UNITED STATES SECURITIES AND EXCHANGE COMMISSION

WASHINGTON, DC 20549

FORM 10-Q

(Mark One)

X

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

For the quarterly period ended June 30, 2021

OR

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

For the transition period from _____ to _

Commission File Number: 001-39219

Revolution Medicines, Inc.

(Exact Name of Registrant as Specified in its Charter)

 Delaware
 47-2029180

 (State or other jurisdiction of incorporation or organization)
 (I.R.S. Employer Identification No.)

 700 Saginaw Drive
 94063

 Redwood City, CA
 94063

 (Address of principal executive offices)
 (Zip Code)

 Registrant's telephone number, including area code: (650) 481-6801

Securities registered pursuant to Section 12(b) of the Act:

Title of each class	Trading Symbol(s)	Name of each exchange on which registered
Common Stock \$0.0001 Par Value per Share	RVMD	The Nasdaq Stock Market LLC
		(Nasdaq Global Select Market)

Indicate by check mark whether the registrant (1) has filed all reports required to be filed by Section 13 or 15(d) of the Securities Exchange Act of 1934 during the preceding 12 months (or for such shorter period that the registrant was required to file such reports), and (2) has been subject to such filing requirements for the past 90 days. Yes 🛛 No 🗆

Indicate by check mark whether the registrant has submitted electronically every Interactive Data File required to be submitted pursuant to Rule 405 of Regulation S-T (§232.405 of this chapter) during the preceding 12 months (or for such shorter period that the registrant was required to submit such files). Yes 🗵 No 🗆

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, a smaller reporting company, or an emerging growth company. See the definitions of "large accelerated filer," "accelerated filer," "smaller reporting company," and "emerging growth company" in Rule 12b-2 of the Exchange Act.

Large accelerated filer		Accelerated filer	
Non-accelerated filer	\boxtimes	Smaller reporting company	
		Emerging growth company	X

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.

Indicate by check mark whether the registrant is a shell company (as defined in Rule 12b-2 of the Exchange Act). Yes 🗌 No 🗵

As of August 6, 2021, the registrant had 73,581,042 shares of common stock, \$0.0001 par value per share, outstanding.

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SPECIAL NOTE REGARDING FORWARD-LOOKING STATEMENTS

This Quarterly Report on Form 10-Q contains forward-looking statements concerning our business, operations and financial performance and condition, as well as our plans, objectives and expectations for our business, operations and financial performance and condition. Any statements contained herein that are not statements of historical facts may be deemed to be forward-looking statements. These statements involve known and unknown risks, uncertainties and other important factors that are in some cases beyond our control and may cause our actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements.

In some cases, you can identify forward-looking statements by terminology such as "aim," "anticipate," "assume," "believe," "contemplate," "continue," "could," "due," "estimate," "expect," "goal," "intend," "may," "objective," "plan," "predict," "potential," "positioned," "seek," "should," "target," "will," "would," and other similar expressions that are predictions of or indicate future events and future trends, or the negative of these terms or other comparable terminology. These forward-looking statements include, but are not limited to, statements about:

- the scope, progress, results and costs of developing our product candidates or any other future product candidates, and conducting preclinical studies and clinical trials, including our RMC-4630 Phase 1/2 clinical program;
- the scope, progress, results and costs related to the research and development of our pipeline;
- the timing of and costs involved in obtaining and maintaining regulatory approval for any of current or future product candidates, and any related restrictions, limitations and/or warnings in the label of an approved product candidate;
- our expectations regarding the potential market size and size of the potential patient populations for RMC-4630, our other product candidates and any
 future product candidates, if approved for commercial use;
- our ability to maintain existing and establish new collaborations, licensing or other arrangements and the financial terms of any such agreements, including our collaboration with Sanofi;
- our commercialization, marketing and manufacturing capabilities and expectations;
- the rate and degree of market acceptance of our product candidates, as well as the pricing and reimbursement of our product candidates, if approved;
- the implementation of our business model and strategic plans for our business, product candidates and technology, including additional indications for which we may pursue;
- the scope of protection we are able to establish and maintain for intellectual property rights covering our product candidates, including the projected term of patent protection;
- estimates of our expenses, future revenue, capital requirements, our needs for additional financing and our ability to obtain additional capital;
- developments and projections relating to our competitors and our industry, including competing therapies and procedures;
- regulatory and legal developments in the United States and foreign countries;
- the performance of our third-party suppliers and manufacturers;
- our ability to attract and retain key scientific or management personnel;
- our expectations regarding the period during which we will qualify as an emerging growth company under the Jumpstart Our Business Startups Act of 2012;
- our expectations regarding our ability to obtain, maintain, enforce and defend our intellectual property protection for our product candidates; and
- other risks and uncertainties, including those listed under the caption "Risk Factors."

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We have based these forward-looking statements largely on management's current expectations, estimates, forecasts and projections about our business and the industry in which we operate and management's beliefs and assumptions and are not guarantees of future performance or development and involve known and unknown risks, uncertainties and other factors that are in some cases beyond our control. These forward-looking statements speak only as of the date of this Quarterly Report on Form 10-Q and are subject to a number of risks, uncertainties and assumptions described in the section titled "Risk Factors" and elsewhere in this Quarterly Report on Form 10-Q. Because forward-looking statements are inherently subject to risks and uncertainties, some of which cannot be predicted or quantified, you should not rely on these forward-looking statements as predictions of future events. The events and circumstances reflected in our forward-looking statements may not be achieved or occur and actual results could differ materially from those projected in the forward-looking statements. Except as required by applicable law, we do not plan to publicly update or revise any forward-looking statements contained herein until after we distribute this Quarterly Report on Form 10-Q, whether as a result of any new information, future events or otherwise.

In addition, statements that "we believe" and similar statements reflect our beliefs and opinions on the relevant subject. These statements are based upon information available to us as of the date of this report, and while we believe such information forms a reasonable basis for such statements, such information may be limited or incomplete, and our statements should not be read to indicate that we have conducted an exhaustive inquiry into, or review of, all potentially available relevant information. These statements are inherently uncertain, and you are cautioned not to unduly rely upon these statements.

Investors and others should note that we may announce material business and financial information to our investors using our investor relations website (https://ir.revmed.com), Securities and Exchange Commission (SEC) filings, webcasts, press releases and conference calls. We use these mediums, including our website, to communicate with our members and public about our company, our products and other issues. It is possible that the information that we make available may be deemed to be material information. We therefore encourage investors and others interested in our company to review the information that we make available on our website.

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PART I—FINANCIAL INFORMATION

REVOLUTION MEDICINES, INC. CONDENSED CONSOLIDATED BALANCE SHEETS (in thousands, except share and per share data) (unaudited)

		June 30,	December 31,		
		2021		2020	
Assets					
Current assets:					
Cash and cash equivalents	\$	195,600	\$	104,268	
Marketable securities		450,722		336,473	
Accounts receivable		6,527		6,393	
Prepaid expenses and other current assets		8,162		6,988	
Total current assets		661,011		454,122	
Property and equipment, net		10,744		8,902	
Operating lease right-of-use asset		25,925		27,435	
Intangible assets, net		60,410		60,945	
Goodwill		14,608		14,608	
Restricted cash		1,084		1,084	
Other noncurrent assets		264		305	
Total assets	\$	774,046	\$	567,401	
Liabilities, redeemable convertible preferred stock and stockholders' equity					
Current liabilities:					
Accounts payable	\$	10,435	\$	12,609	
Accrued expenses and other current liabilities		22,103		18,784	
Operating lease liability, current		4,926		3,672	
Deferred revenue, current		8,337		12,111	
Total current liabilities		45,801		47,176	
Deferred revenue, noncurrent		7,621		8,481	
Deferred tax liability		7,444		7,444	
Operating lease liability, noncurrent		27,566		28,992	
Other noncurrent liabilities		608		632	
Total liabilities	·	89,040		92,725	
Commitments and contingencies (Note 7)					
Stockholders' equity:					
Preferred stock, \$0.0001 par value; 10,000,000 shares authorized at					
June 30, 2021 and December 31, 2020, respectively; zero shares					
issued and outstanding at June 30, 2021 and December 31, 2020, respectively		—		—	
Common stock, \$0.0001 par value; 300,000,000 shares authorized at June 30, 2021 and December 31, 2020, respectively; 73,576,989 and 66,599,748					
shares issued and outstanding at June 30, 2021 and December 31, 2020,					
respectively		8		7	
Additional paid-in capital		1,032,009		740,098	
Accumulated other comprehensive income		9		116	
Accumulated deficit		(347,020)		(265,545)	
Total stockholders' equity		685,006		474,676	
Total liabilities, redeemable convertible preferred stock and stockholders' equity	\$	774,046	\$	567,401	

The accompanying notes are an integral part of these condensed consolidated financial statements.

REVOLUTION MEDICINES, INC. CONDENSED CONSOLIDATED STATEMENTS OF OPERATIONS (in thousands, except share and per share data) (unaudited)

	Three Months Ended June 30,					Six Months Ended June 30,				
		2021		2020		2021		2020		
Revenue:										
Collaboration revenue	\$	8,698	\$	10,025	\$	18,829	\$	21,571		
Total revenue		8,698		10,025		18,829		21,571		
Operating expenses:										
Research and development		45,936		32,918		86,794		60,375		
General and administrative		7,297		5,091		13,967		10,262		
Total operating expenses		53,233		38,009		100,761		70,637		
Loss from operations		(44,535)		(27,984)		(81,932)		(49,066)		
Other income (expense), net:										
Interest income		236		730		469		1,639		
Interest expense		—		(19)		(12)		(40)		
Total other income, net		236		711		457		1,599		
Loss before income taxes		(44,299)		(27,273)		(81,475)		(47,467)		
Benefit from income taxes		_		58		_		733		
Net loss	\$	(44,299)	\$	(27,215)	\$	(81,475)	\$	(46,734)		
Redeemable convertible preferred stock dividends - undeclared and	-				_					
cumulative		_						(2,219)		
Net loss attributable to common stockholders	\$	(44,299)	\$	(27,215)	\$	(81,475)	\$	(48,953)		
Net loss per share attributable to common stockholders - basic and diluted	\$	(0.60)	\$	(0.46)	\$	(1.13)	\$	(1.11)		
Weighted-average common shares used to compute net loss per share, basic and diluted		73,399,714		58,752,494		71,917,508		44,025,372		

The accompanying notes are an integral part of these condensed consolidated financial statements.

REVOLUTION MEDICINES, INC. CONDENSED CONSOLIDATED STATEMENTS OF COMPREHENSIVE LOSS (in thousands) (unaudited)

	Three Months Ended June 30,					Six Months Er	ıded June 30,		
		2021 2020				2021		2020	
Net loss	\$	(44,299)	\$	(27,215)	\$	(81,475)	\$	(46,734)	
Other comprehensive income/(loss)									
Unrealized gain (loss) on investments, net		(54)		255		(107)		217	
Total comprehensive loss	\$	(44,353)	\$	(26,960)	\$	(81,582)	\$	(46,517)	

The accompanying notes are an integral part of these condensed consolidated financial statements.

REVOLUTION MEDICINES, INC. CONDENSED CONSOLIDATED STATEMENTS OF REDEEMABLE CONVERTIBLE PREFERRED STOCK AND STOCKHOLDERS' EQUITY (DEFICIT) (in thousands, except share data) (unaudited)

	Redeemable Preferre		Common Stock Shares Amount		Additional Paid-in	Accumulated other comprehensive	Accumulated	Total Stockholders'
	Shares	Amount			Capital	income	Deficit	Equity/(Deficit)
Balance at December 31, 2020	—	\$ —	66,599,748	\$ 7	\$ 740,098	\$ 116	\$ (265,545)	\$ 474,676
Issuance of common stock upon follow-on offering, net of offering costs of \$18,855	_	_	6,666,666	1	281,144	_	_	281,145
Issuance of common stock pursuant to stock option exercises	_	_	166,897	_	555	_	_	555
Issuance of common stock related to vesting of restricted stock units	_	_	1,798	_	_	_	_	_
Vesting of early exercised stock options	_	_	_	_	46	_	_	46
Stock-based compensation expense	_	_		_	3,387	_	_	3,387
Net unrealized loss on marketable securities	_	_			_	(53)	_	(53)
Net loss	_	—		_	_	_	(37,176)	(37,176)
Balance at March 31, 2021	_	\$ —	73,435,109	\$8	\$ 1,025,230	\$ 63	\$ (302,721)	\$ 722,580
Issuance of common stock pursuant to stock option exercises	_	_	64,862	_	257	_	_	257
Issuance of common stock related to employee stock purchase plan	_	_	46,201	_	1,175	_	_	1,175
Issuance of common stock related to vesting of restricted stock units	_	_	30,817	_	_	_	_	_
Vesting of early exercised stock options	_	_		_	34	_	_	34
Stock-based compensation expense	—	—	_	_	5,313	—	_	5,313
Net unrealized loss on marketable securities		_		_	_	(54)	_	(54)
Net loss							(44,299)	(44,299)
Balance at June 30, 2021		<u>\$ </u>	73,576,989	\$8	\$ 1,032,009	\$ 9	\$ (347,020)	\$ 685,006

The accompanying notes are an integral part of these condensed consolidated financial statements.

REVOLUTION MEDICINES, INC. CONDENSED CONSOLIDATED STATEMENTS OF REDEEMABLE CONVERTIBLE PREFERRED STOCK AND STOCKHOLDERS' EQUITY (DEFICIT) (in thousands, except share data) (unaudited)

	Redeemable C Preferred		Common Stock			Accumulated other comprehensive	Accumulated	Total Stockholders'
	Shares	Amount	Shares	Amount	Capital	income	Deficit	Equity/(Deficit)
Balance at December 31, 2019	39,600,423	\$ 305,109	3,292,124	\$ —	\$ 4,738	\$ 74	\$ (157,386)	\$ (152,574)
Conversion of redeemable convertible preferred stock into common stock	(39,600,423)	(305,109)	39,600,423	4	305,105	_	_	305,109
Issuance of common stock upon initial public offering, net of offering costs of \$23,003	_	_	16,100,000	2	250,695	_	_	250,697
Issuance of common stock pursuant to stock option exercises	_	_	11,097	_	27	_	_	27
Vesting of early exercised stock options	_	_	_		47	_	_	47
Stock-based compensation expense	_	_	_		1,567	_	_	1,567
Net unrealized loss on marketable securities	_	_	_		_	(38)	_	(38)
Net loss	_	_	_	_	_		(19,519)	(19,519)
Balance at March 31, 2020		\$ —	59,003,644	\$ 6	\$ 562,179	\$ 36	\$ (176,905)	\$ 385,316
Issuance of common stock pursuant to stock option								
exercises	—	_	21,578		86	_	—	86
Vesting of early exercised stock options		_	_		39		_	39
Repurchase of common stock	—	_	(9,354)		—	_	—	_
Stock-based compensation expense		—	_		2,017	_	_	2,017
Net unrealized gain on marketable securities	—	_	_	_	—	255	—	255
Net loss		_	_		_		(27,215)	(27,215)
Balance at June 30, 2020		\$ —	59,015,868	\$6	\$ 564,321	\$ 291	\$ (204,120)	\$ 360,498

The accompanying notes are an integral part of these condensed consolidated financial statements.

REVOLUTION MEDICINES, INC. CONDENSED CONSOLIDATED STATEMENTS OF CASH FLOWS (in thousands) (unaudited)

	Six Months Ended June 30,						
		2021		2020			
Cash flows from operating activities							
Net loss	\$	(81,475)	\$	(46,734)			
Adjustments to reconcile net loss to net cash used in operating activities:							
Amortization of intangible assets		535		534			
Stock-based compensation expense		8,700		3,584			
Depreciation and amortization		1,474		1,272			
Net amortization (accretion) of premium (discount) on marketable securities		1,227		(18)			
Amortization of operating lease right-of-use asset		1,510		1,409			
Changes in operating assets and liabilities:							
Accounts receivable		(134)		1,406			
Prepaid expenses and other current assets		(1,134)		(2,541)			
Accounts payable		(523)		(941			
Accrued expenses and other current liabilities		3,243		1,286			
Deferred revenue		(4,634)		(5,660			
Deferred tax liability				(734			
Operating lease liability		(172)		(1,273			
Other noncurrent assets				274			
Other noncurrent liabilities		56		127			
Net cash used in operating activities		(71,327)		(48,009			
Cash flows from investing activities							
Purchases of marketable securities		(392,273)		(305,205)			
Maturities of marketable securities		276,691		107,065			
Sales of marketable securities		—		3,005			
Purchases of property and equipment		(4,891)		(1,077			
Net cash used in investing activities		(120,473)		(196,212			
Cash flows from financing activities							
Proceeds from issuance of common stock, net of issuance costs		281,145		252,300			
Proceeds from issuance of common stock under equity incentive plans		812		113			
Proceeds from issuance of common stock related to employee stock purchase plan		1,175		_			
Repurchases of early exercised stock				(5			
Net cash provided by financing activities		283,132		252,408			
Net increase in cash, cash equivalents and restricted cash		91,332		8,187			
Cash, cash equivalents and restricted cash - beginning of period		105,352		16,873			
Cash, cash equivalents and restricted cash - end of period	\$	196,684	\$	25,060			
Reconciliation of cash, cash equivalents and restricted cash to consolidated balance sheets	<u>.</u>		<u> </u>	,			
Cash and cash equivalents		195,600		23,976			
Restricted cash		1,084		1,084			
Cash, cash equivalents and restricted cash - end of period	\$	196,684	\$	25,060			
Supplemental disclosure of non-cash investing and financing activities			<u> </u>				
Vesting of early exercised options and restricted stock	\$	80	\$	91			
Purchases of property and equipment in accounts payable and accrued expenses and other current liabilities	ψ	238	ψ	530			
Right-of-use assets obtained in exchange for operating lease liabilities		230		21,188			

The accompanying notes are an integral part of these condensed consolidated financial statements.

REVOLUTION MEDICINES, INC. Notes to Unaudited Condensed Consolidated Financial Statements

1. Organization

Revolution Medicines, Inc. (the Company) is a clinical-stage precision oncology company focused on developing targeted therapies to inhibit frontier targets in RAS-addicted cancers. The Company was founded in October 2014 and is headquartered in Redwood City, California.

Liquidity

The Company has incurred net operating losses in each year since inception. As of June 30, 2021, the Company had an accumulated deficit of \$347.0 million. Management believes that its existing cash, cash equivalents and marketable securities will enable the Company to fund its planned operations for at least 12 months following the issuance date of these condensed consolidated financial statements. The Company has been able to fund its operations through the issuance and sale of common stock and redeemable convertible preferred stock in addition to upfront payments and research and development cost reimbursement received under the Company's collaboration agreement with Genzyme Corporation, an affiliate of Sanofi. Future capital requirements will depend on many factors, including the timing and extent of spending on research and development and payments the Company may receive under the Sanofi collaboration agreement or future collaboration agreements, if any. There can be no assurance that, in the event the Company requires additional financing, such financing will be available at terms acceptable to the Company, if at all. Failure to generate sufficient cash flows from operations, raise additional capital and reduce discretionary spending should additional capital not become available could have a material adverse effect on the Company's ability to achieve its business objectives.

Public offerings

In February 2020, the Company closed its initial public offering (IPO), and issued 16,100,000 shares of its common stock (including the exercise in full by the underwriters of their option to purchase an additional 2,100,000 shares of the Company's common stock) at a price to the public of \$17.00 per share for net proceeds of \$250.7 million, after deducting underwriting discounts and commissions of \$19.2 million and expenses of \$3.8 million.

In July 2020, the Company issued 6,900,000 shares of its common stock in an underwritten public offering (including the exercise in full by the underwriters of their option to purchase an additional 900,000 shares of the Company's common stock) at a price of \$26.00 per share for net proceeds of \$167.8 million, after deducting underwriting discounts and commissions of \$10.8 million and expenses of \$0.8 million.

In February 2021, the Company issued 6,666,666 shares of its common stock in an underwritten public offering (including the exercise in full by the underwriters of their option to purchase an additional 869,565 shares of the Company's common stock) at a price of \$45.00 per share for net proceeds of \$281.1 million, after deducting underwriting discounts and commissions of \$18.0 million and expenses of \$0.9 million.

2. Summary of significant accounting policies

Basis of presentation

The condensed consolidated financial statements have been prepared in conformity with generally accepted accounting principles in the United States (GAAP) and applicable rules of the Securities and Exchange Commission (SEC) regarding interim financial reporting and, in the opinion of management, include all normal and recurring adjustments which are necessary to state fairly the Company's financial position and results of operations for the reported periods. The accompanying condensed consolidated financial statements and related financial information should be read in conjunction with the audited consolidated financial statements and the related notes thereto for the year ended December 31, 2020 included in the Company's Annual Report on Form 10-K for the year ended December 31, 2020 filed with the SEC on March 2, 2021. Certain information and note disclosures normally included in the financial statements prepared in accordance with GAAP have been condensed or omitted in accordance with such rules and regulations. The condensed consolidated financial statements for the periods ended June 30, 2021 and June 30, 2020 include the accounts of the Company and its wholly owned subsidiary, Warp Drive Bio, Inc. (Warp Drive). All intercompany balances and transactions have been eliminated in consolidation. The functional and reporting currency of the Company and its subsidiary is the U.S. dollar.

Reverse stock split

On February 7, 2020, the Company amended and restated its amended and restated certificate of incorporation to effect a 1-for-4.8661 reverse stock split of the Company's common stock and redeemable convertible preferred stock. The par value and authorized



shares of the common stock and redeemable convertible preferred stock were not adjusted as a result of the reverse stock split. All issued and outstanding common stock, options to purchase common stock and per share amounts contained in the financial statements have been retroactively adjusted to give effect to the reverse stock split for all periods presented.

Use of estimates

The preparation of consolidated financial statements in conformity with GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities and disclosure of contingent assets and liabilities as of the date of the financial statements and the reported amounts of revenues and expenses during the reporting period. On an ongoing basis, management evaluates its estimates, including those related to revenue recognition, clinical accruals, valuation of in-process research and development and developed technologies, income taxes, useful lives of property and equipment and intangible assets, impairment of goodwill, and stock-based compensation. The extent to which the COVID-19 pandemic may directly or indirectly impact the Company's business, financial condition and results of operations is highly uncertain and subject to change. The Company considered the potential impact of the COVID-19 pandemic on its estimates and assumptions and there was not a material impact to the Company's estimates, and there may be changes to the estimates in future periods.

Concentration of credit risk and other risks and uncertainties

Financial instruments that potentially subject the Company to concentration of credit risk consist of cash, cash equivalents and marketable securities. The Company's cash is held by two financial institutions in the United States, which management believes to be of high credit quality. The Company invests in money market funds, U.S. government debt securities, U.S. government agency bonds, commercial paper and corporate bonds. The Company has not experienced any losses on its deposits of cash and cash equivalents.

The Company is subject to credit risk as its accounts receivable and collaboration revenue are entirely related to its collaboration agreement with Sanofi. See Note 8, "Sanofi collaboration agreement."

The Company's clinical trial sites for its RMC-4630 and RMC-5552 clinical studies may be affected by the COVID-19 outbreak due to prioritization of hospital resources toward the COVID-19 outbreak, travel or quarantine restrictions imposed by governments, and the inability to access sites for initiation and patient monitoring and enrollment. As a result, patient screening, new patient enrollment, monitoring and data collection may be affected or delayed. The Company is aware that several clinical sites involved in its RMC-4630 clinical studies temporarily stopped or delayed enrolling new patients, with exemptions if appropriate, and it is possible that these or other clinical sites may be similarly affected in the future. These developments may delay the Company's clinical trial timelines. Some of the Company's third-party manufacturers which it uses for the supply of materials for product candidates or other materials necessary to manufacture product to conduct preclinical tests and clinical trials and contract research organizations may be impacted by COVID-19, and should they experience disruptions, such as temporary closures or suspension of services, the Company would likely experience delays in advancing clinical trials.

Leases

The Company determines if an arrangement is, or contains, a lease at inception and then classifies the lease as operating or financing based on the underlying terms and conditions of the contract. Leases with terms greater than one year are initially recognized on the balance sheet as right-of-use assets and lease liabilities based on the present value of lease payments over the expected lease term. The interest rate implicit in lease contracts is typically not readily determinable. As such, the Company utilizes the incremental borrowing rate, which is the rate incurred to borrow, on a collateralized basis, an amount equal to the lease payments over a similar term and in a similar economic environment of the applicable country or region. Variable lease payments are excluded from the right-of-use assets and operating lease liabilities and are recognized in the period in which the obligation for those payments is incurred.

Recent accounting pronouncements

From time to time, new accounting pronouncements are issued by the FASB, under its ASC or other standard setting bodies, and adopted by the Company as of the specified effective date, unless otherwise discussed below.

Recently adopted accounting pronouncements

In December 2019, the Financial Accounting Standards Board (FASB) issued ASU 2019-12, *Income Taxes (Topic 740)-Simplifying the Accounting for Income Taxes* (ASU 2019-12). ASU 2019-12 simplifies the accounting for income taxes by removing certain exceptions to the general principles in Topic 740 and clarifying and amending existing guidance. This ASU is effective for the

Company for the fiscal year beginning after December 15, 2021 and interim periods within fiscal years beginning after December 15, 2021. ASU 2019-12 became effective for the Company in the first quarter of 2021 with early adoption permitted. The Company adopted the standard on January 1, 2021 and concluded that adoption of the standard did not have a material impact on its consolidated financial statements.

Recent accounting pronouncements not yet adopted

In June 2016, the FASB issued ASU No. 2016-13, *Financial Instruments—Credit Losses (Topic 326): Measurement of Credit Losses on Financial Instruments* (ASU 2016-13), which requires the measurement and recognition of expected credit losses for financial assets held at amortized cost. ASU 2016-13 replaces the existing incurred loss impairment model with an expected loss model. It also eliminates the concept of other-than-temporary impairment and requires credit losses related to available-for-sale debt securities to be recorded through an allowance for credit losses rather than as a reduction in the amortized cost basis of the securities. These changes will result in earlier recognition of credit losses. In November 2018, the FASB issued ASU No. 2018-19, *Codification Improvements to Topic 326, Financial Instruments—Credit Losses* (ASU 2018-19) which narrowed the scope and changed the effective date for non-public entities for ASU 2016-13. The FASB subsequently issued supplemental guidance within ASU No. 2019-05, *Financial Instruments—Credit Losses* (Topic 326): *Targeted Transition Relief* (ASU 2019-05). ASU 2019-05 provides an option to irrevocably elect the fair value option for certain financial assets previously measured at amortized cost basis. ASU 2016-13 is applicable to the Company for the fiscal year beginning after December 15, 2021. Early adoption is permitted. The Company is currently evaluating the impact the adoption of these ASUs will have on its consolidated financial statements and related disclosures.

In August 2018, the FASB issued ASU No. 2018-15, *Intangibles—Goodwill and Other-Internal-Use Software (Subtopic 350-40): Customer's Accounting for Implementation Costs Incurred in a Cloud Computing Arrangement That Is a Service Contract (ASU 2018-15). ASU 2018-15 aligns the requirements for capitalizing implementation costs incurred in a cloud computing arrangement that is a service contract with the requirements for capitalizing implementation costs incurred to develop or obtain internal-use-software. This ASU is effective for the Company for the fiscal year beginning after December 15, 2020, and interim periods within fiscal years beginning after December 31, 2021. The Company is currently evaluating the impact of this ASU on the Company's consolidated financial statements.*

In October 2020, the FASB issued ASU 2020-08, *Codification Improvements to Subtopic 310-20, Receivables-Nonrefundable Fees and Other Costs* (ASU 2020-08). This ASU clarifies that an entity should reevaluate whether a callable debt security is within the scope of ASC paragraph 310-20-35-33 for each reporting period. The guidance is effective for public business entities for fiscal years, and interim periods within those fiscal years, beginning after December 15, 2021. Early application is not permitted. All entities should apply ASU 2020-08 on a prospective basis as of the beginning of the period of adoption for existing or newly purchased callable debt securities. The Company is currently evaluating the impact of ASU 2020-08 on the Company's consolidated financial statements.

In October 2020, FASB issued ASU 2020-10, *Codification Improvements* (ASU 2020-10). This update contains amendments that improve the consistency of the Codification by including all disclosure guidance in the appropriate Disclosure Section (Section 50). Many of the amendments arose because the Board provided an option to give certain information either on the face of the financial statements or in the notes to financial statements and that option only was included in the Other Presentation Matters Section (Section 45) of the Codification. The option to disclose information in the notes to financial statements should have been codified in the Disclosure Section as well as the Other Presentation Matters Section (or other Section of the Codification in which the option to disclose in the notes to financial statements appears). The amendments in this Update do not change GAAP and, therefore, are not expected to result in a significant change in practice. The amendments are effective for the Company for fiscal years beginning after December 15, 2021, including interim period within those fiscal years. Early adoption is permitted. Adoption shall be applied retrospectively. The Company is currently evaluating the impacts of the provisions of ASU 2020-10 on its consolidated financial statements and related disclosures.

3. Fair value measurements

The carrying amounts of the Company's certain financial instruments, including cash equivalents, accounts payable and accrued expenses and other current liabilities approximate fair value due to their relatively short maturities and market interest rates, if applicable. Refer to Note 4 regarding the fair value of the Company's available-for-sale securities.

Assets and liabilities recorded at fair value on a recurring basis in the consolidated balance sheets are categorized based upon the level of judgment associated with the inputs used to measure their fair values. Fair value is defined as the exchange price that would be received for an asset or an exit price that would be paid to transfer a liability in the principal or most advantageous market for the asset or liability in an orderly transaction between market participants on the measurement date. Valuation techniques used to



measure fair value must maximize the use of observable inputs and minimize the use of unobservable inputs. The authoritative guidance on fair value measurements establishes a three-tier fair value hierarchy for disclosure of fair value measurements as follows:

Level 1—Observable inputs such as unadjusted, quoted prices in active markets for identical assets or liabilities at the measurement date;

Level 2—Inputs (other than quoted prices included in Level 1) are either directly or indirectly observable for the asset or liability. These include quoted prices for similar assets or liabilities in active markets and quoted prices for identical or similar assets or liabilities in markets that are not active; and

Level 3—Unobservable inputs that are supported by little or no market activity and that are significant to the fair value of the assets or liabilities.

The following table presents information about the Company's financial assets that are measured at fair value and indicates the fair value hierarchy of the valuation:

	June 30, 2021							
	Total		Level 1		Level 2			Level 3
	(in thousands)							
Assets:								
Money market funds (1)	\$	192,391	\$	192,391	\$	—	\$	_
Commercial paper (1, 2)		237,884		_		237,884		_
U.S. government and agency securities (2)		114,395		_		114,395		_
Corporate bonds ⁽²⁾		104,442		—		104,442		_
Total	\$	649,112	\$	192,391	\$	456,721	\$	

December 31, 2020							
Total		Level 1		Level 2		evel 3	
(in thousands)							
\$ 16,696	\$	16,696	\$	—	\$		
151,663		—		151,663			
270,520				270,520		_	
3,200		—		3,200		_	
\$ 442,079	\$	16,696	\$	425,383	\$		
\$	\$ 16,696 151,663 270,520 <u>3,200</u>	\$ 16,696 \$ 151,663 270,520 3,200	Total Level 1 (in tho \$ 16,696 \$ 16,696 151,663 270,520 3,200	Total Level 1 (in thousand \$ 16,696 \$ 16,696 \$ 151,663 270,520 3,200	Total Level 1 Level 2 (in thousands) (in thousands) \$ 16,696 \$ — 151,663 — 270,520 — 3,200 —	Total Level 1 Level 2 L (in thousands) (in thousands) (in thousands) \$ 16,696 \$ 16,696 \$ — \$ 151,663 \$ 270,520 — 270,520 270,520 3,200 — 3,200	

(1) (2)

Included in cash and cash equivalents on the consolidated balance sheets.

(2) Included in marketable securities on the consolidated balance sheets.

Money market funds are measured at fair value on a recurring basis using quoted prices. U.S. government debt securities, U.S. government agency bonds, commercial paper and corporate bonds are measured at fair value, which is derived from independent pricing sources based on quoted prices in active markets for similar securities.

There were no transfers between Levels 1, 2 or 3 for any of the periods presented.

4. Available-for-sale securities

The following tables summarize the estimated value of the Company's available-for-sale marketable securities and cash equivalents and the gross unrealized gains and losses:

	June 30, 2021								
	Amortized cost		Gross unrealized gain (in the		Gross unrealized loss housands)			Estimated fair value	
Marketable securities:				(in the	usanu	3)			
Commercial paper	\$	231,886	\$	16	\$	(17)	\$	231,885	
U.S. government and agency securities		114,373		22				114,395	
Corporate bonds		104,454		3		(15)		104,442	
Total marketable securities		450,713		41		(32)		450,722	
Cash equivalents:									
Money market funds		192,391		_				192,391	
Commercial paper		5,999		_				5,999	
Total cash equivalents		198,390				_		198,390	
Total available-for-sale investments	\$	649,103	\$	41	\$	(32)	\$	649,112	

	December 31, 2020							
	A	mortized cost	u	Gross nrealized gain	un	Gross realized loss		Estimated air value
Marketable securities:				(in tho	isanus)			
Commercial paper	\$	69,871	\$		\$	(5)	\$	69,866
U.S. government and agency securities		266,481		131		(5)		266,607
Total marketable securities		336,352		131		(10)		336,473
Cash equivalents:								
Money market funds		16,696		_		_		16,696
Commercial paper		81,800		_		(3)		81,797
U.S. government and agency securities		3,913		—		—		3,913
Corporate bonds		3,202				(2)		3,200
Total cash equivalents		105,611				(5)		105,606
Total available-for-sale investments	\$	441,963	\$	131	\$	(15)	\$	442,079

The amortized cost and estimated fair value of the Company's available-for-sale securities by contractual maturity are summarized below as of June 30, 2021:

		June 30, 2021							
	I	Amortized cost				realized		Estimated fair value	
				(in tho	usands))			
Mature in one year or less	\$	630,984	\$	36	\$	(32)	\$	630,988	
Mature after one year through two years		18,119		5		—		18,124	
Total marketable securities	\$	649,103	\$	41	\$	(32)	\$	649,112	

5. Balance sheet components

Property and equipment, net

Property and equipment, net consists of the following:

		June 30,	De	cember 31,
		2021		2020
)		
Laboratory equipment	\$	10,830	\$	9,978
Leasehold improvements		7,201		3,387
Computer equipment and software		2,163		1,578
Furniture and fixtures		68		48
Construction in progress		9		1,981
		20,271		16,972
Less: accumulated depreciation and amortization		(9,527)		(8,070)
Property and equipment, net	\$	10,744	\$	8,902

Depreciation and amortization expense for property and equipment amounted to \$0.8 million and \$0.6 million for the three months ended June 30, 2021 and 2020, respectively, and \$1.5 and \$1.3 million for the six months ended June 30, 2021 and 2020, respectively.

Accrued expenses and other current liabilities

Accrued expenses and other current liabilities consist of the following:

	_	June 30,	De	ecember 31,
		2021		2020
		(in thousands)		
Accrued compensation	\$	4,719	\$	7,736
Accrued research and development		16,733		10,459
Accrued professional services		552		492
Other		99		97
Total accrued expenses and other current liabilities	\$	22,103	\$	18,784

6. Intangible assets and goodwill

Intangible assets, net

Intangible assets, net consist of the following as of June 30, 2021:

	Gr	oss value	amo	umulated ortization housands)	Net book value	Weighted- average remaining useful life (in years)
In-process research and development — RAS						
Programs	\$	55,800	\$		\$ 55,800	n/a
Developed technology - tri-complex platform		7,480		(2,870)	4,610	4.3
Total	\$	63,280	\$	(2,870)	\$ 60,410	

Amortization expense for the three months ended June 30, 2021 and 2020 was \$0.3 million and \$0.3 million, respectively, and for the six months ended June 30, 2021 and 2020 was \$0.5 and \$0.5 million, respectively.

As of June 30, 2021, future amortization expense is as follows:

	A	mount
	(in t	10usands)
2021 (remaining six months)	\$	533
2022		1,069
2023		1,069
2024		1,069
2025		870
Total	\$	4,610

Intangible assets, net consist of the following as of December 31, 2020:

	Gı	oss value	am	cumulated ortization chousands)	Net book value	Weighted- average remaining useful life (in years)
In-process research and development — RAS			((,		()
Programs	\$	55,800	\$	—	\$ 55,800	n/a
Developed technology - tri-complex platform		7,480		(2,335)	5,145	4.8
Total	\$	63,280	\$	(2,335)	\$ 60,945	

Goodwill

The following summarizes the change in the carrying value of goodwill for the three months ended June 30, 2021:

	Amount
	(in thousands)
Balance at December 31, 2020	\$ 14,608
Adjustment	—
Balance at June 30, 2021	\$ 14,608

No impairment has been recognized as of June 30, 2021. Goodwill recorded is not deductible for income tax purposes.

7. Commitments and contingencies

Leases

In January 2015, as amended in September 2016, the Company entered into an operating lease for approximately 42,000 square feet of office and laboratory space located at 700 Saginaw Drive, Redwood City, California (the 700 Building), with a term through April 2023. In April 2020, the Company amended the lease to lease an additional 19,000 square feet of office, laboratory and research and development space located at 300 Saginaw Drive, Redwood City, California (the 300 Building), beginning on December 15, 2020 and ending December 31, 2030, and to extend the Company's existing lease term for the 700 Building until December 31, 2030. The Company has the option to extend the lease term for the 700 Building and the 300 Building together for an additional ten years after December 31, 2030.

The annual base rent for the lease of the 300 Building is \$1.2 million until December 31, 2021, after which the annual base rent will increase by approximately 3.5% in each subsequent year of the lease term. The annual base rent for the lease of the 700 Building remains unchanged through April 30, 2023, and increases to \$2.8 million for the 12 month period ending April 30, 2024, after which the annual base rent increases by approximately 3.5% in each subsequent year of the lease term.

In connection with the lease amendment, the Company issued a letter of credit for \$0.9 million, which is included in restricted cash on the consolidated balance sheet as of June 30, 2021 and December 31, 2020, respectively.

Through June 30, 2021, the landlord has provided the Company with \$3.4 million in tenant improvement allowances for the 700 Building and \$3.2 million for the 300 building, which were recognized as lease incentives. The lease incentives are being amortized as an offset to rent expense over the lease term in the condensed consolidated statements of operations.

