Prospectus Supplement No. 6 (To Prospectus dated March 21, 2022)

eFFECTOR Therapeutics, Inc.



This prospectus supplement updates, amends and supplements the prospectus dated March 21, 2022 (the "Prospectus"), which forms a part of our Registration Statement on Form S-1 (Registration No. 333-259751). Capitalized terms used in this prospectus supplement and not otherwise defined herein have the meanings specified in the Prospectus.

This prospectus supplement is being filed to update, amend and supplement the information included in the Prospectus with the information contained in our Current Reports on Form 8-K (the "Current Reports"), filed with the SEC on March 28, 2022 and April 4, 2022. Accordingly, we have attached the Current Reports to this prospectus supplement.

This prospectus supplement is not complete without the Prospectus. This prospectus supplement should be read in conjunction with the Prospectus, which is to be delivered with this prospectus supplement, and is qualified by reference thereto, except to the extent that the information in this prospectus supplement updates or supersedes the information contained in the Prospectus.

Our common stock and warrants are listed on the Nasdaq Capital Market under the symbols "EFTR" and "EFTRW." On April 1, 2022, the closing price of our common stock was \$3.82 and the closing price of our warrants was \$0.43.

We are an "emerging growth company" under federal securities laws and are subject to reduced public company reporting requirements. Investing in our securities involves certain risks. See "Risk Factors" beginning on page 6 of the Prospectus.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or determined if the Prospectus or this prospectus supplement is truthful or complete. Any representation to the contrary is a criminal offense.

The date of this prospectus supplement is April 4, 2022.

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

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CURRENT REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934

Date of Report (Date of earliest event reported): March 28, 2022

eFFECTOR Theraneutics Inc

Delaware	001-39866	85-3306396
(State or other jurisdiction of incorporation)	(Commission File Number)	(I.R.S. Employer Identification No.)
142 North Cedros Avenue, Suite B	,	,
Solana Beach, California		92075
(Address of principal executive offices)		(Zip Code)
(Registr	(858) 925-8215 rant's telephone number, including area code)	
(Former nar	ne or former address, if changed since last rep	ort)
Check the appropriate box below if the Form 8-K filing is in following provisions: Written communications pursuant to Rule 425 under the S	, ,	ng obligation of the registrant under any of the
☐ Soliciting material pursuant to Rule 14a-12 under the Exc	· · · · · · · · · · · · · · · · · · ·	
☐ Pre-commencement communications pursuant to Rule 14d	d-2(b) under the Exchange Act (17 CFR	2 240.14d-2(b))
☐ Pre-commencement communications pursuant to Rule 13c	e-4(c) under the Exchange Act (17 CFR	. 240.13e-4(c))
Securities registered pursuant to Section 12(b) of the Act:		
	Trading Symbol(s)	Name of each exchange on which registered
Title of each class	Symbol(s)	
Title of each class Common stock, \$0.0001 par value per share	EFTR	Nasdaq Capital Market

Emerging growth company \boxtimes

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act. □

Item 5.08 Shareholder Director Nominations.

On March 28, 2022, the Board of Directors of eFFECTOR Therapeutics, Inc. (the "Company") established the date of the Company's 2022 Annual Meeting of Stockholders (the "2022 Annual Meeting"). The 2022 Annual Meeting will be held on June 9, 2022, and the record date for the 2022 Annual Meeting will be April 13, 2022. The time of the 2022 Annual Meeting will be set forth in the Company's definitive proxy statement for the 2022 Annual Meeting to be filed with the Securities and Exchange Commission. The Company is holding the 2022 Annual Meeting as a virtual-only meeting, which will be conducted via live audio online webcast.

The Company's bylaws permit stockholders to bring director nominations and other proposals for consideration at the 2022 Annual Meeting, provided they comply with the notice and other requirements set forth in the bylaws and SEC rules. Because the Company did not hold an annual meeting of stockholders during the previous year, all stockholder notices to bring director nominations and other matters before the 2022 Annual Meeting must be received by the Company no later than April 7, 2022, which is the tenth date following this public announcement of the 2022 Annual Meeting date. This is also the date the Company has determined to be a reasonable time before it expects to begin to print and distribute its proxy materials, and therefore, under SEC rules, any stockholder who wishes to include director nominations or other proposals in the Company's proxy materials for the 2022 Annual Meeting must submit proper notice no later than April 7, 2022. Stockholders may submit written notice for any director nomination or other proposal to:

eFFECTOR Therapeutics, Inc., Attn: Corporate Secretary 142 North Cedros Avenue, Suite B Solana Beach, California 92075

Any written notice pursuant to the foregoing must also comply with all other applicable requirements found in the rules of the U.S. Securities and Exchange Commission, including Rule 14a-8, the Delaware General Corporation Law, and the Company's bylaws.

