

OMB APPROVAL	
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Check this box if no longer subject to Section 16. Form 4 or Form 5 obligations may continue. See Instruction 1(b).

STATEMENT OF CHANGES IN BENEFICIAL OWNERSHIP

Filed pursuant to Section 16(a) of the Securities Exchange Act of 1934 or Section 30(h) of the Investment Company Act of 1940

1. Name and Address of Reporting Person* <u>ARCH Venture Partners X, LLC</u> (Last) (First) (Middle) 8755 WEST HIGGINS ROAD SUITE 1025 (Street) CHICAGO IL 60631 (City) (State) (Zip)	2. Issuer Name and Ticker or Trading Symbol <u>Sienna Biopharmaceuticals, Inc. [SNNA]</u>	5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director <input checked="" type="checkbox"/> 10% Owner Officer (give title below) Other (specify below)
	3. Date of Earliest Transaction (Month/Day/Year) 02/22/2019	
4. If Amendment, Date of Original Filed (Month/Day/Year)		

Table I - Non-Derivative Securities Acquired, Disposed of, or Beneficially Owned

1. Title of Security (Instr. 3)	2. Transaction Date (Month/Day/Year)	2A. Deemed Execution Date, if any (Month/Day/Year)	3. Transaction Code (Instr. 8)		4. Securities Acquired (A) or Disposed Of (D) (Instr. 3, 4 and 5)			5. Amount of Securities Beneficially Owned Following Reported Transaction(s) (Instr. 3 and 4)	6. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)	7. Nature of Indirect Beneficial Ownership (Instr. 4)
			Code	V	Amount	(A) or (D)	Price			
Common Stock	02/22/2019		P		150,000	A	\$2.5	2,900,575	I	See footnote ⁽¹⁾⁽²⁾
Common Stock	02/22/2019		P		150,000	A	\$2.5	1,032,493	I	See footnote ⁽¹⁾⁽²⁾
Common Stock	02/22/2019		P		1,865,800	A	\$2.5	1,865,800	I	See footnote ⁽¹⁾⁽³⁾

Table II - Derivative Securities Acquired, Disposed of, or Beneficially Owned (e.g., puts, calls, warrants, options, convertible securities)

1. Title of Derivative Security (Instr. 3)	2. Conversion or Exercise Price of Derivative Security	3. Transaction Date (Month/Day/Year)	3A. Deemed Execution Date, if any (Month/Day/Year)	4. Transaction Code (Instr. 8)		5. Number of Derivative Securities Acquired (A) or Disposed of (D) (Instr. 3, 4 and 5)		6. Date Exercisable and Expiration Date (Month/Day/Year)		7. Title and Amount of Securities Underlying Derivative Security (Instr. 3 and 4)	8. Price of Derivative Security (Instr. 5)	9. Number of derivative Securities Beneficially Owned Following Reported Transaction(s) (Instr. 4)	10. Ownership Form: Direct (D) or Indirect (I) (Instr. 4)	11. Nature of Indirect Beneficial Ownership (Instr. 4)
				Code	V	(A)	(D)	Date Exercisable	Expiration Date					

1. Name and Address of Reporting Person* <u>ARCH Venture Partners X, LLC</u> (Last) (First) (Middle) 8755 WEST HIGGINS ROAD SUITE 1025 (Street) CHICAGO IL 60631 (City) (State) (Zip)		
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1. Name and Address of Reporting Person*

[ARCH Venture Fund X Overage, L.P.](#)

(Last) (First) (Middle)

8755 WEST HIGGINS ROAD
SUITE 1025

(Street)

CHICAGO IL 60631

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[ARCH Venture Partners X Overage, L.P.](#)

(Last) (First) (Middle)

8755 WEST HIGGINS ROAD
SUITE 1025

(Street)

CHICAGO IL 60631

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[CRANDELL KEITH](#)

(Last) (First) (Middle)

8755 WEST HIGGINS ROAD
SUITE 1025

(Street)

CHICAGO IL 60631

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[NELSEN ROBERT](#)

(Last) (First) (Middle)

8755 WEST HIGGINS ROAD
SUITE 1025

(Street)