Upon the execution of the lease amendment, which was deemed to be a lease modification, the Company re-evaluated the assumptions used during the adoption of ASC 842 for the lease. The Company determined the amendment consists of two separate contracts under ASC 842. One contract is related to a new right-of-use asset for the 300 Building, which is being accounted for as an operating lease, and the other is related to the modification of the original lease term for the 700 Building. As a result, the Company recorded a right-of-use asset of \$6.4 million and a lease liability of \$9.0 million for the 300 Building and an increase of \$14.8 million to the right-of-use asset and lease liability for the 700 Building upon execution of the lease amendment. The Company is recognizing rent expense for both buildings on a straight-line basis through the remaining extended term of the respective leases.

As part of the Warp Drive acquisition in October 2018, the Company assumed an operating lease for approximately 22,000 square feet of office and laboratory space located in Cambridge, Massachusetts (the Cambridge Lease), which expires in February 2023, with an option to extend the term through February 2028, subject to certain conditions. In March 2019, the Company fully subleased the Cambridge Lease to Casma Therapeutics, Inc. (Casma), a related party, on financial terms substantially the same as the original lease. The sublease term with Casma is through the remainder of the Cambridge Lease term. The sublease by Casma and related sublease payments by Casma to the Company are fully guaranteed by Third Rock Ventures, LLC, a related party. In conjunction with the Cambridge Lease, the Company issued a letter of credit for \$0.2 million, which is included in restricted cash on the condensed consolidated balance sheets as of June 30, 2021 and December 31, 2020.

The balance sheet classification of the Company's lease liabilities was as follows:

]	4,926 27,566	De	cember 31,
		2021		2020
		(in tho	usands)	
Operating lease liabilities:				
Operating lease liability – current	\$	4,926	\$	3,672
Operating lease liability – noncurrent		27,566		28,992
Total operating lease liabilities		32,492		32,664
Financing lease liabilities:				
Accrued expenses and other current liabilities				19
Total financing lease liabilities		_		19
Total lease liabilities	\$	32,492	\$	32,683

For the three months ended June 30, 2021 and 2020, operating lease cost was \$0.5 million and \$0.7 million, respectively, net of sublease income of \$0.5 million and \$0.5 million, respectively, and tenant improvement allowance credits of \$0.1 million and \$0.1 million, respectively. For the six months ended June 30, 2021 and 2020, operating lease cost was \$1.3 and \$1.1 million, respectively, net of sublease income was \$1.1 and \$0.9 million, respectively, and tenant improvement allowance credits of \$0.5 million and \$(0.5) million for the three months ended June 30, 2021 and 2020, respectively. The operating cash flows provided by (used in) operating leases were \$0.5 million and \$(0.5) million for the three months ended June 30, 2021 and 2020, respectively. The operating cash flows used in operating leases were \$0.2 and \$1.3 million for the six months ended June 30, 2021 and 2020, respectively. Short-term lease costs were immaterial for the three and six months ended June 30, 2021 and 2020, respectively.

As of June 30, 2021, the maturities of the Company's operating lease liabilities were as follows (in thousands):

2021 (remaining six months)	\$ 2,508
2022	5,144
2023	4,173
2024	4,215
2025	4,363
Thereafter	24,213
Total undiscounted lease payments	\$ 44,616
Less: Imputed interest	(12,124)
Total operating lease liabilities	\$ 32,492

The amounts reflected in the table above include the Company's lease payments for the Cambridge lease, but do not reflect any offset for the sublease payments the Company is entitled to receive from Casma.



Operating lease liabilities are based on the net present value of the remaining lease payments over the remaining lease term. In determining the present value of lease payments, the Company uses its incremental borrowing rate. The weighted-average discount rate used to determine the operating lease liability was 7.1%. As of June 30, 2021 and December 31, 2020, the weighted-average remaining lease term was 8.8 years and 9.1 years, respectively.

Legal matters

From time to time, the Company may be involved in litigation related to claims that arise in the ordinary course of its business activities. The Company accrues for these matters when it is probable that losses will be incurred and these losses can be reasonably estimated. As of June 30, 2021 and December 31, 2020, respectively, the Company does not believe that any such matters, individually or in the aggregate, will have a material adverse effect on the Company's financial position, results of operations or cash flows.

Indemnification

The Company enters into standard indemnification arrangements in the ordinary course of business. Pursuant to these arrangements, the Company indemnifies, holds harmless and agrees to reimburse the indemnified parties for losses suffered or incurred by the indemnified party, in connection with any trade secret, copyright, patent or other intellectual property infringement claim by any third party with respect to its technology. The term of these indemnification agreements is generally perpetual any time after the execution of the agreement. The maximum potential amount of future payments the Company could be required to make under these arrangements is not determinable. The Company has not incurred costs to defend lawsuits or settle claims related to these indemnification agreements. As a result, the Company believes the fair value of these agreements is minimal.

8. Sanofi collaboration agreement

In June 2018, the Company entered into a collaborative research, development and commercialization agreement (the Sanofi Agreement) with Aventis, Inc. (an affiliate of Sanofi) to research and develop SHP2 inhibitors, including RMC-4630, for any indications. The Sanofi Agreement was assigned to Genzyme Corporation, a Sanofi affiliate, in December 2018. Pursuant to the Sanofi Agreement, the Company granted Sanofi a worldwide, exclusive, sublicensable (subject to the Company's consent in certain circumstances) license under certain of the Company's patents and know-how to research, develop, manufacture, use, sell, offer for sale, import and otherwise commercialize SHP2 inhibitors, including RMC-4630, for any and all uses, subject to the Company's exercise of rights and performance obligations under the Sanofi Agreement.

In October 2018, the Company acquired Warp Drive in exchange for issuance of shares of the Company's Series B redeemable convertible preferred stock and payment of cash. Sanofi was a stockholder of Warp Drive and received the Company's Series B redeemable convertible preferred stock during the transaction and accordingly became an investor and related party of the Company. As a result of the Company's underwritten offering of common stock in February 2021, Sanofi's ownership percentage in the Company decreased and Sanofi is no longer considered a related party.

Under the Sanofi Agreement, the Company received a non-refundable, upfront cash payment of \$50 million in July 2018 and could also receive up to \$520 million in development and regulatory milestone payments, including up to \$235 million upon the achievement of specified development milestones and up to \$285 million upon the achievement of certain marketing approval milestones.

The Company has primary responsibility for early clinical development and for the manufacture of SHP2 inhibitors for Phase 1 and nonregistrational Phase 2 clinical trials of RMC-4630 pursuant to an initial development plan. A joint research and development committee is responsible for preparing development plans for other SHP2 inhibitors approved by this committee for development, if any. Sanofi is responsible to reimburse the Company all internal and external costs and expenses to perform the Company's activities under approved development plans, except for costs and expenses related to studies designated in the Sanofi Agreement as RevMed Studies, for which the Company will bear all costs and expenses, and for the planned RMC-4630-03 study, for which Sanofi will reimburse the Company for 50% of the costs and expenses. Unreimbursed costs incurred by the Company for any RevMed Studies or for the RMC-4630-03 study are subject to future reimbursement by Sanofi through a buy-in payment pursuant to the terms of the Sanofi Agreement if Sanofi uses the data from a RevMed Study or the RMC-4630-03 study in support of a marketing approval application. There currently are no active RevMed Studies.

The Company is also primarily responsible for performing preclinical research on SHP2 inhibitors, pursuant to a research plan that is currently approved through 2021. The research plan and budget beyond 2021 will be determined by a joint research and development committee, over which Sanofi has final decision-making power subject to certain exceptions. Sanofi is responsible to reimburse the Company for all internal and external costs and expenses incurred to perform activities under the approved research plans, with the exception of internal and external research costs and expenses under the approved research plans for 2019 and 2020,

for which Sanofi was obligated to reimburse the Company for 80% of such costs. The Company was responsible for 20% of all internal and external research costs incurred under the research plans for 2019 and 2020. Sanofi is responsible to reimburse the Company for all internal and external costs and expenses incurred under the research plan for 2021.

In the United States, the Company will share equally with Sanofi the profits and losses applicable to commercialization of SHP2 inhibitor products, pursuant to a profit/loss share agreement that the parties will negotiate based on key terms agreed in the Sanofi Agreement. On a product-by-product basis, Sanofi will also be required to pay the Company tiered royalties on annual net sales of each product outside the United States ranging from high single digit to mid-teen percentages.

Unless terminated earlier, the Sanofi Agreement will continue in effect until the later of the expiration of all of Sanofi's milestone and royalty payment obligations and the expiration of the profit/loss share agreement. Sanofi may terminate the Sanofi Agreement in its entirety or on a country-by-country or product-by-product basis for any reason or for significant safety concerns, upon prior notice to the Company. Sanofi may terminate the Sanofi Agreement if an undisputed material breach by the other party is not cured within a defined period of time, or immediately upon notice for insolvency-related events of the other party. The Company may terminate the Sanofi Agreement after a certain number of years if Sanofi develops a competing program without commencing a registrational clinical trial for a SHP2 inhibitor product candidate, and subject to certain other conditions. The Company may also terminate the Sanofi Agreement at any time, if Sanofi ceases certain critical activities for SHP2 inhibitor product candidates for more than a specified period of time, provided that such cessations of critical activity were not a result of certain specified factors, and subject to certain other conditions. Upon any termination of the Sanofi Agreement with respect to any product or country, all licenses to Sanofi with respect to such product or country shall automatically terminate and all rights generally revert back to the Company.

The Company identified the following promises in the agreement (1) the license related to SHP2 inhibitors, (2) the performance of research and development services for Phase 1 clinical studies and Phase 2 clinical trials that are non-registrational clinical trials and (3) the performance of manufacturing services for the non-registrational clinical trials. The Company determined that the license is not distinct from the services within the context of the agreement because the research, development and manufacturing significantly increase the utility of the intellectual property. The intellectual property (IP) related to SHP2 inhibitors, which is proprietary to the Company, is the foundation for the research and development activities. The manufacturing services are a necessary and integral part of the research and development services as they could only be conducted utilizing the outcomes of these services. Given the research and development services under the Sanofi Agreement are expected to involve significant further development of the initial IP, the Company has concluded that the research, development and manufacturing services are not distinct from the license, and thus the license, research and development services are combined into a single performance obligation.

For revenue recognition purposes, the Company determined that the duration of the contract begins on the effective date of the Sanofi Agreement in July 2018 and ends upon completion of the non-registrational clinical trials. The contract duration is defined as the period in which parties to the contract have present enforceable rights and obligations. The Company analyzed the impact of Sanofi terminating the agreement prior to the completion of these trials and determined that there were significant economic costs to Sanofi for doing so.

The Company determined that the transaction price of the Sanofi Agreement was \$196.1 million as of June 30, 2021. In order to determine the transaction price, the Company evaluated all the payments to be received during the duration of the contract. The Company determined that the \$50.0 million upfront payment and \$146.1 million of estimated variable consideration for expense reimbursements from Sanofi for agreed upon research and development services as of June 30, 2021 constituted consideration to be included in the transaction price, which is to be allocated to the combined performance obligation. Development and regulatory milestones under the Sanofi Agreement were considered but not included in the transaction price, as it is probable that a significant revenue reversal could occur if they were included. The Company will re-evaluate the transaction price in each reporting period and as uncertain events are resolved or other changes in circumstances occur.

The license, research, development and manufacturing services are combined as one performance obligation that will be performed over the duration of the contract, which is from the effective date of the Sanofi Agreement through to the completion of studies. The Company concluded that it would utilize a cost-based input method to measure proportional performance and to calculate the corresponding amount of revenue to recognize. In applying the cost-based input method of revenue recognition, the Company uses actual costs incurred relative to estimated costs to fulfill the combined performance obligation. These costs consist primarily of internal full-time equivalent efforts and third-party costs. Revenue is recognized based on actual costs incurred as a percentage of total estimated costs as the Company completes its performance obligations. The cumulative effect of revisions to estimated costs to complete the Company's performance obligations will be recorded in the period in which changes are identified and amounts can be reasonably estimated.

During the three months ended June 30, 2021 and 2020, the Company recognized \$8.7 million and \$10.0 million of collaboration revenue associated with this agreement, respectively, and \$18.8 million and \$21.6 million for the six months ended June 30, 2021 and 2020, respectively.

As of June 30, 2021 and December 31, 2020, \$8.3 million and \$12.1 million of deferred revenue is classified as current and \$7.6 million and \$8.5 million is classified as noncurrent.

9. Redeemable convertible preferred stock

Upon the closing of the IPO in February 2020, all shares of redeemable convertible preferred stock then outstanding converted into 39,600,423 shares of common stock. There were no shares of redeemable convertible preferred stock outstanding as of June 30, 2021 and December 31, 2020, respectively.

10. Common stock

As of June 30, 2021 and December 31, 2020, the Company's certificate of incorporation authorized the Company to issue 300,000,000 shares of common stock, respectively, at a par value of \$0.0001 per share. Each share of common stock is entitled to one vote. The holders of common stock are also entitled to receive dividends whenever funds are legally available and when declared by the Board of Directors, subject to prior rights of the redeemable convertible preferred stockholders. As of June 30, 2021, no dividends have been declared to date.

The Company has reserved shares of common stock, on an as-converted basis, for future issuance as follows:

	June 30,	December 31,
	2021	2020
Outstanding options to purchase common stock	6,012,477	5,118,979
Unvested restricted stock units of common stock	383,768	85,639
Available for future issuance under the 2020 Incentive Award Plan	6,680,902	4,806,916
Available for issuance under the 2020 Employee Stock Purchase Plan	1,119,518	499,722
Total	14,196,665	10,511,256

11. Stock-based compensation

2020 Incentive Award Plan

In February 2020, the Company adopted the 2020 Equity Incentive Plan (2020 Plan). The 2020 Plan became effective on February 11, 2020. The 2020 Plan provides for a variety of stock-based compensation awards, including stock options, stock appreciation rights, restricted stock awards, restricted stock unit awards, performance bonus awards, performance stock unit awards, dividend equivalents, or other stock or cash based awards. Under the 2020 Plan, the Company generally grants stock-based awards with service-based vesting conditions only. Options and restricted stock unit awards granted typically vest over a four-year period, but may be granted with different vesting terms.

Following the effectiveness of the 2020 Plan, the Company will not make any further grants under the 2014 Equity Incentive Plan (2014 Plan). However, the 2014 Plan will continue to govern the terms and conditions of the outstanding awards granted under it. Shares of common stock subject to awards granted under the 2014 Plan that are forfeited or lapse unexercised and which following the effective date of the 2020 Plan are not issued under the 2014 Plan will be available for issuance under the 2020 Plan. As of June 30, 2021, there were 6,680,902 shares of common stock reserved for issuance pursuant to the 2020 Plan.

2014 Equity Incentive Plan

In December 2014, the Company adopted the 2014 Plan which provided for the Company to issue restricted common stock, or to grant incentive stock options or nonqualified stock options for the purchase of common stock, to employees, members of the Board of Directors and consultants of the Company under terms and provisions established by the Board of Directors. The Company generally granted stock-based awards with service-based vesting conditions only. Options granted typically vest over a four-year period but may be granted with different vesting terms.

The Company allows its employees, non-employees and directors to exercise options granted under the 2014 Plan prior to vesting. The shares related to early exercised stock options are subject to the Company's lapsing repurchase right upon termination of employment at the original purchase price. In order to vest, the holders are required to provide continued service to the Company. The proceeds are initially recorded in other noncurrent liabilities and are reclassified to common stock and additional paid-in capital as the repurchase right lapses. As of June 30, 2021 and December 31, 2020, there were 71,549 and 130,793 shares, respectively, and \$0.2 million and \$0.2 million, respectively, recorded in other noncurrent liabilities, related to early exercised shares that were subject to repurchase.

2020 Employee Stock Purchase Plan

In February 2020, the Company adopted the 2020 Employee Stock Purchase Plan (2020 ESPP). Under the 2020 ESPP, employees have the ability to purchase shares of the Company's common stock through payroll deductions at a discount during a series of offering periods of 24 months, each comprised of four six-month purchase periods. The purchase price will be the lower of 85% of the closing trading price per share of the Company's common stock on the first day of an offering period in which an employee is enrolled or 85% of the closing trading price per share on the purchase date, which will occur on the last trading day of each purchase period.

As of June 30, 2021, there have been 75,438 shares purchased under the 2020 ESPP and a total of 1,119,518 shares of common stock were available for future issuance under the ESPP. As of June 30, 2021, there was \$2.4 million of unrecognized compensation cost related to the ESPP.

Stock options

The following summarizes option activity under both the 2020 Plan and the 2014 Plan:

	Number of Shares underlying options	Weighted- average exercise price		Weighted- average remaining contractual <u>term</u> (in years)	 Aggregate intrinsic value i thousands)
Balance, December 31, 2020	5,118,979	\$	7.37	8.25	\$ 165,088
Options granted	1,203,786		40.62		
Options exercised	(231,759)		3.84		
Options cancelled	(78,529)		26.68		
Balance, June 30, 2021	6,012,477	\$	13.91	8.15	\$ 119,111
Options vested and expected to vest as of June 30, 2021	6,012,477	\$	13.91	8.15	\$ 119,111
Options vested and exercisable as of June 30, 2021	2,475,472	\$	6.20	7.46	\$ 64,138

As of June 30, 2021, there was \$46.2 million of unrecognized stock-based compensation expense related to unvested stock options that is expected to be recognized over a weighted-average period of 2.5 years.

Restricted stock units

Restricted stock units (RSUs) have been granted to employees and directors. The value of an RSU award is based on the Company's stock price on the date of grant. The shares underlying the RSU awards are not issued until the RSUs vest. Upon vesting, each RSU converts into one share of the Company's common stock. The Company has granted RSUs pursuant to the 2020 Plan.

Activity under the 2020 Plan with respect to the Company's RSUs during the three months ended June 30, 2021 was as follows:

Number of Shares	av grant	erage date fair	weighted- average remaining contractual <u>term</u> (in years)	i	ggregate ntrinsic value thousands)
85,639	\$	35.13	1.73	\$	3,390
343,797		40.63			
(32,615)		39.27			
(13,053)		35.29			
383,768	\$	39.70	1.87	\$	12,181
383,768	\$	39.70	1.87	\$	12,181
	Shares 85,639 343,797 (32,615) (13,053) 383,768	Number of Shares average grant value 85,639 \$ 343,797 (32,615) (13,053) (3383,768)	Shares value per share 85,639 \$ 35.13 343,797 40.63 (32,615) 39.27 (13,053) 35.29 383,768 \$ 39.70	Weighted-average average grant date fair value per shareaverage remaining contractual term (in years)85,639\$ 35.131.7385,639\$ 35.131.73343,79740.63(32,615)39.27(13,053)35.29383,768\$ 39.701.87	Weighted-average average grant date fair value per shareaverage remaining contractual termA i i85,639\$ 35.131.73\$343,79740.63\$(32,615)39.27\$(13,053)35.29\$383,768\$ 39.701.87\$

The number of RSUs vested includes shares of common stock that the Company withheld to satisfy the minimum statutory tax withholding requirements. As of June 30, 2021, there was \$14.8 million of total unrecognized compensation cost related to RSUs that is expected to be recognized over a weighted average period of 3.5 years.

Stock-based compensation expense

Total stock-based compensation expense related to stock options, RSUs and the 2020 ESPP by function was as follows:

	Th	ree Month 3	s En 0,	led June	Six Months Ended June 30,				
		2021	2020		2021	2020			
		(in tho	isano	ls)		(in tho	usands)		
Research and development	\$	3,113	\$	1,170	\$	4,989	\$	2,063	
General and administrative		2,200		847		3,711		1,521	
Total	\$	5,313	\$	2,017	\$	8,700	\$	3,584	

Stock-based compensation related to options granted to non-employees was \$0.1 million and \$0.2 million for the three months ended June 30, 2021 and 2020, respectively, and \$0.3 million and \$0.4 million for the six months ended June 30, 2021 and 2020, respectively.

12. Net loss per share attributable to common stockholders

The following table sets forth the computation of basic and diluted net loss per share attributable to common stockholders:

		Three Months l	l June 30,		Six Months E	June 30,			
		2021 2020				2021	2020		
	(in t	(in thousands, except share and per share data)			(in	thousands, except dat			
Numerator:									
Net loss	\$	(44,299)	\$	(27,215)	\$	(81,475)	\$	(46,734)	
Redeemable convertible preferred stock dividends-									
undeclared and cumulative								(2,219)	
Net loss attributable to common stockholders	\$	\$ (44,299)		(27,215)	\$	(81,475)	\$	(48,953)	
Denominator:									
Weighted-average shares outstanding		73,480,020		59,013,236		72,005,252		44,314,960	
Less: Weighted-average unvested restricted shares and									
shares subject to repurchase		(80,306)		(260,742)		(87,744)		(289,588)	
Weighted-average shares used to compute net loss per share									
attributable to common stockholders-basic and diluted		73,399,714		58,752,494		71,917,508		44,025,372	
Net loss per share attributable to common stockholders-basic									
and diluted	\$	(0.60)	\$	(0.46)	\$	(1.13)	\$	(1.11)	
	20								

The following outstanding potentially dilutive shares have been excluded from the calculation of diluted net loss per share for the periods presented due to their anti-dilutive effect:

	As of Ju	ne 30,
	2021	2020
Options to purchase common stock	6,012,477	5,691,470
Options early exercised subject to future vesting	71,549	233,361
Unvested restricted stock units of common stock	383,768	48,660
Expected shares to be purchased under ESPP	146,961	110,341
Total	6,614,755	6,083,832

13. Related party relationships

Following the Company's acquisition of Warp Drive, in January 2019, the Company entered into a sublease agreement with Casma to sublease the Cambridge Lease. The sublease by Casma and related sublease payments by Casma to the Company are fully guaranteed by an affiliate of Third Rock Ventures.

14. Subsequent Events

In July 2021, the Company and Sanofi determined to deprioritize the RMC-4630-02 study and are no longer enrolling patients in the study.

In August 2021, the Company entered into a letter agreement with Sanofi, adding the RMC-4630-03 study to the development plan under the Sanofi Agreement. Under the letter agreement, Sanofi will reimburse the Company for 50% of the costs and expenses of the RMC-4630-03 study. Unreimbursed costs incurred by the Company for the RMC-4630-03 study are subject to future reimbursement by Sanofi through a buy-in payment pursuant to the terms of the Sanofi Agreement, if Sanofi uses the data from the study in support of a marketing approval application.

Item 2. Management's Discussion and Analysis of Financial Condition and Results of Operations.

You should read the following discussion and analysis of our financial condition and results of operations in conjunction with the condensed consolidated financial statements and the related notes included elsewhere in this Quarterly Report on Form 10-Q. In addition to historical financial information, this discussion contains forward-looking statements based upon current expectations that involve risks and uncertainties. As a result of many factors, including those factors set forth in the "Risk Factors" section of this report, our actual results could differ materially from the results described or implied by the forward-looking statements contained in the following discussion and analysis.

Overview

We are a clinical-stage precision oncology company focused on developing targeted therapies to inhibit frontier targets in RAS-addicted cancers. We possess sophisticated structure-based drug discovery capabilities built upon deep chemical biology and cancer pharmacology know-how and innovative, proprietary technologies that enable the creation of small molecules tailored to unconventional binding sites. Our understanding of genetic drivers and adaptive resistance mechanisms in cancer, coupled with robust drug discovery and medicinal chemistry capabilities, has guided us to establish a deep pipeline targeting critical signaling nodes within the RAS pathway and associated pathways. This cohesive approach underpins our clinical strategy of exploring mechanism-based dosing paradigms and in-pathway combinations to optimize treatment for cancer patients.

Our research and development pipeline comprises RAS(ON) inhibitors that bind directly to RAS variants, which we refer to as RAS(ON) Inhibitors, and RAS companion inhibitors that target key nodes in the RAS pathway or associated pathways, which we refer to as RAS Companion Inhibitors. Our RAS Companion Inhibitors (e.g., SHP2, mTORC1 and SOS1 inhibitors) are designed primarily for combination treatment strategies involving one or more therapeutic agents, which may include our RAS(ON) Inhibitors. Our long-term goal is to combine selected RAS Companion Inhibitors with RAS(ON) Inhibitors on behalf of patients based on molecular tumor features.

RAS(ON) Inhibitors

Our RAS(ON) Inhibitors are based on our proprietary tri-complex technology platform, which enables a highly differentiated approach to inhibiting the active, GTP-bound form of RAS (RAS(ON)). We are developing a portfolio of compounds that we believe are the first and only RAS(ON) inhibitors to use this mechanism of action. RMC-6291, our inhibitor targeting KRAS^{G12C}/NRAS^{G12C}(ON), and RMC-6236, our inhibitor of multiple RAS variants, which we refer to as RASMULTI(ON), are each in IND-enabling preclinical development. We currently plan to submit INDs for RMC-6291 and RMC-6236 in the first half of 2022. In addition, we have inhibitors targeting KRAS^{G13C}(ON) and KRAS^{G12D}(ON) in the lead optimization stage of preclinical development. We are focused on developing additional mutant-selective inhibitors and plan to nominate a third RAS(ON) Inhibitor as a development candidate in the second half of 2021.

RAS Companion Inhibitors

RMC-4630

Our RAS Companion Inhibitor RMC-4630 is designed as a potent and selective inhibitor of SHP2, a central node in the RAS signaling pathway. In collaboration with Sanofi, we are evaluating RMC-4630 in a multi-cohort Phase 1/2 clinical program.

Our highest priority hypothesis regarding RMC-4630 is that it can deliver additional clinical benefit to patients with RAS-addicted tumors treated with a RAS inhibitor. There are currently two active or planned clinical studies of RMC-4630 in combination with RAS inhibitors:

(1) an active Amgen-sponsored Phase 1b study of RMC-4630 in combination with Amgen's KRAS^{G12C}(OFF) inhibitor, sotorasib (LumakrasTM); and

(2) RMC-4630-03, a planned Phase 2 study of RMC-4630 in combination with sotorasib that we are sponsoring under a clinical trial collaboration and supply agreement with Amgen under our global partnership with Sanofi.

Amgen is currently evaluating RMC-4630 in an active Phase 1b study in combination with Amgen's KRASG12C(OFF) agent sotorasib in Amgen's CodeBreaK 101c study. This study is currently dosing patients at the RMC-4630 target dose of 200 mg on a D1D2 weekly schedule (i.e., the full dose used by us in monotherapy) in combination with sotorasib 960 mg on a daily schedule. We look forward to selection of a combination dose for this study in the second half of 2021.



As a complement to the CodeBreaK 101c study, we are sponsoring RMC-4630-03, which is an additional planned global Phase 2 study of RMC-4630 in combination with sotorasib. RMC-4630-03 will enroll patients with non-small cell lung cancer (NSCLC) carrying a KRAS^{G12C} mutation who have failed prior standard therapy and who have not previously been treated with a RAS inhibitor. We currently expect enrollment in this study to begin in the second half of 2021 and to have preliminary findings by the end of 2022.

AstraZeneca has informed us that, for reasons unrelated to RMC-4630, they have made a strategic decision not to move forward with a clinical combination study incorporating RMC-4630. Therefore, we no longer expect to provide RMC-4630 for this study.

There are also four other active, planned or recently deprioritized clinical studies of RMC-4630 that do not involve combinations with RAS inhibitors:

- (1) RMC-4630-01, an active Phase 1 study of RMC-4630 as monotherapy;
- (2) a Sanofi-sponsored active Phase 1/2 study of RMC-4630 in combination with the PD-1 inhibitor pembrolizumab (Keytruda®);
- (3) a combination of RMC-4630 with an ERK inhibitor in patients with pancreatic cancer as part of an investigator sponsored study planned by Netherlands Cancer Institute; and
- (4) the recently deprioritized RMC-4630-02 study, which is a Phase 1b/2 study that included an arm studying RMC-4630 in combination with the MEK inhibitor cobimetinib (Cotellic®) and an arm studying RMC-4630 in combination with the EGFR inhibitor osimertinib (Tagrisso®).

RMC-4630-01 is an ongoing Phase 1 study in patients with advanced cancers, including those with tumors harboring genetically defined mutations in the RAS signaling pathway. This study is evaluating the safety, pharmacokinetics and pharmacodynamic effects of RMC-4630 as a single agent at the recommended phase 2 dose and schedule (RP2DS) of 200 mg administered on Day 1 and Day 2 (D1D2) weekly. We are evaluating this dose and schedule in an expansion cohort of the RMC-4630-01 study, consisting of patients with gynecologic tumors harboring NF1^{LOF} mutations.

RMC-4630 is being evaluated by Sanofi in a global Phase 1/2 study in combination with the PD-1 inhibitor pembrolizumab (Keytruda®). Sanofi has selected a RP2DS and preparation is underway for Phase 2 expansion focused on evaluating this combination as front-line treatment for NSCLC patients whose tumors express PD-L1.

RMC-4630-02 is a Phase 1b/2 study of RMC-4630 that includes an arm that studied RMC-4630 in combination with the MEK inhibitor cobimetinib in patients with advanced cancers that harbor mutations in the RAS signaling pathway. This study evaluated the safety, tolerability, anti-tumor activity, pharmacokinetic and pharmacodynamic profiles of RMC-4630 and cobimetinib at the RP2DS of RMC-4630 140 mg and cobimetinib 40 mg, administered on a D1D2 weekly schedule, in expansion cohorts of patients with colorectal cancer (CRC) harboring KRAS^{G12V} or KRAS^{G12D} mutations and others drawing from a broader set of histotypes (including NSCLC) and RAS pathway genotypes. In a group of NSCLC patients in this arm of the study treated with the drug combination using the RP2DS we had described in 2020, we had 11 efficacy evaluable subjects. We observed acceptable tolerability in this group, and one patient with a KRASG12V tumor mutation and gene amplification exhibited a confirmed partial response with 44.7% tumor volume reduction. In a group of CRC patients, we had 25 efficacy evaluable subjects. In this group, we observed acceptable tolerability, and the best clinical response was stable disease. Although we will observe and support patients who are part of the study, we and Sanofi determined that these data showed insufficient clinical benefit to warrant enrolling new patients in this study and the study has been deprioritized.

The EGFR inhibitor arm of RMC-4630-02 evaluated RMC-4630 in combination with the EGFR inhibitor osimertinib (Tagrisso®) in patients with EGFR-positive NSCLC. The arm was designed to evaluate the safety, tolerability, and pharmacokinetics of RMC-4630 and osimertinib with different doses of RMC-4630, administered on a D1D2 weekly schedule, with the approved dose of osimertinib. Overlapping on-pathway toxicities with the combination of RMC-4630 and osimertinib were anticipated and observed. As with MEK inhibitors, many EGFR inhibitors are less selective for a mutated cancer driver than are targeted inhibitors for other drivers, such as sotorasib for KRASG^{12C}, and thus the combination of EGFR antagonists with SHP2 inhibitors presents a potential clinical development challenge. We and Sanofi did not identify a combination dose and schedule with acceptable tolerability, indicating that the combined suppression of RAS signaling in normal tissues caused by these two agents presents too big a hurdle to justify advancing this approach. Based on these data, no further patients will be enrolled in this study and the study has been deprioritized.

RMC-5552

Our RAS Companion Inhibitor RMC-5552 is designed as a selective inhibitor of hyperactivated mTORC1 signaling in tumors. We are evaluating RMC-5552 first as a monotherapy in a Phase 1 study (RMC-5552-01), and plan to evaluate RMC-5552 in combination with RAS inhibitors for patients with cancers harboring a RAS mutation and co-occurring mutations in the mTOR signaling pathway. The RMC-5552-01 study is ongoing, with the first patient dosed in April 2021. We expect initial safety, pharmacokinetic and single agent activity data for this compound in 2022.

RMC-5845

Our RAS Companion Inhibitor RMC-5845 targets SOS1, a protein that plays a key role in converting RAS(OFF) to RAS(ON) in cells. RMC-5845 is in the IND-enabling stage of preclinical development and is intended for select combination therapies for certain genetically-defined tumors. We currently expect RMC-5845 to be IND-ready in the second half of 2021, but because we have prioritized other activities, we no longer expect to submit an IND for this compound in 2021 and will continue to evaluate this potential IND along with our other assets.

Collaboration agreement with Sanofi

In June 2018, we entered into a collaborative research, development and commercialization agreement (the Sanofi Agreement) with Aventis, Inc. (an affiliate of Sanofi), to research and develop SHP2 inhibitors, including RMC-4630, for any indications. The Sanofi Agreement was assigned to Genzyme Corporation, a Sanofi affiliate, in December 2018. For the purposes of this discussion, we refer to Genzyme Corporation as Sanofi. Pursuant to the Sanofi Agreement, we granted Sanofi a worldwide, exclusive, sublicensable (subject to our consent in certain circumstances) license under certain of our patents and know-how to research, develop, manufacture, use, sell, offer for sale, import and otherwise commercialize SHP2 inhibitors, including RMC-4630, for any and all uses, subject to our exercise of rights and performance of obligations under the Sanofi Agreement. Such intellectual property exclusively licensed to Sanofi includes our interest under any of our solely-owned or jointly-owned inventions arising out of activities undertaken pursuant to the development of SHP2 inhibitor product candidates under the Sanofi Agreement.

Under the Sanofi Agreement, we have primary responsibility for early clinical development of RMC-4630 pursuant to a development plan that is currently approved through 2021. The development plan and budget beyond 2021 will be determined by a joint research and development committee, over which Sanofi has final decision-making power subject to certain exceptions. Sanofi is



responsible to reimburse us all internal and external costs and expenses to perform our activities under approved development plans, except for costs and expenses related to studies designated in the Sanofi Agreement as RevMed Studies, for which we will bear all costs and expenses, and for the planned RMC-4630-03 study, for which we have agreed that Sanofi will reimburse us for 50% of the costs and expenses. Unreimbursed costs borne by us for any RevMed Studies and for our 50% share of the RMC-4630-03 collaboration study are subject to future reimbursement by Sanofi through a buy-in payment pursuant to the terms of the Sanofi Agreement if they use data from a RevMed Study or the RMC-4630-03 study in support of a marketing approval application. There currently are no active RevMed Studies.

We are responsible for the manufacture of SHP2 inhibitors for Phase 1 and non-registrational Phase 2 clinical trials at Sanofi's cost, while Sanofi is responsible for manufacturing SHP2 inhibitors for all other clinical trials and commercial supply. Sanofi has the sole right and responsibility to perform all regulatory activities under the Sanofi Agreement, except with respect to certain trials conducted by us or otherwise conducted under our IND, including our current clinical trials evaluating RMC-4630. Once we have completed all clinical trials for a product candidate that are assigned to us under a development plan, all regulatory approvals for such product candidate are automatically assigned to Sanofi.

We are also primarily responsible for performing preclinical research on SHP2 inhibitors, pursuant to a research plan that is currently approved through 2021. The research plan and budget beyond 2021 will be determined by a joint research and development committee, over which Sanofi has final decision-making power subject to certain exceptions. Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred to perform activities under approved research plans, with the exception of internal and external research costs and expenses under approved research plans for 2019 and 2020, for which Sanofi was obligated to reimburse us for 80% of such costs. We were responsible for 20% of all internal and external research costs incurred under the research plans for 2019 and 2020. Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred under the research plans for 2019 and 2020. Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred under the research plans for 2019 and 2020. Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred under the research plans for 2019 and 2020. Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred under the research plan for 2021.

Unless otherwise delegated to us by the joint commercialization committee, Sanofi also has the sole right and responsibility for all aspects of the commercialization of SHP2 inhibitors in the world for any and all uses, at its expense, subject to our right to elect to co-promote SHP2 inhibitors in the United States. Sanofi is obligated to use commercially reasonable efforts to seek marketing approval for at least one SHP2 inhibitor product candidate in certain major market countries. Sanofi agrees to provide us, and we agree to provide Sanofi, with research, development and commercialization updates through the joint committees.

During the term of the Sanofi Agreement, we may not, alone or with any affiliate or third party, conduct certain research activities with respect to, or develop or commercialize, any product that contains a SHP2 inhibitor outside of the Sanofi Agreement.

Pursuant to the Sanofi Agreement, we received an upfront payment of \$50 million from Sanofi in July 2018. Upon the achievement of specified development and regulatory milestones, Sanofi will be obligated to pay us up to \$520 million in the aggregate, including up to \$235 million upon the achievement of specified development milestones and up to \$285 million upon achievement of certain marketing approval milestones. In the United States, we will share equally with Sanofi the profits and losses applicable to commercialization of SHP2 inhibitor products, pursuant to a profit/loss share agreement that the parties will negotiate based on key terms agreed in the Sanofi Agreement. On a product-by-product basis, Sanofi will also be required to pay us tiered royalties on annual net sales of each product outside the United States ranging from high single digit to mid-teen percentages. The royalty payments are subject to reduction under specified conditions set forth in the Sanofi Agreement. Subject to certain exceptions, the royalties are payable on a product-by-product and country-by-country basis until the latest of the expiration of all valid claims covering such product in such country contained in the patents licensed to Sanofi under the Sanofi Agreement and the expiration of regulatory exclusivity for such product in such country.

Sanofi has the sole and exclusive right to file, prosecute and maintain any patents licensed to it pursuant to the Sanofi Agreement, as well as to enforce infringement of or defend claims against such patents that relate to SHP2 inhibitor products.

Unless terminated earlier, the Sanofi Agreement will continue in effect until the later of the expiration of all of Sanofi's milestone and royalty payment obligations and the expiration of the profit/loss share agreement. Upon expiration of the Sanofi Agreement, the licenses granted to Sanofi thereunder shall become fully paid-up, royalty-free, perpetual and irrevocable. Sanofi may terminate the Sanofi Agreement in its entirety or on a country-by-country or product-by-product basis for any reason or for significant safety concerns, upon prior notice to us within certain specified time periods. Sanofi may terminate the Sanofi Agreement in its entirety upon our change of control, with prior notice. Either party may terminate the Sanofi Agreement if an undisputed material breach by the other party is not cured within a defined period of time, or immediately upon notice for insolvency-related events of the other party. We may terminate the Sanofi Agreement after a certain number of years if Sanofi develops a competing program without commencing a registrational clinical trial for a SHP2 inhibitor product candidate, and subject to certain other conditions. We may also terminate the Sanofi Agreement at any time, if Sanofi ceases certain critical activities for SHP2 inhibitor product candidates for more than a specified period of time, provided that such cessations of critical activity were not a result of certain specified factors, and subject to certain other conditions. Upon any termination of the Sanofi Agreement with respect to any product or country, all licenses to Sanofi with respect to such product, other than by us for Sanofi's material breach or insolvency, we may be required to pay Sanofi royalties on worldwide net sales of reverted products up to mid-single digit percentages based on the development and regulatory status of such reverted products, in each case subject to reductions in accordance with the terms of the Sanofi Agreement.

