PROSPECTUS SUPPLEMENT NO. 7 (to Prospectus dated August 12, 2021)



Sema4 Holdings Corp.

236,223,401 Shares of Common Stock

7,236,667 Warrants to Purchase Shares of Common Stock

21,995,000 Shares of Common Stock Underlying Warrants

This prospectus supplement supplements the prospectus dated August 12, 2021 (the "Prospectus"), which forms a part of our registration statement on Form S-1 (No. 333-258467). This prospectus supplement is being filed to update and supplement the information in the Prospectus with the information contained in our current report on Form 8-K, filed with the Securities and Exchange Commission (the "SEC") on January 31, 2022 (the "Current Report"). Accordingly, we have attached the Current Report to this prospectus supplement.

The Prospectus and this prospectus supplement relate to the offer and sale from time to time by the selling securityholders named in the Prospectus (the "Selling Securityholders") of (A) up to 236,223,401 shares of our Class A common stock, par value \$0.0001 per share ("Class A common stock" or "common stock"), consisting of (i) up to 35,000,000 shares of our Class A common stock issued in a private placement pursuant to subscription agreements each entered into on February 9, 2021; (ii) up to 11,068,750 shares of our Class A common stock issued in connection with the consummation of the Business Combination (as defined in the Prospectus), in exchange for shares of our Class B common stock originally issued in a private placement to CMLS Holdings LLC (the "Sponsor"); (iii) up to 182,917,984 shares of our Class A common stock issued or issuable to certain former stockholders and equity award holders of Sema4 (the "Sema4 equity holders") in connection with or as a result of the consummation of the Business Combination, consisting of (a) up to 149,856,840 shares of our Class A common stock; (b) up to 14,039,568 shares of our Class A common stock issuable upon the exercise or vesting of certain equity awards; and (c) up to 19,021,576 shares of Class A common stock that certain Sema4 equity holders have the contingent right to receive upon the achievement of certain vesting conditions; and (iv) up to 7,236,667 shares of our Class A common stock issuable upon the exercise of the private placement warrants (as defined below); and (B) up to 7,236,667 warrants (the "private placement warrants") originally issued in a private placement to the Sponsor and certain of the other Initial Stockholders (as defined in the Prospectus).

In addition, the Prospectus and this prospectus supplement relate to the offer and sale of: (i) up to 14,758,333 shares of our Class A common stock that are issuable by us upon the exercise of 14,758,333 warrants (the "public warrants") originally issued in our initial public offering; and (ii) up to 7,236,667 shares of our Class A common stock that are issuable by us upon the exercise of the private placement warrants following the public resale of the private placement warrants by the Selling Securityholders pursuant to the Prospectus and this prospectus supplement.

Our common stock and public warrants are listed on the Nasdaq Global Select Market (the "Nasdaq") under the symbols "SMFR" and "SMFRW," respectively. On January 28, 2022, the last reported sales price of our common stock was \$3.07 per share and the last reported sales price of our public warrants was \$0.68 per warrant.

This prospectus supplement updates and supplements the information in the Prospectus and is not complete without, and may not be delivered or utilized except in combination with, the Prospectus, including any amendments or supplements thereto. This prospectus supplement should be read in conjunction with the Prospectus and if there is any inconsistency between the information in the Prospectus and this prospectus supplement, you should rely on the information in this prospectus supplement.

Investing in our securities involves risks. See the section entitled "Risk Factors" beginning on page 10 of the Prospectus to read about factors you should consider before buying our securities.

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

The date of this prospectus supplement is January 31, 2022

UNITED STATES

SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

FORM 8-K

CURRENT REPORT
Pursuant to Section 13 OR 15(d) of The Securities Exchange Act of 1934

Date of Report (date of earliest event reported): January 31, 2022 (January 25, 2022)

Sema4 Holdings Corp.

(Exact name of registrant as specified in its charter)		
Delaware	001-39482	85-1966622
(State or other jurisdiction of incorporation or organization)	(Commission File Number)	(I.R.S. Employer Identification No.)
333 Ludlow Street, North Tower, 8th Floor		
Stamford, Connecticut		06902
(Address of Principal Executive Offices)		(Zip Code)
	(800) 298-6470 s telephone number, including are r former address, if changed since	
Check the appropriate box below if the Form 8-K filing is intenfollowing provisions (see General Instruction A.2. below):	ded to simultaneously satisfy the	filing obligation of the registrant under any of the
\square Written communications pursuant to Rule 425 under the Sec	curities Act (17 CFR 230.425)	
☐ Soliciting material pursuant to Rule 14a-12 under the Excha	ange Act (17 CFR 240.14a-12)	
☐ Pre-commencement communications pursuant to Rule 14d-	2(b) under the Exchange Act (17 (CFR 240.14d-2(b))
☐ Pre-commencement communications pursuant to Rule 13e-	4(c) under the Exchange Act (17 C	CFR 240.13e-4(c))
Securities registered pursuant to Section 12(b) of the Act:		
Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Class A common stock, par value \$0.0001 per share	SMFR	The Nasdaq Global Select Market
Warrants to purchase one share of Class A common stock, each at an exercise price of \$11.50 per share	ch SMFRW	The Nasdaq Global Select Market
Indicate by check mark whether the registrant is an emerging grapher) or Rule 12b-2 of the Securities Exchange Act of 1934		405 of the Securities Act of 1933 (§230.405 of this

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.□

Emerging growth company ⊠

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

(b) Departure of Certain Officers.

James Coffin, Ph.D., the President and Chief Operating Officer of Sema4 Holdings Corp. (the "Company"), will leave the Company for personal reasons, effective February 1, 2022.

(e) Compensatory Arrangements with Certain Officers.

In connection with Mr. Coffin's departure, the Company and Mr. Coffin have entered into a Separation Agreement (the "Separation Agreement"), pursuant to which Mr. Coffin will receive the following severance benefits in exchange for a customary release of claims against the Company:

- a severance payment equal to 12 months of Mr. Coffin's annual base salary;
- a discretionary 2021 annual bonus payment;
- 12 months of reimbursement of COBRA continuation benefits;
- accelerated vesting of 25,425 stock options that were otherwise scheduled to vest on February 2, 2022; and
- an extended period to exercise certain of Mr. Coffin's vested stock options through May 30, 2022.

The foregoing description of the Separation Agreement does not purport to be complete and is qualified in its entirety by reference to the full text of the Separation Agreement, a copy of which is attached as Exhibit 10.1 and is incorporated herein by reference.

Item 9.01 Financial Statements and Exhibits.

(d) Exhibits.

<u>Exhibit</u>	
<u>Number</u>	<u>Description</u>
10.1	Separation Agreement with James Coffin dated as of January 25, 2022.
104	Cover Page Interactive Data File (embedded within the Inline XBRL Document)

SIGNATURES

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.

Sema4 Holdings Corp.

Date: January 31, 2022 By: /s/ Eric Schadt

Name: Eric Schadt

Title: Chief Executive Officer