UNITED STATES SECURITIES AND EXCHANGE COMMISSION

WASHINGTON, D.C. 20549

SCHEDULE 13D

Under the Securities Exchange Act of 1934 (Amendment No. 1)*

Spruce Biosciences, Inc.

(Name of Issuer)

Common Stock, par value \$0.0001 per share (Title of Class of Securities)

85209E 109 (CUSIP Number)

Peter Haahr Novo Holdings A/S Tuborg Havnevej 19 Hellerup, Denmark DK-2900 +45 3527 6592

Copy to:

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(Name, Address and Telephone Number of Person Authorized to Receive Notices and Communications)

May 19, 2021 (Date of Event which Requires Filing of this Statement)

If the filing person has previously filed a statement on Schedule 13G to report the acquisition which is subject of this Schedule 13D, and is filing
his statement because of Rule 13d-1(e), 13d-1(f) or 13d-1(g), check the following box. \Box

Note: Schedules filed in paper format shall include a signed original and five copies of the schedule, including all exhibits. See Rule 13d-7(b) for other parties to whom copies are to be sent.

* The remainder of this cover page shall be filled out for a reporting person's initial filing on this form with respect to the subject class of securities, and for any subsequent amendment containing information which would alter disclosures provided in a prior cover page.

The information required on the remainder of this cover page shall not be deemed to be "filed" for the purpose of Section 18 of the Securities Exchange Act of 1934 (the "Act") or otherwise subject to the liabilities of that section of the Act but shall be subject to all other provisions of the Act (however, see the Notes).

CUSII	P No.: 8520	9E 10	9			
1.	1. Name of Reporting Person:					
Novo Holdings A/S						
2. Check the Appropriate Box if a Member of Group (See Instructions):						
(a) \Box (b) \Box						
3.	SEC Use	Only	:			
4.	Source o	f Fun	ds:			
	WC					
5.		Discl	osure of Legal Proceedings is Required Pursuant to <u>Items 2(d)</u> or <u>2(e):</u>			
6.	Citizens	nip or	Place of Organization:			
	Denmark					
	Demina					
		7.	Sole Voting Power:			
	umber of	8.	4,514,336			
	Shares Beneficially		Shared Voting Power:			
O,	Owned By		0			
Each Reporting		9.	Sole Dispositive Power:			
	Person With:		4,514,336			
			Shared Dispositive Power:			
			0			
11.	Aggregat	e Amo	ount Beneficially Owned by Each Reporting Person:			
12	4,514,336					
12.	Check if the Aggregate Amount in Row (11) Excludes Certain Shares:					

(1) Based upon 23,330,135 shares of the Issuer's Common Stock outstanding as of May 11, 2021, as reported in the Issuer's 10-Q filed with the Securities and Exchange Commission ("SEC") on May 12, 2021.

13.

14.

19.4%(1)

CO

Type of Reporting Person:

Percent of Class Represented By Amount In Row (11):

This amendment ("Amendment No. 1"), amends the Schedule 13D originally filed with the SEC on October 16, 2020, (the "Schedule"), to report a decrease in beneficial ownership of Common Stock of the Issuer held by Novo Holdings A/S resulting from the sale of shares by Novo Holdings A/S. All capitalized terms contained herein but not otherwise defined shall have the meaning ascribed to such terms in the Schedule.

Item 2. Identity and Background

Item 2 is amended and replaced in its entirety as follows:

- (a) Novo Holdings A/S is a Danish limited liability company that is wholly owned by Novo Nordisk Foundation (the "Foundation"), a Danish commercial foundation. Novo Holdings A/S is the holding company in the group of Novo companies (currently comprised of Novo Nordisk A/S and Novozymes A/S) and is responsible for managing the Foundation's assets, including its financial assets. Based on the governance structure of Novo Holdings A/S and the Foundation, the Foundation is not deemed to have any beneficial ownership of the securities of the Issuer held by Novo Holdings A/S. Tiba Aynechi, Ph.D. is employed as a senior partner at Novo Ventures (US), Inc. and was designated to the board of directors of the Issuer by Novo Holdings A/S in May 2016. Dr. Aynechi is not deemed to be a beneficial owner of the securities held by Novo Holdings A/S.
 - The name of each director and executive officer of both Novo Holdings A/S and the Foundation is set forth on <u>Schedule I</u> to this Schedule 13D.
- (b) The business address of both Novo Holdings A/S and the Foundation is Tuborg Havnevej 19, 2900 Hellerup, Denmark.
 The residence or business address of each director and executive officer of both Novo Holdings A/S and the Foundation is set forth on Schedule I to this Schedule 13D.
- (c) Novo Holdings A/S, a holding company that is responsible for managing the Foundation's assets, provides seed and venture capital to development stage companies and invests in well-established companies within the life science and biotechnology sector.
 - The Foundation is a Danish self-governing and profit-making foundation, whose objectives are to provide a stable basis for commercial and research activities undertaken by the group of Novo companies and to support scientific, humanitarian and social purposes through grants.
- (d) Within the last five years, neither Novo Holdings A/S, the Foundation, nor any person named in <u>Schedule I</u> has been convicted in any criminal proceedings.
- (e) Within the last five years, neither Novo Holdings A/S, the Foundation, nor any person named in <u>Schedule I</u> was a party to a civil proceeding of a judicial or administrative body of competent jurisdiction and as a result of such proceeding was or is subject to a judgment, decree or final order enjoining future violations of, or prohibiting or mandating activities subject to, federal or state securities laws or finding any violation with respect to such laws.