Through June 30, 2021, we have received an aggregate of \$140.7 million from Sanofi, including the upfront payment and research and development expense reimbursements.

Financial Operations Overview

Collaboration revenue

Collaboration revenue consists of revenue under the Sanofi Agreement for our SHP2 program. We entered into the Sanofi Agreement in June 2018. We received a \$50.0 million upfront payment from Sanofi in July 2018, receive reimbursement for research and development services, and are entitled to future potential development and regulatory milestones.

For further information on our revenue recognition policies, see "Note 2. Summary of significant accounting policies" in the "Notes to Consolidated Financial Statements" contained in Part II, Item 8 of our 2020 Annual Report on Form 10-K.

Research and development expenses

We substantially rely on third parties to conduct our preclinical studies, clinical trials and manufacturing. We estimate research and development expenses based on estimates of services performed, and rely on third party contractors and vendors to provide us with timely and accurate estimates of expenses of services performed to assist us in these estimates. Research and development expenses consist primarily of costs incurred for the development of our product candidates and costs associated with identifying compounds through our discovery platform, which include:

- expenses incurred under agreements with third-party contract organizations, investigative clinical trial sites that conduct research and development activities on our behalf, and consultants;
- costs related to production of clinical and preclinical materials, including fees paid to contract manufacturers;
- laboratory and vendor expenses related to the execution of discovery programs, preclinical and clinical trials;
- employee-related expenses, which include salaries, benefits and stock-based compensation; and
- facilities and other expenses, which include allocated expenses for rent and maintenance of facilities, depreciation and amortization expense, information technology and other supplies.

We expense all research and development costs in the periods in which they are incurred. Costs for certain development activities are recognized based on an evaluation of the progress to completion of specific tasks using information and data provided to us by our vendors, collaborators and third-party service providers. Nonrefundable advance payments for goods or services to be received in future periods for use in research and development activities are deferred and recorded as prepaid assets. The prepaid amounts are then expensed as the related goods are delivered or as services are performed.

Sanofi is responsible to reimburse us all internal and external costs and expenses to perform our activities under approved development plans, except for 50% of the RMC-4630-03 study and any future RevMed Study. Additionally, Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred to perform activities under approved research plans for RMC-4630, with the exception of internal and external research costs and expenses under the approved research plans for 2019 and 2020,

for which Sanofi was obligated to reimburse us for 80% of such costs. We were responsible for 20% of all internal and external research costs incurred under the research plans for 2019 and 2020. Sanofi is responsible to reimburse us for all internal and external costs and expenses incurred under the research plan for 2021. These reimbursements from Sanofi are recorded as collaboration revenue.

We expect our research and development expenses to increase for the foreseeable future as we continue to invest in discovering and developing product candidates and advancing product candidates into later stages of development, which may include conducting larger clinical trials. The process of conducting the necessary research and development and clinical trials to seek regulatory approval for product candidates is costly and time-consuming, and the successful development of our product candidates is highly uncertain. As a result, we are unable to determine the duration and completion costs of our research and development projects or clinical trials or if and to what extent we will generate revenue from the commercialization and sale of any of our product candidates.

General and administrative expenses

General and administrative expenses consist primarily of personnel-related costs, consultants and professional services expenses, including legal, audit, accounting and human resources services, insurance, allocated facilities and information technology costs, and other general operating expenses not otherwise classified as research and development expenses. Personnel-related costs consist of salaries, benefits and stock-based compensation. Facilities costs consist of rent, utilities and maintenance of facilities. We expect our general and administrative expenses to increase for the foreseeable future due to anticipated increases in headcount and as a result of operating as a public company, including expenses related to compliance with the rules and regulations of the Securities and Exchange Commission, the Nasdaq Global Select Market, additional insurance expenses, investor relations activities and other administrative and professional services.

Interest income

Interest income primarily consists of interest earned on our cash equivalents and marketable securities.

Interest expense

Interest expense primarily consists of interest related to our capital lease and interest on other outstanding obligations.

Benefit from income taxes

Benefit from income taxes relates to net changes in the deferred tax liability associated with our Warp Drive acquisition resulting from changes in the effective state tax rate and changes in our valuation allowance.

Results of operations

Comparison of the three and six months ended June 30, 2021 and 2020

June 30, 30,	
Increase/ 2021 2020 (decrease) 2021 2020	Increase/ (decrease)
(in thousands) (in thousands)	
Revenue:	
Collaboration revenue \$ 8,698 \$ 10,025 \$ (1,327) \$ 18,829 \$ 21,571 \$	\$ (2,742)
Total revenue 8,698 10,025 (1,327) 18,829 21,571	(2,742)
Operating expenses:	
Research and development 45,936 32,918 13,018 86,794 60,375	26,419
General and administrative 7,297 5,091 2,206 13,967 10,262	3,705
Total operating expenses 53,233 38,009 15,224 100,761 70,637	30,124
Loss from operations (44,535) (27,984) (16,551) (81,932) (49,066)	(32,866)
Other income (expense), net:	
Interest income 236 730 (494) 469 1,639	(1,170)
Interest expense — (19) 19 (12) (40)	28
Total other income, net 236 711 (475) 457 1,599	(1,142)
Loss before income taxes (44,299) (27,273) (17,026) (81,475) (47,467)	(34,008)
Benefit from income taxes — 58 (58) — 733	(733)
Net loss \$ (44,299) \$ (27,215) \$ (17,084) \$ (81,475) \$ (46,734)	\$ (34,741)

Collaboration revenue

Collaboration revenue consists of revenue under the Sanofi Agreement, which was entered into in June 2018. Collaboration revenue decreased by \$1.3 million, or 13%, during the three months ended June 30, 2021 compared to the same period in 2020. The decrease in collaboration revenue during the three months ended June 30, 2021 was primarily due to lower reimbursed research and development costs incurred by us for our SHP2 program under the Sanofi Agreement resulting from lower clinical trial costs.

Collaboration revenue decreased by \$2.7 million, or 13%, during the six months ended June 30, 2021 compared to the same period in 2020. The decrease in collaboration revenue during the six months ended June 30, 2021 was primarily due to lower reimbursed research and development costs incurred by us for our SHP2 program under the Sanofi Agreement resulting from lower clinical trial and manufacturing costs.

Research and development expenses

Research and development expenses increased by \$13.0 million, or 40%, during the three months ended June 30, 2021 compared to the same period in 2020. The increase in research and development expenses during the three months ended June 30, 2021 was primarily due to a \$8.0 million increase in third party costs for our preclinical research portfolio, primarily driven by higher chemistry contract research organization, material sourcing and manufacturing costs; a \$2.4 million increase in salaries and other employee-related expenses due to increased headcount to support our research and development programs; and a \$1.9 million increase in stock-based compensation.

Research and development expenses increased by \$26.4 million, or 44%, during the six months ended June 30, 2021 compared to the same period in 2020. The increase in research and development expenses during the six months ended June 30, 2021 was primarily due to a \$18.6 million increase in third party costs for our preclinical research portfolio, primarily driven by higher chemistry contract research organization, material sourcing and manufacturing costs; a \$4.2 million increase in salaries and other employee-related expenses due to increased headcount to support our research and development programs; and a \$2.9 million increase in stock-based compensation.

General and administrative expenses

General and administrative expenses increased by \$2.2 million, or 43%, during the three months ended June 30, 2021 compared to the same period in 2020. The increase in general and administrative expenses during the three months ended June 30, 2021 was primarily due to an increase of \$1.4 million in stock-based compensation expense and an increase of \$0.7 million in salaries and other employee-related expenses due to increased headcount.

General and administrative expenses increased by \$3.7 million, or 36%, during the six months ended June 30, 2021 compared to the same period in 2020. The increase in general and administrative expenses during the six months ended June 30, 2021 was primarily due to an increase of \$2.2 million in stock-based compensation expense and an increase of \$1.2 million in salaries and other employee-related expenses due to increased headcount.

Interest income

Interest income decreased by \$0.5 million and \$1.2 million during the three and six months ended June 30, 2021, respectively, compared to the same periods in 2020 due to lower interest rates.

Interest expense

Interest expense was less than \$0.1 million for the three and six months ended June 30, 2021 and 2020, respectively.

Benefit from income taxes

Benefit from income taxes was zero and \$0.1 million for the three months ended June 30, 2021 and 2020, respectively, and zero and \$0.7 million for the six months ended June 2021 and 2020, respectively. Benefit from income taxes for the three and six months ended June 30, 2020, respectively, relates to a reduction in our effective state tax rate and the resulting impact on the deferred tax liabilities from the Warp Drive acquisition.

On March 27, 2020, the United States enacted the Coronavirus Aid, Relief, and Economic Security Act (the CARES Act) as a result of the Coronavirus pandemic, which contains among other things, numerous income tax provisions. Some of these tax provisions are expected to be effective retroactively for years ending before the date of enactment. We have evaluated the current legislation and at this time, we do not anticipate the CARES Act to have a material impact on our financial statements.

Liquidity and capital resources

Liquidity

In February 2020, we closed our IPO and issued 16,100,000 shares of our common stock at a price to the public of \$17.00 per share for net proceeds of approximately \$250.7 million, after deducting underwriting discounts and commissions of \$19.2 million and expenses of \$3.8 million.

In July 2020, we issued 6,900,000 shares of our common stock in an underwritten public offering at a price of \$26.00 per share for net proceeds of \$167.8 million, after deducting underwriting discounts and commissions of \$10.8 million and offering expenses of \$0.8 million.

In February 2021, we issued 6,666,666 shares of our common stock in an underwritten public offering at a price of \$45.00 per share for net proceeds of \$281.1 million, after deducting underwriting discounts and commissions of \$18.0 million and offering expenses of \$0.9 million.

Our operations have been financed primarily by our public offerings of common stock, net proceeds of \$230.6 million from the issuance of our preferred stock and \$140.7 million received under the Sanofi Agreement for upfront payments and for research and development cost reimbursement.

As of June 30, 2021, we had \$646.3 million in cash, cash equivalents and marketable securities.

As of June 30, 2021, we had an accumulated deficit of \$347.0 million. Our primary use of cash is to fund operating expenses, which consist primarily of research and development expenditures related to our product candidates and our pre-clinical research portfolio, and to a lesser extent, general and administrative expenditures. We expect our expenses to continue to increase in connection with our ongoing activities, particularly as we continue to advance our product candidates and pre-clinical research portfolio.

We believe that our existing cash, cash equivalents and marketable securities will enable us to fund our planned operations for at least 12 months following the date of this report.

The timing and amount of our future funding requirements depends on many factors, including:

- the scope, progress, results and costs of researching and developing our product candidates and programs, and of conducting preclinical studies and clinical trials;
- the timing of, and the costs involved in, obtaining marketing approvals for product candidates we develop if clinical trials are successful;
- the success of our collaboration with Sanofi, including the continued reimbursement by Sanofi of substantially all of our research costs and all of our development costs for our SHP2 program under the Sanofi Agreement (other than for the RMC-4630-03 study or any RevMed Studies);
- whether we achieve certain clinical and regulatory milestones under the Sanofi Agreement, each of which would trigger additional payments to us;
- the cost of commercialization activities for RMC-4630, to the extent not borne by Sanofi, and any other future product candidates we develop, whether alone or in collaboration, including marketing, sales and distribution costs if RMC-4630 or any other product candidate we develop is approved for sale;
- the cost of manufacturing our current and future product candidates for clinical trials in preparation for marketing approval and in preparation for commercialization;
- our ability to establish and maintain strategic licenses or other arrangements and the financial terms of such agreements;
- the costs involved in preparing, filing, prosecuting, maintaining, expanding, defending and enforcing patent claims, including litigation costs and the outcome of such litigation;
- the timing, receipt and amount of sales of, profit share or royalties on, our future products, if any;
- the emergence of competing cancer therapies or other adverse market developments; and
- any plans to acquire or in-license other programs or technologies.

We expect to need to obtain substantial additional funding in the future for our research and development activities and continuing operations. Sanofi reimburses us for almost all of our research and development expenses associated with our SHP2 program, however Sanofi has the right to terminate the Sanofi Agreement for any reason, upon prior notice to us within certain specified time periods and upon any such termination by Sanofi with respect to any product or country, all licenses to Sanofi with respect to such product or country shall automatically terminate and all rights generally revert back to us. If we need to raise additional capital to fund our operations, funding may not be available to us on acceptable terms, or at all. If we are unable to obtain adequate financing when needed, we may have to delay, reduce the scope of or suspend one or more of our clinical trials, research and development programs or commercialization efforts. We may seek to raise any necessary additional capital through a combination of public or private equity offerings, debt financings and collaborations or licensing arrangements. If we do raise additional capital through public or private equity offerings, the ownership interest of our existing stockholders will be diluted, and the terms of these securities may include liquidation or other preferences that adversely affect our stockholders' rights. If we raise additional capital through debt financing, we may be subject to covenants limiting or restricting our ability to take specific actions, such as incurring additional debt, making capital expenditures or declaring dividends, and if the debt is convertible into our common stock, the ownership interest of our stockholders may be diluted. If we are unable to raise capital, we may need to delay, reduce or terminate planned activities to reduce costs. Doing so will likely harm our ability to execute our business plans.

Cash flows

The following table summarizes our consolidated cash flows for the periods indicated:

	 Six Months Ended June 30,						
	 2021	2020					
	(in thousands)						
Net cash provided by (used in) provided by:							
Operating activities	\$ (71,327)	\$	(48,009)				
Investing activities	(120,473)		(196,212)				
Financing activities	283,132		252,408				
Net change in cash and cash equivalents	\$ 91,332	\$	8,187				

Cash used in operating activities

During the six months ended June 30, 2021, cash used in operating activities of \$71.3 million was attributable to a net loss of \$81.5 million and a net change of \$3.3 million in our operating assets and liabilities, partially offset by \$13.4 million in non-cash charges. The non-cash charges primarily consisted of stock-based compensation expense of \$8.7 million, depreciation and amortization of \$1.5 million, amortization of operating lease right-of-use asset of \$1.5 million and net amortization of premium on marketable securities of \$1.2 million. The change in operating assets and liabilities was primarily due to a \$4.6 million decrease in deferred revenue associated with the Sanofi Agreement, and a \$1.1 million increase in prepaid expenses and other current assets primarily resulting from the timing of prepayments made for research and development activities and insurance, partially offset by an increase in accrued expenses and other current liabilities of \$3.2 million.

During the six months ended June 30, 2020, cash used in operating activities of \$48.0 million was attributable to a net loss of \$46.7 million and a net change of \$8.1 million in our operating assets and liabilities, partially offset by a net change of \$6.8 million in non-cash charges. The non-cash charges primarily consisted of stock-based compensation expense of \$3.6 million and depreciation and amortization of \$1.8 million. The change in operating assets and liabilities was primarily due to a \$5.7 million decrease in deferred revenue associated with the Sanofi Agreement and \$2.5 million increase in prepaid expenses and other current assets primarily resulting from the timing of prepayments made for research and development activities.

Cash used in investing activities

During the six months ended June 30, 2021, cash used in investing activities of \$120.5 million, was primarily comprised of purchases of marketable securities of \$392.3 million and purchases of property and equipment of \$4.9 million partially offset by maturities of marketable securities of \$276.7 million.

During the six months ended June 30, 2020, cash used in investing activities of \$196.2 million, was primarily comprised of purchases of marketable securities of \$305.2 million and purchases of property and equipment of \$1.0 million, partially offset by cash provided by maturities of marketable securities of \$107.1 million and sale of marketable securities of \$3.0 million.

Cash provided by financing activities

During the six months ended June 30, 2021, cash provided by financing activities of \$283.1 million was comprised of \$281.1 million in net proceeds from the issuance of common stock related to our public offering in February 2021, \$1.2 million in proceeds from issuance of common stock related to our employee stock purchase plan and \$0.8 million in proceeds from the issuance of common stock upon the exercise of stock options.

During the six months ended June 30, 2020, cash provided by financing activities of \$252.4 million was comprised primarily of proceeds from issuance of common stock, net of offering costs related to our IPO in February 2020.

Contractual obligations and commitments

The following table summarizes our commitments and contractual obligations as of June 30, 2021:

	Payments Due by Period									
	Less than Total 1 year 1-3 years 3-5 years									ore than 5 years
					(in t	housands)				
Operating lease obligations	\$	44,617	\$	5,066	\$	8,851	\$	8,727	\$	21,973
Total contractual obligations	\$	44,617	\$	5,066	\$	8,851	\$	8,727	\$	21,973

Our contractual obligations reflect our minimum payments due for office and laboratory space leases in Redwood City, California and Cambridge, Massachusetts, which are our operating leases, and our equipment leases, which are our financing leases.

Our primary Redwood City lease commenced in January 2015 and ends in December 2030. In April 2020, we amended our Redwood City lease to lease an additional 19,483 square feet of office, laboratory and research and development space located at 300 Saginaw Drive, Redwood City, California beginning on December 15, 2020 and ending December 31, 2030. Under the amendment, our existing lease term for the premises located at 700 Saginaw Drive, Redwood City, California was extended until December 31, 2030.

As part of the Warp Drive acquisition, we assumed Warp Drive's office and laboratory space lease in Cambridge, which ends in February 2023. In March 2019, we fully subleased the Cambridge lease to Casma Therapeutics, Inc. (Casma) on financial terms substantially the same as the original lease. The amounts reflected in the table above include our lease payments for the Cambridge lease, but do not reflect any offset for the sublease payments we are entitled to receive from Casma. The sublease by Casma and related sublease payments by Casma to us are fully guaranteed by Third Rock Ventures, LLC.

We enter into agreements in the normal course of business with contract research organizations for clinical trials, contract manufacturing organizations to provide clinical trial materials and with vendors for preclinical studies and other services and products for operating purposes which are generally cancelable at any time by us upon 30 to 90 days prior written notice. These payments are not included in this table of contractual obligations.

Off-balance sheet arrangements

We have not entered into any off-balance sheet arrangements, as defined in Item 303 of Regulation S-K.

Indemnification agreements

We enter into standard indemnification arrangements in the ordinary course of business. Pursuant to these arrangements, we indemnify, hold harmless and agree to reimburse the indemnified parties for losses suffered or incurred by the indemnified party, in connection with any trade secret, copyright, patent or other intellectual property infringement claim by any third party with respect to its technology. The term of these indemnification agreements is generally perpetual any time after the execution of the agreement. The maximum potential amount of future payments we could be required to make under these arrangements is not determinable. We have never incurred costs to defend lawsuits or settle claims related to these indemnification agreements. As a result, we believe the fair value of these agreements is minimal.

Critical accounting policies, significant judgments and use of estimate

Our management's discussion and analysis of our financial condition and results of operations is based on our consolidated financial statements, which have been prepared in accordance with United States generally accepted accounting principles (U.S. GAAP). The preparation of these consolidated financial statements requires us to make estimates and assumptions that affect the reported amounts of assets and liabilities and the disclosure of contingent assets and liabilities at the date of the consolidated financial statements, as well as the reported expenses incurred during the reporting periods. Our estimates are based on our historical experience and on various other factors that we believe are reasonable under the circumstances, the results of which form the basis for making judgments about the carrying value of assets and liabilities that are not readily apparent from other sources. Actual results may differ from these estimates under different assumptions or conditions.

For a discussion of our critical accounting estimates, see Part II, Item 7. Management's Discussion and Analysis of Financial Condition and Results of Operations in our 2020 Annual Report on Form 10-K for the year ended December 31, 2020, as filed with the SEC on March 2, 2021 (our 2020 Form 10-K). There have been no material changes to these critical accounting estimates since our 2020 Form 10-K.

Emerging growth company status

In April 2012, the Jumpstart Our Business Startups Act of 2012 (the JOBS Act) was enacted. Section 107 of the JOBS Act provides that an "emerging growth company," or an EGC, can take advantage of the extended transition period provided in Section 7(a)(2)(B) of the Securities Act of 1933, as amended (the Securities Act) for complying with new or revised accounting standards. Thus, an EGC can delay the adoption of certain accounting standards until those standards would otherwise apply to private companies. We early adopted ASC 606 and ASC 842 as the JOBS Act does not preclude an EGC from early adopting a new or revised accounting standard earlier than the time that such standard applies to private companies. We have elected to use the extended transition period for any other new or revised accounting standards during the period in which we remain an emerging growth company; however, we may adopt certain new or revised accounting standards early.

For further information regarding our significant accounting policies, see "Note 2. Summary of significant accounting policies" in the "Notes to Unaudited Condensed Consolidated Financial Statements" contained in Part I, Item 1 of this report and "Note 2. Summary of significant accounting policies" in the "Notes to Consolidated Financial Statements" contained in Part II, Item 8 of our 2020 Form 10-K. There have been no material changes to these critical accounting policies since our 2020 Form 10-K.

Recent accounting pronouncements

For a description of the expected impact of recent accounting pronouncements, see "Note 2. Summary of significant accounting policies" in the "Notes to Unaudited Condensed Consolidated Financial Statements" contained in Part I, Item 1 of this report.

Item 3. Quantitative and Qualitative Disclosures About Market Risk.

Interest rate risk

We are exposed to market risks in the ordinary course of our business. These risks primarily include interest rate sensitivities. The primary objective of our investment activities is to preserve capital to fund our operations. We also seek to maximize income from our investments without assuming significant risk. To achieve our objectives, we maintain a portfolio of investments in a variety of securities of high credit quality and short-term duration, invested in compliance with our policy.

We held cash, cash equivalents and marketable securities of \$646.3 million and \$440.7 million as of June 30, 2021 and December 31, 2020, respectively, which consisted of bank deposits, money market funds, U.S. government debt securities, U.S. government agency bonds, commercial paper and corporate bonds. Such interest-earning instruments carry a degree of interest rate risk; however, historical fluctuations in interest income have not been significant for us. Due to the short-term maturities of our cash equivalents, an immediate one percent change in interest rates would not have a material effect on the fair value of our cash equivalents and marketable securities.

Foreign currency risk

Our expenses are generally denominated in U.S. dollars. However, we have entered into a limited number of contracts with vendors for research and development services with payments denominated in foreign currencies, including the Euro, British Pound and Chinese Yuan. We are subject to foreign currency transaction gains or losses on our contracts denominated in foreign currencies. To date, foreign currency transaction gains and losses have not been material to our consolidated financial statements, and we have not had a formal hedging program with respect to foreign currency. A 10% increase or decrease in current exchange rates would not have a material effect on our financial results.

Item 4. Controls and Procedures.

Evaluation of disclosure controls and procedures

Our management, with the participation of our President and Chief Executive Officer and Director and our Senior Vice President, Finance and Principal Accounting Officer, our principal executive officer and principal financial officer, respectively, have evaluated our disclosure controls and procedures (as defined in Rules 13a-15(e) and 15d-15(e) under the Securities Exchange Act of 1934, as amended) as of June 30, 2021. Based on the evaluation, our President and Chief Executive Officer and Director and our Senior Vice President, Finance and Principal Accounting Officer have concluded that, as of June 30, 2021, our disclosure controls and procedures were, in design and operation, effective to the reasonable assurance level.

Changes in internal control over financial reporting

There were no changes in our internal controls over financial reporting identified in connection with the evaluation required by Rule 13a-15(d) and 15d-15(d) of the Exchange Act that occurred during the three months ended June 30, 2021 that have materially affected, or are reasonably likely to materially affect, our internal control over financial reporting.

Inherent limitation on the effectiveness over financial reporting

The effectiveness of any system of internal control over financial reporting, including ours, is subject to inherent limitations, including the exercise of judgment in designing, implementing, operating, and evaluating the controls and procedures, and the inability to eliminate misconduct completely. Accordingly, any system of internal control over financial reporting, including ours, no matter how well designed and operated, can only provide reasonable, not absolute assurances. In addition, projections of any evaluation of effectiveness to future periods are subject to the risk that controls may become inadequate because of changes in conditions, or that the degree of compliance with the policies or procedures may deteriorate. We intend to continue to monitor and upgrade our internal controls as necessary or appropriate for our business, but there can be no assurance that such improvements will be sufficient to provide us with effective internal control over financial reporting.

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PART II—OTHER INFORMATION

Item 1. Legal Proceedings.

From time to time, we may become involved in litigation or other legal proceedings. We are not currently a party to any litigation or legal proceedings that, in the opinion of our management, are likely to have a material adverse effect on our business. Regardless of outcome, litigation can have an adverse impact on our business, financial condition, results of operations and prospects because of defense and settlement costs, diversion of management resources and other factors.

Item 1A. Risk Factors.

Summary of Material Risks Associated with Our Business

The principal risks and uncertainties affecting our business include the following:

- The COVID-19 pandemic, or other epidemic and pandemic diseases or governmental or other actions taken in response to them, could significantly disrupt our business.
- We are a clinical-stage precision oncology company with a limited operating history and no products approved for commercial sale. We have incurred significant losses since inception. We expect to incur losses for at least the next several years and may never achieve or maintain profitability, which, together with our limited operating history, makes it difficult to assess our future viability.
- We have never generated revenue from product sales and may never be profitable.
- We will require substantial additional financing to achieve our goals, which may not be available on acceptable terms, or at all. A failure to obtain this
 necessary capital when needed could force us to delay, limit, reduce or terminate our product development or commercialization efforts.
- We are early in our development efforts. Our business is dependent on the successful development of our current and future product candidates. If we are unable to advance our current or future product candidates through clinical trials, obtain marketing approval and ultimately commercialize any product candidates we develop, or experience significant delays in doing so, our business will be materially harmed.
- Preclinical development is uncertain. Our preclinical programs may experience delays or may never advance to clinical trials, which would adversely
 affect our ability to obtain regulatory approvals or commercialize our product candidates on a timely basis or at all, which would have an adverse
 effect on our business.
- Historically, direct inhibition of any RAS protein has been challenging due to a lack of tractable, or "druggable," binding pockets and we are not aware of any programs in clinical development that have successfully targeted any RAS(ON) protein. Given this approach is unproven, it may not be successful.
- The results of preclinical studies and early-stage clinical trials may not be predictive of future results.
- If we encounter difficulties enrolling patients in our clinical trials, our clinical development activities could be delayed or otherwise be adversely affected.
- We are dependent on our collaboration with Sanofi for the development of RMC-4630 and may depend on Sanofi for the development and commercialization of any other future SHP2 inhibitor product candidates. Under certain circumstances, Sanofi may unilaterally terminate the collaboration for convenience, which would materially and adversely affect our business.
- We are currently developing and may in the future, develop RMC-4630 and other product candidates in combination with other therapies, which exposes us to additional risks.
- We face significant competition, and if our competitors develop and market products that are more effective, safer or less expensive than the product candidates we develop, our commercial opportunities will be negatively impacted.
- If we and our collaborators are unable to obtain and maintain sufficient patent and other intellectual property protection for our product candidates and technology, our competitors could develop and commercialize products and technology similar or identical to ours, and we may not be able to compete effectively in our market or successfully commercialize any product candidates we may develop.

The summary risk factors described above should be read together with the text of the full risk factors below in the section entitled "Risk Factors" and the other information set forth in this Quarterly Report on Form 10-Q, including our condensed consolidated financial statements and the related notes, as well as in other documents that we file with the U.S. Securities and Exchange



Commission. The risks summarized above or described in full below are not the only risks that we face. Additional risks and uncertainties not precisely known to us or that we currently deem to be immaterial may also materially adversely affect our business, financial condition, results of operations, and future growth prospects.

Investing in our common stock involves a high degree of risk. You should carefully consider the risks described below, as well as the other information in this Quarterly Report on Form 10-Q, including our financial statements and the related notes and the section titled "Management's Discussion and Analysis of Financial Condition and Results of Operations" before deciding whether to invest in our common stock. The occurrence of any of the events or developments described below could harm our business, competitive position, financial condition, results of operations, cash flows and prospects. In such an event, the market price of our common stock could decline, and you may lose all or part of your investment. Many of the following risks and uncertainties and those contained in the documents incorporated by reference herein are, and will be, exacerbated by the COVID-19 pandemic and any worsening of the global business and economic environment as a result. Additional risks and uncertainties not presently known to us or that we currently deem immaterial also may impair our business operations and the market price of our common stock.

Risks related to the COVID-19 pandemic

The COVID-19 pandemic, or other epidemic and pandemic diseases or governmental or other actions taken in response to them, could significantly disrupt our business.

Outbreaks of epidemic, pandemic or contagious diseases, such as the recent SARS-CoV-2 virus, or coronavirus, which causes coronavirus disease 2019 (COVID-19) or, historically, the Ebola virus, Middle East Respiratory Syndrome, Severe Acute Respiratory Syndrome or the H1N1 virus, could significantly disrupt our business. These outbreaks pose the risk that we or our employees, contractors, suppliers, and other partners may be prevented from conducting business activities for an indefinite period of time due to spread of the disease within these groups, or due to restrictions that may be requested or mandated by governmental authorities. Business disruptions could include disruptions or restrictions on our ability to travel, as well as temporary closures of all or part of our facilities and the facilities of our partners, clinical trial sites, service providers, suppliers or contract manufacturers. As the COVID-19 pandemic evolves and spreads, both across the United States and through the world, we continue to actively monitor the impact that COVID-19 is having and may have on our business. The pandemic and the measures taken by governmental authorities could disrupt and delay our ongoing clinical trials, our preclinical activities, the manufacture or shipment of both drug substance and finished drug product for our product candidates for preclinical testing and clinical trials and otherwise significantly disrupt our business.

As a result of the COVID-19 pandemic, the state of California, where our corporate offices are located, and many counties where our offices are located or our employees reside, have issued and may in the future issue orders for all residents to remain at home, except as needed for essential activities, and have placed restrictions on the scope and conduct of business activities. We have taken steps to ensure the safety of our patients and employees, while working to ensure the sustainability of our business operations as this unprecedented situation continues to evolve. As a result, we have implemented policies that require or permit many of our employees to work remotely. Some of these policies may continue for an indefinite period and may become more restrictive in response to developments related to the pandemic and the associated governmental responses. We continue to evaluate the impact of COVID-19 on the healthcare system and work with healthcare providers supporting our clinical studies to mitigate risk to patients while taking into account regulatory, institutional, and government guidance and policies.

Our clinical trial sites for our RMC-4630 and RMC-5552 clinical studies may be affected by the COVID-19 outbreak due to prioritization of hospital resources toward the COVID-19 outbreak, travel, quarantine or other restrictions imposed by governments, and the inability to access sites for initiation and patient monitoring and enrollment. As a result, patient screening, new patient enrollment, monitoring and data collection may be affected or delayed. We are aware that several clinical sites involved in our RMC-4630 clinical studies temporarily stopped or delayed enrolling new patients, with exemptions if appropriate, and it is possible that these or other clinical sites may be similarly affected in the future. These developments may delay our clinical trial timelines. Our clinical trials currently permit patients to receive COVID-19 vaccines while they are on study. The potential impact of our candidates on the safety and efficacy of COVID-19 vaccines, and the potential impact on of COVID-19 vaccines on the safety and efficacy of our candidates is unknown at this time, but it possible that adverse impacts will negatively affect our clinical trials.

Although we are currently not aware of any material impacts on our supply chain of our current or potential product candidates as a result of the COVID-19 pandemic, some of our third-party manufacturers which we use for the supply of materials for product candidates or other materials necessary to manufacture product to conduct preclinical tests and clinical trials and contract research organizations that we may utilize may be impacted by COVID-19, and should they experience continued disruptions, such as temporary closures or suspension of services, we would likely experience delays in advancing clinical trials. Furthermore, the spread of the virus may affect the operations of key governmental agencies, such as the U.S. Food and Drug Administration (the FDA), which may delay the development of our product candidates. The spread of an infectious disease, including COVID-19, may also result in the inability of our suppliers to deliver components or raw materials on a timely basis or at all. Such events may result in a period of business disruption, and in reduced operations, or doctors and medical providers may be unwilling to participate in our clinical trials.



In addition, a significant outbreak of epidemic, pandemic or contagious diseases in the human population, such as the global COVID-19 pandemic, could result in a widespread health crisis and adversely affect the economies and financial markets of many countries, resulting in an economic downturn that could affect demand for our current or future products.

While the potential economic impact brought by, and the duration of, COVID-19 may be difficult to assess or predict, a continuing widespread pandemic could result in significant disruption of global financial markets, reducing our ability to access capital, which could in the future negatively affect our liquidity. In addition, a recession or market correction resulting from the spread of COVID-19 could materially affect the value of our common stock.

Risks related to our limited operating history, financial position and need for additional capital

We are a clinical-stage precision oncology company with a limited operating history and no products approved for commercial sale. We have incurred significant losses since inception. We expect to incur losses for at least the next several years and may never achieve or maintain profitability, which, together with our limited operating history, makes it difficult to assess our future viability.

Biopharmaceutical product development is a highly speculative undertaking and involves a substantial degree of risk. We are a clinical-stage precision oncology company, and we have only a limited operating history upon which you can evaluate our business and prospects. We currently have no products approved for commercial sale, have not generated any revenue from sales of products and have incurred losses in each year since our inception in October 2014. In addition, we have limited experience as a company and have not yet demonstrated an ability to successfully overcome many of the risks and uncertainties frequently encountered by companies in new and rapidly evolving fields, particularly in the biopharmaceutical industry.

Since inception, we have incurred significant net losses. Our net losses were \$108.2 million, \$47.7 million and \$41.8 million, for the years ended December 31, 2020, 2019 and 2018, respectively. Our net loss was \$44.3 million and \$81.5 million for the three and six months ended June 30, 2021. As of June 30, 2021, we had an accumulated deficit of \$347.0 million. We have funded our operations to date primarily with proceeds from the sale of common stock and preferred stock and upfront payments and research and development cost reimbursement received under our collaboration agreement with Genzyme Corporation, an affiliate of Sanofi, or the Sanofi Agreement. To date, we have devoted substantially all of our resources to organizing and staffing our company, business planning, raising capital, acquiring and discovering development programs, securing intellectual property rights and conducting discovery, research and development activities for our programs. We have not yet demonstrated our ability to successfully complete any clinical trials, including pivotal clinical trials, obtain marketing approvals, manufacture a commercial-scale product, or arrange for a third party to do so on our behalf, or conduct sales and marketing activities necessary for successful product commercialization. Our product candidates will require substantial additional development time and resources before we will be able to apply for or receive regulatory approvals and, if approved, begin generating revenue from product sales. We expect to continue to incur significant expenses and operating losses for the foreseeable future.

We have never generated revenue from product sales and may never be profitable.

Our ability to generate revenue from product sales and achieve profitability depends on our ability, alone or with our collaboration partners, to successfully complete the development of, and obtain the regulatory approvals necessary to commercialize, our development programs. We do not anticipate generating revenue from product sales for the next several years, if ever. Our ability to generate future revenue from product sales depends heavily on our, Sanofi's, and any potential future collaborators' success in:

- completing clinical and preclinical development of product candidates and programs and identifying and developing new product candidates;
- seeking and obtaining marketing approvals for any product candidates that we develop;
- launching and commercializing product candidates for which we obtain marketing approval by establishing a sales force, marketing, medical affairs
 and distribution infrastructure or, alternatively, collaborating with a commercialization partner;
- achieving adequate coverage and reimbursement by third-party payors for product candidates that we develop;
- establishing and maintaining supply and manufacturing relationships with third parties that can provide adequate, in both amount and quality, products
 and services to support clinical development and the market demand for product candidates that we develop, if approved;
- obtaining market acceptance of product candidates that we develop as viable treatment options;
- addressing any competing technological and market developments;

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- negotiating favorable terms in any collaboration, licensing or other arrangements into which we may enter and performing our obligations in such collaborations;
- maintaining, protecting, enforcing and expanding our portfolio of intellectual property rights, including patents, trade secrets and know-how;
- · defending against third-party interference, infringement or other intellectual property-related claims, if any; and
- attracting, hiring and retaining qualified personnel.

Even if one or more of the product candidates that we develop is approved for commercial sale, we anticipate incurring significant costs associated with commercializing any approved product candidate. Our expenses could increase beyond expectations if we are required by the FDA, the European Medicines Agency (the EMA) or other regulatory agencies to perform clinical trials or studies in addition to those that we currently anticipate. Even if we are able to generate revenue from the sale of any approved products, we may not become profitable and may need to obtain additional funding to continue operations.

We will require substantial additional financing to achieve our goals, which may not be available on acceptable terms, or at all. A failure to obtain this necessary capital when needed could force us to delay, limit, reduce or terminate our product development or commercialization efforts.

Our operations have consumed substantial amounts of cash since inception. Since our inception, we have invested a significant portion of our efforts and financial resources in research and development activities for our initial preclinical and clinical product candidates. Preclinical studies and clinical trials and additional research and development activities will require substantial funds to complete. As of June 30, 2021, we had cash, cash equivalents and marketable securities of \$646.3 million. In February 2020, we raised \$250.7 million upon the completion of our initial public offering (IPO), net of underwriting discounts and commissions and offering expenses. In July 2020, we raised \$167.8 million upon the completion of a follow-on public offering, net of underwriting discounts and commissions and offering expenses. In February 2021, we raised \$281.1 million upon the completion of a follow-on public offering, net of underwriting discounts and commissions and offering expenses. We expect to continue to spend substantial amounts to continue the preclinical and clinical development of our current and future programs. If we are able to gain marketing approval for product candidates that we develop, we will require significant additional amounts of cash in order to launch and commercialize our product candidates to the extent that their launch and commercialization are not the responsibility of Sanofi or another collaborator that we may contract with in the future. In addition, other unanticipated costs may arise. Because the design and outcome of our current, planned and potential future clinical trials is highly uncertain, we cannot reasonably estimate the actual amounts necessary to successfully complete the development and commercialization of any product candidate we develop.