SIGNATURE

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

eFFECTOR Therapeutics, Inc.

Date: March 28, 2022 By: /s/ Michael Byrnes

Name: Michael Byrnes
Title: Chief Financial Officer

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

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CURRENT REPORT PURSUANT TO SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934

Date of Report (Date of earliest event reported): April 1, 2022

eFFECTOR Therapeutics, Inc. (Exact name of registrant as specified in its charter)

001-39866

85-3306396

Delaware

	e or other jurisdiction of incorporation)	(Commission File Number)	(I.K.S. Employer Identification No.)		
	142 North Cedros Avenue, Suite B				
Solana Beach, California			92075		
	(Address of principal executive offices)		(Zip Code)		
	(Registr	(858) 925-8215 rant's telephone number, including area code)			
	(Former nar	me or former address, if changed since last repo	rt)		
Check the appropri		tended to simultaneously satisfy the filin	ng obligation of the registrant under any of the		
□ Written comr	Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)				
☐ Soliciting ma	terial pursuant to Rule 14a-12 under the l	Exchange Act (17 CFR 240.14a-12)			
□ Pre-commend	cement communications pursuant to Rule	14d-2(b) under the Exchange Act (17 C	FR 240.14d-2(b))		
□ Pre-commend	cement communications pursuant to Rule	13e-4(c) under the Exchange Act (17 C	FR 240.13e-4(c))		
Securities registere	d pursuant to Section 12(b) of the Act:				
7	Fitle of each class	Trading Symbol(s)	Name of each exchange on which registered		
Common stock,	\$0.0001 par value per share	EFTR	Nasdaq Capital Market		
Warrants to purchase common stock		EFTRW	Nasdaq Capital Market		
	nark whether the registrant is an emerging bb-2 of the Securities Exchange Act of 19		5 of the Securities Act of 1933 (Sec.230.405 of this		
Emerging growth c	ompany 🗆				
~ ~ ~ ~	wth company, indicate by check mark if the notal accounting standards provided pursion	e	stended transition period for complying with any		

Item 5.02 Departure of Directors or Certain Officers; Election of Directors; Appointment of Certain Officers; Compensatory Arrangements of Certain Officers.

On April 1, 2022, Premal Patel, M.D., Ph.D., Chief Medical Officer of eFFECTOR Therapeutics, Inc. (the "Company"), resigned from the Company in order to pursue a new career opportunity. Mr. Patel's duties and responsibilities have been reassigned to existing executive officers and other management personnel at the Company. Dr. Patel will continue to serve the company as a consultant to the company until Sept. 30, 2022. Dr. Patel's resignation from the Company was not due to any disagreement with the Company or its management on any matter relating to the Company's operations, policies or practices.

In connection with Dr. Patel's termination of employment with the Company on April 1, 2022 (the "Effective Date"), Dr. Patel executed a general release of claims (the "Release"), pursuant to which (i) Dr. Patel's vested Company stock options will remain eligible to become exercisable until April 1, 2023, and (ii) the Company will continue to pay the employer portion of the healthcare coverage for Mr. Patel and his eligible dependents, in each case, until December 31, 2022. Pursuant to a Consulting Agreement (the "Consulting Agreement") entered into as of the Effective Date, Dr. Patel will receive an hourly consulting fee of \$600, and continued vesting of his outstanding equity awards during the term of his consulting services.

The foregoing description of the Release and Consulting Agreement is a summary only and is subject to and qualified in its entirety by the terms of the Release and Consulting Agreement, copies of which will be filed as exhibits to the Company's Quarterly Report on Form 10-Q for the fiscal quarter ending June 30, 2022.

SIGNATURE

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

eFFECTOR Therapeutics, Inc.

Date: April 4, 2022 By: /s/ Michael Byrnes

Name: Michael Byrnes

Title: Chief Financial Officer