Item 5. Interest in Securities of the Issuer

Item 5 of the Schedule is amended and replaced in its entirety as follows:

(a) Novo Holdings A/S beneficially owns 4,514,336 shares of Common Stock (the "Novo Shares") representing approximately 19.4% of the Issuer's outstanding shares of Common Stock, based upon 23,330,135 shares of the Issuer's Common Stock outstanding as of May 11, 2021, as reported in the Issuer's 10-Q filed with the SEC on May 12, 2021.

- (b) Novo Holdings A/S is a Danish limited liability company wholly owned by the Novo Nordisk Foundation. Novo Holdings A/S, through its Board of Directors (the "Novo Board"), has the sole power to vote and dispose of the Novo Shares. The Novo Board may exercise voting and dispositive control over the Novo Shares with approval by a majority of the Novo Board. As such, no individual member of the Novo Board is deemed to hold any beneficial ownership or reportable pecuniary interest in the Novo Shares. Except as described in this Amendment No. 1, neither the Foundation nor any person listed on Schedule I has the power to direct the vote as to, or the disposition of, the Novo Shares.
- (c) On May 19, 2021, Novo Holdings A/S sold 475,000 shares at \$13.90 per share in a block sale broker transaction. No other transactions in the Issuer's Common Stock were effected by Novo Holdings A/S nor the Foundation nor any person listed on <u>Schedule I</u> to the Schedule in the Issuer's Common Stock since the date of the most recent Schedule 13D filing.
- (d) Novo Holdings A/S does not know of any other person having the right to receive or the power to direct the receipt of dividends from, or the proceeds from the sale of, the Novo Shares.
 - (e) Not applicable.

SIGNATURE

After reasonable inquiry and to the best of my knowledge and belief, I certify that the information set forth in this statement is true, complete and correct.

Dated: May 21, 2021 Novo Holdings A/S

/s/ Peter Haahr

By: Peter Haahr

ts: Chief Financial Officer

Schedule I

Information regarding each director and executive officer of both Novo Holdings A/S and the Novo Nordisk Foundation is set forth below.

Novo Holdings A/S

Name, Title	Address	Principal Occupation	Citizenship
Lars Rebien Sørensen, Chairman of the Board	Christianholms Tværvej 27, 2930 Klampenborg Denmark	Professional Board Director	Denmark
Steen Riisgaard, Vice Chairman of the Board	Hestetangsvej 155, 3520 Farum, Denmark	Professional Board Director	Denmark
Jean-Luc Butel, Director	235 Arcadia Road unit # 10-3 289843 Singapore	Global Healthcare Advisor, President, K8 Global Pte Ltd.	Singapore
Jeppe Christiansen, Director	c/o Kasper Fonager Christiansen Classensgade 59, 5. th. 2100 Kobenhavn Ø Denmark	Chief Executive Officer, Fondsmaeglerselskabet Maj Invest A/S	Denmark
Francis Michael Cyprian Cuss, Director	111 Rippling Brook Way, Bernardsville, NJ 07924 USA	Former Executive Vice President and Chief Scientific Officer of Bristol-Myers Squibb	United Kingdom
Viviane Monges, Director	Chemin de Craivavers 32, 1012 Lausanne, Switzerland	Professional Board Director	France
Henrik Poulsen, Director	Emiliekildevej 36 2930 Klampenborg Denmark	Professional Board Director and Senior Advisor, A.P. Møller Holding A/S,	Denmark
Poul Carsten Stendevad, Director	3220 Idaho Ave NW Washington, DC 20016 USA	Co-Chief Investment Officer for Sustainability, Bridgewater Associates	Denmark
Kasim Kutay, Chief Executive Officer of Novo Holdings A/S	Bredgade 65, 3.tv. 1260 Copenhagen K. Denmark	Chief Executive Officer of Novo Holdings A/S	United Kingdom
Peter Haahr, Chief Financial Officer of Novo Holdings A/S	Ordrup Have 21 2920 Charlottenlund Denmark	Chief Financial Officer of Novo Holdings A/S	Denmark

Novo Nordisk Foundation							
Name, Title	Address	Principal Occupation	Citizenship				
Lars Rebien Sørensen, Chairman of the Board	Christianholms Tværvej 27 2930 Klampenborg Denmark	Professional Board Director	Denmark				
Marianne Philip, Vice Chairman of the Board	Annasvej 28 2900 Hellerup Denmark	Attorney	Denmark				
Steen Riisgaard, Director	Hestetangsvej 155 3520 Farum Denmark	Professional Board Director	Denmark				
Mads Krogsgaard Thomsen, Chief Executive Officer	Præstevejen 38 3230 Græsted Denmark	Chief Executive Officer, Novo Nordisk Foundation	Denmark				
Anne Marie Kverneland, Director	Nybrovej 216 2800 Kgs. Lyngby Denmark	Laboratory technician, Novo Nordisk A/S	Denmark				
Lars Bo Køppler, Director	Anemonevej 7 3550 Slangerup Denmark	Technician, Novozymes A/S	Denmark				
Lars Henrik Fugger, Director	72 Staunton Road, Headington Great Britain	Professor, John Radcliffe Hospital, University of Oxford, Oxford, Great Britain	Denmark				
Lars Henrik Munch, Director	Galionsvej 46 1437 Copenhagen K Denmark	Professional Board Director	Denmark				
Mads Boritz Grøn, Director	Horsevænget 4 3400 Hillerød Denmark	Senior Lead Auditor	Denmark				
Liselotte Højgaard, Director	Grønningen 21 1270 Copenhagen K Denmark	Professor	Denmark				