The timing and amount of our future funding requirements depend on many factors, including:

- the scope, progress, results and costs of researching and developing our product candidates and programs, and of conducting preclinical studies and clinical trials;
- the timing of, and the costs involved in, obtaining marketing approvals for product candidates we develop if clinical trials are successful;
- the success of our collaboration with Sanofi, including the continued reimbursement by Sanofi of substantially all of our research costs and all of our development costs for our SHP2 program (other than for the RMC-4630-03 study or any RevMed Studies) under the Sanofi Agreement;
- whether we achieve certain clinical and regulatory milestones under the Sanofi Agreement, each of which would trigger additional payments to us;
- the cost of commercialization activities for RMC-4630, to the extent not borne by Sanofi, and any other future product candidates we develop, whether alone or in collaboration, including marketing, sales and distribution costs if RMC-4630 or any other product candidate we develop is approved for sale;
- the cost of manufacturing our current and future product candidates for clinical trials in preparation for marketing approval and in preparation for commercialization;
- our ability to establish and maintain strategic licenses or other arrangements and the financial terms of such agreements;
- the costs involved in preparing, filing, prosecuting, maintaining, expanding, defending and enforcing patent claims, including litigation costs and the
 outcome of such litigation;
- the timing, receipt and amount of sales of, profit share or royalties on, our future products, if any;
- the emergence of competing cancer therapies or other adverse market developments; and



• any plans to acquire or in-license other programs or technologies.

Other than our Sanofi collaboration on SHP2 inhibitors, including RMC-4630, we do not have any committed external source of funds or other support for our development efforts. We expect to finance our cash needs through a combination of public or private equity offerings, debt financings, credit or loan facilities, collaborations, strategic alliances, licensing arrangements and other marketing or distribution arrangements. In addition, we may seek additional capital due to favorable market conditions or strategic considerations even if we believe we have sufficient funds for our current or future operating plans.

Our ability to raise additional funds will depend on financial, economic and other factors, many of which are beyond our control. Additional funds may not be available when we need them, on terms that are acceptable to us, or at all. If adequate funds are not available to us on a timely basis, we may be required to:

- delay, limit, reduce or terminate preclinical studies, clinical trials or other research and development activities or eliminate one or more of our development programs altogether; or
- delay, limit, reduce or terminate our efforts to establish manufacturing and sales and marketing capabilities or other activities that may be necessary to commercialize any future approved products, or reduce our flexibility in developing or maintaining our sales and marketing strategy.

Our operating results may fluctuate significantly, which will make our future results difficult to predict and could cause our results to fall below expectations.

Our quarterly and annual operating results may fluctuate significantly, which will make it difficult for us to predict our future results. These fluctuations may occur due to a variety of factors, many of which are outside of our control and may be difficult to predict, including:

- the timing and cost of, and level of investment in, research, development and commercialization activities, which may change from time to time;
- the timing and status of enrollment for our clinical trials;
- the timing of regulatory approvals, if any, in the United States and internationally;
- the timing of expanding our operational, financial and management systems and personnel, including personnel to support our clinical development, quality control, manufacturing and commercialization efforts and our operations as a public company;
- the cost of manufacturing, as well as building out our supply chain, which may vary depending on the quantity of productions, and the terms of any
 agreements we enter into with third-party suppliers;
- timing and amount of any milestone, royalty or other payments due under any current or future collaboration or license agreement, including the Sanofi Agreement;
- coverage and reimbursement policies with respect to any future approved products, and potential future drugs that compete with our products;
- the timing and cost to establish a sales, marketing, medical affairs and distribution infrastructure to commercialize any products for which we may obtain marketing approval and intend to commercialize on our own or jointly with Sanofi;
- expenditures that we may incur to acquire, develop or commercialize additional products and technologies;
- the level of demand for any future approved products, which may vary significantly over time;
- future accounting pronouncements or changes in our accounting policies; and
- the timing and success or failure of preclinical studies and clinical trials for our product candidates or competing product candidates, or any other change in the competitive landscape of our industry, including consolidation among our competitors or collaboration partners.

The cumulative effects of these factors could result in large fluctuations and unpredictability in our quarterly and annual operating results. As a result, comparing our operating results on a period-to-period basis may not be meaningful. Investors should not rely on our past results as an indication of our future performance.



This variability and unpredictability could also result in our failing to meet the expectations of industry or financial analysts or investors for any period. If our revenue or operating results fall below the expectations of analysts or investors or below any forecasts we may provide to the market, or if the forecasts we provide to the market are below the expectations of analysts or investors, the price of our common stock could decline substantially. Such a stock price decline could occur even when we have met any previously publicly stated revenue or operating guidance we may provide.

Risks related to product development and regulatory process

We are early in our development efforts. Our business is dependent on the successful development of our current and future product candidates. If we are unable to advance our current or future product candidates through clinical trials, obtain marketing approval and ultimately commercialize any product candidates we develop, or experience significant delays in doing so, our business will be materially harmed.

We are early in our development efforts. Our most advanced product candidates, RMC-4630 and RMC-5552, are currently being evaluated in clinical trials. Our other programs are in the preclinical stage. We have invested substantially all of our efforts and financial resources in the identification of targets and preclinical development of small molecules to treat cancer.

The success of our business, including our ability to finance our company and generate revenue from products in the future, which we do not expect will occur for several years, if ever, will depend heavily on the successful development and eventual commercialization of the product candidates we develop, which may never occur. Our current product candidates, and any future product candidates we develop, will require additional preclinical and clinical development, management of clinical, preclinical and manufacturing activities, marketing approval in the United States and other markets, demonstrating effectiveness to pricing and reimbursement authorities, obtaining sufficient manufacturing supply for both clinical development and commercial production, building of a commercial organization, and substantial investment and significant marketing efforts before we generate any revenues from product sales.

We have not previously submitted a new drug application (NDA) to the FDA or similar approval filings to a comparable foreign regulatory authority, for any product candidate. An NDA or other relevant regulatory filing must include extensive preclinical and clinical data and supporting information to establish that the product candidate is safe and effective for each desired indication. The NDA or other relevant regulatory filing must also include significant information regarding the chemistry, manufacturing and controls for the product. We cannot be certain that our current or future product candidates will be successful in clinical trials or receive regulatory approval. For example, we recently determined to deprioritize our RMC-4630-02 study and are no longer enrolling patients in this study. Further, even if they are successful in clinical trials, our product candidates or any future product candidates may not receive regulatory approval. If we do not receive regulatory approvals for current or future product candidates, we may not be able to continue our operations. Even if we successfully obtain regulatory approval to market a product candidate, our revenue will depend, in part, upon the size of the markets in the territories for which we gain regulatory approval and have commercial rights, as well as the availability of competitive products, whether there is sufficient third-party reimbursement and adoption by physicians.

We plan to seek regulatory approval to commercialize our product candidates both in the United States and in select foreign countries. While the scope of regulatory approval generally is similar in other countries, in order to obtain separate regulatory approval in other countries we must comply with numerous and varying regulatory requirements of such countries regarding safety and efficacy. Other countries also have their own regulations governing, among other things, clinical trials and commercial sales, as well as pricing and distribution of drugs, and we may be required to expend significant resources to obtain regulatory approval and to comply with ongoing regulations in these jurisdictions.

The success of our current and future product candidates will depend on several factors, including the following:

- successful completion of clinical trials and preclinical studies;
- sufficiency of our financial and other resources to complete the necessary preclinical studies and clinical trials;
- acceptance of investigational new drug applications (INDs) for our planned clinical trials or future clinical trials;
- successful enrollment and completion of clinical trials, particularly where competitors may also be recruiting patients;
- data from our clinical programs that supports an acceptable risk-benefit profile of our product candidates in the intended populations;
- receipt and maintenance of marketing approvals from applicable regulatory authorities;

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- establishing agreements with third-party manufacturers for clinical supply for our clinical trials and commercial manufacturing, if one of our product candidates is approved;
- entry into collaborations to further the development of our product candidates;
- · obtaining and maintaining our portfolio of intellectual property rights, including patents, trade secrets and know-how;
- enforcing and defending intellectual property rights and claims;
- obtaining and maintaining regulatory exclusivity for our product candidates;
- successfully launching commercial sales of our product candidates, if approved;
- acceptance of the product candidate's benefits and uses, if approved, by patients, the medical community and third-party payors;
- the prevalence, duration and severity of potential side effects or other safety issues experienced with our product candidates following approval;
- effectively competing with other therapies; and
- obtaining and maintaining healthcare coverage and adequate reimbursement from third-party payors.

If we are not successful with respect to one or more of these factors in a timely manner or at all, we could experience significant delays or an inability to successfully commercialize the product candidates we develop, which would materially harm our business. If we do not receive marketing approvals for any product candidate we develop, we may not be able to continue our operations.

Preclinical development is uncertain. Our preclinical programs may experience delays or may never advance to clinical trials, which would adversely affect our ability to obtain regulatory approvals or commercialize our product candidates on a timely basis or at all, which would have an adverse effect on our business.

In order to obtain approval from the FDA or comparable foreign authorities to market a new small molecule product, we must demonstrate proof of safety and efficacy in humans. To meet these requirements, we will have to conduct adequate and well-controlled clinical trials. Before we can commence clinical trials for a product candidate, we must complete extensive preclinical studies that support our planned INDs in the United States. We have two product candidates in clinical development and the rest of our programs are in preclinical research or development. We cannot be certain of the timely completion or outcome of our preclinical studies and cannot predict if the FDA or foreign authorities will accept our proposed clinical programs or if the outcome of our preclinical studies will ultimately support further development of our programs. As a result, we cannot be sure that we will be able to submit INDs or similar applications on the timelines we expect, if at all, and we cannot be sure that submission of INDs or similar applications will result in the FDA or other regulatory authorities allowing additional clinical trials to begin.

Conducting preclinical testing is a lengthy, time-consuming and expensive process. The length of time may vary substantially according to the type, complexity and novelty of the program, and often can be several years or more per program. Delays associated with programs for which we are directly conducting preclinical studies may cause us to incur additional operating expenses. Moreover, we may be affected by delays or decisions to discontinue development associated with the studies of certain programs that are the responsibility of Sanofi or our potential future partners over which we have no control. For example, AstraZeneca has informed us that, for reasons unrelated to RMC-4630, they have made a strategic decision not to move forward with a clinical combination study incorporating RMC-4630. The commencement and rate of completion of preclinical studies and clinical trials for a product candidate may be delayed by many factors, including, for example:

- inability to generate sufficient preclinical or other in vivo or in vitro data to support the initiation of clinical studies;
- delays in reaching a consensus with regulatory agencies on study design and obtaining regulatory authorization to commence clinical trials;
- obtaining sufficient quantities of starting materials, intermediate materials and our product candidates for use in preclinical studies and clinical trials from third-party suppliers on a timely basis; and
- delays due to the COVID-19 pandemic, including the introduction of remote work policies, or reduced workforce resulting from illness, or delays at our third-party contract research organizations throughout the world, due to similar restrictions imposed by governments or reduced workforce resulting from illness.

Moreover, even if clinical trials do begin for our preclinical programs, our development efforts may not be successful, and clinical trials that we conduct or that third parties conduct on our behalf may not demonstrate sufficient safety or efficacy to obtain the requisite regulatory approvals for any product candidates we develop. Even if we obtain positive results from preclinical studies or initial clinical trials, we may not achieve the same success in future trials.

Some of our programs focus on the discovery and development of "Beyond Rule of 5" small molecules. Such molecules can be associated with longer development timelines and greater costs compared to traditional small molecule drugs. Our "Beyond Rule of 5" product candidates may take longer to develop and/or manufacture relative to traditional small molecules, and we may not be able to formulate "Beyond Rule of 5" candidates for certain routes of administration.

We enlist various technologies and capabilities that give us chemical access to challenging sites on target proteins that generally are not accessible using conventional small molecule drug discovery approaches. For each target, we consider the specific structural, physico-chemical, functional and dynamic properties of the target and deploy the approach or approaches that appear most likely to yield viable development candidates. The "Rule of 5" is a set of criteria used in pharmaceutical drug development to determine whether chemical compounds have certain physico-chemical properties that make them likely to be orally active drugs in humans. In some instances, the compounds we discover and develop are traditional small molecules (i.e. less than 500 daltons) with properties that generally satisfy conventional pharmaceutical "Rule of 5" criteria, while in other cases, they are larger (i.e. more than 500 daltons) "Beyond Rule of 5" (BRo5) compounds that do not satisfy these criteria. For example, our mTORC1 program and our RAS(ON) Inhibitors each include pursuit of BRo5 compounds.

BRo5 compounds have been successfully pursued by many pharmaceutical companies. Examples of BRo5 compounds include natural products and semisynthetic derivatives, peptidomimetics, macrocycles and degraders. However, larger molecular weight small molecules often cannot be formulated into orally absorbed drugs and also often face solubility, potency, bioavailability and stability challenges, among others. In addition, many of the commonly used predictive and other drug development tools are designed specifically for traditional Rule of 5 small molecule drugs rather than BRo5 molecules, contributing to the difficulty and uncertainty of development of BRo5 compounds.

Due to their size and complexity, drug development of our BRo5 compounds may be slower and/or more expensive than drug development of traditional "Rule of 5" compounds, resulting in program delays, increased costs or failure to obtain regulatory approval in a commercially reasonable timeframe, if at all. Our competitors developing traditional small molecules in areas where we are developing BRo5 compounds could obtain regulatory approval and reach the market before we do. Even if we succeed in generating an approved drug from a BRo5 compound, it may be less convenient to administer, have higher grade and/or more frequent side effects or be more costly to manufacture and formulate than competing products on the market. The discovery and development of BRo5 small molecules may pose risks to us such as:

- BRo5 small molecules may present difficult synthetic chemistry and manufacturing challenges, including with any scale-up of our product candidates in sufficient quality and quantity;
- BRo5 small molecules may be challenging to purify, including with any scale-up of our product candidates in sufficient quality and quantity;
- BRo5 small molecules may present solubility challenges;
- BRo5 small molecules may present oral absorption challenges due to low passive permeability, and may not achieve acceptable oral bioavailability for development and may result in poor pharmaceutical properties for formulation development;
- BRo5 small molecules may present cell permeability challenges, especially with regards to lipophilicity, hydrogen bond donor and rotatable bond count, and high topological polar surface area;
- BRo5 small molecules may have a propensity to be substrates for efflux proteins such as the adenosine triphosphate (ATP) binding cassette (ABC) transporter protein family, including multidrug resistance protein 1. Cancer cells may overexpress these transporter proteins causing an increase in expulsion of BRo5 small molecules from the cell. For example, as the site of action of our RAS(ON) Inhibitors is inside the cell, expulsion by these transporter proteins may decrease the effective concentration in the cell sufficiently to reduce target inhibition and thereby render a RAS-dependent tumor less susceptible to the inhibitory activity of a BRo5 small molecule, such as our product candidates;
- BRo5 small molecules may present central nervous system (CNS) penetration challenges due to low passive permeability and/or interaction with efflux transporters at the blood-brain barrier and this could limit sensitivity of CNS tumors to BRo5 small molecules;

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- BRo5 small molecules may present formulation vehicle challenges for administration, such as intravenous and subcutaneous administration, due to
 aspects such as solubility and hydrophobicity;
- BRo5 small molecules may present stability and shelf-life limitations due to the incorporation of labile functionality in their scaffolds, including for example in the development of RMC-5552 which currently requires a cold chain storage of zero degrees Celsius; and
- BRo5 small molecules may present off-target toxicities due to physico-chemical properties such as lipophilicity, which is the ability to dissolve fats, oils and lipids, the presence of off-target pharmacophores in the molecule that can interact with other cellular proteins, or other characteristics that have not been fully characterized within a novel chemical scaffold or platform.

These and other risks related to our research and development of BRo5 small molecules may result in delays in development, an increase in development costs and/or the failure to develop any BRo5 small molecule to approval. As a result, our competitors may develop products more rapidly and cost effectively than we do if they are able to target the same indications as our product candidates using conventional small molecules. In particular, competitors may develop and commercialize a product that competes with a RAS(ON) Inhibitor product candidate we may develop, as some of our competitors in this area are pursuing conventional small molecules directed to other forms of RAS, such as RAS(OFF), and are further along in development than we currently are.

The regulatory approval processes of the FDA, the EMA and comparable foreign authorities are lengthy, time-consuming and inherently unpredictable, and if we are ultimately unable to obtain regulatory approval for our product candidates, our business will be substantially harmed.

The time required to obtain approval by the FDA, the EMA and comparable foreign authorities is unpredictable but typically takes many years following the commencement of clinical trials and depends upon numerous factors, including the substantial discretion of the regulatory authorities. In addition, approval policies, regulations, or the type and amount of clinical data necessary to gain approval may change during the course of a product candidate's clinical development and may vary among jurisdictions. We have not obtained regulatory approval for any product candidate and it is possible that none of our current or future product candidates will ever obtain regulatory approval.

Our current and future product candidates could fail to receive regulatory approval for many reasons, including the following:

- the FDA, the EMA or comparable foreign regulatory authorities may disagree with the design or implementation of our clinical trials;
- we may be unable to demonstrate to the satisfaction of the FDA, the EMA or comparable foreign regulatory authorities that a product candidate is safe or effective for its proposed indication or indications;
- the results of clinical trials may not meet the level of statistical significance required by the FDA, the EMA or comparable foreign regulatory authorities for approval;
- we may be unable to demonstrate that a product candidate's clinical and other benefits outweigh its safety risks;
- the FDA, the EMA or comparable foreign regulatory authorities may disagree with our interpretation of data from clinical trials or preclinical studies;
- the data collected from clinical trials of our product candidates may not be sufficient to support the submission of an NDA to the FDA or other submission or to obtain regulatory approval in the United States, the European Union or elsewhere;
- the FDA, the EMA or comparable foreign regulatory authorities may find deficiencies with or fail to approve the manufacturing processes or facilities of third-party manufacturers with which we contract for clinical and commercial supplies; and
- the approval policies or regulations of the FDA, the EMA or comparable foreign regulatory authorities may significantly change in a manner rendering our clinical data insufficient for approval.

This lengthy approval process as well as the unpredictability of clinical trial results may result in our failing to obtain regulatory approval to market any product candidate we develop. The FDA, the EMA and other comparable foreign authorities have substantial discretion in the approval process, and determining when or whether regulatory approval will be obtained for any product candidate that we develop. Even if we believe the data collected from future clinical trials of our product candidates are promising, this data may not be sufficient to support approval by the FDA, the EMA or any other regulatory authority.

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In addition, even if we were to obtain approval, regulatory authorities may approve any of our product candidates for fewer or more limited indications than we request, may not approve the price we may desire to charge for our products, may grant approval contingent on the performance of costly postmarketing clinical trials, or may approve a product candidate with a label that does not include the labeling claims necessary or desirable for the successful commercialization of that product candidate. Any of the foregoing scenarios could materially harm the commercial prospects for our product candidates.

Further, we have not previously submitted an NDA to the FDA, or a Marketing Authorization Application (MAA) to the EMA. We cannot be certain that any of our programs will be successful in clinical trials or receive regulatory approval. Further, product candidates we develop may not receive regulatory approval even if they are successful in clinical trials. If we do not receive regulatory approvals for our product candidates, we may not be able to continue our operations.

Clinical product development involves a lengthy and expensive process, with uncertain outcomes. We may experience delays in completing, or ultimately be unable to complete, the development and commercialization of our current and future product candidates.

To obtain the requisite regulatory approvals to commercialize any of our product candidates, we must demonstrate through extensive preclinical studies and clinical trials that our products are safe or effective in humans. Clinical testing is expensive and can take many years to complete, and its outcome is inherently uncertain. Failure can occur at any time during the clinical trial process and our future clinical trial results may not be successful. For example, we recently determined to deprioritize our RMC-4630-02 study and are no longer enrolling patients in this study.

We may experience delays in completing our clinical trials or preclinical studies and initiating or completing additional clinical trials. We may also experience numerous unforeseen events during our clinical trials that could delay or prevent our ability to receive marketing approval or commercialize the product candidates we develop, including:

- actions by regulators, Institutional Review Boards (IRBs) or ethics committees may cause us or our investigators to not commence or conduct a clinical trial at a prospective trial site or at all sites and cause us to pause or stop an in-process clinical trial;
- we may experience delays in reaching, or fail to reach, agreement on acceptable terms with prospective trial sites and prospective contract research organizations (CROs);
- the number of patients required for clinical trials may be larger than we anticipate;
- it may be difficult to enroll a sufficient number of patients for our clinical trials or enrollment in these clinical trials may be slower than we anticipate, including in both cases because appropriate patients must have the relevant mutations in the signaling pathways our therapies are designed to target;
- participants may drop out of these clinical trials or fail to return for post-treatment follow-up at a higher rate than we anticipate;
- patients may not comply with our clinical trial protocols, particularly with respect to intermittent dosing, which we are evaluating for our product candidates;
- our third-party contractors may fail to comply with regulatory requirements or meet their contractual obligations to us in a timely manner, or at all, or
 may deviate from the clinical trial protocol or drop out of the trial, which may require that we add new clinical trial sites or investigators;
- the supply or quality of materials for product candidates we develop or other materials necessary to conduct clinical trials may be insufficient or inadequate; and
- our collaborators may delay the development process by waiting to take action or focusing on other priorities.

We could encounter delays if a clinical trial is suspended or terminated by us, by the IRBs or ethics committees of the institutions in which such trials are being conducted, by the Data Safety Monitoring Board (DSMB) for such trial or by the FDA or other regulatory authorities. Such authorities may impose such a suspension or termination due to a number of factors, including failure to conduct the clinical trial in accordance with regulatory requirements or our clinical protocols, inspection of the clinical trial operations or trial site by the FDA or other regulatory authorities resulting in the imposition of a clinical hold, unforeseen safety issues or adverse side effects, failure to demonstrate a benefit from using a product, changes in governmental regulations or administrative actions or lack of adequate funding to continue the clinical trial. Many of the factors that cause, or lead to, a delay in the commencement or completion of clinical trials may also ultimately lead to the denial of marketing approval of our product candidates.

If we experience delays in the completion of, or termination of, any clinical trial of our product candidates, the commercial prospects of our product candidates will be harmed, and our ability to generate product revenues from any of these product candidates will be



delayed. In addition, any delays in completing our clinical trials will increase our costs, slow down our product candidate development and approval process and jeopardize our ability to commence product sales and generate revenues. Clinical trial delays could also allow our competitors to bring products to market before we do or shorten any periods during which we have the exclusive right to commercialize our product candidates and impair our ability to commercialize our product candidates.

Our clinical trial sites may be affected by the COVID-19 outbreak due to prioritization of hospital resources toward the COVID-19 outbreak, travel or quarantine restrictions imposed by governments, and the inability to access sites for initiation and patient monitoring and enrollment. As a result, patient screening, new patient enrollment, monitoring and data collection may be affected or delayed. We are aware that several clinical sites involved in our RMC-4630 clinical studies temporarily stopped or delayed enrolling new patients, with exemptions if appropriate, and it is possible that these or other clinical sites may be similarly affected in the future. These developments may delay our clinical trial timelines. Our clinical trials currently permit patients to receive COVID-19 vaccines while they are on study. The potential impact of our candidates on the safety and efficacy of COVID-19 vaccines, and the potential impact on of COVID-19 vaccines on the safety and efficacy of our candidates is unknown at this time, but it possible that adverse impacts will negatively affect our clinical trials.

Some of our third-party manufacturers which we use for the supply of materials for product candidates or other materials necessary to manufacture product to conduct clinical trials and contract research organizations that we may utilize may be impacted by COVID-19, and should they experience disruptions, such as temporary closures or suspension of services, we would likely experience delays in advancing these trials.

Many of the factors described above that cause, or lead to, a delay in the commencement or completion of clinical trials may also ultimately lead to the denial of regulatory approval of our product candidates or result in the development of our product candidates being stopped early.

Historically, direct inhibition of any RAS protein has been challenging due to a lack of tractable, or "druggable," binding pockets and we are not aware of any programs in clinical development that have successfully targeted any RAS(ON) protein. Given this approach is unproven, it may not be successful.

Historically, direct inhibition of any RAS protein has been challenging due to a lack of tractable, or "druggable," binding pockets. Our tri-complex technology has enabled us to design potent, cell-active inhibitors of multiple mutant RAS(ON) proteins. We are not aware of any programs in clinical development that have successfully targeted any RAS(ON) protein. We cannot be certain that our approach will lead to the development of approvable or marketable products, alone or in combination with other therapies.

The results of preclinical studies and early-stage clinical trials may not be predictive of future results.

The results of preclinical studies may not be predictive of the results of clinical trials, and the results of any early-stage clinical trials we commence may not be predictive of the results of the later-stage clinical trials. Product candidates in later stages of clinical trials may fail to show the desired safety and efficacy despite having progressed through preclinical studies and initial clinical trials. For example, historically, targeted therapies have been susceptible to resistance mutations in cancer cells that facilitate escape from anti-tumor response. Should such resistance mutations arise in patients being treated with our product candidates, the clinical benefit associated with those candidates may be compromised.

There can be no assurance that any of our current or future clinical trials will ultimately be successful or support further clinical development of any of our product candidates. There is a high failure rate for drugs proceeding through clinical trials. A number of companies in the pharmaceutical and biotechnology industries have suffered significant setbacks in clinical development even after achieving promising results in earlier studies. Even if our clinical trials are completed, the results may not be sufficient to obtain regulatory approval of any products.

Interim, "topline" and preliminary data from our clinical trials may differ materially from the final data.

From time to time, we may disclose interim data from our clinical trials. For example, we have reported interim Phase 1 clinical data for RMC-4630 as a single agent and interim Phase 1b/2 clinical data for RMC-4630 in combination with the MEK inhibitor cobimetinib. In each case, this interim data included a limited number of patients and time of exposure to the study drug. Interim data from clinical trials are subject to the risk that one or more of the clinical outcomes may materially change as patient enrollment continues and more data on existing patients become available. Our clinical trial program is ongoing, and the final results may be materially different from those reflected in any interim data we report.

From time to time, we may also publicly disclose preliminary or "topline" data from our clinical trials, which are based on a preliminary analysis of thenavailable data, and the results and related findings and conclusions are subject to change following a more comprehensive review of the data related to the particular trial. We also make assumptions, estimations, calculations and conclusions as part of our analyses of data, and we may not have received or had the opportunity to fully and carefully evaluate all data. As a result, the topline results that we report may differ from future results of the same clinical trials, or different conclusions or considerations may qualify such topline results once additional data have been received and fully evaluated. Topline data also remain subject to audit and verification procedures that may result in the final data being materially different from the preliminary data we previously published. As a result, topline data should be viewed with caution until the final data are available.

Further, others, including regulatory agencies, may not accept or agree with our assumptions, estimates, calculations, conclusions or analyses or may interpret or weigh the importance of data differently, which could impact the value of the particular program, the approvability or commercialization of the particular product candidate or product and the value of our company in general. In addition, the information we choose to publicly disclose regarding a particular study or clinical trial is typically a summary of extensive information, and you or others may not agree with what we determine is the material or otherwise appropriate information to include in our disclosure, and any information we determine not to disclose may ultimately be deemed significant with respect to future decisions, conclusions, views, activities or otherwise regarding a particular product, product candidate or our business. If the topline data that we report differ from actual results, or if others, including regulatory authorities, disagree with the conclusions reached, our ability to obtain approval for, and commercialize, our product candidates may be harmed.

If we encounter difficulties enrolling patients in our clinical trials, our clinical development activities could be delayed or otherwise be adversely affected.

The timely completion of clinical trials in accordance with their protocols depends, among other things, on our ability to enroll a sufficient number of patients who remain in the trial until its conclusion. We may experience difficulties in patient enrollment in our clinical trials for a variety of reasons. The enrollment of patients depends on many factors, including:

- the patient eligibility criteria defined in the protocol;
- our ability to enroll a sufficient number of patients with mutations in the signaling pathways our therapies are designed to target;
- the size of the patient population required for analysis of the trial's primary endpoints;
- the proximity of patients to study sites;
- the design of the trial;
- our ability to recruit clinical trial investigators with the appropriate competencies and experience;
- clinicians' and patients' perceptions as to the potential advantages of the product candidate being studied in relation to other available therapies, including any new products that may be approved for the indications we are investigating;
- our ability to obtain and maintain patient consents for participation in our clinical trials and, where appropriate, biopsies for future patient enrichment efforts;
- the risk that patients enrolled in clinical trials will not remain on the trial through the completion of evaluation; and
- the ability of our clinical trial investigators to enroll patients in cases of outbreak of disease, including COVID-19, or other natural disasters.

In addition, our clinical trials will compete with approved therapies, including sotorasib, as well as other clinical trials for product candidates that are in the same therapeutic areas (and that seek to evaluate patients with cancer cells having the same mutations, particularly with patients having KRASG^{12C} mutations) as our current and potential future product candidates. This competition and competition with approved therapies, including sotorasib, will reduce the number and types of patients available to us, because some patients who might have opted to enroll in our trials may instead opt to pursue a treatment regime using an approved therapy or enroll in a trial conducted by one of our competitors. Since the number of qualified clinical investigators is limited, we conduct some of our clinical trials at the same clinical trial sites that some of our competitors use, which will reduce the number of patients who are available for our clinical trials at such sites. Moreover, because our current and potential future product candidates may represent a departure from more commonly used methods for cancer treatment, potential patients and their doctors may be inclined to use conventional therapies, such as chemotherapy, rather than enroll patients in our ongoing or any future clinical trials.

Our clinical trial sites for our RMC-4630 and RMC-5552 clinical studies may be affected by the COVID-19 outbreak due to prioritization of hospital resources toward the COVID-19 outbreak, travel or quarantine restrictions imposed by governments, and the inability to access sites for initiation and patient monitoring and enrollment. As a result, patient screening, new patient enrollment, monitoring and data collection may be affected or delayed. We are aware that several clinical sites involved in our RMC-4630 clinical studies temporarily stopped or delayed enrolling new patients, with exemptions if appropriate, and it is possible that these or other clinical sites may be similarly affected in the future. These developments may delay our clinical trial timelines. Some of our third-party manufacturers which we use for the supply of materials for product candidates or other materials necessary to manufacture product to conduct clinical trials and contract research organizations that we may utilize may be impacted by COVID-19, and should



they experience disruptions, such as temporary closures or suspension of services, we would likely experience delays in advancing these trials.

Delays in patient enrollment may result in increased costs or may affect the timing or outcome of clinical trials, which could prevent completion of these trials and adversely affect our ability to advance the development of the product candidates we develop.

Our current or future product candidates may cause undesirable side effects or have other properties when used alone or in combination with other approved products or investigational new drugs that could delay or halt their clinical development, prevent their marketing approval, limit their commercial potential or result in significant negative consequences.

Undesirable or clinically unmanageable side effects could occur and cause us or regulatory authorities to interrupt, delay or halt clinical trials and could result in a more restrictive label or the delay or denial of marketing approval by the FDA or comparable foreign regulatory authorities. Results of our trials could reveal a high and unacceptable severity and prevalence of side effects or unexpected characteristics.

For example, the safety data we have released in 2020 for both our RMC-4630-01 and RMC-4630-02 trials included both serious adverse events (SAEs) and other adverse events (AEs). Our recent decision to deprioritize the cobimetinib arm of our RMC-4630-02 study and to stop enrolling patients in this arm of the study was based on part on the safety data we generated.

Although our current and future product candidates will undergo safety testing to the extent possible and, where applicable, under such conditions discussed with regulatory authorities, not all adverse effects of drugs can be predicted or anticipated. Unforeseen side effects could arise either during clinical development or, if such side effects are more rare, after our products have been approved by regulatory authorities and the approved product has been marketed, resulting in the exposure of additional patients. So far, we have not demonstrated that RMC-4630 or any other product candidate is safe in humans, and we cannot predict if ongoing or future clinical trials will do so.

Furthermore, certain of our product candidates, such as RMC-4630, are currently being, and may in the future be, co-administered with approved or experimental therapies, such as Roche's MEK inhibitor cobimetinib, Amgen's KRASG^{12C}(OFF) inhibitor sotorasib, Merck's PD-1 inhibitor pembrolizumab, AstraZeneca's EGFR inhibitor osimertinib, or Lilly's investigational ERK inhibitor LY3214996, which is the subject of a potential investigator sponsored trial with the Netherlands Cancer Institute. These combinations may have additional side effects. For example, overlapping on-pathway toxicities with the combination of RMC-4630 and osimertinib were anticipated and observed and our recent decision to deprioritize the osimertinib arm of our RMC-4630-02 study and to stop enrolling patients in this arm of the study was based in part on what we believe were on-pathway toxicities. The uncertainty resulting from the use of our product candidates in combination with other therapies may make it difficult to accurately predict side effects in future clinical trials.

If any of our product candidates receives marketing approval and we or others later identify undesirable side effects caused by such products, a number of potentially significant negative consequences could result, including:

- regulatory authorities may withdraw their approval of the product;
- we may be required to recall a product or change the way such product is administered to patients;
- additional restrictions may be imposed on the marketing of the particular product or the manufacturing processes for the product or any component thereof;
- regulatory authorities may require the addition of labeling statements, such as a "black box" warning or a contraindication;
- we may be required to implement a Risk Evaluation and Mitigation Strategy (REMS) or create a Medication Guide outlining the risks of such side effects for distribution to patients;
- we could be sued and held liable for harm caused to patients;
- the product may become less competitive; and
- our reputation may suffer.

Any of the foregoing events could prevent us from achieving or maintaining market acceptance of the particular product candidate, if approved. In addition, if one or more of our product candidates prove to be unsafe, our entire technology platform and pipeline could be affected.



Even if we complete the necessary preclinical studies and clinical trials, the marketing approval process is expensive, time-consuming and uncertain and may prevent us or any of our existing or potential future collaboration partners from obtaining approvals for the commercialization of any product candidate we develop.

Any current or future product candidate we may develop and the activities associated with their development and commercialization, including their design, testing, manufacture, safety, efficacy, recordkeeping, labeling, storage, approval, advertising, promotion, sale, and distribution, are subject to comprehensive regulation by the FDA and other regulatory authorities in the United States and by comparable authorities in other countries. Failure to obtain marketing approval for a product candidate will prevent us from commercializing the product candidate in a given jurisdiction. We have not received approval to market any product candidates from regulatory authorities in any jurisdiction and it is possible that none of our current or future product candidates will ever obtain regulatory approval. We have no experience in filing and supporting the applications necessary to gain marketing approvals and expect to rely on third-party CROs or regulatory consultants to assist us in this process. Securing regulatory approval requires the submission of extensive preclinical and clinical data and supporting information to the various regulatory authorities for each therapeutic indication to establish the product candidate's safety and efficacy. Securing regulatory approval also requires the submission of information about the product manufacturing process to, and inspection of manufacturing facilities by, the relevant regulatory authority. Any product candidates we develop may not be effective, may be only moderately effective, or may prove to have undesirable or unintended side effects, toxicities or other characteristics that may preclude our obtaining marketing approval or prevent or limit commercial use.

The process of obtaining marketing approvals, both in the United States and abroad, is expensive, may take many years if additional clinical trials are required, if approval is obtained at all, and can vary substantially based upon a variety of factors, including the type, complexity, and novelty of the product candidates involved. Changes in marketing approval policies during the development period, changes in or the enactment of additional statutes or regulations, or changes in regulatory review for each submitted product application, may cause delays in the approval or rejection of an application. The FDA and comparable authorities in other countries have substantial discretion in the approval process and may refuse to accept any application or may decide that our data are insufficient for approval and require additional preclinical, clinical or other studies. In addition, varying interpretations of the data obtained from preclinical and clinical testing could delay, limit, or prevent marketing approval of a product candidate. Any marketing approval we ultimately obtain may be limited or subject to restrictions or post-approval commitments that render the approved product not commercially viable.

If we experience delays in obtaining approval or if we fail to obtain approval of any current or future product candidates we may develop, the commercial prospects for those product candidates may be harmed.

Obtaining and maintaining marketing approval of our current and future product candidates in one jurisdiction does not mean that we will be successful in obtaining marketing approval of our current and future product candidates in other jurisdictions.

Obtaining and maintaining marketing approval of our current and future product candidates in one jurisdiction does not guarantee that we will be able to obtain or maintain marketing approval in any other jurisdiction, while a failure or delay in obtaining marketing approval in one jurisdiction may have a negative effect on the marketing approval process in others. For example, even if the FDA grants marketing approval of a product candidate, comparable regulatory authorities in foreign jurisdictions must also approve the manufacturing, marketing and promotion of the product candidate in those countries. Approval procedures vary among jurisdictions and can involve requirements and administrative review periods different from, and greater than, those in the United States, including additional preclinical studies or clinical trials as clinical studies conducted in one jurisdiction may not be accepted by regulatory authorities in other jurisdictions. In many jurisdictions outside the United States, a product candidate must be approved for reimbursement before it can be approved for sale in that jurisdiction. In some cases, the price that we may charge for our products is also subject to approval.

We may also submit marketing applications in other countries. Regulatory authorities in jurisdictions outside of the United States have requirements for approval of product candidates with which we must comply prior to marketing in those jurisdictions. Obtaining foreign marketing approvals and compliance with foreign regulatory requirements could result in significant delays, difficulties and costs for us and could delay or prevent the introduction of our products in certain countries. If we fail to comply with the regulatory requirements in international markets or receive applicable marketing approvals, our target market will be reduced and our ability to realize the full market potential of our product candidates will be harmed.

Adverse events in the field of oncology or the biopharmaceutical industry could damage public perception of our current or future product candidates and negatively affect our business.

The commercial success of our products will depend in part on public acceptance of the use of targeted cancer therapies. While a number of targeted cancer therapies have received regulatory approval and are being commercialized, our approach to targeting cancer cells carrying tumor causing mutations, including oncogenic RAS(ON) pathway mutations, is novel and unproven. Adverse events in



clinical trials of our product candidates, or post-marketing activities, or in clinical trials of others developing similar products or that are related to approved targeted therapies, particularly those targeting oncogenic RAS pathway mutations, including sotorasib, and the resulting publicity, as well as any other adverse events in the field of oncology that may occur in the future, could result in a decrease in demand for any product that we may develop. If public perception is influenced by claims that the use of cancer therapies is unsafe, whether related to our therapies or those of our competitors, our products may not be accepted by the general public or the medical community.

Future adverse events in oncology or the biopharmaceutical industry could also result in greater governmental regulation, stricter labeling requirements and potential regulatory delays in the testing or approvals of our products. Any increased scrutiny could delay or increase the costs of obtaining marketing approval for the product candidates we develop.

