United States SECURITIES AND EXCHANGE COMMISSION

WASHINGTON, D.C. 20549

SCHEDULE 13G

Under the Securities Exchange Act of 1934 (Amendment No. 8)*

IONIS PHARMACEUTICALS, INC. (formerly Isis Pharmaceuticals, Inc.)

(Name of Issuer)

Common Stock (Title of Class of Securities)

462222100 (CUSIP Number)

December 31, 2018 (Date of Event Which Requires Filing This Statement)

Check the appropriate box to designate the rule pursuant to which this Schedule is filed:

☐ Rule 13d-1(b)

☐ Rule 13d-1(c)

☐ Rule 13d-1(d)

* 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 the disclosures provided in a prior cover page.

The information required in 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 ("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).

CUSIP No. 462222100

	- 101						
1.	Names of Reporting Persons						
	BB Bio	otec	h AG				
2.	Check th	ne A	ppropriate Box if a Member of a Group (See Instructions)				
	(a) 🗵	(1	b) \square				
3.	SEC Use	e On	ly				
4.	Citizens	hip (or Place of Organization				
	Switze	rlai	nd				
	SWILZE	5.	Sole Voting Power				
		٥.	Sole voting rower				
			0				
	mber of Shares	6.	Shared Voting Power				
	neficially		č				
	vned by		8,741,334				
	Each	7.	Sole Dispositive Power				
Re	porting						
	Person		0				
	with:	8.	Shared Dispositive Power				
			8,741,334				
9.	Aggrega	te A	mount Beneficially Owned by Each Reporting Person				
10	8,741,3		A				
10.	Check if	the	Aggregate Amount in Row (9) Excludes Certain Shares □				
11.	Percent	of C	lass Represented by amount in Row (9)				
11.	1. refeelt of Class Represented by affiount in Row (9)						
	6.4%						
12.							
1	HC. CO						

CUSIP No. **462222100**

CODII	110. 102		vov					
1.	1. Names of Reporting Persons							
	Biotech Target N.V.							
	I.R.S. Identification Nos. of above persons (entities only):							
	N/A							
2.	. Check the Appropriate Box if a Member of a Group (See Instructions) (a) ⊠ (b) □							
3.	. SEC Use Only							
4.	Citizens	hip (or Place of Organization					
	Curaç	ao						
		5.	Sole Voting Power					
Nu	mber of		0					
S	Shares	6.	Shared Voting Power					
	neficially wned by		8,741,334					
	Each	7.						
F	eporting Person		0					
	with:	8.	Shared Dispositive Power					
			8,741,334					
9.	Aggrega	ite A	mount Beneficially Owned by Each Reporting Person					
	8,741,334							
10.	Check i	f the	Aggregate Amount in Row (9) Excludes Certain Shares □					
11.	11. Percent of Class Represented by amount in Row (9)							
	6.4%							
12.	2. Type of Reporting Person (See Instructions)							
	CO							

Item	2					
<u> Targ</u>	2(a) <u>set")</u>	Name of Person Filing: BB Biotech AG ("BB Biotech") on behalf of its wholly-owned subsidiary, Biotech Target N.V. ("Biotech				
	2(b)	Address of Principal Business Office or, if none, Residence:				
		BB Biotech AG: Schwertstrasse 6, CH-8200 Schaffhausen, Switzerland				
		Biotech Target N.V.: Ara Hill Top Building, Unit A-5, Pletterijweg Oost 1, Curaçao				
	2(c)	Citizenship: BB Biotech AG: Switzerland				
		Biotech Target N.V.: Curação				
	2(d)	Title of Class of Securities <u>Common Stock, \$0.001 par value</u>				
	2(e)	CUSIP Number <u>462222100</u>				
Item 4. Ownership						
1.	Provide the following information regarding the aggregate number and percentage of the class of securities of the issuer identified in I					
	(a)	Amount beneficially owned: <u>8,741,334</u>				
	(b)	Percent of class: 6.4%				
	(c)	Number of shares as to which the person has:				
	(i)	Sole power to vote or to direct the vote <u>0</u>				
	(ii)	Shared power to vote or to direct the vote 8,741,334				

Item 10. Certification

(iii) Sole power to dispose or to direct the disposition of <u>0</u>

(iv) Shared power to dispose or to direct the disposition of 8,741,334

By signing below I certify that, to the best of my knowledge and belief, the securities referred to above were not acquired and are not held for the purpose of or with the effect of changing or influencing the control of the issuer of the securities and were not acquired and are not held in connection with or as a participant in any transaction having that purpose or effect.

SIGNATURES

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

BB Biotech AG

Date: February 7, 2019	By:	/s/ Michael Hutter	
		Signatory Authority	
	Name:	Michael Hutter	
	Title:	Signatory Authority	
Date: February 7, 2019	Ву:	/s/ Ivo Betschart	
		Signatory Authority	
	Name:	Ivo Betschart	
	Title:	Signatory Authority	
Biotech Target N.V.			
Date: February 7, 2019	By:	/s/ Michael Hutter	
		Signatory Authority	
	Name:	Michael Hutter	
	Title:	Signatory Authority	
	_		
Date: <u>February 7, 2019</u>	Ву:	/s/ Ivo Betschart	
		Signatory Authority	
	Name:	Ivo Betschart	
	Title:	Signatory Authority	

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Exhibit Index

Exhibit A: Agreement by and between BB Biotech AG and Biotech Target N.V. with respect to the filing of this disclosure statement.*

* Previously filed as an exhibit to BB Biotech AG and Biotech Target N.V.'s Schedule 13G filed with the Securities and Exchange Commission on July 20, 2011.

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