: Frank E. Thomas Title: Attorney-in-Fact Pursuant to the requirements of the Securities Act of 1933, as amended, the undersigned, the duly authorized representative in the United States of Orchard Therapeutics plc has signed this Pre-Effective Amendment No. 1 to Registration Statement on Form F-6 to be signed on October 29, 2018.

By: /s/ Tais Laureano

Name:Tais LaureanoTitle:Assistant Secretary on behalf of Cogency Global Inc.

<u>Exhibit</u>

<u>Document</u>

Sequentially <u>Numbered Page</u>