Even if we receive marketing approval of a product candidate, we will be subject to ongoing regulatory obligations and continued regulatory review, which may result in significant additional expense and we may be subject to penalties if we fail to comply with regulatory requirements or experience unanticipated problems with our products, if approved.

Any marketing approvals that we receive for any current or future product candidate may be subject to limitations on the approved indicated uses for which the product may be marketed or the conditions of approval, or contain requirements for potentially costly post-market testing and surveillance to monitor the safety and efficacy of the product candidate. The FDA may also require REMS as a condition of approval of any product candidate, which could include requirements for a medication guide, physician communication plans or additional elements to ensure safe use, such as restricted distribution methods, patient registries and other risk minimization tools. In addition, if the FDA or a comparable foreign regulatory authority approves a product candidate, the manufacturing processes, labeling, packaging, distribution, adverse event reporting, storage, advertising, promotion, import and export and record keeping for the product candidate will be subject to extensive and ongoing regulatory requirements. These requirements include submissions of safety and other post-marketing information and reports, registration, as well as continued compliance with current Good Manufacturing Practice (cGMP) and Good Clinical Practice (GCP) for any clinical trials that we conduct post-approval. Later discovery of previously unknown problems with any approved candidate, including adverse events of unanticipated severity or frequency, or with our third-party manufacturers or manufacturing processes, or failure to comply with regulatory requirements, may result in, among other things:

- restrictions on the marketing or manufacturing of the product, withdrawal of the product from the market, or product recalls;
- fines, untitled and warning letters, or holds on clinical trials;
- refusal by the FDA to approve pending applications or supplements to approved applications we filed or suspension or revocation of license approvals;
- · product seizure or detention, or refusal to permit the import or export of the product; and
- injunctions or the imposition of civil or criminal penalties.

The FDA's and other regulatory authorities' policies may change, and additional government regulations may be enacted that could prevent, limit or delay marketing approval of a product. We cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative action, either in the United States or abroad. If we are slow or unable to adapt to changes in existing requirements or the adoption of new requirements or policies, or if we are not able to maintain regulatory compliance, we may lose any marketing approval that we may have obtained.

Even if a current or future product candidate receives marketing approval, it may fail to achieve the degree of market acceptance by physicians, patients, third-party payors and others in the medical community necessary for commercial success.

If any current or future product candidate we develop receives marketing approval, whether as a single agent or in combination with other therapies, it may nonetheless fail to gain sufficient market acceptance by physicians, patients, third-party payors, and others in the medical community to be a viable product. For example, current approved immunotherapies, and other cancer treatments like chemotherapy and radiation therapy, are well established in the medical community, and doctors may continue to rely on these therapies. The degree of market acceptance of any product candidate, if approved for commercial sale, will depend on a number of factors, including:

- efficacy and potential advantages compared to alternative treatments;
- the ability to offer our products, if approved, for sale at competitive prices;
- convenience and ease of administration compared to alternative treatments;
- the willingness of the target patient population to try new therapies and of physicians to prescribe these therapies;

- the strength of marketing and distribution support;
- the ability to obtain sufficient third-party coverage and adequate reimbursement, including with respect to the use of the approved product as a combination therapy;
- adoption of a companion diagnostic and/or complementary diagnostic (if any); and
- the prevalence and severity of any side effects.

The market opportunities for any current or future product candidate we develop, if and when approved, may be limited to those patients who are ineligible for established therapies or for whom prior therapies have failed, and may be small.

Cancer therapies are sometimes characterized as first-line, second-line or third-line, and the FDA often approves new therapies initially only for third-line use. When cancer is detected early enough, first-line therapy, usually chemotherapy, hormone therapy, surgery, radiation therapy or a combination of these, is sometimes adequate to cure the cancer or prolong life without a cure. Second- and third-line therapies are administered to patients when prior therapy is not effective. We expect to initially seek approval of RMC-4630 and any other product candidates we develop as a therapy for patients who have received one or more prior treatments. Subsequently, for those products that prove to be sufficiently beneficial, if any, we would expect to seek approval potentially as a first-line therapy, but there is no guarantee that product candidates we develop, even if approved, would be approved for first-line therapy, and, prior to any such approvals, we may have to conduct additional clinical trials.

The number of patients who have the cancers we are targeting, including those with the necessary mutations, may turn out to be lower than expected. Additionally, the potentially addressable patient population for our current programs or future product candidates may be limited, if and when approved. Even if we obtain significant market share for any product candidate, if and when approved, if the potential target populations are small, we may never achieve commercial success without obtaining marketing approval for additional indications, including to be used as first- or second-line therapy.

We are currently developing and may in the future, develop RMC-4630 and other product candidates in combination with other therapies, which exposes us to additional risks.

We are developing RMC-4630 in combination with Amgen's KRASG12C(OFF) inhibitor sotorasib, and Merck's PD-1 inhibitor pembrolizumab, and may in the future, develop RMC-4630 and other product candidates in combination with one or more approved cancer therapies. Even if any product candidate we develop were to receive marketing approval or be commercialized for use in combination with other existing therapies, we would continue to be subject to the risks that the FDA or similar regulatory authorities outside of the United States could revoke approval of the therapy used in combination with our product candidate or that safety, efficacy, manufacturing or supply issues could arise with these existing therapies. Combination therapies are commonly used for the treatment of cancer, and we would be subject to similar risks if we develop any of our product candidates for use in combination with other drugs or for indications other than cancer. This could result in our own products being removed from the market or being less successful commercially.

We or our collaborators may also evaluate RMC-4630 or any other current or future product candidates in combination with one or more other cancer therapies that have not yet been approved for marketing by the FDA or similar regulatory authorities outside of the United States or with approved cancer therapies at an unapproved dose and/or schedule, and/or with approved cancer therapies in unapproved indications. For example, we plan to provide RMC-4630 to the Netherlands Cancer Institute to support their evaluation of RMC-4630 in combination with Lilly's ERK inhibitor LY3214996. We will not be able to market and sell RMC-4630, or any product candidate we develop in combination with any such cancer therapies, outside existing approved labels that do not ultimately obtain marketing approval.

If the FDA or similar regulatory authorities outside of the United States do not approve the drugs we choose to evaluate in combination with or any product candidate we develop or revoke their approval of, or if safety, efficacy, manufacturing, or supply issues arise with, these drugs, we may be unable to obtain approval of or market or any product candidate we develop.

In addition, Sanofi primarily controls the research and development activities of our SHP2 inhibitors, including RMC-4630, pursuant to the terms of the Sanofi Agreement, and may disagree with us regarding which other therapies should be evaluated in combination with RMC-4630. As a result of any such disagreement, our completion of a trial in combination with our preferred combination product candidate may be delayed or prevented.

We face significant competition, and if our competitors develop and market products that are more effective, safer or less expensive than the product candidates we develop, our commercial opportunities will be negatively impacted.

The life sciences industry is highly competitive. We are currently developing therapies that will compete, if approved, with other products and therapies that currently exist or are being developed. Products we may develop in the future are also likely to face competition from other products and therapies, some of which we may not currently be aware. We have competitors both in the United States and internationally, including major multinational pharmaceutical companies, established biotechnology companies, specialty pharmaceutical companies, universities and other research institutions. Many of our competitors have significantly greater financial, manufacturing, marketing, product development, technical and human resources than we do. Large pharmaceutical companies, in particular, have extensive experience in clinical testing, obtaining marketing approvals, recruiting patients and manufacturing pharmaceutical products. These companies also have significantly greater research and marketing capabilities than we do and may also have products that have been approved or are in late stages of development, and collaborative arrangements in our target markets with leading companies and research institutions. Established pharmaceutical companies may also invest heavily to accelerate discovery and development of novel compounds or to in-license novel compounds that could make the product candidates that we develop obsolete. Mergers and acquisitions in the pharmaceutical and biotechnology industries may result in even more resources being concentrated among a smaller number of our competitors. As a result of all of these factors, our competitors may succeed in obtaining patent protection and/or marketing approval or discovering, developing and commercializing products in our field before we do.

There is a large number of companies developing or marketing treatments for cancer, including many major pharmaceutical and biotechnology companies. These treatments consist of small molecule drug products, biologics, cell-based therapies and traditional chemotherapy. Smaller and other early stage companies may also prove to be significant competitors. In addition, academic research departments and public and private research institutions may be conducting research on compounds that could prove to be competitive.

There are also several programs in development targeting SHP2, including those clinical programs run by Novartis AG, Jacobio Pharmaceuticals Co. Ltd. (licensed to AbbVie Inc.), Relay Therapeutics Inc. (licensed to Roche), Erasca, Inc., Navire Pharma, Inc. and Genhouse Bio Co. Ltd. There are several RAS pathway mutations programs, including those directed at KRAS^{G12C} (OFF) and KRAS^{G12D} mutations, including clinical programs directed at KRAS^{G12C} (OFF) being conducted by Amgen Inc., Mirati Therapeutics, Inc., Roche, Jacobio Pharmaceuticals Co. Ltd., Betta Pharmaceuticals Co., Ltd., Novartis AG, InventisBio, Eli Lilly, and Genhouse Bio Co. Ltd. Other clinical programs directed at mutant RAS are being conducted by Merck & Co./Moderna Therapeutics, Boehringer Ingelheim and Gilead Sciences, Inc. Smaller and other early-stage companies may also prove to be significant competitors. In addition, academic research departments and public and private research institutions may be conducting research on compounds that could prove to be competitive.

Our commercial opportunity could be reduced or eliminated if our competitors develop and commercialize products that are safer, more effective, have fewer or less severe effects, are more convenient, have a broader label, are marketed more effectively, are reimbursed or are less expensive than any products that we may develop. Our competitors also may obtain FDA, EMA or other marketing approval for their products more rapidly than we may obtain approval for ours, which could result in our competitors establishing a strong market position before we are able to enter the market. Even if the product candidates we develop achieve marketing approval, they may be priced at a significant premium over competitive products if any have been approved by then, resulting in reduced competitiveness.

Third parties compete with us in recruiting and retaining qualified scientific and management personnel, establishing clinical trial sites and patient registration for clinical trials, as well as in acquiring technologies complementary to, or necessary for, our programs. In addition, the biopharmaceutical industry is characterized by rapid technological change. If we fail to stay at the forefront of technological change, we may be unable to compete effectively. Technological advances or products developed by our competitors may render our product candidates obsolete, less competitive or not economical.

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Even if we are able to commercialize any product candidates, such products may become subject to unfavorable pricing regulations or third-party coverage and reimbursement policies, which would harm our business.

The regulations that govern marketing approvals, pricing and reimbursement for new products vary widely from country to country. Some countries require approval of the sale price of a product before it can be marketed. In many countries, the pricing review period begins after marketing approval is granted. In some foreign markets, prescription pharmaceutical pricing remains subject to continuing governmental control even after initial approval is granted. As a result, we might obtain marketing approval for a product candidate in a particular country, but then be subject to price regulations that delay our commercial launch of the product candidate, possibly for lengthy time periods, and negatively impact the revenues we are able to generate from the sale of the product candidate in that country. Adverse pricing limitations may hinder our ability to recoup our investment in one or more product candidates, even if our product candidates obtain marketing approval.

Our ability to commercialize any product candidates, whether as a single agent or combination therapy, successfully also will depend in part on the extent to which coverage and reimbursement for these product candidates and related treatments will be available from government authorities, private health insurers and other organizations. Government authorities and third-party payors, such as private health insurers and health maintenance organizations, decide which medications they will pay for and establish reimbursement levels. It is difficult to predict at this time what government authorities and thirdparty payors will decide with respect to coverage and reimbursement for our programs.

A primary trend in the U.S. healthcare industry and elsewhere is cost containment. Government authorities and third-party payors have attempted to control costs by limiting coverage and the amount of reimbursement for particular products and requiring substitutions of generic products and/or biosimilars. Increasingly, third-party payors are requiring that drug companies provide them with predetermined discounts from list prices and are challenging the prices charged for drugs. We cannot be sure that coverage will be available for any product candidate that we commercialize and, if coverage is available, the level of reimbursement. These third-party payors are also examining the cost-effectiveness of drugs in addition to their safety and efficacy.

Reimbursement may impact the demand for, or the price of, any product candidate for which we obtain marketing approval. If reimbursement is not available or is available only to limited levels, we may not be able to successfully commercialize any product candidate for which we obtain marketing approval.

There may be significant delays in obtaining coverage and reimbursement for newly approved drugs, as the process is time-consuming and costly, and coverage may be more limited than the purposes for which the drug is approved by the FDA or comparable foreign regulatory authorities. Additionally, no uniform policy requirement for coverage and reimbursement for drug products exists among third-party payors in the United States, which may result in coverage and reimbursement for drug products that can differ significantly from payor to payor. Moreover, eligibility for reimbursement does not imply that any drug will be paid for in all cases or at a rate that covers our costs, including research, development, manufacture, sale and distribution. Interim reimbursement levels for new drugs, if applicable, may also not be sufficient to cover our costs and may not be made permanent. Reimbursement rates may vary according to the use of the drug and the clinical setting in which it is used, may be based on reimbursement levels already set for lower cost drugs and may be incorporated into existing payments for other services. Net prices for drugs may be reduced by mandatory discounts or rebates required by government healthcare programs or private payors and by any future relaxation of laws that presently restrict imports of drugs from countries where they may be sold at lower prices than in the United States. Third-party payors often rely upon Medicare coverage policy and payment limitations in setting their own reimbursement policies.

We may fail to select or capitalize on the most scientifically, clinically and commercially promising or profitable mutant RAS(ON) targets.

We have limited technical, managerial and financial resources to determine which of our lead generation stage RAS(ON) Inhibitors should be advanced into further preclinical development, initial clinical trials, later-stage clinical development and potential commercialization. From our RAS(ON) Inhibitors that have completed our lead optimization phase of preclinical development, we have selected RMC-6291, our inhibitor targeting KRASG12C/NRASG12C(ON), and RMC-6236, our inhibitor of multiple RAS variants, which we refer to as RASMULTI(ON) for IND-enabling preclinical development. In selecting these or other development candidates, we may make incorrect determinations. Our decisions to allocate our research and development, management and financial resources toward particular development candidates or therapeutic areas may not lead to the development of viable commercial products and may divert resources from better opportunities. Similarly, our decisions to delay or terminate development programs may also be incorrect and could cause us to miss valuable opportunities.



We may not be successful in our efforts to identify or discover other product candidates and may fail to capitalize on programs or product candidates that may present a greater commercial opportunity or for which there is a greater likelihood of success.

The success of our business depends upon our ability to identify, develop and commercialize product candidates. Research programs to identify new product candidates require substantial technical, financial and human resources, and we may fail to identify potential product candidates for numerous reasons.

Additionally, because we have limited resources beyond those provided by Sanofi on SHP2 and RMC-4630, we may forego or delay pursuit of opportunities with certain programs or product candidates or for indications that later prove to have greater commercial potential. However, the advancement of this product candidate may ultimately prove to be unsuccessful or less successful than another program in our pipeline that we might have chosen to pursue on a less aggressive basis. Our estimates regarding the potential market for our product candidates could be inaccurate, and our spending on current and future research and development programs may not yield any commercially viable products. If we do not accurately evaluate the commercial potential for a particular product candidate, we may relinquish valuable rights to that product candidate through collaboration, licensing or other arrangements in cases in which it would have been more advantageous for us to retain sole development and commercialization rights to such product candidate. For example, we licensed worldwide development and commercialization rights with respect to RMC-4630 to Sanofi and will receive only milestone payments, an equal share of profits and losses in the United States and royalties on annual net sales of each product outside the United States. Alternatively, we may allocate internal resources to a product candidate in a therapeutic area in which it would have been more advantageous to enter into a partnering arrangement.

If any of these events occur, we may be forced to abandon or delay our development efforts with respect to a particular product candidate or fail to develop a potentially successful product candidate.

We may need to use existing commercial diagnostic tests or develop, or enter into a collaboration or partnership to develop, novel complementary diagnostics and/or novel companion diagnostics for some of our current or future product candidates. If we or our future partners are unable to successfully develop these companion diagnostics or complementary diagnostics, or experience significant delays in doing so, we may not realize the full commercial potential of our future product candidates.

As one of the key elements of our product development strategy, we seek to identify cancer patient populations that may derive meaningful benefit from our current or future product candidates. Because predictive biomarkers may be used to identify the right patients for our programs and our current or future product candidates, we believe that our success may depend, in part, on our ability to use existing diagnostic tests (such as Foundation Medicine's FoundationOne[®] CDX), or develop novel complementary diagnostics and/or novel companion diagnostics in collaboration with partners.

In the event that novel tests will need to be developed, we have little experience in the development of diagnostics. As such, we expect to rely on future partners in developing appropriate diagnostics to pair with our current or future product candidates. We may be unsuccessful in entering into collaborations for the development of companion diagnostics for our programs and our current or future product candidates.

Complementary diagnostics and/or companion diagnostics are subject to regulation by the FDA and similar regulatory authorities outside the United States as medical devices and require separate regulatory approval or clearance prior to commercialization. If we, our partners, or any third parties that we engage to assist us, are unable to successfully develop complementary diagnostics and/or companion diagnostics for our product candidates and any future product candidates, or experience delays in doing so:

- the development of our product candidates and any other future product candidates may be adversely affected if we are unable to appropriately select patients for enrollment in our clinical trials; and
- we may not realize the full commercial potential of our product candidates and any other future product candidates that receive marketing approval if, among other reasons, we are unable to appropriately identify, or it takes us longer to identify, patients who are likely to benefit from therapy with our products, if approved.

We may seek and fail to obtain fast track or breakthrough therapy designations for our current or future product candidates. If we are successful, these programs may not lead to a faster development or regulatory review process, and they do not guarantee we will receive approval for any product candidate. We may also seek to obtain accelerated approval for one or more of our product candidates but the FDA or foreign regulators may disagree that we have met the requirements for such approval.

If a product is intended for the treatment of a serious or life-threatening condition and preclinical or clinical data demonstrate the potential to address an unmet medical need for this condition, the product sponsor may apply for fast track designation. The FDA has broad discretion whether or not to grant this designation, so even if we believe a particular product candidate is eligible for this designation, the FDA may reach a different conclusion and not grant it. Even if we do receive fast track designation, we may not



experience a faster development process, review or approval compared to conventional FDA procedures. The FDA may rescind the fast track designation if it believes that the designation is no longer supported by data from our clinical development program.

We may also seek breakthrough therapy designation for any product candidate that we develop. A breakthrough therapy is defined as a drug that is intended, alone or in combination with one or more other drugs, to treat a serious or life-threatening disease or condition, and preliminary clinical evidence indicates that the drug may demonstrate substantial improvement over currently approved therapies on one or more clinically significant endpoints, such as substantial treatment effects observed early in clinical development. Like fast track designation, breakthrough therapy designation is within the discretion of the FDA. Accordingly, even if we believe a product candidate we develop meets the criteria for designation as a breakthrough therapy, the FDA may disagree and instead determine not to make such designation. In any event, the receipt of breakthrough therapy designation for a product candidate may not result in a faster development process, review or approval compared to drugs considered for approval under conventional FDA procedures and does not assure ultimate approval by the FDA. In addition, even if a product candidate we develop qualifies as a breakthrough therapy, the FDA may later decide that the drug no longer meets the conditions for qualification and rescind the designation.

Drugs designated as fast track products or breakthrough therapies by the FDA are also eligible for accelerated approval if the product has an effect on a surrogate endpoint that is reasonably likely to predict clinical benefit, or on a clinical endpoint that can be measured earlier than irreversible morbidity or mortality, that is reasonably likely to predict an effect on irreversible morbidity or mortality or other clinical benefit, taking into account the severity, rarity, or prevalence of the condition and the availability or lack of alternative treatments. As a condition of accelerated approval, the FDA will generally require the sponsor to perform adequate and well-controlled post-marketing clinical studies to verify and describe the anticipated effect on irreversible morbidity or mortality or other clinical benefit. In addition, the FDA requires pre-approval of promotional materials for accelerated approval products, once approved. We cannot guarantee that the FDA will agree any of our product candidates has met the criteria to receive accelerated approval, which would require us to conduct additional clinical testing prior to seeking FDA approval. Even if any of our product candidates received approval through this pathway, the product may fail required post-approval confirmatory clinical trials, and we may be required to remove the product from the market or amend the product label in a way that adversely impacts its marketing.

Jurisdictions where we may seek to pursue product candidates outside of the United States have processes similar to the breakthrough designation and fast track processes described above, and to the extent we desire to enter these markets, we will face similar risks and challenges as those described in the United States.

We may seek Orphan Drug Designation for product candidates we develop, and we may be unsuccessful or may be unable to maintain the benefits associated with Orphan Drug Designation, including the potential for market exclusivity.

As part of our business strategy, we may seek Orphan Drug Designation for product candidates we develop. Regulatory authorities in some jurisdictions, including the United States and Europe, may designate drugs for relatively small patient populations as orphan drugs. Under the Orphan Drug Act, the FDA may designate a drug as an orphan drug if it is a drug intended to treat a rare disease or condition, which is generally defined as a patient population of fewer than 200,000 individuals annually in the United States, or a patient population greater than 200,000 in the United States where there is no reasonable expectation that the cost of developing the drug will be recovered from sales in the United States. In the United States, Orphan Drug Designation entitles a party to financial incentives such as opportunities for grant funding towards clinical trial costs, tax advantages and user-fee waivers.

Similarly, in Europe, the European Commission grants Orphan Drug Designation after receiving the opinion of the EMA Committee for Orphan Medicinal Products on an Orphan Drug Designation application. Orphan Drug Designation is intended to promote the development of drugs that are intended for the diagnosis, prevention or treatment of life-threatening or chronically debilitating conditions affecting not more than 5 in 10,000 persons in Europe and for which no satisfactory method of diagnosis, prevention, or treatment has been authorized (or the product would be a significant benefit to those affected). Additionally, designation is granted for drugs intended for the diagnosis, prevention, or treatment of a life-threatening, seriously debilitating or serious and chronic condition and when, without incentives, it is unlikely that sales of the drug in Europe would be sufficient to justify the necessary investment in developing the drug. In Europe, Orphan Drug Designation entitles a party to a number of incentives, such as protocol assistance and scientific advice specifically for designated orphan medicines, and potential fee reductions depending on the status of the sponsor.

Generally, if a drug with an Orphan Drug Designation subsequently receives the first marketing approval for the indication for which it has such designation, the drug is entitled to a period of marketing exclusivity, which precludes the EMA or the FDA from approving another marketing application for the same drug and indication for that time period, except in limited circumstances. The applicable period is seven years in the United States and ten years in Europe. The European exclusivity period can be reduced to six years if a drug no longer meets the criteria for Orphan Drug Designation or if the drug is sufficiently profitable such that market exclusivity is no longer justified.

We may be unsuccessful in obtaining Orphan Drug Designation for our product candidates. In addition, even if we obtain orphan drug exclusivity for a product candidate, that exclusivity may not effectively protect the product candidate from competition because



different therapies can be approved for the same condition. Even after an orphan drug is approved, the FDA can subsequently approve the same drug for the same condition if the FDA concludes that the later drug is clinically superior in that it is shown to be safer, more effective or makes a major contribution to patient care. In addition, a designated orphan drug may not receive orphan drug exclusivity if it is approved for a use that is broader than the indication for which it received orphan designation. Moreover, orphan drug exclusive marketing rights in the United States may be lost if the FDA later determines that the request for designation was materially defective or if the manufacturer is unable to assure sufficient quantity of the drug to meet the needs of patients with the rare disease or condition. Orphan Drug Designation neither shortens the development time or regulatory review time of a drug nor gives the drug any advantage in the regulatory review or approval process. While we may seek Orphan Drug Designation for applicable indications for our current and any future product candidates, we may never receive such designations. Even if we do receive such designations, there is no guarantee that we will enjoy the benefits of those designations.

If product liability lawsuits are brought against us, we may incur substantial liabilities and may be required to limit commercialization of any approved products.

We face an inherent risk of product liability as a result of the clinical testing of product candidates and will face an even greater risk if we commercialize any products. For example, we may be sued if any product candidate we develop causes or is perceived to cause injury or is found to be otherwise unsuitable during clinical testing, manufacturing, marketing or sale. Any such product liability claims may include allegations of defects in manufacturing, defects in design, a failure to warn of dangers inherent in the product, negligence, strict liability or a breach of warranties. Claims could also be asserted under state consumer protection acts. If we cannot successfully defend ourselves against product liability claims, we may incur substantial liabilities or be required to limit commercialization of any approved products. Even successful defense would require significant financial and management resources. Regardless of the merits or eventual outcome, liability claims may result in:

- decreased demand for any approved product;
- injury to our reputation;
- withdrawal of clinical trial participants;
- initiation of investigations by regulators;
- costs to defend the related litigation;
- a diversion of management's time and our resources;
- substantial monetary awards to trial participants or patients;
- product recalls, withdrawals or labeling, marketing or promotional restrictions;
- exhaustion of any available insurance and our capital resources and potential increase in our insurance premiums and/or retention amounts; and
- the inability to commercialize any product candidate.

Our inability to obtain sufficient product liability insurance at an acceptable cost to protect against potential product liability claims could prevent or inhibit the commercialization of products we develop, alone or with collaboration partners.

Insurance coverage is increasingly expensive. We may not be able to maintain insurance at a reasonable cost or in an amount adequate to satisfy any liability that may arise, if at all. Our insurance policy contains various exclusions, and we may be subject to a product liability claim for which we have no coverage. We may have to pay any amounts awarded by a court or negotiated in a settlement that exceed our coverage limitations or that are not covered by our insurance, and we may not have, or be able to obtain, sufficient capital to pay such amounts. Even if our agreements with Sanofi or any future collaborator entitle us to indemnification against losses, such indemnification is limited and may not be available or adequate should any claim arise.

Healthcare legislative reform measures may have a material adverse effect on our business and results of operations.

In the United States, there have been and continue to be a number of legislative initiatives to contain healthcare costs. For example, in March 2010, the Affordable Care Act (the ACA) was passed, which substantially changes the way healthcare is financed by both governmental and private insurers, and significantly impacts the U.S. pharmaceutical industry. The ACA, among other things, increases the minimum Medicaid rebates owed by manufacturers under the Medicaid Drug Rebate Program and extends the rebate program to individuals enrolled in Medicaid managed care organizations, establishes annual fees and taxes on manufacturers of certain branded prescription drugs, and creates a new Medicare Part D coverage gap discount program, in which manufacturers must agree to offer 70% point-of-sale discounts off negotiated prices of applicable brand drugs to eligible beneficiaries during their coverage gap period, as a condition for the manufacturer's outpatient drugs to be covered under Medicare Part D.

Since its enactment, there have been judicial, executive and Congressional challenges to certain aspects of the ACA. On June 17, 2021, the U.S. Supreme Court dismissed the most recent judicial challenge to the ACA brought by several states without specifically ruling on the constitutionality of the ACA. Prior to the Supreme Court's decision, President Biden issued an executive order initiating a special enrollment period from February 15, 2021 through August 15, 2021 for purposes of obtaining health insurance coverage through the ACA marketplace. The executive order also instructed certain governmental agencies to review and reconsider their existing policies and rules that limit access to healthcare. It is unclear how healthcare reform measures enacted by Congress or implemented by the Biden administration or other challenges to the ACA, if any, will impact the ACA or our business.

Other legislative changes have been proposed and adopted in the United States since the ACA was enacted. On August 2, 2011, the Budget Control Act of 2011, among other things, included aggregate reductions of Medicare payments to providers of 2% per fiscal year. These reductions went into effect on April 1, 2013 and, due to subsequent legislative amendments to the statute, will remain in effect through 2030, with the exception of a temporary suspension from May 1, 2020 through December 31, 2021 unless additional congressional action is taken. In addition, in January 2013, the American Taxpayer Relief Act of 2012 was signed into law, which, among other things, further reduced Medicare payments to several types of providers.

Moreover, payment methodologies may be subject to changes in healthcare legislation and regulatory initiatives. We expect that additional state and federal healthcare reform measures will be adopted in the future, any of which could limit the amounts that federal and state governments will pay for healthcare products and services, which could result in reduced demand for any product candidate we develop or complementary diagnostics or companion diagnostics or additional pricing pressures.

Additionally, there has been increasing legislative and enforcement interest in the United States with respect to specialty drug pricing practices. Specifically, there have been several recent U.S. Congressional inquiries and proposed and enacted federal and state legislation designed to, among other things, bring more transparency to drug pricing, reduce the cost of prescription drugs under Medicare, review the relationship between pricing and manufacturer patient programs, and reform government program reimbursement methodologies for drugs.

In addition, FDA regulations and guidance may be revised or reinterpreted by the FDA in ways that may significantly affect our business. For example, the results of the 2020 presidential election may impact our business and industry. The previous presidential administration took several executive actions, including the issuance of a number of Executive Orders, that could impose significant burdens on, or otherwise materially delay, the FDA's ability to engage in routine oversight activities such as implementing statutes through rulemaking, issuance of guidance, and review and approval of marketing applications. It is difficult to predict whether or how these requirements will be interpreted and implemented, or whether they will be rescinded and replaced under the Biden administration. The policies and priorities of an incoming administration are unknown and could materially impact the regulations governing our product candidates. Any new regulations or guidance, or revisions or reinterpretations of existing regulations or guidance, may impose additional costs or lengthen FDA review times for our product candidates. We cannot determine how changes in regulations, statutes, policies, or interpretations when and if issued, enacted or adopted, may affect our business in the future.

Disruptions at the FDA and other government agencies caused by funding shortages or global health concerns could hinder their ability to hire and retain key leadership and other personnel, prevent new products and services from being developed or commercialized in a timely manner or otherwise prevent those agencies from performing normal business functions on which the operation of our business may rely, which could negatively impact our business.

The ability of the FDA to review and approve new products can be affected by a variety of factors, including government budget and funding levels, ability to hire and retain key personnel and accept the payment of user fees, and statutory, regulatory, and policy changes. Average review times at the agency have fluctuated in recent years as a result.

Disruptions at the FDA and other agencies may also slow the time necessary for new drugs to be reviewed and/or approved by necessary government agencies, which would adversely affect our business. For example, over the last several years the U.S. government has shut down several times and certain regulatory agencies, such as the FDA, have had to furlough critical FDA and other government employees and stop critical activities. If a prolonged government shutdown occurs, it could significantly impact the ability of the FDA to timely review and process our regulatory submissions, which could have a material adverse effect on our business.

Separately, in response to the COVID-19 pandemic, on March 10, 2020, the FDA announced its intention to postpone most inspections of foreign manufacturing facilities and products and subsequently, on March 18, 2020, the FDA temporarily postponed routine surveillance inspections of domestic manufacturing facilities. Subsequently, on July 10, 2020, the FDA announced its intention to resume certain on-site inspections of domestic manufacturing facilities subject to a risk-based prioritization system. The FDA has announced that it intends to use this risk-based assessment system to identify the categories of regulatory activity that can occur within a given geographic area, ranging from mission critical inspections to resumption of all regulatory activities. Additionally, on April 15, 2021, the FDA issued a guidance document in which the FDA described its plans to conduct voluntary remote interactive evaluations of certain drug manufacturing facilities and clinical research sites. According to this guidance, the FDA intends to request such remote interactive evaluations in situations where an in-person inspection would not be prioritized, deemed mission-critical, or where direct



inspection is otherwise limited by travel restrictions, but where the FDA determines that remote evaluation would still be appropriate. Regulatory authorities outside the U.S. may adopt similar restrictions or other policy measures in response to the COVID-19 pandemic and may experience delays in their regulatory activities. If a prolonged government shutdown occurs, or if global health concerns continue to prevent the FDA or other regulatory authorities from conducting business as usual or conducting inspections, reviews or other regulatory activities, it could significantly impact the ability of the FDA to timely review and process our regulatory submissions.

We are subject to stringent privacy laws, information security policies and contractual obligations governing the use, processing and transfer of personal information.

We receive, generate and store significant and increasing volumes of sensitive information, such as health information, insurance information and other potentially personally identifiable information. We face a number of risks relative to protecting this critical information, including loss of access risk, inappropriate use or disclosure, inappropriate modification and the risk of our being unable to adequately monitor, audit and modify our controls over our critical information. This risk extends to the third-party vendors we use to manage this sensitive data.

We are subject to a variety of local, state, national and international laws, directives and regulations that apply to the collection, use, retention, protection, disclosure, transfer and other processing of personal data in the different jurisdictions in which we operate, including comprehensive regulatory systems in the U.S. and Europe. Further, various states have implemented certain data privacy and security laws and regulations that impose restrictive requirements regulating the use and disclosure of health information and other personally identifiable information. California enacted the California Consumer Privacy Act (CCPA), which creates individual privacy rights for California consumers and increases the privacy and security obligations of entities handling certain personal data. The CCPA went into effect on January 1, 2020, and became enforceable by the California Attorney General on July 1, 2020. The CCPA has been amended from time to time, and, further a new privacy law, the California Privacy Rights Act (CPRA) was approved by California voters in the November 3, 2020 election. Effective starting January 1, 2023, the CPRA will significantly modify the CCPA, including by expanding consumers' rights with respect to certain sensitive personal information. The CPRA also creates a new state agency that will be vested with authority to implement and enforce the CCPA and the CPRA. It remains unclear what, if any, further modifications will be made to the CCPA or CPRA, or how such legislation will be interpreted. This may potentially result in further uncertainty and require us to incur additional costs and expenses in efforts to comply. Certain other state laws impose similar privacy obligations and all 50 states have laws including obligations to provide notification of security breaches of computer databases that contain personal information to affected individuals, state officers and others. For example, the CCPA has prompted a number of proposals for new federal and state-level privacy legislation. This legislation may add additional complexity, variation in requirements, restrictions and potential legal risk, require additional investment of resources in compliance programs, impact strategies and the availability of previously useful data and could result in increased compliance costs and/or changes in business practices and policies.

State laws and regulations are not necessarily preempted by federal laws and regulations, such as the Health Insurance Portability and Accountability Act of 1996 (HIPAA), particularly if a state affords greater protection to individuals than federal law. Where state laws are more protective, we have to comply with the stricter provisions. In addition to fines and penalties imposed upon violators, some of these state laws also afford private rights of action to individuals who believe their personal information has been misused. The interplay of federal and state laws may be subject to varying interpretations by courts and government agencies, creating complex compliance issues for us and data we receive, use and share, potentially exposing us to additional expense, adverse publicity and liability. Legal requirements relating to the collection, storage, handling, and transfer of personal information and personal data continue to evolve and may result in increased public scrutiny and escalating levels of enforcement, sanctions and increased costs of compliance.

The collection and use of personal data in the European Union (EU) and the European Economic Area (EEA), are governed by the General Data Protection Regulation (GDPR). The GDPR imposes stringent requirements for controllers and processors of personal data, including, for example, more robust disclosures to individuals and a strengthened individual data rights regime, shortened timelines for data breach notifications, limitations on retention of information, increased requirements pertaining to special categories of data, such as health data, and additional obligations when we contract with third-party processors in connection with the processing of the personal data. The GDPR also imposes strict rules on the transfer of personal data out of the EU and the EEA to the United States and other third countries. In July 2020, the Court of Justice of the European Union issued a decision that struck down the EU-U.S. Privacy Shield framework, which provided companies with a mechanism to comply with data protection requirements when transferring personal data from the EU to the United States and additionally called into question the validity of the European Commission's Standard Contractual Clauses, on which U.S. companies rely to transfer personal data from Europe to the United States and elsewhere. In September 2020, the Swiss Federal Data Protection and Information Commissioner issued an opinion that stated it no longer considers the Swiss-U.S. Privacy Shield adequate for the purposes of personal data transfers from Switzerland to the United States. These developments may result in European data protection regulators applying differing standards for, and requiring ad hoc verification of, transfers of personal data from Europe to the United States. To the extent that we engage in such transfers, including through third-party vendors, if we are unable to implement safeguards to ensure that our transfers are lawful or if any safeguards upon which we rely are invalidated, we will face increased exposure to litigation, reg



processing. If we are unable to engage in such transfers because there is no lawful mechanism to do so, the functionality or effectiveness of our products and services may decrease and our marketing efforts, plans and activities may be adversely impacted. In addition, the GDPR provides that EU and EEA member states may make their own further laws and regulations limiting the processing of personal data, including biometric or health data.

The GDPR applies extraterritorially, and we may be subject to the GDPR because of our data processing activities that involve the personal data of individuals located in the EEA, such as in connection with any EEA clinical trials. GDPR regulations may impose additional responsibility and liability in relation to the personal data that we process and we may be required to put in place additional mechanisms to ensure compliance with the new data protection rules. This may be onerous and may interrupt or delay our development activities.

Other jurisdictions outside the EEA are similarly introducing or enhancing privacy and data security laws, rules and regulations, which could increase our compliance costs and the risks associated with non-compliance. We cannot guarantee that we or our vendors may be in compliance with all applicable international regulations as they are enforced now or as they evolve. For example, our privacy and cybersecurity policies may be insufficient to protect any personal information we collect, or may not comply with applicable laws, in which case we may be subject to regulatory enforcement actions, lawsuits or reputational damage. If we or our vendors fail to comply with the GDPR and the applicable national data protection laws of the EU or EEA member states, or if regulators assert we have failed to comply with these laws, it may lead to regulatory enforcement actions, which can result in monetary penalties of up to €20,000,000 or up to 4% of the total worldwide annual turnover of the preceding financial year, whichever is higher, and other administrative penalties. Further, beginning on January 1, 2021, companies have to comply with both the GDPR and the United Kingdom GDPR (the UK GDPR), which together with the UK Data Protection Act 2018, retains the GDPR in United Kingdom national law. The UK GDPR mirrors the fines under the GDPR, i.e. fines up to the greater of £17.5 million or 4% of global turnover. The relationship between the United Kingdom and the EU in relation to certain aspects of data protection law remains unclear, and it is unclear how United Kingdom data protection laws and regulations will develop over time, and how data transfers to and from the United Kingdom will be regulated. These changes could lead to additional costs and increase our overall risk exposure. Currently there is a four- to six-month grace period agreed in the EU and United Kingdom Trade and Cooperation Agreement, ending June 30, 2021 at the latest, whilst the parties discuss an adequacy decision. The European Commission published a draft adequacy decision on February 19, 2021. If adopted, the decision will enable data transfers from EU member states to the United Kingdom for a four-year period, subject to subsequent extensions. We may incur liabilities, expenses, costs and other operational losses under the GDPR and privacy laws of the applicable EU and EEA Member States and the United Kingdom in connection with any measures we take to comply with them.

Compliance with U.S. and international data protection laws and regulations could cause us to incur substantial costs or require us to change our business practices and compliance procedures in a manner adverse to our business. Penalties for violations of these laws vary and may be significant. Moreover, complying with these various laws could require us to take on more onerous obligations in our contracts, restrict our ability to collect, use and disclose data, or in some cases, impact our ability to operate in certain jurisdictions. In addition, we rely on third-party vendors to collect, process and store data on our behalf and we cannot guarantee that such vendors are in compliance with all applicable data protection laws and regulations. Our or our vendors' failure to comply with U.S. and international data protection laws and regulations could result in government enforcement actions (which could include civil or criminal penalties), private litigation and adverse publicity. Claims that we have violated individuals' privacy rights, failed to comply with data protection laws, or breached our contractual obligations, even if we are not found liable, could be expensive and time consuming to defend and could result in adverse publicity.

Risks related to reliance on third parties

We are dependent on our collaboration with Sanofi for the development of RMC-4630 and may depend on Sanofi for the development and commercialization of any other future SHP2 inhibitor product candidates. Under certain circumstances, Sanofi may unilaterally terminate the collaboration for convenience, which would materially and adversely affect our business.

In June 2018, we entered into a collaborative research, development and commercialization agreement with Sanofi (the Sanofi Agreement), focused on researching, developing and commercializing SHP2 inhibitors as cancer therapies and potentially other indications. Sanofi primarily controls the research and development activities pursuant to the terms of the Sanofi Agreement, and our lack of control over these activities, including with respect to RMC-4630, could result in delays or other difficulties in the development and commercialization of product candidates, which may prevent completion of intended NDA filings in a timely fashion, if at all. Because of the allocation of responsibilities under the Sanofi Agreement, we are wholly dependent on Sanofi for the success of the RMC-4630 program. Any dispute with Sanofi may result in the delay or termination of the research, development or commercialization of RMC-4630 or other SHP2 inhibitor product candidates, and may result in costly litigation that diverts management attention and resources away from our day-to-day activities. For example, we plan to evaluate RMC-4630 in combination with other therapies (which may include product candidates from our pipeline), and Sanofi may disagree with us regarding which other therapies should be evaluated in combination with RMC-4630. As a result of this disagreement, our completion of a trial in combination with our preferred combination product candidate may be delayed or prevented.



In addition, Sanofi can terminate the Sanofi Agreement (including for convenience), and in the event Sanofi terminates the Sanofi Agreement, we would be prevented from receiving any research and development funding, milestone payments, profit share payments, royalty payments and other benefits under that agreement. Termination of the Sanofi Agreement could require us to seek additional funding in order to avoid delaying, reducing the scope of, or suspending, one or more of our research and development programs or clinical trials. In addition, any decision by Sanofi to terminate the Sanofi Agreement may negatively impact public perception of RMC-4630, or all of the SHP2 program covered by the Sanofi Agreement. We cannot provide any assurance with respect to the success of the Sanofi collaboration.

In addition to our collaboration with Sanofi, we may depend on collaborations with other third parties for the development and commercialization of our product candidates in the future. If our collaborations are not successful, we may not be able to capitalize on the market potential of these product candidates.

In the future, we may form or seek other strategic alliances, joint ventures, or collaborations, or enter into additional licensing arrangements with third parties that we believe will complement or augment our development and commercialization efforts with respect to product candidates we develop.

Collaborations involving our current and future product candidates, including our collaborations with Sanofi and Amgen may pose the following risks to us:

- collaborators have significant discretion in determining the efforts and resources that they will apply to these collaborations and may have incentives that are different than ours;
- collaborators may delay clinical trials, provide insufficient funding for a clinical trial program, stop a clinical trial, abandon a product candidate, repeat
 or conduct new clinical trials or require a new formulation of a product candidate for clinical testing;
- collaborators could independently develop, or develop with third parties, products that compete directly or indirectly with our products or product candidates;
- a collaborator with marketing, manufacturing and distribution rights to one or more products may not commit sufficient resources to or otherwise not perform satisfactorily in carrying out these activities;
- collaborators may not properly prosecute, maintain, enforce or defend our intellectual property rights or may use our proprietary information in a way
 that gives rise to actual or threatened litigation that could jeopardize or invalidate our intellectual property or proprietary information or expose us to
 potential litigation, or other intellectual property proceedings;
- collaborators may own or co-own intellectual property covering products that result from our collaboration with them, and in such cases, we may not have the exclusive right to develop, license or commercialize this intellectual property;
- disputes may arise with respect to ownership of any intellectual property developed pursuant to our collaborations;
- disputes may arise between a collaborator and us that cause the delay or termination of the research, development or commercialization of the product candidate, or that result in costly litigation or arbitration that diverts management attention and resources; and
- if a present or future collaborator of ours were to be involved in a business combination, the continued pursuit and emphasis on our product development or commercialization program under such collaboration could be delayed, diminished or terminated, including if the partner in such a business combination has products that compete with ours.

As a result, if we enter into additional collaboration agreements and strategic partnerships or license our intellectual property, products or businesses, we may not be able to realize the benefit of such transactions if we are unable to successfully integrate them with our existing operations, which could delay our timelines or otherwise adversely affect our business. We also cannot be certain that, following a strategic transaction or license, we will achieve the revenue or specific net income that justifies such transaction. Any delays in entering into new collaborations or strategic partnership agreements related to any product candidate we develop could delay the development and commercialization of our product candidates, which would harm our business prospects, financial condition, and results of operations.

We may seek to establish additional collaborations, and, if we are not able to establish them on commercially reasonable terms, we may have to alter our development and commercialization plans.

The advancement of our product candidates and development programs and the potential commercialization of our current and future product candidates will require substantial additional cash to fund expenses. For some of our programs, we may decide to collaborate with additional pharmaceutical and biotechnology companies with respect to development and potential commercialization. Any of these relationships may require us to incur non-recurring and other charges, increase our near and long term expenditures, issue securities that dilute our existing stockholders, or disrupt our management and business.

We face significant competition in seeking appropriate strategic partners and the negotiation process is time-consuming and complex. Whether we reach a definitive agreement for other collaborations will depend, among other things, upon our assessment of the collaborator's resources and expertise, the terms and conditions of the proposed collaboration and the proposed collaborator's



evaluation of a number of factors. Those factors may include the design or results of clinical trials, the progress of our clinical trials, the likelihood of approval by the FDA or similar regulatory authorities outside the United States, the potential market for the subject product candidate, the costs and complexities of manufacturing and delivering such product candidate to patients, the potential of competing products, the existence of uncertainty with respect to our ownership of technology, which can exist if there is a challenge to such ownership without regard to the merits of the challenge and industry and market conditions generally. The collaborator may also consider alternative product candidates or technologies for similar indications that may be available to collaborate on and whether such a collaboration could be more attractive than the one with us for our product candidate. Further, we may not be successful in our efforts to establish a strategic partnership or other alternative arrangements for future product candidates because they may be deemed to be at too early of a stage of development for collaborative effort and third parties may not view them as having the requisite potential to demonstrate safety and efficacy.

We may also be restricted under existing collaboration agreements from entering into future agreements on certain terms with potential collaborators. For example, under the Sanofi Agreement, we have granted worldwide exclusive rights under our intellectual property to Sanofi for SHP2 inhibitors, and during the term of the agreement we will be restricted from granting similar rights to other parties. This exclusivity could limit our ability to enter into collaborators.

In addition, there have been a significant number of recent business combinations among large pharmaceutical companies that have resulted in a reduced number of potential future collaborators.

We may not be able to negotiate collaborations on a timely basis, on acceptable terms, or at all. If we are unable to do so, we may have to curtail the development of the product candidate for which we are seeking to collaborate, reduce or delay its development program or one or more of our other development programs, delay its potential commercialization or reduce the scope of any sales or marketing activities, or increase our expenditures and undertake development or commercialization activities at our own expense. If we elect to increase our expenditures to fund development or commercialization activities on our own, we may need to obtain additional capital, which may not be available to us on acceptable terms or at all. If we do not have sufficient funds, we may not be able to further develop our product candidates or bring them to market and generate product revenue.

If conflicts arise between us and our collaborators or strategic partners, these parties may act in a manner adverse to us and could limit our ability to implement our strategies.

If conflicts arise between our corporate or academic collaborators or strategic partners and us, the other party may act in a manner adverse to us and could limit our ability to implement our strategies. Sanofi, Amgen or future collaborators or strategic partners may develop, either alone or with others, products in related fields that are competitive with the products or potential products that are the subject of these collaborations. Competing products, either developed by the collaborators or strategic partners or to which the collaborators or strategic partners have rights, may result in the withdrawal of partner support for our product candidates. Our current or future collaborators or strategic partners may preclude us from entering into collaborations with their competitors, fail to obtain timely regulatory approvals, terminate their agreements with us prematurely, or fail to devote sufficient resources to the development and commercialization of products. Any of these developments could harm our product development efforts.

We rely on third parties to conduct our ongoing and planned clinical trials for RMC-4630, RMC-5552 and any other product candidates we develop. If these third parties do not successfully carry out their contractual duties, comply with regulatory requirements or meet expected deadlines, we may not be able to obtain marketing approval for or commercialize RMC-4630, RMC-5552 or any other product candidates we develop.

We do not have the ability to independently conduct clinical trials. We rely on medical institutions, clinical investigators, CROs, contract laboratories, and other third parties, including collaboration partners, to conduct or otherwise support ongoing or planned clinical trials for RMC-4630, RMC-5552 and other product candidates. We rely heavily on these parties for execution of clinical trials and control only certain aspects of their activities. Nevertheless, we are responsible for ensuring that each of our clinical trials is conducted in accordance with the applicable protocol, legal and regulatory requirements and scientific standards, and our reliance on CROs will not relieve us of our regulatory responsibilities. For any violations of laws and regulations during the conduct of our clinical trials, we could be subject to untitled and warning letters or enforcement action that may include civil penalties up to and including criminal prosecution.

We and third parties are required to comply with regulations and requirements, including GCP, for conducting, monitoring, recording and reporting the results of clinical trials to ensure that the data and results are scientifically credible and accurate, and that the trial patients are adequately informed of the potential risks of participating in clinical trials and their rights are protected. These regulations are enforced by the FDA, the Competent Authorities of the Member States of the EEA and comparable foreign regulatory authorities for any drugs in clinical development. The FDA enforces GCP requirements through periodic inspections of clinical trial sponsors, principal investigators and trial sites. If we or third parties fail to comply with applicable GCP, the clinical data generated in our clinical trials may be deemed unreliable and the FDA or comparable foreign regulatory authorities may require us to perform additional clinical trials before approving our marketing applications. Upon inspection, the FDA may determine that any of our current or future clinical trials do not comply with GCP. In addition, our clinical trials must be conducted with product candidates produced under cGMP regulations. Our failure or the failure of third parties to comply with these regulations may require us to repeat clinical trials, which would delay the marketing approval process and could also subject us to enforcement action. We also are required to



register certain ongoing clinical trials and provide certain information, including information relating to the trial's protocol, on a government-sponsored database, ClinicalTrials.gov, within specific timeframes. Failure to do so can result in fines, adverse publicity and civil and criminal sanctions.

Although we intend to design the clinical trials for our product candidates, or be involved in the design when other parties sponsor the trials, third parties conduct all or a portion of the clinical trials involving our product candidates. For example, Amgen is conducting the Phase 1b trial evaluating the combination of RMC-4630 and the KRASG^{12C}(OFF) inhibitor sotorasib in Amgen's CodeBreaK 101c study and Sanofi is conducting the Phase 1/2 trial evaluating the combination of RMC-4630 and Merck's PD-1 inhibitor pembrolizumab. In addition, in March 2020, the Pancreatic Cancer Collective (a strategic partnership between Lustgarten Foundation and Stand Up To Cancer) announced that it had awarded funding to the Netherlands Cancer Institute for its study using RMC-4630 in combination with an investigational ERK inhibitor (LY3214996) in patients with pancreatic cancer. We plan to provide RMC-4630 to support this investigator sponsored study. As a result, many important aspects of our clinical development, including their conduct and timing, will be outside of our direct control. Our reliance on third parties to conduct future clinical trials will also result in less direct control over the management of data developed through clinical trials than would be the case if we were relying entirely upon our own staff. Communicating with outside parties can also be challenging, potentially leading to mistakes as well as difficulties in coordinating activities. Outside parties may:

- have staffing difficulties;
- fail to comply with contractual obligations;
- experience regulatory compliance issues;
- have incentives that are different than ours;
- undergo changes in priorities or become financially distressed; or
- form relationships with other entities, some of which may be our competitors.

These factors may materially adversely affect the willingness or ability of third parties to conduct our clinical trials and may subject us to unexpected cost increases that are beyond our control. If the CROs do not perform clinical trials in a satisfactory manner, breach their obligations to us or fail to comply with regulatory requirements, the development, marketing approval and commercialization of our product candidates may be delayed, we may not be able to obtain marketing approval and commercialize our product candidates, or our development program may be materially and irreversibly harmed. If we are unable to rely on clinical data collected by our CROs, we could be required to repeat, extend the duration of, or increase the size of any clinical trials we conduct and this could significantly delay commercialization and require significantly greater expenditures.

If any of our relationships with these third-party CROs terminate, we may not be able to enter into arrangements with alternative CROs on commercially reasonable terms, or at all. If CROs do not successfully carry out their contractual duties or obligations or meet expected deadlines, if they need to be replaced or if the quality or accuracy of the clinical data they obtain are compromised due to the failure to adhere to our clinical protocols, regulatory requirements or for other reasons, any clinical trials such CROs are associated with may be extended, delayed or terminated, and we may not be able to obtain marketing approval for or successfully commercialize our product candidates.

Our clinical trial sites for our RMC-4630 and RMC-5552 clinical studies may be affected by the COVID-19 outbreak due to prioritization of hospital resources toward the COVID-19 outbreak, travel or quarantine restrictions imposed by governments, and the inability to access sites for initiation and patient monitoring and enrollment. As a result, patient screening, new patient enrollment, monitoring and data collection may be affected or delayed. We are aware that several clinical sites involved in our RMC-4630 clinical studies temporarily stopped or delayed enrolling new patients, with exemptions if appropriate, and it is possible that these or other clinical sites may be similarly affected in the future. These developments may delay our clinical trial timelines. Some of our third-party manufacturers which we use for the supply of materials for product candidates or other materials necessary to manufacture product to conduct clinical trials and contract research organizations that we may utilize may be impacted by COVID-19, and should they experience disruptions, such as temporary closures or suspension of services, we would likely experience delays in advancing these trials.

We rely on third parties to manufacture preclinical and clinical drug supplies, and we intend to rely on third parties to produce commercial supplies of any approved product, which increases the risk that we will not have sufficient quantities of these product candidates or products or such quantities at an acceptable cost, which could delay, prevent or impair our development or commercialization efforts.

We do not own or operate manufacturing facilities for the production of preclinical, clinical or commercial supplies of the product candidates that we are developing or evaluating in our development programs. We have limited personnel with experience in drug manufacturing and lack the resources and the capabilities to manufacture any of our product candidates on a preclinical, clinical or commercial scale. We rely on third parties for supply of our preclinical and clinical drug supplies (including key starting and intermediate materials), and our strategy is to outsource all manufacturing of our product candidates and products to third parties, including Sanofi.

In order to conduct clinical trials of product candidates, we will need to have them manufactured in potentially large quantities. Our third-party manufacturers may be unable to successfully increase the manufacturing capacity for any of our clinical drug supplies (including key starting and intermediate materials) in a timely or cost-effective manner, or at all. In addition, quality issues may arise during scale-up activities and at any other time. For example, ongoing data on the stability of our product candidates may shorten the expiry of our product candidates and lead to clinical trial material supply shortages, and potentially clinical trial delays. If these third-party manufacturers are unable to successfully scale up the manufacture of our product candidates in sufficient quality and quantity, the development, testing and clinical trials of that product candidate may be delayed or infeasible, and regulatory approval or commercial launch of that product candidate may be delayed or not obtained.

Our use of new third-party manufacturers increases the risk of delays in production or insufficient supplies of our product candidates (and the key starting and intermediate materials for such product candidates) as we transfer our manufacturing technology to these manufacturers and as they gain experience manufacturing our product candidates (and the key starting and intermediate materials for such product candidates).

Even after a third-party manufacturer has gained significant experience in manufacturing our product candidates (or the key starting and intermediate materials for such product candidates) or even if we believe we have succeeded in optimizing the manufacturing process, there can be no assurance that such manufacturer will produce sufficient quantities of our product candidates (or the key starting and intermediate materials for such product candidates) in a timely manner or continuously over time, or at all.

We may be delayed if we need to change the manufacturing process used by a third party. Further, if we change an approved manufacturing process, then we may be delayed if the FDA or a comparable foreign authority needs to review the new manufacturing process before it may be used.

We do not currently have any agreements with third-party manufacturers for long-term commercial supply. In the future, we may be unable to enter into agreements with third-party manufacturers for commercial supplies of any product candidate that we develop, or may be unable to do so on acceptable terms. Even if we are able to establish and maintain arrangements with third-party manufacturers, reliance on third-party manufacturers entails risks, including:

- reliance on the third party for regulatory compliance and quality assurance;
- the possible breach of the manufacturing agreement by the third party;
- the possible misappropriation of our proprietary information, including our trade secrets and know-how; and
- the possible termination or non-renewal of the agreement by the third party at a time that is costly or inconvenient for us.

Third-party manufacturers may not be able to comply with cGMP requirements or similar regulatory requirements outside the United States. Our failure, or the failure of our third-party manufacturers, to comply with applicable requirements could result in sanctions being imposed on us, including fines, injunctions, civil penalties, delays, suspension or withdrawal of approvals, license revocation, seizures or recalls of product candidates or products, operating restrictions and/or criminal prosecutions, any of which could significantly and adversely affect supplies of our product candidates.

Our future product candidates and any products that we may develop may compete with other product candidates and products for access to manufacturing facilities. There are a limited number of manufacturers that operate under cGMP requirements particularly for the development of monoclonal antibodies, and that might be capable of manufacturing for us.

If the third parties that we engage to supply any materials or manufacture product for our preclinical tests and clinical trials should cease to continue to do so for any reason, we likely would experience delays in advancing these tests and trials while we identify and qualify replacement suppliers or manufacturers and we may be unable to obtain replacement supplies on terms that are favorable to us. In addition, if we are not able to obtain adequate supplies of our product candidates or the substances used to manufacture them, it will be more difficult for us to develop our product candidates and compete effectively.

Some of our third-party manufacturers which we use for the supply of materials for product candidates or other materials necessary to manufacture product to conduct clinical trials are located in may be affected by COVID-19, and should they experience disruptions, such as temporary closures or suspension of services, we would likely experience delays in advancing these trials.

Our current and anticipated future dependence upon others for the manufacture of our product candidates (or the key starting and intermediate materials for such product candidates) may adversely affect our future profit margins and our ability to develop product candidates and commercialize any products that receive marketing approval on a timely and competitive basis.

Our future relationships with customers and third-party payors in the United States and elsewhere may be subject, directly or indirectly, to applicable anti-kickback, fraud and abuse, false claims, transparency, health information privacy and security and other healthcare laws and regulations, which could expose us to criminal sanctions, civil penalties, contractual damages, reputational harm, administrative burdens and diminished profits and future earnings.

Healthcare providers, physicians and third-party payors in the United States and elsewhere will play a primary role in the recommendation and prescription of any product candidates for which we obtain marketing approval. Our future arrangements with

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third-party payors and customers may expose us to broadly applicable fraud and abuse and other healthcare laws and regulations, including, without limitation, the federal Anti- Kickback Statute and the federal False Claims Act (FCA), which may constrain the business or financial arrangements and relationships through which we sell, market and distribute any products for which we obtain marketing approval. In addition, we may be subject to transparency laws and patient privacy regulation by the U.S. federal and state governments and by governments in foreign jurisdictions in which we conduct our business. The applicable federal, state and foreign healthcare laws and regulations that may affect our ability to operate include:

- the federal Anti-Kickback Statute, which prohibits, among other things, knowingly and willfully soliciting, receiving, offering or paying any
 remuneration (including any kickback, bribe, or rebate), directly or indirectly, overtly or covertly, in cash or in kind, to induce, or in return for, either
 the referral of an individual, or the purchase, lease, order or recommendation of any good, facility, item or service for which payment may be made, in
 whole or in part, under the Medicare and Medicaid programs or other federal healthcare programs. A person or entity can be found guilty of violating
 the statute without actual knowledge of the statute or specific intent to violate it. The Anti-Kickback Statute has been interpreted to apply to
 arrangements between pharmaceutical manufacturers on the one hand and prescribers, purchasers, and formulary managers on the other. There are a
 number of statutory exceptions and regulatory safe harbors protecting some common activities from prosecution;
- the federal civil and criminal false claims laws, including the FCA, and civil monetary penalty laws, which prohibit any person or entity from, among
 other things, knowingly presenting, or causing to be presented, a false, fictitious or fraudulent claim for payment to, or approval by, the federal
 government or knowingly making, using or causing to be made or used a false record or statement material to a false or fraudulent claim to the federal
 government. In addition, the government may assert that a claim including items or services resulting from a violation of the federal Anti-Kickback
 Statute constitutes a false or fraudulent claim for purposes of the FCA;
- the federal Health Insurance Portability and Accountability Act of 1996 (HIPAA), which created federal criminal statutes that prohibit knowingly and willfully executing, or attempting to execute, a scheme to defraud any healthcare benefit program or obtain, by means of false or fraudulent pretenses, representations, or promises, any of the money or property owned by, or under the custody or control of, any healthcare benefit program, regardless of the payor (e.g., public or private) and knowingly and willfully falsifying, concealing or covering up by any trick or device a material fact or making any materially false statements in connection with the delivery of, or payment for, healthcare benefits, items or services relating to healthcare matters. Similar to the federal Anti-Kickback Statute, a person or entity can be found guilty of violating HIPAA without actual knowledge of the statutes or specific intent to violate them;
- HIPAA, as amended by the Health Information Technology for Economic and Clinical Health Act of 2009 (HITECH) and their respective
 implementing regulations, which impose, among other things, specified requirements relating to the privacy, security and transmission of individually
 identifiable health information held by covered entities and their business associates. HITECH also created new tiers of civil monetary penalties,
 amended HIPAA to make civil and criminal penalties directly applicable to business associates, and gave state attorneys general new authority to file
 civil actions for damages or injunctions in federal courts to enforce the federal HIPAA laws and seek attorneys' fees and costs associated with
 pursuing federal civil actions;
- the Physician Payments Sunshine Act, created under the ACA, and its implementing regulations, which requires manufacturers of drugs, devices, biologics and medical supplies for which payment is available under Medicare, Medicaid or the Children's Health Insurance Program (with certain exceptions) to report annually to CMS information related to payments or other transfers of value made to physicians (defined to include doctors, dentists, optometrists, podiatrists and chiropractors) and teaching hospitals, as well as ownership and investment interests held by physicians and their immediate family members. Beginning in 2022, applicable manufacturers also will be required to report such information regarding payments and transfers of value provided during the previous year to physician assistants, nurse practitioners, clinical nurse specialists, certified nurse anesthetists, anesthesiologist assistants and certified nurse midwives;
- federal consumer protection and unfair competition laws, which broadly regulate marketplace activities and activities that potentially harm consumers; and
- analogous or related foreign, state or local laws and regulations, including anti-kickback and false claims laws, which may apply to sales or marketing arrangements and claims involving healthcare items or services reimbursed by non-governmental third-party payors, including private insurers; laws that require pharmaceutical companies to comply with the pharmaceutical industry's voluntary compliance guidelines and the relevant compliance guidance promulgated by the federal government or otherwise restrict payments that may be made to healthcare providers; laws that require drug manufacturers to report information related to payments and other transfers of value to physicians and other healthcare providers or marketing expenditures; and healthcare and data protection laws in the European Union and other jurisdictions, including reporting requirements detailing interactions with and payments to healthcare providers and laws governing the privacy and security of certain protected information, such as the California Consumer Privacy Act (CCPA), which creates individual privacy rights for California consumers and increases the privacy and security obligations of entities handling certain personal data, and General Data Protection Regulation (GDPR), which imposes obligations and restrictions on the collection and use of personal data relating to individuals located in the European Economic Area (the EEA) and the United Kingdom (including health data).

Because of the breadth of the laws described above and the narrowness of the statutory exceptions and regulatory safe harbors available under them, it is possible that some of our business activities could be subject to challenge under one or more of these laws. The scope and enforcement of each of these laws is uncertain and subject to rapid change in the current environment of healthcare reform, especially in light of the lack of applicable precedent and regulations. Federal and state enforcement bodies have recently increased their scrutiny of interactions between healthcare companies and healthcare providers, which has led to a number of investigations, prosecutions, convictions and settlements in the healthcare industry. Ensuring that our business arrangements with third parties comply with applicable healthcare laws, as well as responding to investigations by government authorities, can be time- and resource-consuming and can divert management's attention from the business.

If our operations are found to be in violation of any of the laws described above or any other government regulations that apply to us, we may be subject to penalties, including civil, criminal and administrative penalties, damages, fines, disgorgement, individual imprisonment, possible exclusion from participation in federal and state funded healthcare programs, contractual damages and the curtailment or restricting of our operations, as well as additional reporting obligations and oversight if we become subject to a corporate integrity agreement or other agreement to resolve allegations of non-compliance with these laws, any of which could harm our ability to operate our business and our financial results. Further, if the physicians or other providers or entities with whom we expect to do business are found not to be in compliance with applicable laws, they may be subject to criminal, civil or administrative sanctions, including exclusions from government funded healthcare programs. In addition, the approval and commercialization of any product candidate we develop outside the United States will also likely subject us to foreign equivalents of the healthcare laws mentioned above, among other foreign laws.

Risks related to intellectual property

If we and our collaborators are unable to obtain and maintain sufficient patent and other intellectual property protection for our product candidates and technology, our competitors could develop and commercialize products and technology similar or identical to ours, and we may not be able to compete effectively in our market or successfully commercialize any product candidates we may develop.

Our success depends in significant part on our ability and the ability of our collaborators to obtain, maintain, enforce and defend patents and other intellectual property rights with respect to our product candidates and technology and to operate our business without infringing, misappropriating, or otherwise violating the intellectual property rights of others. If we and our collaborators are unable to obtain and maintain sufficient intellectual property protection for our product candidates or the product candidates that we may identify, or if the scope of the intellectual property protection obtained is not sufficiently broad, our competitors and other third parties could develop and commercialize product candidates similar or identical to ours, and our ability (and the ability of our collaborators) to successfully commercialize the product candidates that we (and our collaborators) may pursue may be impaired. Our patent coverage with respect to our clinical and preclinical programs is limited, and we can provide no assurance that any of our current or future patent applications will result in issued patents or that any issued patents will provide us with any competitive advantage. Failure to obtain such issued patents could have a material adverse effect on our and Sanofi's ability to develop and commercialize SHP2 inhibitor products, including RMC-4630, and on our ability to receive milestone, royalty or other payments from Sanofi pursuant to the Sanofi Agreement or our ability to develop or commercialize any of our our ability to develop.

We seek to protect our proprietary positions by, among other things, filing patent applications in the United States and abroad related to our current product candidates and the product candidates that we may identify. Obtaining, maintaining, defending and enforcing pharmaceutical patents is costly, time consuming and complex, and we may not be able to file and prosecute all necessary or desirable patent applications, or maintain, enforce and license any patents that may issue from such patent applications, at a reasonable cost or in a timely manner. It is also possible that we will fail to identify patentable aspects of our research and development output before it is too late to obtain patent protection. We may not have the right to control the preparation, filing, prosecution and maintenance of patent applications, or to maintain the rights to patents licensed to or from third parties.

Although we enter into confidentiality agreements with parties who have access to confidential or patentable aspects of our research and development output, such as our employees, collaborators, CROs, contract manufacturers, consultants, advisors and other third parties, any of these parties may breach the agreements and disclose such output before a patent application is filed, thereby jeopardizing our ability to seek patent protection. Further, we may not be aware of all third-party intellectual property rights potentially relating to our product candidates. Publications of discoveries in the scientific literature often lag behind the actual discoveries, and patent applications in the United States and other jurisdictions are typically not published until 18 months after filing or, in some cases, not at all. Therefore, we cannot know with certainty whether we were the first to make the inventions claimed in our patents or pending patent applications, or that we were the first to file for patent protection of such inventions.

The patent position of pharmaceutical companies generally is highly uncertain, involves complex legal, technological and factual questions and has, in recent years, been the subject of much debate and litigation throughout the world. In addition, the laws of foreign countries may not protect our rights to the same extent as the laws of the United States, or vice versa. As a result, the issuance, scope, validity, enforceability, and commercial value of our patent rights are highly uncertain. The subject matter claimed in a patent application can be significantly reduced before the patent is issued, and its scope can be reinterpreted after issuance. Therefore, our pending and future patent applications may not result in patents being issued in relevant jurisdictions that protect our product

candidates, in whole or in part, or which effectively prevent others from commercializing competitive product candidates, and even if our patent applications issue as patents in relevant jurisdictions, they may not issue in a form that will provide us with any meaningful protection for our product candidates or technology, prevent competitors from competing with us or otherwise provide us with any competitive advantage. Additionally, our competitors may be able to circumvent our patents by developing similar or alternative product candidates or technologies in a non-infringing manner.

The issuance of a patent is not conclusive as to its inventorship, scope, validity or enforceability, and our patents may be challenged in the courts or patent offices in the United States and abroad. We may be subject to a third-party preissuance submission of prior art to the United States Patent and Trademark Office (the USPTO) or become involved in opposition, derivation, revocation, reexamination, *inter partes* review, post-grant review or interference proceedings challenging our patent rights or the patent rights of others, or other proceedings in the USPTO or applicable foreign offices that challenge priority of invention or other features of patentability. An adverse determination in any such submission, proceeding or litigation could result in loss of exclusivity or freedom to operate, patent claims being narrowed, invalidated or held unenforceable, in whole or in part, or limits of the scope or duration of the patent protection of our product candidates, all of which could limit our ability to stop others from using or commercializing similar or identical product candidates or technology to compete directly with us, without payment to us, or result in our inability to manufacture or commercialize products without infringing third-party patent rights. In addition, if the breadth or strength of protection provided by our patents and patent applications is threatened, regardless of the outcome, it could dissuade companies from collaborating with us to license, develop or commercialize current or future product candidates, or could have a material adverse effect on our ability to raise funds necessary to continue our research programs or clinical trials. Such proceedings also may result in substantial cost and require significant time from our scientists and management, even if the eventual outcome is favorable to us.

In addition, given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such candidates might expire before or shortly after such candidates are commercialized. As a result, our patent portfolio may not provide us with sufficient rights to exclude others from commercializing products or technology similar or identical to ours for a meaningful amount of time, or at all. Moreover, some of our owned or licensed patents and patent applications are, and may in the future be, co-owned with third parties. If we are unable to obtain exclusive licenses to any such co-owners' interest in such patents or patent applications, such co-owners may be able to license their rights to other third parties, including our competitors, and our competitors could market competing products and technology. In addition, we may need the cooperation of any such co-owners in order to enforce such patents against third parties, and such cooperation may not be provided to us. Any of the foregoing could harm our competitive position, business, financial condition, results of operations and prospects.

We have entered into licensing agreements with third parties. If we or a third party fail to comply with the obligations in the agreements under which we license intellectual property rights to or from third parties, or these agreements are terminated, or we otherwise experience disruptions to business relationships with our licensors or licensees, competitive position, business, financial condition, results of operations and prospects could be harmed.

In addition to patent and other intellectual property rights we own or co-own, we have licensed, and may in the future license, patent and other intellectual property rights to and from other parties. Licenses may not provide us with exclusive rights to use the applicable intellectual property and technology in all relevant fields of use and in all territories in which we may wish to develop or commercialize our products and technology in the future. As a result, we may not be able to prevent competitors from developing and commercializing competitive products or technologies.

In addition, in some circumstances, we may not have the right to control the preparation, filing and prosecution of patent applications or to maintain, defend and enforce the patents that we license to or from third parties, and we may have to rely on our partners to fulfill these responsibilities. For example, in June 2018, we entered into the Sanofi Agreement, wherein we exclusively licensed the worldwide rights in our SHP2 inhibitor program, including RMC-4630, to Sanofi. Although we have review and comment rights regarding patent prosecution decisions, Sanofi retains ultimate decision-making control, as well as the sole and exclusive right to enforce infringement of or defend claims against patents that relate to SHP2 inhibitor products licensed to it pursuant to the Sanofi Agreement. Consequently, any such licensed patents and applications may not be prepared, filed, prosecuted, maintained, enforced, and defended in a manner consistent with the best interests of our business. If our current or future licensors, licensees or collaborators fail to prepare, file, prosecute, maintain, enforce, and defend licensed patents and other intellectual property rights, such rights may be reduced or eliminated, and our right to develop and commercialize any of our product candidates or technology that are the subject of such licensed rights could be adversely affected. In addition, our licensors may own or control intellectual property that has not been licensed to us and, as a result, we may be subject to claims, regardless of their merit, that we are infringing or otherwise violating the licensor's rights.

If we fail to comply with our obligations in the agreements under which we license intellectual property rights from third parties, the licensor may have the right to terminate the license. If these agreements are terminated, the underlying patents fail to provide the intended exclusivity or we otherwise experience disruptions to our business relationships with our licensors, we could lose intellectual property rights that are important to our business or be prevented from developing and commercializing our product candidates, and competitors could have the freedom to seek regulatory approval of, and to market, products identical to ours. Termination of these agreements or reduction or elimination of our rights under these agreements may also result in our having to negotiate new or reinstated agreements with less favorable terms, cause us to lose our rights under these agreements, including our rights to important



intellectual property or technology, or impede, delay or prohibit the further development or commercialization of one or more product candidates that rely on such agreements. It is possible that we may be unable to obtain any additional licenses at a reasonable cost or on reasonable terms, if at all. In that event, we may be required to expend significant time and resources to redesign our product candidates or the methods for manufacturing them or to develop or license replacement technology, all of which may not be feasible on a technical or commercial basis.

In addition, the research resulting in certain of our owned and in-licensed patent rights and technology was funded in part by the U.S. federal or state governments. As a result, the government may have certain rights, including march-in rights, to such patent rights and technology. When new technologies are developed with government funding, the government generally obtains certain rights in any resulting patents, including a non-exclusive license authorizing the government to use the invention for noncommercial purposes. These rights may permit the government to disclose our confidential information to third parties or allow third parties to use our licensed technology. The government can exercise its march-in rights if it determines that action is necessary because we fail to achieve practical application of the government-funded technology, because action is necessary to alleviate health or safety needs, to meet requirements of federal regulations, or to give preference to U.S. industry. In addition, our rights in such inventions may be subject to certain requirements to manufacture products embodying such inventions in the United States. Any of the foregoing could harm our competitive position, business, financial condition, results of operations and prospects.

Licensing of intellectual property is of critical importance to our business and involves complex legal, business and scientific issues and certain provisions in intellectual property license agreements may be susceptible to multiple interpretations. Disputes may arise between us and our licensing partners regarding intellectual property subject to a license agreement, including:

- the scope of rights granted under the license agreement and other interpretation-related issues;
- whether and the extent to which technology and processes of one party infringe on intellectual property of the other party that are not subject to the licensing agreement;
- rights to sublicense patent and other rights to third parties;
- any diligence obligations with respect to the use of the licensed technology in relation to development and commercialization of our product candidates, and what activities satisfy those diligence obligations;
- the ownership of inventions and know-how resulting from the joint creation or use of intellectual property;
- rights to transfer or assign the license; and
- the effects of termination.

The resolution of any contract interpretation disagreement that may arise could narrow what we believe to be the scope of our rights to the relevant intellectual property or technology, or increase what we believe to be our financial or other obligations under the relevant agreement, either of which could harm our business, financial condition, results of operations and prospects. Moreover, if disputes over intellectual property that we have licensed prevent or impair our ability to maintain our current licensing arrangements on acceptable terms, we may be unable to successfully develop and commercialize the affected product candidates.

In addition, if our licensors or licensees fail to abide by the terms of the license, if the licensors or licensees fail to prevent infringement by third parties or if the licensed patents or other rights are found to be invalid or unenforceable, our business, competitive position, financial condition, results of operations and prospects could be materially harmed.

If we are unable to obtain licenses from third parties on commercially reasonable terms or at all, our business could be harmed.

It may be necessary for us to use the patented or proprietary technology of third parties to commercialize our products, in which case we would be required to obtain a license from these third parties. The licensing of third-party intellectual property rights is a competitive area, and more established companies may pursue strategies to license or acquire third-party intellectual property rights that we may consider attractive or necessary. More established companies may have a competitive advantage over us due to their size, capital resources and greater clinical development and commercialization capabilities. In addition, companies that perceive us to be a competitor may be unwilling to assign or license rights to us. We also may be unable to license or acquire third party intellectual property rights on terms that would allow us to make an appropriate return on our investment or at all. If we are unable to license needed technology, or if we are forced to license this technology on unfavorable terms, our business could be materially harmed. If we are unable to obtain a necessary license, we may be unable to develop or commercialize the affected product candidates, and the third parties owning such intellectual property rights could seek either an injunction prohibiting our sales, or, with respect to our sales, an obligation on our part to pay royalties or other forms of compensation. Even if we are able to obtain a license, it may be non-exclusive, thereby giving our competitors access to the same technologies licensed to us.

We may not identify relevant third-party patents or may incorrectly interpret the relevance, scope or expiration of a third-party patent, which might subject us to infringement claims or adversely affect our ability to develop and market our product candidates.

We cannot guarantee that any of our or our licensors' patent searches or analyses, including the identification of relevant patents, the scope of patent claims or the expiration of relevant patents, are complete or thorough, nor can we be certain that we have identified



each and every third-party patent and pending patent application in the United States and abroad that is relevant to or necessary for the commercialization of our product candidates in any jurisdiction. For example, U.S. patent applications filed before November 29, 2000 and certain U.S. patent applications filed after that date that will not be filed outside the United States remain confidential until patents issue. Patent applications in the United States and elsewhere are published approximately 18 months after the earliest filing for which priority is claimed, with the earliest filing date being commonly referred to as the priority date. Therefore, patent applications covering our product candidates could have been filed by third parties without our knowledge. Additionally, pending patent applications that have been published can, subject to certain limitations, be later amended in a manner that could cover our product candidates or the use of our product candidates. The scope of a patent claim is determined by an interpretation of the law, the written disclosure in a patent and the patent's prosecution history. Our interpretation of the relevance or the scope of a patent or a pending application may be incorrect, which may negatively impact our ability to market our product candidates. We may incorrectly determine that our product candidates are not covered by a thirdparty patent or may incorrectly predict whether a third party's pending application will issue with claims of relevant scope. Our determination of the expiration date of any patent in the United States or abroad that we consider relevant may be incorrect, which may negatively impact our ability to develop and market our product candidates. Our failure to identify and correctly interpret relevant patents may negatively impact our ability to develop and market our product candidates.

If we fail to identify and correctly interpret relevant patents, we may be subject to infringement claims. We cannot guarantee that we will be able to successfully settle or otherwise resolve any infringement claims. If we fail in any of these disputes, in addition to being forced to pay damages, which may be significant, we may be temporarily or permanently prohibited from commercializing any of our product candidates that are held to be infringing. We might, if possible, also be forced to redesign product candidates so that we no longer infringe the third-party intellectual property rights. Any of these events, even if we were ultimately to prevail, could require us to divert substantial financial and management resources that we would otherwise be able to devote to our business.

Patent terms may be inadequate to protect our competitive position on our product candidates for an adequate amount of time.

Patents have a limited lifespan. In the United States, if all maintenance fees are timely paid, the natural expiration of a patent is generally 20 years from its earliest U.S. non-provisional filing date. Various extensions may be available, but the life of a patent, and the protection it affords, is limited. Even if patents covering our product candidates are obtained, once the patent life has expired for a product candidate, we may be open to competition from competitive medications, including generic medications. Given the amount of time required for the development, testing and regulatory review of new product candidates, patents protecting such product candidates might expire before or shortly after such product candidates are commercialized. As a result, our owned and licensed patent portfolio may not provide us with sufficient rights to exclude others from commercializing product candidates similar or identical to ours for a meaningful amount of time, or at all.

Depending upon the timing, duration and conditions of any FDA marketing approval of our product candidates, one or more of our owned or licensed U.S. patents may be eligible for limited patent term extension under the Drug Price Competition and Patent Term Restoration Act of 1984, referred to as the Hatch-Waxman Amendments, and similar legislation in the European Union and certain other countries. The Hatch-Waxman Amendments permit a patent term extension of up to five years for a patent covering an approved product as compensation for effective patent term lost during product development and the FDA regulatory review process. However, we may not receive an extension if we fail to exercise due diligence during the testing phase or regulatory review process, fail to apply within applicable deadlines, fail to apply prior to expiration of relevant patents or otherwise fail to satisfy applicable requirements. Moreover, the length of the extension could be less than we request. Only one patent per approved product can be extended, the extension cannot extend the total patent term beyond 14 years from approval and only those claims covering the approved drug, a method for using it or a method for manufacturing it may be extended. If we are unable to obtain patent term extension or the term of any such extension is less than we request, the period during which we can enforce our patent rights for the applicable products could be reduced. Further, if this occurs, our competitors may take advantage of our investment in development and trials by referencing our clinical and preclinical data and launch their product earlier than might otherwise be the case.

Also, there are detailed rules and requirements regarding the patents that may be submitted to the FDA for listing in the Approved Drug Products with Therapeutic Equivalence Evaluations (the Orange Book). We may be unable to obtain patents covering our product candidates that contain one or more claims that satisfy the requirements for listing in the Orange Book. Even if we submit a patent for listing in the Orange Book, the FDA may decline to list the patent, or a manufacturer of generic drugs may challenge the listing. If one of our product candidates is approved and a patent covering that product candidate is not listed in the Orange Book, a manufacturer of generic drugs would not have to provide advance notice to us of any abbreviated new drug application filed with the FDA to obtain permission to sell a generic version of such product candidate.

We may not be able to protect our intellectual property rights throughout the world.

Filing, prosecuting, maintaining, defending and enforcing patents on our product candidates in all countries throughout the world would be prohibitively expensive, and our intellectual property rights in some countries outside the United States can be less extensive than those in the United States. In addition, the laws and enforcement practices of some foreign countries do not protect intellectual



property rights to the same extent as federal and state laws in the United States. Consequently, we may not be able to prevent third parties from practicing our inventions in all countries outside the United States, or from selling or importing products made using our inventions in and into the United States or other jurisdictions. Competitors may use our technologies in jurisdictions where we have not obtained patent protection to develop their own products and may export otherwise infringing products to territories where we have patent protection, but enforcement rights are not as strong as those in the United States. These products may compete with our product candidates and our patents or other intellectual property rights may not be effective or sufficient to prevent them from competing.

Many companies have encountered significant problems in protecting and defending intellectual property rights in foreign jurisdictions. The legal systems of some countries do not favor the enforcement or protection of patents, trade secrets and other intellectual property, which could make it difficult for us to stop the infringement of our patents or marketing of competing products in violation of our intellectual property and proprietary rights generally. Proceedings to enforce our intellectual property rights in foreign jurisdictions could result in substantial costs and divert our efforts and attention from other aspects of our business, could put our patents at risk of being invalidated or interpreted narrowly and our patent applications at risk of not issuing and could provoke third parties to assert claims against us. We may not prevail in any lawsuits that we initiate, and the damages or other remedies awarded, if any, may not be commercially meaningful.

Many foreign countries, including some European Union countries, India, Japan and China, have compulsory licensing laws under which a patent owner may be compelled under specified circumstances to grant licenses to third parties. In addition, many countries limit the enforceability of patents against government agencies or government contractors. In those countries, we may have limited remedies if patents are infringed or if we are compelled to grant a license to a third party, which could materially diminish the value of the applicable patents. This could limit our potential revenue opportunities. Accordingly, our efforts to enforce our intellectual property rights around the world may be inadequate to obtain a significant commercial advantage from the intellectual property that we develop or license.

Changes in patent law could diminish the value of patents in general, thereby impairing our ability to protect our product candidates.

Obtaining and enforcing patents in the pharmaceutical industry is inherently uncertain, due in part to ongoing changes in the patent laws. For example, in the United States, depending on decisions by Congress, the federal courts, and the USPTO, the laws and regulations governing patents, and interpretation thereof, could change in unpredictable ways that could weaken our and our licensors' or collaborators' ability to obtain new patents or to enforce existing or future patents. For example, the U.S. Supreme Court has ruled on several patent cases in recent years, either narrowing the scope of patent protection available in certain circumstances or weakening the rights of patent owners in certain situations. Therefore, there is increased uncertainty with regard to our and our licensors' or collaborators' ability to obtain patents once obtained.

Patent reform legislation could increase the uncertainties and costs surrounding the prosecution of our and our licensors' or collaborators' patent applications and the enforcement or defense of our or our licensors' or collaborators' issued patents. For example, assuming that other requirements for patentability are met, prior to March 2013, in the United States, the first to invent the claimed invention was entitled to the patent, while outside the United States, the first to file a patent application was entitled to the patent. After March 2013, under the Leahy-Smith America Invents Act (the Leahy-Smith Act) enacted in September 2011, the United States transitioned to a first inventor to file system in which, assuming that other requirements for patentability are met, the first inventor to file a patent application will be entitled to the patent on an invention regardless of whether a third party was the first to invent the claimed invention. The Leahy-Smith Act also includes a number of significant changes that affect the way patent applications are prosecuted and may also affect patent litigation. These include allowing third-party submission of prior art to the USPTO during patent prosecution and additional procedures to challenge the validity of a patent by USPTO-administered post-grant proceedings, including post-grant review, *inter partes* review and derivation proceedings. The USPTO has developed regulations and procedures to govern administration of the Leahy-Smith Act, and many of the substantive changes to patent awasociated with the Leahy-Smith Act, particularly the first inventor-to-file provisions. Accordingly, it is not clear what, if any, impact the Leahy-Smith Act will have on the operation of our business. However, the Leahy-Smith Act and its implementation could increase the uncertainties and costs surrounding the prosecution of our or our licensors' patent applications and the enforcement or defense of our or our licensors' patent applications and the enforcement or defense of our or our

Obtaining and maintaining our patent protection depends on compliance with various procedural, document submission, fee payment and other requirements imposed by governmental patent agencies, and our patent protection could be reduced or eliminated if we fail to comply with these requirements.

Periodic maintenance fees, renewal fees, annuity fees, and various other fees are required to be paid to the USPTO and foreign patent agencies in several stages over the lifetime of a patent. In certain circumstances, we rely on our licensors and collaborators to pay these fees. The USPTO and various foreign patent agencies also require compliance with a number of procedural, documentary, fee payment and other similar requirements during the patent application and prosecution process. Non-compliance events that could result in abandonment or lapse of a patent or patent application include failure to respond to official communications within prescribed



time limits, non-payment of fees and failure to properly legalize and submit formal documents. While an inadvertent lapse can in some cases be cured by payment of a late fee or by other means in accordance with the applicable rules, there are situations in which non-compliance can result in irrevocable abandonment or lapse of the patent or patent application, resulting in partial or complete loss of patent rights in the relevant jurisdiction. If we or our licensors or collaborators fail to maintain the patents and patent applications covering our product candidates, our competitors might be able to enter the market with similar or identical products or technology, which would harm our business, financial condition, results of operations and prospects.

We may become involved in lawsuits to protect or enforce our patents or other intellectual property, which could be expensive, time-consuming and unsuccessful, and issued patents covering our technology and product candidates could be found invalid or unenforceable if challenged.

Competitors and other third parties may infringe or otherwise violate our issued patents or other intellectual property or the patents or other intellectual property of our licensors. In addition, our patents or the patents of our licensors may become involved in inventorship or priority disputes. Our pending patent applications cannot be enforced against third parties practicing the technology claimed in such applications unless and until a patent issues from such applications. To counter infringement or other unauthorized use, we may be required to file infringement claims, which can be expensive and time-consuming. Our ability to enforce patent rights also depends on our ability to detect infringement. It may be difficult to detect infringers who do not advertise the components or methods that are used in connection with their products and services. Moreover, it may be difficult or impossible to obtain evidence of infringement in a competitor's or potential competitor's product or service. Any claims we assert against perceived infringers could provoke these parties to assert counterclaims against us alleging that we infringe their patents or that our patents are invalid or unenforceable. In a patent infringement proceeding, a court may decide that a patent of ours is invalid or unenforceable, in whole or in part, construe the patent's claims narrowly or refuse to stop the other party from using the technology at issue on the grounds that our patents do not cover the technology. An adverse result in any litigation proceeding could put one or more of our owned or licensed patents at risk of being invalidated, held unenforceable or interpreted narrowly. We may find it impractical or undesirable to enforce our intellectual property against some third parties.

If we were to initiate legal proceedings against a third party to enforce a patent directed to our product candidates, or one of our future product candidates, the defendant could counterclaim that our patent is invalid or unenforceable. In patent litigation in the United States, defendant counterclaims alleging invalidity or unenforceability are commonplace. Grounds for a validity challenge could be an alleged failure to meet any of several statutory requirements, including lack of novelty, obviousness, non-enablement or insufficient written description. Grounds for an unenforceability assertion could be an allegation that someone connected with prosecution of the patent withheld relevant information from the USPTO or made a misleading statement during prosecution. Third parties may also raise similar claims before the USPTO or an equivalent foreign body, even outside the context of litigation. Potential proceedings include re-examination, post-grant review, inter partes review, interference proceedings, derivation proceedings and equivalent proceedings in foreign jurisdictions (e.g., opposition proceedings). Such proceedings could result in the revocation of, cancellation of, or amendment to our patents in such a way that they no longer cover our technology or any product candidates that we may develop. The outcome following legal assertions of invalidity and unenforceability is unpredictable. With respect to the validity question, for example, we cannot be certain that there is no invalidating prior art of which we and the patent examiner were unaware during prosecution. If a defendant were to prevail on a legal assertion of invalidity or unenforceability, we would lose at least part, and perhaps all, of the patent protection on the applicable product candidates or technology covered by the patent rendered invalid or unenforceabile.

Interference proceedings provoked by third parties or brought by us or declared by the USPTO may be necessary to determine the priority of inventions with respect to our patents or patent applications. An unfavorable outcome could require us to cease using the related technology or to attempt to license rights to it from the prevailing party. Our business could be materially harmed if the prevailing party does not offer us a license on commercially reasonable terms.

Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during this type of litigation.

Some of our competitors are larger than we are and have substantially greater resources. They are, therefore, likely to be able to sustain the costs of complex patent litigation or proceedings more effectively than we can because of their greater financial resources and more mature and developed intellectual property portfolios. Accordingly, despite our efforts, we may not be able to prevent third parties from infringing upon, misappropriating or otherwise violating our intellectual property. Even if resolved in our favor, litigation or other legal proceedings relating to intellectual property claims could result in substantial costs and diversion of management resources, which could harm our business. In addition, the uncertainties associated with litigation could compromise our ability to raise the funds necessary to continue our clinical trials, continue our internal research programs, or in-license needed technology or other product candidates. There could also be public announcements of the results of the hearing, motions, or other interim proceedings or developments. If securities analysts or investors perceive those results to be negative, it could cause the price of shares of our common stock to decline.

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Third parties may initiate legal proceedings alleging that we are infringing, misappropriating or otherwise violating their intellectual property rights, the outcome of which would be uncertain and could negatively impact the success of our business.

Our commercial success depends upon our ability to develop, manufacture, market and sell our product candidates and use our proprietary technologies without infringing, misappropriating or otherwise violating the intellectual property and other proprietary rights of third parties. There is considerable intellectual property litigation in the pharmaceutical industry.

We may become party to, or threatened with, future adversarial proceedings or litigation regarding intellectual property rights with respect to our product candidates and their manufacture and our other technology, including re-examination, interference, post-grant review, inter partes review or derivation proceedings before the USPTO or an equivalent foreign body. Numerous U.S. and foreign issued patents and pending patent applications owned by third parties exist in the fields in which we are developing our product candidates. Third parties may assert infringement claims against us based on existing patents or patents that may be granted in the future, regardless of their merit.

Even if we believe third-party intellectual property claims are without merit, there is no assurance that a court would find in our favor on questions of infringement, validity, enforceability or priority. A court of competent jurisdiction could hold that third-party patents asserted against us are valid, enforceable and infringed, which could materially and adversely affect our ability to commercialize any product candidates we may develop and any other product candidates or technologies covered by the asserted third-party patents. In order to successfully challenge the validity of a U.S. patent in federal court, we would need to overcome a presumption of validity. As this burden is a high one requiring us to present clear and convincing evidence as to the invalidity of a U.S. patent claim, there is no assurance that a court of competent jurisdiction would invalidate the claims of any such U.S. patent. If we are found to infringe, misappropriate or otherwise violate a third party's intellectual property rights, and we are unsuccessful in demonstrating that these rights are invalid or unenforceable, we could be required to obtain a license from such a third party in order to continue developing and marketing our products and technology. However, we may not be able to obtain any required license on commercially reasonable terms or at all. Even if we were able to obtain a license, it could be non-exclusive, thereby giving our competitors access to the same technologies licensed to us. We could be forced, including by court order, to cease commercializing the infringing technology or product. A finding of infringement could prevent us from commercializing our product candidates or force us to cease some of our business operations. In the event of a successful claim of infringement against us, we may have to pay substantial damages, including treble damages and attorneys' fees for willful infringement, pay royalties and other fees, redesign our infringing product candidate or obtain one or more licenses from third

We may be subject to claims by third parties asserting that we or our employees have infringed upon, misappropriated or otherwise violated their intellectual property rights, or claiming ownership of what we regard as our own intellectual property.

Many of our employees were previously employed at other biotechnology or pharmaceutical companies and our consultants and advisors may work for other biotechnology or pharmaceutical companies in addition to us. Although we try to ensure that our employees, consultants and advisors do not use the proprietary information or know-how of others in their work for us, we may be subject to claims that we or these individuals have used or disclosed intellectual property, including trade secrets or other proprietary information, of any of these individuals' former or concurrent employers or clients. We may also be subject to claims that patents and applications we have filed to protect inventions of our employees, consultants and advisors, even those related to one or more of our product candidates, are rightfully owned by their former or concurrent employer. Litigation may be necessary to defend against these claims.

If we fail in prosecuting or defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights or personnel. Even if we are successful in prosecuting or defending against these claims, litigation could result in substantial costs, delay development of our product candidates and be a distraction to management.

We may be subject to claims challenging the inventorship of our patents and other intellectual property.

We or our licensors may be subject to claims that former employees, collaborators or other third parties have an interest in our owned or in-licensed patents, trade secrets, or other intellectual property as an inventor or co-inventor. For example, we or our licensors may have inventorship disputes arise from conflicting obligations of employees, consultants or others who are involved in developing our product candidates. While it is our policy to require our employees and contractors who may be involved in the development of intellectual property to execute agreements assigning this intellectual property to us, we may be unsuccessful in executing such an agreement with each party who in fact develops intellectual property that we regard as our own. Our and their assignment agreements may not be self-executing or may be breached, and litigation may be necessary to defend against these and other claims challenging inventorship or our or our licensors' ownership of our owned or in-licensed patents, trade secrets or other intellectual property. If we or our licensors fail in defending any such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights, such as exclusive ownership of, or right to use, intellectual property that is important to our product candidates. Even if we are successful in defending against such claims, litigation could result in substantial costs and be a distraction to management and other employees.



If we are unable to protect the confidentiality of our trade secrets, our business and competitive position would be harmed.

We rely on trade secrets and confidentiality agreements to protect our unpatented know-how, technology and other proprietary information (including unpatented know-how associated with Warp Drive) and to maintain our competitive position. Trade secrets and know-how can be difficult to protect. We seek to protect these trade secrets and other proprietary technology, in part, by entering into non-disclosure and confidentiality agreements with parties who have access to them, such as our employees, collaborators, CROs, contract manufacturers, consultants, advisors and other third parties. We also enter into confidentiality and invention or patent assignment agreements with our employees and consultants. We cannot guarantee that we have entered into these agreements with each party that may have or has had access to our trade secrets or proprietary technology and processes. Despite our efforts, any of these parties may breach the agreements and disclose our proprietary information, including our trade secrets, and we may not be able to obtain adequate remedies for such breaches. Unauthorized parties may also attempt to copy or reverse engineer certain aspects of our products that we consider proprietary. Monitoring unauthorized uses and disclosures is difficult, and we do not know whether the steps we have taken to protect our proprietary information will be effective.

We also seek to preserve the integrity and confidentiality of our confidential proprietary information by maintaining physical security of our premises and physical and electronic security of our information technology systems, but it is possible that these security measures could be breached. Enforcing a claim that a party illegally disclosed or misappropriated a trade secret is difficult, expensive and time-consuming, and the outcome is unpredictable. In addition, some courts inside and outside the United States are less willing or unwilling to protect trade secrets. If any of our trade secrets were to be lawfully obtained or independently developed by a competitor or other third party, we would have no right to prevent them from using that technology or information to compete with us. If any of our trade secrets were to be disclosed to, or independently developed by, a competitor or other third party, our competitive position would be materially and adversely harmed.

If our trademarks and trade names are not adequately protected, then we may not be able to build name recognition in our markets of interest and our business may be adversely affected.

Our registered or unregistered trademarks or trade names may be challenged, infringed, circumvented or declared generic or determined to be infringing on other marks. We may not be able to protect our rights to these trademarks and trade names, which we need to build name recognition among potential collaborators or customers in our markets of interest. At times, competitors may adopt trade names or trademarks similar to ours, thereby impeding our ability to build brand identity and possibly leading to market confusion. In addition, there could be potential trade name or trademarks or trade names. Over the long term, if we are unable to establish name recognition based on our trademarks and trade names, then we may not be able to compete effectively and our business may be adversely affected. We may license our trademarks and trade names to third parties, such as distributors. Though these license agreements may provide guidelines for how our trademarks and trade names may be used, a breach of these agreements or misuse of our trademarks and tradenames by our licensees may jeopardize our rights in or diminish the goodwill associated with our trademarks and trade names. Our efforts to enforce or protect our proprietary rights related to trademarks, trade names, trade secrets, domain names, copyrights or other intellectual property may be ineffective and could result in substantial costs and diversion of resources.

Intellectual property rights do not necessarily address all potential threats.

The degree of future protection afforded by our intellectual property rights is uncertain because intellectual property rights have limitations and may not adequately protect our business or permit us to maintain our competitive advantage. For example:

- others may be able to make products that are similar to any product candidates we may develop or utilize similar technology but that are not covered by the claims of our patents or the patents that we license or may own in the future;
- we, or our current or future licensors, might not have been the first to make the inventions covered by an issued patent or pending patent application that we license or may own in the future;
- we, or our current or future licensors might not have been the first to file patent applications covering certain of our or their inventions;
- others may independently develop similar or alternative technologies or duplicate any of our technologies without infringing our owned or licensed intellectual property rights;
- it is possible that our pending owned or licensed patent applications or those that we may own or license in the future will not lead to issued patents;
- issued patents that we hold rights to may be held invalid or unenforceable, including as a result of legal challenges by our competitors;
- our competitors might conduct research and development activities in countries where we do not have patent rights and then use the information learned from such activities to develop competitive products for sale in our major commercial markets;
- we may not develop additional proprietary technologies that are patentable;

- the patents of others may harm our business; and
- we may choose not to file a patent in order to maintain certain trade secrets or know-how, and a third party may subsequently file a patent covering such intellectual property.

Risks related to employee matters and managing our growth

We are highly dependent on our key personnel, and if we are not successful in attracting, motivating and retaining highly qualified personnel, we may not be able to successfully implement our business strategy.

We are highly dependent on members of our executive team. The loss of the services of any of them may adversely impact the achievement of our objectives. Any of our executive officers could leave our employment at any time, as all of our employees are "at-will" employees. We currently do not have "key person" insurance on any of our employees. The loss of the services of one or more of our key personnel might impede the achievement of our research, development and commercialization objectives.

Recruiting and retaining qualified employees, consultants and advisors for our business, including scientific and technical personnel, is critical to our success. Competition for skilled personnel is intense and the turnover rate can be high. We may not be able to attract and retain personnel on acceptable terms given the competition among numerous pharmaceutical and biotechnology companies and academic institutions for skilled individuals. In addition, failure to succeed in preclinical studies, clinical trials or applications for marketing approval may make it more challenging to recruit and retain qualified personnel. The inability to recruit, or the loss of services of certain executives, key employees, consultants or advisors, may impede the progress of our research, development and commercialization objectives.

We currently have no sales organization. If we are unable to establish sales capabilities on our own or through third parties, we may not be able to market and sell any products effectively, if approved, or generate product revenue.

We currently do not have a marketing or sales organization. In order to commercialize any product, if approved, in the United States and foreign jurisdictions, we must build our marketing, sales, distribution, managerial and other non-technical capabilities or make arrangements with third parties to perform these services, and we may not be successful in doing so. In advance of any of our product candidates receiving regulatory approval, we expect to establish a sales organization with technical expertise and supporting distribution capabilities to commercialize each such product candidate, which will be expensive and time-consuming. We have no prior experience in the marketing, sale and distribution of pharmaceutical products, and there are significant risks involved in building and managing a sales organization, including our ability to hire, retain, and incentivize qualified individuals, generate sufficient sales leads, provide adequate training to sales and marketing personnel, and effectively manage a geographically dispersed sales and marketing team. Any failure or delay in the development of our internal sales, marketing and distribution capabilities would adversely impact the commercialization of these products. We may choose to collaborate with third parties that have direct sales forces and established distribution systems, either to augment our own sales force and distribution systems. If we are unable to enter into such arrangements on acceptable terms or at all, we may not be able to successfully commercialize our product candidates.

We will need to grow the size of our organization, and we may experience difficulties in managing this growth.

As of June 30, 2021, we had 150 full-time employees, including 122 employees engaged in research and development. As our development and commercialization plans and strategies develop, and as we operate as a public company, we expect to need additional managerial, research and development, operational, sales, marketing, financial and other personnel. Future growth would impose significant added responsibilities on members of management, including:

- identifying, recruiting, integrating, maintaining and motivating additional employees;
- managing our internal development efforts effectively, including the clinical and FDA review process for RMC-4630, RMC-5552 and any other
 product candidate we develop, while complying with our contractual obligations to contractors and other third parties; and
- improving our operational, financial and management controls, reporting systems and procedures.

Our future financial performance and our ability to advance development of and, if approved, commercialize RMC-4630, RMC-5552 and any other product candidate we develop will depend, in part, on our ability to effectively manage any future growth, and our management may also have to divert a disproportionate amount of its attention away from day-to-day activities in order to devote a substantial amount of time to managing these growth activities.

We currently rely, and for the foreseeable future will continue to rely, in substantial part on certain independent organizations, advisors and consultants to provide certain services, including substantially all aspects of marketing approval, clinical management, and manufacturing. We cannot assure you that the services of independent organizations, advisors and consultants will continue to be available to us on a timely basis when needed, or that we can find qualified replacements. In addition, if we are unable to effectively manage our outsourced activities or if the quality or accuracy of the services provided by consultants is compromised for any reason, our clinical trials may be extended, delayed or terminated, and we may not be able to obtain marketing approval of any current or



future product candidates or otherwise advance our business. We cannot assure you that we will be able to manage our existing consultants or find other competent outside contractors and consultants on economically reasonable terms, or at all.

If we are not able to effectively expand our organization by hiring new employees and expanding our groups of consultants and contractors, we may not be able to successfully implement the tasks necessary to further develop and commercialize RMC-4630, RMC-5552 and any future product candidates we develop and, accordingly, may not achieve our research, development and commercialization goals.

We have in the past engaged and may in the future engage in strategic transactions; these transactions could affect our liquidity, dilute our existing stockholders, increase our expenses and present significant challenges in focus and energy to our management or prove not to be successful.

From time to time, we may consider strategic transactions, such as acquisitions of companies, asset purchases and out-licensing or in-licensing of intellectual property, products or technologies. For example, in October 2018, we acquired all of the outstanding shares of Warp Drive Bio, which became our direct wholly-owned subsidiary.

Additional potential transactions that we may consider in the future include a variety of business arrangements, including spin-offs, strategic partnerships, joint ventures, restructurings, divestitures, business combinations and investments. Any future transactions could result in potentially dilutive issuances of our equity securities, including our common stock, or the incurrence of debt, contingent liabilities, amortization expenses or acquired in-process research and development expenses, any of which could affect our financial condition, liquidity and results of operations. Future acquisitions may also require us to obtain additional financing, which may not be available on favorable terms or at all. These transactions may never be successful and may require significant time and attention of management. In addition, the integration of any business that we may acquire in the future may disrupt our existing business and may be a complex, risky and costly endeavor for which we may never realize the full benefits of the acquisition.

If we fail to comply with environmental, health and safety laws and regulations, we could become subject to fines or penalties or incur costs that could have a material adverse effect on the success of our business.

We are subject to numerous environmental, health and safety laws and regulations, including those governing laboratory procedures and the handling, use, storage, treatment and disposal of hazardous materials and wastes. Our operations involve the use of hazardous and flammable materials, including chemicals and biological and radioactive materials. Our operations also produce hazardous waste products. We generally contract with third parties for the disposal of these materials and wastes. We cannot eliminate the risk of contamination or injury from these materials. In the event of contamination or injury resulting from our use of hazardous materials, we could be held liable for any resulting damages, and any liability could exceed our resources. We also could incur significant costs associated with civil or criminal fines and penalties.

Although we maintain workers' compensation insurance to cover us for costs and expenses, we may incur due to injuries to our employees resulting from the use of hazardous materials, this insurance may not provide adequate coverage against potential liabilities. We do not maintain insurance for environmental liability or toxic tort claims that may be asserted against us in connection with our storage or disposal of biological, hazardous or radioactive materials.

We or the third parties upon whom we depend may be adversely affected by earthquakes, outbreak of disease, or other natural disasters and our business continuity and disaster recovery plans may not adequately protect us from a serious disaster.

Our corporate headquarters and other facilities are located in the San Francisco Bay Area, which in the past has experienced both severe earthquakes and wildfires. We do not carry earthquake insurance. Earthquakes, wildfires or other natural disasters could severely disrupt our operations, and have a material adverse effect on our business, results of operations, financial condition and prospects.

If a natural disaster, power outage, outbreak of disease, or other event occurred that prevented us from using all or a significant portion of our headquarters, that damaged critical infrastructure, such as our enterprise financial systems or manufacturing resource planning and enterprise quality systems, or that otherwise disrupted operations, it may be difficult or, in certain cases, impossible, for us to continue our business for a substantial period of time. As an example, our operations are currently limited as a result of the COVID-19 pandemic, as described in the risk factor entitled *"The COVID-19 pandemic, or other epidemic and pandemic diseases or governmental or other actions taken in response to them, could significantly disrupt our business."* The disaster recovery and business continuity plans we have in place currently are limited and are unlikely to prove adequate in the event of a serious disaster or similar event. We may incur substantial expenses as a result of the limited nature of our disaster recovery and business continuity plans, which, particularly when taken together with our lack of earthquake insurance, could have a material adverse effect on our business.

Furthermore, integral parties used in our preclinical activities and in our supply chain are similarly vulnerable to natural disasters, outbreak of disease, or other sudden, unforeseen and severe adverse events. If such an event were to affect our preclinical activities or our supply chain, it could have a material adverse effect on our business.



Our internal computer systems, or those used by our CROs or other contractors or consultants, may fail or suffer security breaches.

Despite the implementation of security measures, our internal computer systems and those of our future CROs and other contractors and consultants are vulnerable to damage from computer viruses and unauthorized access. While we have not to our knowledge experienced any such material system failure or security breach to date, if such an event were to occur and cause interruptions in our operations, it could result in a material disruption of our development programs and our business operations. For example, the loss of clinical trial data from completed or future clinical trials could result in delays in our marketing approval efforts and significantly increase our costs to recover or reproduce the data. Likewise, we rely on third parties for the manufacture of our product candidates and to conduct clinical trials, and similar events relating to their computer systems could also have a material adverse effect on our business. To the extent that any disruption or security breach were to result in a loss of, or damage to, our data or applications, or inappropriate disclosure of confidential or proprietary information, we could incur liability and the further development and commercialization of our future product candidates could be delayed.

Our employees, independent contractors, vendors, principal investigators, CROs and consultants may engage in misconduct or other improper activities, including non-compliance with regulatory standards and requirements and insider trading.

We are exposed to the risk that our employees, independent contractors, vendors, principal investigators, CROs and consultants may engage in fraudulent conduct or other illegal activity. Misconduct by these parties could include intentional, reckless or negligent conduct or disclosure of unauthorized activities to us that violate the regulations of the FDA or comparable foreign regulatory authorities, including those laws requiring the reporting of true, complete and accurate information to such authorities; healthcare fraud and abuse laws and regulations in the United States and abroad; or laws that require the reporting of financial information or data accurately. In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, misconduct, kickbacks, self-dealing and other abusive practices. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Activities subject to these laws also involve the improper use of information obtained in the course of clinical trials or creating fraudulent data in our preclinical studies or clinical trials, which could result in regulatory sanctions and cause serious harm to our reputation. It is not always possible to identify and deter misconduct by employees and other third parties, and the precautions we take to detect and prevent this activity may not be effective in controlling unknown or unmanaged risks or losses or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to comply with these laws or regulations. Additionally, we are subject to the risk that a person could allege fraud or other misconduct, even if none occurred. If any such actions are instituted against us, and we are not successful in defending ourselves or asserting our rights, those actions could have a significant impact on our

Risks related to our common stock

The price of our common stock is volatile and fluctuates substantially, which could result in substantial losses for investors.

Our stock price is highly volatile. The stock market in general, and the market for biopharmaceutical companies in particular, have experienced extreme volatility that has often been unrelated to the operating performance of particular companies. The COVID-19 pandemic, actions taken by governments in response to this pandemic and the related economic impacts, have exacerbated this volatility, particularly as it relates to stocks of biopharmaceutical companies like ours.

The market price for our common stock may be influenced by many factors, including:

- our research and development efforts and our ability to discover and develop product candidates;
- results of our clinical trials and preclinical studies or those of our competitors;
- the success of competitive products or technologies;
- regulatory or legal developments in the United States and other countries;
- · developments or disputes concerning patent applications, issued patents or other proprietary rights;
- the recruitment or departure of key personnel;
- the level of expenses related to our product candidates or clinical development programs;
- the results of our efforts to discover, develop, acquire or in-license product candidates or companion diagnostics;
- actual or anticipated changes in estimates as to financial results, development timelines or recommendations by securities analysts;
- variations in our financial results or those of companies that are perceived to be similar to us;
- changes in the structure of healthcare payment systems;

- market conditions in the pharmaceutical and biotechnology sectors; and
- general economic, industry and market conditions.

An active and liquid market for our common stock may not be sustained.

Our common stock is currently listed on the Nasdaq Global Select Market under the symbol "RVMD". The price for our common stock may vary and an active and liquid market in our common stock may not be sustained. The lack of an active market may impair the value of your shares, your ability to sell your shares at the time you wish to sell them and the prices that you may obtain for your shares. An inactive market may also impair our ability to raise capital by selling our common stock and our ability to acquire other companies, products or technologies by using our common stock as consideration.

We do not intend to pay dividends on our common stock so any returns will be limited to the value of our stock.

We do not currently intend to pay any cash dividends on our common stock for the foreseeable future. We currently intend to invest our future earnings, if any, to fund our growth. Therefore, stockholders are not likely to receive any dividends on their common stock for the foreseeable future. Since we do not intend to pay dividends, stockholders' ability to receive a return on their investment will depend on any future appreciation in the market value of our common stock. There is no guarantee that our common stock will appreciate or even maintain the price at which our holders have purchased it.

We are an emerging growth company, and we cannot be certain if the reduced reporting requirements applicable to emerging growth companies will make our common stock less attractive to investors.

We are an emerging growth company, as defined in the JOBS Act. For as long as we continue to be an emerging growth company, we may take advantage of exemptions from various reporting requirements, that are applicable to other public companies that are not emerging growth companies, including not being required to comply with the auditor attestation requirements of Section 404 of the Sarbanes-Oxley Act, reduced disclosure obligations regarding executive compensation and exemptions from the requirements of holding nonbinding advisory votes on executive compensation and stockholder approval of any golden parachute payments not previously approved. We will remain an emerging growth company until the earlier of (1) December 31, 2025, (2) the last day of the fiscal year in which we have total annual gross revenue of at least \$1.07 billion, (3) the last day of the fiscal year in which we are deemed to be a large accelerated filer, which requires that we be subject to the reporting requirements of Section 13(a) or 15(d) of the Exchange Act for a period of at least 12 calendar months and that the market value of our common stock that is held by non-affiliates to exceed \$700.0 million as of the prior June 30th, or (4) the date on which we have issued more than \$1.0 billion in non-convertible debt during the prior three-year period. We currently expect that we will no longer be an emerging growth company on December 31, 2021. We cannot predict if investors will find our common stock less attractive because we may rely on these exemptions. If some investors find our common stock less attractive as a result, our stock price may be depressed, there may be a less active trading market for our common stock and our stock price may be more volatile.

Under the JOBS Act, emerging growth companies can also delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have elected to use the extended transition period for any other new or revised accounting standards during the period in which we remain an emerging growth company; however, we may adopt certain new or revised accounting standards early. As a result, changes in rules of U.S. generally accepted accounting principles or their interpretation, the adoption of new guidance or the application of existing guidance to changes in our business could significantly affect our financial position and results of operations.

Our executive officers, directors and their affiliates have significant influence over our company, which will limit your ability to influence corporate matters and could delay or prevent a change in corporate control.

As of June 30, 2021, our executive officers, directors and their affiliates beneficially owned, in the aggregate, approximately 17% of our outstanding common stock. As a result, these stockholders, if they act together, will be able to influence our management and affairs and the outcome of matters submitted to our stockholders for approval, including the election of directors and any sale, merger, consolidation or sale of all or substantially all of our assets. In addition, this concentration of ownership might adversely affect the market price of our common stock by:

- delaying, deferring or preventing a change of control of us;
- impeding a merger, consolidation, takeover or other business combination involving us; or
- discouraging a potential acquirer from making a tender offer or otherwise attempting to obtain control of us.

Sales of a substantial number of shares of our common stock in the public market could cause our stock price to fall.

If our existing stockholders sell, or indicate an intention to sell, substantial amounts of our common stock in the public market, the market price of our common stock could decline.

As of June 30, 2021, 14.2 million shares of common stock that are either subject to outstanding options or reserved for future issuance under our equity incentive plans are eligible for sale in the public market to the extent permitted by the provisions of various vesting



schedules, lock-up agreements and Rule 144 and Rule 701 under the Securities Act. If these additional shares of common stock are sold, or if it is perceived that they will be sold, in the public market, the market price of our common stock could decline.

In addition, as of June 30, 2021, holders of approximately 14.7 million shares of our common stock are entitled to rights with respect to the registration of their shares under the Securities Act. Registration of these shares under the Securities Act would result in the shares becoming freely tradable without restriction under the Securities Act, except for shares purchased by affiliates. Any sales of securities by these stockholders could have a material adverse effect on the market price of our common stock.

Our ability to utilize our net operating loss carryforwards and certain other tax attributes has been limited by "ownership changes" and may be further limited.

Under Sections 382 and 383 of the Internal Revenue Code of 1986, as amended (the Code), and corresponding provisions of state law, if a corporation undergoes an "ownership change" (generally defined as a greater than 50 percentage point change (by value) in its equity ownership over a rolling threeyear period), the corporation's ability to use its pre-change net operating loss carryforwards and other pre-change tax attributes to offset its post-change income may be limited. We have experienced ownership changes in the past, and we may experience ownership changes in the future as a result of our public offerings or other changes in our stock ownership (some of which are not in our control). Use of our federal and state net operating loss carryforwards have been limited as a result of ownership changes and could be further limited if we experience additional ownership changes.

Provisions in our charter documents and under Delaware law could discourage a takeover that stockholders may consider favorable and may lead to entrenchment of management.

Our amended and restated certificate of incorporation and amended and restated bylaws contain provisions that could delay or prevent changes in control or changes in our management without the consent of our board of directors. These provisions include the following:

- a classified board of directors with three-year staggered terms, which may delay the ability of stockholders to change the membership of our board of directors;
- no cumulative voting in the election of directors, which limits the ability of minority stockholders to elect director candidates;
- the exclusive right of our board of directors to appoint a director to fill a vacancy created by the expansion of the board of directors or the resignation, death or removal of a director, which prevents stockholders from being able to fill vacancies on our board of directors;
- the ability of our board of directors to authorize the issuance of shares of preferred stock and to determine the price and other terms of those shares, including preferences and voting rights, without stockholder approval, which could be used to significantly dilute the ownership of a hostile acquiror;
- the ability of our board of directors to alter our amended and restated bylaws without obtaining stockholder approval;
- the required approval of at least 66 2/3% of the shares entitled to vote at an election of directors to adopt, amend or repeal our amended and restated bylaws or repeal the provisions of our amended and restated certificate of incorporation regarding the election and removal of directors;
- a prohibition on stockholder action by written consent, which forces stockholder action to be taken at an annual or special meeting of our stockholders;
- the requirement that a special meeting of stockholders may be called only by our chief executive officer or president or by the board of directors, which may delay the ability of our stockholders to force consideration of a proposal or to take action, including the removal of directors; and
- advance notice procedures that stockholders must comply with in order to nominate candidates to our board of directors or to propose matters to be
 acted upon at a stockholders' meeting, which may discourage or deter a potential acquiror from conducting a solicitation of proxies to elect the
 acquiror's own slate of directors or otherwise attempting to obtain control of us.

We are also subject to the anti-takeover provisions contained in Section 203 of the Delaware General Corporation Law. Under Section 203, a corporation may not, in general, engage in a business combination with any holder of 15% or more of its capital stock unless the holder has held the stock for three years or, among other exceptions, the board of directors has approved the transaction.

Claims for indemnification by our directors and officers may reduce our available funds to satisfy successful third-party claims against us and may reduce the amount of money available to us.

Our amended and restated certificate of incorporation and amended and restated bylaws provide that we will indemnify our directors and officers, in each case to the fullest extent permitted by Delaware law.



In addition, as permitted by Section 145 of the Delaware General Corporation Law, our amended and restated by laws and our indemnification agreements that we have entered into with our directors and officers provide that:

- we will indemnify our directors and officers for serving us in those capacities or for serving other business enterprises at our request, to the fullest extent permitted by Delaware law. Delaware law provides that a corporation may indemnify such person if such person acted in good faith and in a manner such person reasonably believed to be in or not opposed to the best interests of the registrant and, with respect to any criminal proceeding, had no reasonable cause to believe such person's conduct was unlawful;
- we may, in our discretion, indemnify employees and agents in those circumstances where indemnification is permitted by applicable law;
- we are required to advance expenses, as incurred, to our directors and officers in connection with defending a proceeding, except that such directors or
 officers shall undertake to repay such advances if it is ultimately determined that such person is not entitled to indemnification;
- we are not obligated pursuant to our amended and restated bylaws to indemnify a person with respect to proceedings initiated by that person against us or our other indemnitees, except with respect to proceedings authorized by our board of directors or brought to enforce a right to indemnification;
- the rights conferred in our amended and restated bylaws are not exclusive, and we are authorized to enter into indemnification agreements with our directors, officers, employees and agents and to obtain insurance to indemnify such persons; and
- we may not retroactively amend our amended and restated bylaw provisions to reduce our indemnification obligations to directors, officers, employees
 and agents.

Our amended and restated certificate of incorporation and amended and restated bylaws provide for an exclusive forum in the Court of Chancery of the State of Delaware for certain disputes between us and our stockholders, which could limit our stockholders' ability to obtain a favorable judicial forum for disputes with us or our directors, officers or employees.

Our amended and restated certificate of incorporation and amended and restated bylaws provide that the Court of Chancery of the State of Delaware is the exclusive forum for any state law derivative action or proceeding brought on our behalf, any action asserting a breach of fiduciary duty, any action asserting a claim against us arising pursuant to the Delaware General Corporation Law, any action to interpret, apply, enforce, or determine the validity of our amended and restated certificate of incorporation or our amended and restated bylaws, or any action asserting a claim against us that is governed by the internal affairs doctrine; provided that, the exclusive forum provision will not apply to suits brought to enforce any liability or duty created by the Exchange Act or any other claim for which the federal courts have exclusive jurisdiction; and provided further that, if and only if the Court of Chancery of the State of Delaware. Our amended and restated bylaws also provide that the federal district courts of the United States of America will be the exclusive forum for the resolution of any complaint asserting a cause or causes of action under the Securities Act. Such provision is intended to benefit and may be enforced by us, our officers and directors, the underwriters to any offering giving rise to such compliant and any other professional or entity whose profession gives authority to a statement made by that person or entity and who has prepared or certified any part of the documents underlying the offering. Nothing in our amended and restated certificate of incorporation or amended and restated bylaws preclude stockholders that assert claims under the Exchange Act from bringing such claims in state or federal court, subject to applicable law.

We believe these provisions may benefit us by providing increased consistency in the application of Delaware law and federal securities laws by chancellors and judges, as applicable, particularly experienced in resolving corporate disputes, efficient administration of cases on a more expedited schedule relative to other forums and protection against the burdens of multi-forum litigation. This choice of forum provision may limit a stockholder's ability to bring a claim in a judicial forum that it finds favorable for disputes with us or any of our directors, officers, other employees or stockholders, which may discourage lawsuits with respect to such claims, although our stockholders will not be deemed to have waived our compliance with federal securities laws and the rules and regulations thereunder. Furthermore, the enforceability of similar choice of forum provisions in other companies' certificates of incorporation has been challenged in legal proceedings, and it is possible that a court could find these types of provisions to be inapplicable or unenforceable. While the Delaware courts have determined that such choice of forum provisions are facially valid, a stockholder may nevertheless seek to bring a claim in a venue other than those designated in the exclusive-forum provisions, and there can be no assurance that such provisions will be enforced by a court in those other jurisdictions. If a court were to find the choice of forum provision in our amended and restated certificate of incorporation or amended and restated bylaws to be inapplicable or unenforceable in an action, we may incur additional costs associated with resolving such action in other jurisdictions, which could adversely affect our business, financial condition, results of operations and prospects.

General risk factors



Raising additional capital may cause dilution to our stockholders, restrict our operations or require us to relinquish rights to our technologies.

To date, we have primarily financed our operations through the sale of preferred stock and common stock and upfront payments and research and development cost reimbursement received in connection with our collaboration with Sanofi. We will be required to seek additional funding in the future and may do so through a combination of public or private equity offerings, debt financings, credit or loan facilities, collaborations, strategic alliances, licensing arrangements and other marketing or distribution arrangements. If we raise additional capital through marketing and distribution arrangements or other collaborations, strategic alliances or licensing arrangements with third parties, we may have to relinquish certain valuable rights to our product candidates, technologies, future revenue streams or research programs or grant licenses on terms that may not be favorable to us. If we raise additional funds by issuing equity securities, our stockholders may suffer dilution and the terms of any financing may adversely affect the rights of our stockholders. In addition, as a condition to providing additional funds to us, future investors may demand, and may be granted, rights superior to those of existing stockholders. Debt financing, if available, is likely to involve restrictive covenants limiting our flexibility in conducting future business activities, and, in the event of insolvency, debt holders would be repaid before holders of our equity securities would receive any distribution of our corporate assets. Attempting to secure additional financing may also divert our management from our day-to-day activities, which may adversely affect our ability to develop our product candidates.

We are subject to certain U.S. and foreign anti-corruption, anti-money laundering, export control, sanctions, and other trade laws and regulations. We can face serious consequences for violations.

Among other matters, U.S. and foreign anti-corruption, anti-money laundering, export control, sanctions, and other trade laws and regulations, which are collectively referred to as Trade Laws, prohibit companies and their employees, agents, clinical research organizations, legal counsel, accountants, consultants, contractors, and other partners from authorizing, promising, offering, providing, soliciting, or receiving directly or indirectly, corrupt or improper payments or anything else of value to or from recipients in the public or private sector. Violations of Trade Laws can result in substantial criminal fines and civil penalties, imprisonment, the loss of trade privileges, debarment, tax reassessments, breach of contract and fraud litigation, reputational harm, and other consequences. We have direct or indirect interactions with officials and employees of government agencies or government-affiliated hospitals, universities, and other organizations. We also expect our non-U.S. activities to increase in time. We plan to engage third parties for clinical trials and/or to obtain necessary permits, licenses, patent registrations, and other regulatory approvals and we can be held liable for the corrupt or other illegal activities of our personnel, agents, or partners, even if we do not explicitly authorize or have prior knowledge of such activities.

Intellectual property litigation could cause us to spend substantial resources and distract our personnel from their normal responsibilities.

Even if resolved in our favor, litigation or other legal proceedings relating to intellectual property claims may cause us to incur significant expenses, and could distract our technical and management personnel from their normal responsibilities. In addition, there could be public announcements of the results of hearings, motions or other interim proceedings or developments and if securities analysts or investors perceive these results to be negative, it could have a substantial adverse effect on the price of our common stock. Such litigation or proceedings could substantially increase our operating losses and reduce the resources available for development activities or any future sales, marketing or distribution activities. We may not have sufficient financial or other resources to conduct such litigation or proceedings adequately. As noted above, some of our competitors may be able to sustain the costs of such litigation or proceedings more effectively than we can because of their greater financial resources. Uncertainties resulting from the initiation and continuation of patent litigation or other proceedings could compromise our ability to compete in the marketplace, including compromising our ability to raise the funds necessary to continue our clinical trials, continue our research programs, license necessary technology from third parties, or enter into development collaborations that would help us commercialize our product candidates, if approved.

We incur significantly increased costs as a result of operating as a public company, and our management devotes substantial time to new compliance initiatives. We may fail to comply with the rules that apply to public companies, including Section 404 of the Sarbanes-Oxley Act of 2002, which could result in sanctions or other penalties that would harm our business.

We incur significant legal, accounting and other expenses as a public company, including costs resulting from public company reporting obligations under the Securities Exchange Act of 1934, as amended (the Exchange Act), and regulations regarding corporate governance practices. The listing requirements of the Nasdaq Global Select Market and the rules of the Securities and Exchange Commission (SEC) require that we satisfy certain corporate governance requirements relating to director independence, filing annual and interim reports, stockholder meetings, approvals and voting, soliciting proxies, conflicts of interest and a code of conduct. Our management and other personnel devote a substantial amount of time to ensure that we comply with all of these requirements.



Moreover, the reporting requirements, rules and regulations will increase our legal and financial compliance costs and make some activities more timeconsuming and costly. Any changes we make to comply with these obligations may not be sufficient to allow us to satisfy our obligations as a public company on a timely basis, or at all. These reporting requirements, rules and regulations, coupled with the increase in potential litigation exposure associated with being a public company, could also make it more difficult for us to attract and retain qualified persons to serve on our board of directors or board committees or to serve as executive officers, or to obtain certain types of insurance, including directors' and officers' insurance, on acceptable terms. Stockholder activism, the current political environment and the current high level of government intervention and regulatory reform may lead to substantial new regulations and disclosure obligations, which may lead to additional compliance costs and impact the manner in which we operate our business in ways we cannot currently anticipate.

As a public company, we are subject to Section 404 of the Sarbanes-Oxley Act of 2002 (Section 404) and the related rules of the SEC, which generally require our management and independent registered public accounting firm to report on the effectiveness of our internal control over financial reporting. However, for so long as we remain an emerging growth company as defined in the JOBS Act, we intend to take advantage of certain exemptions from various reporting requirements that are applicable to public companies that are not emerging growth companies, including, but not limited to, not being required to comply with the auditor attestation requirements of Section 404. Once we are no longer an emerging growth company or, if prior to such date, we opt to no longer take advantage of the applicable exemption, we will be required to include an opinion from our independent registered public accounting firm on the effectiveness of our internal control over financial reporting. We currently expect that we will no longer be an emerging growth company on December 31, 2021.

In order to provide the reports required by these rules we must conduct reviews and testing of our internal controls. During the course of our review and testing, we may identify deficiencies and be unable to remediate them before we must provide the required reports. Furthermore, if we have a material weakness in our internal controls over financial reporting, we may not detect errors on a timely basis and our financial statements may be materially misstated. We or our independent registered public accounting firm may not be able to conclude on an ongoing basis that we have effective internal control over financial reporting, which could harm our operating results, cause investors to lose confidence in our reported financial information and cause the trading price of our stock to fall. In addition, as a public company we are required to file accurate and timely quarterly and annual reports with the SEC under the Exchange Act. In order to report our results of operations and financial statements on an accurate and timely basis, we will depend on third party vendors to provide timely and accurate notice of their costs to us. Any failure to report our financial results on an accurate and timely basis could result in sanctions, lawsuits, delisting of our shares from the Nasdaq Global Select Market or other adverse consequences that would materially harm to our business.

If we sell shares of our common stock in future financings, stockholders may experience immediate dilution and, as a result, our stock price may decline.

We may from time to time issue additional shares of common stock at a discount from the current trading price of our common stock. As a result, our stockholders would experience immediate dilution upon the purchase of any shares of our common stock sold at such discount. In addition, as opportunities present themselves, we may enter into financing or similar arrangements in the future, including the issuance of debt securities, preferred stock or common stock. If we issue common stock or securities convertible into common stock, our common stockholders would experience additional dilution and, as a result, our stock price may decline.

If securities analysts do not continue to publish research or reports about our business or if they publish negative evaluations of our stock, the price of our stock could decline.

The trading market for our common stock relies, in part, on the research and reports that industry or financial analysts publish about us or our business. If few analysts publish research or reports about us, the trading price of our stock would likely decrease. If one or more of the analysts covering our business downgrade their evaluations of our stock, the price of our stock could decline. If one or more of these analysts cease to cover our stock, we could lose visibility in the market for our stock, which, in turn, could cause our stock price to decline.

If we fail to maintain proper and effective internal control over financial reporting, our ability to produce accurate and timely financial statements could be impaired, investors may lose confidence in our financial reporting and the trading price of our common stock may decline.

Pursuant to Section 404 of Sarbanes-Oxley, our management is required to report upon the effectiveness of our internal control over financial reporting. When we lose our status as an "emerging growth company," our independent registered public accounting firm will be required to attest to the effectiveness of our internal control over financial reporting. We currently expect that we will no longer be an emerging growth company on December 31, 2021. The rules governing the standards that must be met for management



to assess our internal control over financial reporting are complex and require significant documentation, testing and possible remediation.

There may be material weaknesses in our internal control over financial reporting in the future. Any failure to implement and maintain internal control over financial reporting could severely inhibit our ability to accurately report our financial condition, results of operations or cash flows. If we are unable to conclude that our internal control over financial reporting is effective, or if our independent registered public accounting firm determines we have a material weakness in our internal control over financial reporting, investors may lose confidence in the accuracy and completeness of our financial reports, the market price of our common stock could decline, and we could be subject to sanctions or investigations by Nasdaq, the SEC or other regulatory authorities. Failure to remedy any material weakness in our internal control over financial control over financial reporting over financial reporting, or to implement or maintain other effective control systems required of public companies, could also restrict our future access to the capital markets.

Item 2. Unregistered Sales of Equity Securities and Use of Proceeds.

Unregistered Sales of Equity Securities

None.

Use of Proceeds from the Sale of Registered Securities

In February 2020, our registration statement on Form S-1 (File No. 333-235968) relating to our IPO of shares of common stock became effective. The IPO closed on February 18, 2020 at which time we issued 16,100,000 shares of common stock (including the exercise in full by the underwriters of their option to purchase an additional 2,100,000 shares of common stock) at a public offering price of \$17.00 per share. We received net proceeds from the IPO of approximately \$250.7 million, after deducting the underwriting discounts and commissions of \$19.2 million and expenses of \$3.8 million. None of the expenses associated with the IPO were paid to directors, officers, persons owning 10% or more of any class of equity securities, or to our affiliates. J.P. Morgan Securities LLC, Cowen and Company, LLC, SVB Leerink LLC and Guggenheim Securities, LLC acted as joint book-running managers for the offering.

There has been no material change in the planned use of proceeds from our IPO as described in our final prospectus filed with the SEC on February 13, 2020, pursuant to Rule 424(b)(4). We invested the funds received in interest-bearing, investment-grade securities consisting of government securities, corporate bonds and commercial paper.

Issuer Purchases of Equity Securities

None.

Item 3. Defaults Upon Senior Securities.

None.

Item 4. Mine Safety Disclosures.

None.

Item 5. Other Information.

None.

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Item 6. Exhibits.

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Exhibit Number	Exhibit Description	Incorporated by Reference			Provided
		Form	Date	Number	Herewith
2.1	<u>Agreement and Plan of Merger, dated as of October 15, 2018,</u>	S-1	1/17/2020	2.1	
	by and among Revolution Medicines, Inc., Trotsky Merger				
	Sub, Inc., Warp Drive Bio, Inc., and Fortis Advisors LLC.				
3.1	Amended and Restated Certificate of Incorporation.	8-K	2/18/2020	3.1	
3.2	Amended and Restated Bylaws.	8-K	3/8/2021	3.1	
10.1#	Non-Employee Director Compensation Program.				Х
10.2†	Letter Agreement and Amendment, dated as of August 5, 2021				
	by and between Revolution Medicines, Inc. and Genzyme				Х
	Corporation.				
31.1	Certification of Principal Executive Officer Pursuant to Rules				
	<u>13a-14(a) and 15d-14(a) under the Securities Exchange Act of</u>				Х
	<u>1934, as Adopted Pursuant to Section 302 of the Sarbanes-</u>				Λ
	Oxley Act of 2002.				
31.2	Certification of Principal Financial Officer Pursuant to Rules				
	<u>13a-14(a) and 15d-14(a) under the Securities Exchange Act of</u>				Х
	<u>1934, as Adopted Pursuant to Section 302 of the Sarbanes-</u>				Λ
	Oxley Act of 2002.				
32.1*	Certification of Principal Executive Officer Pursuant to 18				
	U.S.C. Section 1350, as Adopted Pursuant to Section 906 of the				Х
	Sarbanes-Oxley Act of 2002.				
32.2*	Certification of Principal Financial Officer Pursuant to 18				
	U.S.C. Section 1350, as Adopted Pursuant to Section 906 of the				Х
	Sarbanes-Oxley Act of 2002.				
101.INS	XBRL Instance Document.				Х
101.SCH	XBRL Taxonomy Extension Schema Document.				Х
101.CAL	XBRL Taxonomy Extension Calculation Linkbase Document.				Х
101.DEF	XBRL Taxonomy Extension Definition Linkbase Document.				Х
101.LAB	XBRL Taxonomy Extension Label Linkbase Document.				Х
101.PRE	XBRL Taxonomy Extension Presentation Linkbase Document.				Х
104	The cover page from the Company's Quarterly Report on Form				
	10-Q for the quarter ended June 30, 2021 has been formatted in				Х
	Inline XBRL.				

Certain portions of this document that constitute confidential information have been redacted in accordance with Regulation S-K, Item 601(b)(10).

Indicates management contract or compensatory plan.

The certifications attached as Exhibits 32.1 and 32.2 that accompany this Quarterly Report on Form 10-Q, are deemed furnished and not filed with the Securities and Exchange Commission and are not to be incorporated by reference into any filing of Revolution Medicines, Inc. under the Securities Act of 1933, as amended, or the Securities Exchange Act of 1934, as amended, whether made before or after the date of this Quarterly Report on Form 10-Q, irrespective of any general incorporation language contained in such filing.

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 thereunto duly authorized.

Revolution Medicines, Inc.

By: /s/ Mark A. Goldsmith Mark A. Goldsmith, M.D., Ph.D. Chief Executive Officer (Principal Executive Officer)

Revolution Medicines, Inc.

By: /s/ Jack Anders

Jack Anders Senior Vice President, Finance and Principal Accounting Officer (Principal Financial and Accounting Officer)

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Date: August 11, 2021

Date: August 11, 2021

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REVOLUTION MEDICINES, INC. NON-EMPLOYEE DIRECTOR COMPENSATION PROGRAM

As amended, June 22, 2021

This Revolution Medicines, Inc. (the "*Company*") Non-Employee Director Compensation Program (this "*Program*") has been adopted under the Company's 2020 Incentive Award Plan (the "*Plan*"). The cash and equity compensation described in this Program shall be paid or made, as applicable, automatically and without further action of the board of directors of the Company (the "*Board*") to each member of the Board who is not an employee of the Company or any parent or subsidiary of the Company (each, a "*Non-Employee Director*") who is entitled to receive such cash or equity compensation, unless such Non-Employee Director declines the receipt of such cash or equity compensation by written notice to the Company. Capitalized terms not otherwise defined herein shall have the meaning ascribed in the Plan.

Cash Compensation

Annual retainers will be paid in the following amounts to Non-Employee Directors:

Non-Employee Director:	\$40,000
Lead Independent Director	\$25,000
Non-Executive Chair:	\$30,000
Audit Committee Chair:	\$15,000
Compensation Committee Chair:	\$12,000
Nominating and Corporate Governance Committee Chair:	\$10,000
Research and Development Committee Chair:	\$10,000
Audit Committee Member (non-Chair):	\$7,500
Compensation Committee Member (non-Chair):	\$6,000
Nominating and Corporate Governance Committee Member (non-Chair):	\$5,000
Research and Development Committee Member (non-Chair):	\$5,000

All annual retainers will be paid in cash quarterly in arrears promptly following the end of the applicable calendar quarter, but in no event more than 30 days after the end of such quarter. In the event a Non-Employee Director does not serve as a Non-Employee Director, or in the applicable positions described above, for an entire calendar quarter, the retainer paid to such Non-Employee Director shall be prorated for the portion of such calendar quarter actually served as a Non-Employee Director, or in such position, as applicable.

Equity Compensation

Each Non-Employee Director who is initially elected or appointed to serve on the Board shall be granted (i) an Option under the Plan or any other applicable Company equity incentive plan then-maintained by the Company to purchase 25,318 shares of Common Stock (the "*Initial Option*") and (ii) 7,234 restricted stock units under the Plan or any other applicable Company equity incentive plan then-maintained by the Company (the "*Initial RSUs*" and together with the Initial Option, the "*Initial Awards*").

The Initial Option will be automatically granted on the date on which such Non-Employee Director commences service on the Board, and will vest as to 1/36th of the shares subject thereto on each monthly anniversary of the applicable date of grant such that the shares subject to the Initial Option are fully vested on the third anniversary of the grant, subject to the Non-Employee Director continuing in service through each such vesting date.

The Initial RSUs will be automatically granted on the date on which such Non-Employee Director commences service on the Board, and, subject to the Non-Employee Director continuing in service through each such vesting date, will vest in 12 substantially equal quarterly installments commencing on the first Quarterly Vesting Date (as defined below) following the grant date. For the purposes of this Program, a "*Quarterly Vesting Date*" shall mean March 15, June 15, September 15 or December 15.

Each Non-Employee Director who has been serving on the Board since prior to April 1 of the calendar year in which an annual meeting of the Company's stockholders (each, an "*Annual Meeting*") occurs and will continue to serve as a Non-Employee Director immediately following such meeting, shall be granted (i) an Option under the Plan or any other applicable Company equity incentive plan thenmaintained by the Company to purchase 12,659 shares of Common Stock (the "*Annual Option*") and (ii) 3,617 restricted stock units under the Plan or any other applicable Company equity incentive plan then-maintained by the Company equity incentive plan then-maintained by the Company (the "*Annual RSUs*" and together with the Annual Option, the "*Annual Awards*").

Notwithstanding the foregoing, the number of shares subject to each Annual Award granted to any Non-Employee Director who joined the Board after the previous Annual Meeting, but prior to April 1 of the calendar year in which the applicable Annual Meeting occurs will be multiplied by a fraction, the numerator of which is the number of days elapsed between the date of such Non-Employee Director's appointment and the applicable Annual Meeting and the denominator of which is 365, and rounded down to the nearest whole share.

For the avoidance of doubt, any Non-Employee Director who joins the Board after April 1 of a calendar year shall not be eligible for Annual Awards granted at the first Annual Meeting that occurs after their appointment in the calendar year of their appointment.

The Annual Option will be automatically granted on the date of the applicable Annual Meeting, and will vest in full on the earlier of (i) the first anniversary of the date of grant and (ii) immediately prior to the Annual Meeting following the date of grant, subject to the Non-Employee Director continuing in service through such vesting date.

The Annual RSUs will be automatically granted on the date of the applicable Annual Meeting, and will vest in full on the earlier of (i) the first anniversary of the first Quarterly Vesting Date following the date of grant and (ii) immediately prior to the Annual Meeting following the date of grant, subject to the Non-Employee Director continuing in service through such vesting date.

Annual Awards:

The per share exercise price of each Option granted to a Non-Employee Director shall equal the Fair Market Value of a share of Common Stock on the date the Option is granted.

The term of each Option granted to a Non-Employee Director shall be ten years from the date the Option is granted.

No portion of the Initial Awards or Annual Awards which is unvested or unexercisable at the time of a Non-Employee Director's termination of service shall become vested and exercisable thereafter.

Members of the Board who are employees of the Company or any parent or subsidiary of the Company who subsequently terminate their service with the Company and any parent or subsidiary of the Company and remain on the Board will not receive Initial Awards, but to the extent that they are otherwise eligible, will be eligible to receive, after termination from service with the Company and any parent or subsidiary of the Company. Annual Awards as described above.

Change in Control

3

Upon a Change in Control of the Company, all outstanding equity awards granted under the Plan and any other equity incentive plan maintained by the Company that are held by a Non-Employee Director shall become fully vested and/or exercisable, irrespective of any other provisions of the Non-Employee Director's Award Agreement.

Reimbursements

The Company shall reimburse each Non-Employee Director for all reasonable, documented, out-of-pocket travel and other business expenses incurred by such Non-Employee Director in the performance of his or her duties to the Company in accordance with the Company's applicable expense reimbursement policies and procedures as in effect from time to time.

Miscellaneous

The other provisions of the Plan shall apply to the Awards granted automatically pursuant to this Program, except to the extent such other provisions are inconsistent with this Program. All applicable terms of the Plan apply to this Program as if fully set forth herein, and all grants of Awards hereby are subject in all respects to the terms of the Plan (including Section 5.5 of the Plan limiting the sum of the grant date fair value of all equity-based Awards and the maximum amount that may become payable pursuant to all cash-based Awards that may be granted to a Service Provider (as defined in the Plan) as compensation for services as a Non-Employee Director during any calendar year to \$1,000,000). If the Company anticipates that the limit in Section 5.5 of the Plan will be exceeded, the Board will reduce compensation to a level deemed appropriate in its sole discretion. The grant of any Award under this Program shall be made solely by and subject to the terms set forth in a written agreement in a form to be approved by the Board and duly executed by an executive officer of the Company.

* * * * *

I hereby certify that the foregoing Program was adopted by the Board of Directors of Revolution Medicines, Inc. on June 22, 2021.

* * * * *

Executed on June 22, 2021.

<u>Margaret A.</u> <u>Horn</u> Corporate Secretary

EXECUTION VERSION

[***] Certain information in this document has been excluded pursuant to Regulation S-K, Item 601(b)(10). Such excluded information is not material and would likely cause competitive harm to the registrant if publicly disclosed.



August 5, 2021

Revolution Medicines, Inc. 700 Saginaw Dr. Redwood City, CA 94063

To Whom It May Concern:

Reference is hereby made to that certain Collaborative Research, Development and Commercialization Agreement (as amended from time to time, the "<u>Agreement</u>"), dated as of June 8, 2018, by and between Revolution Medicines, Inc. ("<u>RevMed</u>") and Aventis, Inc. ("<u>Aventis</u>"), as amended by that certain letter agreement dated as of August 24, 2018, and as assigned by Aventis to Genzyme Corporation ("<u>Sanofi</u>") pursuant to that certain Assignment and Assumption Agreement dated as of December 20, 2018. Except as otherwise specifically indicated herein, capitalized terms used but not defined in this letter agreement (this "<u>Letter</u>") shall have the meanings assigned to them in the Agreement.

In consideration of our ongoing Collaboration, and the mutual covenants contained herein, RevMed and Sanofi hereby acknowledge and agree as follows:

1. RMC-4630-03 – Amgen Doublet.

a. In accordance with the terms of the Agreement, the JRDC has reviewed and approved the proposed RMC-4630-03 study, consisting of a combination study of RMC-4630 and Amgen's LumakrasTM (sotorasib), in each case, as set forth in the protocol and the budget separately agreed to by the Parties through the JRDC and JSC, as applicable (such study, "<u>RMC-4630-03</u>" and such budget, the "<u>RMC-4630-03 Budget</u>"), and has added RMC-4630-03 to the Development Plan. For clarity, notwithstanding any prior agreement by the Parties to the contrary, RMC-4630-03 shall be considered a study under the Development Plan (and not a RevMed Study) and part of the Collaboration subject to the terms of the Agreement as modified by this Letter.

b. Notwithstanding anything in the Agreement to the contrary, Sanofi and RevMed shall each be responsible for fifty percent (50%) of the RevMed R&D Costs for RMC-4630-03, for clarity, including but not limited to RevMed's Manufacturing Costs of any Product required for RMC-4630-03 and costs of supply of Lumakras[™] for the US sites in RMC-4630-03 (provided that such RevMed R&D Costs are incurred per the RMC-4630-03 Budget, as the same may be updated by the Parties from time to time in accordance with the Agreement, and do not exceed [***]% of the applicable amounts set forth in the RMC-4630-03 Budget for the particular Calendar Quarter). The RevMed R&D Costs to be shared by the Parties shall initially be borne by RevMed. Promptly following the end of each Calendar Quarter during which RMC-4630-03 is ongoing or RevMed R&D Costs for RMC-4630-03 are incurred, but in no event later than [***] following the end of such Calendar Quarter, RevMed ⁵⁰ Binney Street, Cambridge, MA02142

Tel: 617.252.7500 - Fax: 617.252.7600

Execution Copy Page 1 of 3

will provide to the JRDC a detailed expense report in form approved by the JRDC with respect to such RevMed R&D Costs incurred by or on behalf of RevMed during such Calendar Quarter consistent with this <u>Section 1(b)</u> (including, if requested by Sanofi in writing, copies of receipts or invoices from Third Parties for all RevMed R&D Out-of-Pocket Costs). RevMed shall deliver an invoice to Sanofi for an amount of cash sufficient to reconcile to Sanofi's agreed percentage of such RevMed R&D Costs. Sanofi will reimburse RevMed in Dollars all undisputed amounts within such expense reports under this <u>Section 1(b)</u> within [***] following receipt of the invoice therefor.

c. Sanofi shall have rights to use, at no additional cost, any data arising from RMC-4630-03 ("<u>RMC-4630-03</u> <u>Study Data</u>") in its performance of its obligations and exercise of its rights under the Collaboration except in connection with filing of MAAs for the Indication and Product Treatment Regimen that were the subject of RMC-4630-03. If Sanofi wishes to use or actually uses, RMC-4630-03 Study Data in support of filing a MAA for the Indication and Product Treatment Regimen that were the subject of RMC-4630-03, it shall notify RevMed in writing and shall make a buy-in payment to RevMed in Dollars equal to [***], such payment to be made within [***] after the date that Sanofi receives an invoice from RevMed setting forth such amount.

2. **No Other Modifications**. This Letter constitutes an Ancillary Agreement, as defined in, and pursuant to, the Agreement. This Letter amends the Agreement and shall be deemed to be a part of and incorporated into the Agreement. In the event of a conflict between this Letter and the Agreement, this Letter shall control. Except as expressly set forth in this Letter, all of the terms and conditions of the Agreement shall remain unchanged and are ratified and confirmed in all respects and remain in full force and effect.

3. **Counterparts**. This Letter may be executed in one (1) or more counterparts, each of which will be deemed an original, but all of which together will constitute one and the same instrument.

4. **Governing Law**. This letter agreement shall be governed by and construed in accordance with the laws of the State of New York without reference to any rules of conflict of laws.

[Signature Page Follows]

50 Binney Street, Cambridge, MA02142 Tel: 617.252.7500 - Fax: 617.252.7600

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Please acknowledge RevMed's agreement with this Letter by returning a signed copy of this Letter.

Sincerely,

Genzyme Corporation

By: <u>/s/ Peter Adamson</u> Name: <u>Peter Adamson</u> Title: Global Head, Oncology Development & Pediatric Innovation

Acknowledged and Agreed:

Revolution Medicines, Inc.

By:___/s/ Margaret A. Horn_____ Name: Margaret A. Horn, J.D. Title: Chief Operating Officer

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CERTIFICATION PURSUANT TO RULES 13a-14(a) AND 15d-14(a) UNDER THE SECURITIES EXCHANGE ACT OF 1934, AS ADOPTED PURSUANT TO SECTION 302 OF THE SARBANES-OXLEY ACT OF 2002

I, Mark A. Goldsmith, certify that:

- 1. I have reviewed this Quarterly Report on Form 10-Q of Revolution Medicines, Inc.;
- 2. Based on my knowledge, this report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this report;
- 3. Based on my knowledge, the financial statements, and other financial information included in this report, fairly present in all material respects the financial condition, results of operations and cash flows of the registrant as of, and for, the periods presented in this report;
- 4. The registrant's other certifying officer(s) and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) and internal control over financial reporting (as defined in Exchange Act Rules 13a-15(f) and 15d-15(f)) for the registrant and have:
 - (a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under our supervision, to ensure that material information relating to the registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this report is being prepared;
 - (b) Designed such internal control over financial reporting, or caused such internal control over financial reporting to be designed under our supervision, to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles;
 - (c) Evaluated the effectiveness of the registrant's disclosure controls and procedures and presented in this report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this report based on such evaluation; and
 - (d) Disclosed in this report any change in the registrant's internal control over financial reporting that occurred during the registrant's most recent fiscal quarter (the registrant's fourth fiscal quarter in the case of an annual report) that has materially affected, or is reasonably likely to materially affect, the registrant's internal control over financial reporting; and
- 5. The registrant's other certifying officer(s) and I have disclosed, based on our most recent evaluation of internal control over financial reporting, to the registrant's auditors and the audit committee of the registrant's board of directors (or persons performing the equivalent functions):
 - (a) All significant deficiencies and material weaknesses in the design or operation of internal control over financial reporting which are reasonably likely to adversely affect the registrant's ability to record, process, summarize and report financial information; and
 - (b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the registrant's internal control over financial reporting.

Date: August 11, 2021

Ву:

/s/ Mark A. Goldsmith Mark A. Goldsmith, M.D., Ph.D. President and Chief Executive Officer (Principal Executive Officer)

CERTIFICATION PURSUANT TO RULES 13a-14(a) AND 15d-14(a) UNDER THE SECURITIES EXCHANGE ACT OF 1934, AS ADOPTED PURSUANT TO SECTION 302 OF THE SARBANES-OXLEY ACT OF 2002

I, Jack Anders, certify that:

- 1. I have reviewed this Quarterly Report on Form 10-Q of Revolution Medicines, Inc.;
- 2. Based on my knowledge, this report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this report;
- 3. Based on my knowledge, the financial statements, and other financial information included in this report, fairly present in all material respects the financial condition, results of operations and cash flows of the registrant as of, and for, the periods presented in this report;
- 4. The registrant's other certifying officer(s) and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) and internal control over financial reporting (as defined in Exchange Act Rules 13a-15(f) and 15d-15(f)) for the registrant and have:
 - (a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under our supervision, to ensure that material information relating to the registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this report is being prepared;
 - (b) Designed such internal control over financial reporting, or caused such internal control over financial reporting to be designed under our supervision, to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles;
 - (c) Evaluated the effectiveness of the registrant's disclosure controls and procedures and presented in this report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this report based on such evaluation; and
 - (d) Disclosed in this report any change in the registrant's internal control over financial reporting that occurred during the registrant's most recent fiscal quarter (the registrant's fourth fiscal quarter in the case of an annual report) that has materially affected, or is reasonably likely to materially affect, the registrant's internal control over financial reporting; and
- 5. The registrant's other certifying officer(s) and I have disclosed, based on our most recent evaluation of internal control over financial reporting, to the registrant's auditors and the audit committee of the registrant's board of directors (or persons performing the equivalent functions):
 - (a) All significant deficiencies and material weaknesses in the design or operation of internal control over financial reporting which are reasonably likely to adversely affect the registrant's ability to record, process, summarize and report financial information; and
 - (b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the registrant's internal control over financial reporting.

Date: August 11, 2021

By:

/s/ Jack Anders Jack Anders Senior Vice President, Finance and Principal Accounting Officer (Principal Financial and Accounting Officer)

CERTIFICATION PURSUANT TO 18 U.S.C. SECTION 1350, AS ADOPTED PURSUANT TO SECTION 906 OF THE SARBANES-OXLEY ACT OF 2002

In connection with the Quarterly Report of Revolution Medicines, Inc. (the "Company") on Form 10-Q for the period ending June 30, 2021 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I certify, pursuant to 18 U.S.C. § 1350, as adopted pursuant to § 906 of the Sarbanes-Oxley Act of 2002, that:

- (1) The Report fully complies with the requirements of section 13(a) or 15(d) of the Securities Exchange Act of 1934; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and result of operations of the Company.

Date: August 11, 2021

Ву:

/s/ Mark A. Goldsmith Mark A. Goldsmith, M.D., Ph.D. Chief Executive Officer (Principal Executive Officer)

CERTIFICATION PURSUANT TO 18 U.S.C. SECTION 1350, AS ADOPTED PURSUANT TO SECTION 906 OF THE SARBANES-OXLEY ACT OF 2002

In connection with the Quarterly Report of Revolution Medicines, Inc. (the "Company") on Form 10-Q for the period ending June 30, 2021 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I certify, pursuant to 18 U.S.C. § 1350, as adopted pursuant to § 906 of the Sarbanes-Oxley Act of 2002, that:

- (1) The Report fully complies with the requirements of section 13(a) or 15(d) of the Securities Exchange Act of 1934; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and result of operations of the Company.

Date: August 11, 2021

By:

Jack Anders Senior Vice President, Finance and Principal Accounting Officer (Principal Financial and Accounting Officer)

/s/ Jack Anders