CHICAGO IL 60631

(City) (State) (Zip)

1. Name and Address of Reporting Person*

[GILLIS STEVEN](#)

(Last) (First) (Middle)

8755 WEST HIGGINS ROAD
SUITE 1025

(Street)

CHICAGO IL 60631

(City) (State) (Zip)

Explanation of Responses:

1. Following the transactions reported herein, includes 1,032,493 shares of common stock held of record by ARCH Venture Fund VIII Overage, L.P. ("ARCH Fund Overage"), 2,900,575 shares of common stock held of record by ARCH Venture Fund VIII, L.P. ("ARCH Fund VIII"), and 1,865,800 shares of common stock held of record by ARCH Venture Fund X Overage, L.P. ("ARCH X Overage").

2. The sole general partner of ARCH Fund VIII is ARCH Venture Partners VIII, L.P. ("ARCH Partners VIII"). The sole general partner of ARCH Partners VIII and ARCH Fund Overage is ARCH Venture Partners VIII, LLC ("ARCH VIII LLC"). ARCH Partners VIII may therefore be deemed to beneficially own the securities held by ARCH Fund VIII and ARCH VIII LLC may be deemed to beneficially own the securities held by ARCH Fund VIII and ARCH Fund Overage. ARCH Partners VIII and ARCH VIII LLC disclaim beneficial ownership of such securities, except to the extent of any pecuniary interest therein. The managing directors of ARCH VIII LLC are Keith L. Crandell, Clinton Bybee and Robert Nelsen, and they may be deemed to beneficially own the shares held by ARCH Fund VIII and ARCH Fund Overage. Messrs. Crandell, Bybee and Nelsen disclaim beneficial ownership of such shares, except to the extent of any pecuniary interest therein.

3. The sole general Partner of ARCH X Overage is ARCH Venture Partners X Overage, L.P. ("AVP X Over GP"). The sole general partner of AVP X Over GP is ARCH Venture Partners X, LLC ("AVP X LLC"). AVP X Over LP and AVP X LLC may therefore be deemed to beneficially own the securities held by ARCH X Overage. AVP X Over GP and AVP X LLC disclaim beneficial ownership of such securities, except to the extent of any pecuniary interest therein. Keith L. Crandell, Robert Nelsen and Steven Gillis are managing directors of AVP X LLC, and they may be deemed to beneficially own the shares held by ARCH X Overage. Messrs. Crandell, Nelsen and Gillis disclaim beneficial ownership of such shares, except to the extent of any pecuniary interest therein.

Remarks:

The Reporting Persons are jointly filing this Form 4 pursuant to Rule 16a-3(j) under the Exchange Act. Because no more than 10 reporting persons can file any one Form 4 through the Securities and Exchange Commission's EDGAR system, ARCH Venture Fund VIII Overage, L.P., ARCH Venture Fund VIII, L.P., ARCH Venture Partners VIII, L.P., ARCH Venture Partners VIII, LLC and certain other affiliates of the Reporting Persons have filed a separate Form 4, in which the direct and indirect transactions of ARCH Venture Fund X Overage, L.P., ARCH Venture Partners X Overage, L.P., ARCH Venture Partners X Overage, LLC are also reported, in addition to being reported on this Form 4.

<p><u>/s/ Mark McDonnell,</u> <u>Attorney-in-Fact for Keith</u> <u>Crandell, Managing Director</u> <u>of ARCH Venture Partners X,</u> <u>LLC, General Partner of</u> <u>ARCH Venture Partners X</u> <u>Overage, L.P., General Partner</u> <u>of ARCH Venture Fund X</u> <u>Overage, L.P.</u></p>	<p><u>02/26/2019</u></p>
<p><u>/s/ Mark McDonnell,</u> <u>Attorney-in-Fact for Keith</u> <u>Crandell, Managing Director</u> <u>of ARCH Venture Partners X,</u> <u>LLC, General Partner of</u> <u>ARCH Venture Fund X</u> <u>Overage, L.P.</u></p>	<p><u>02/26/2019</u></p>
<p><u>/s/ Mark McDonnell,</u> <u>Attorney-in-Fact for Keith</u> <u>Crandell, Managing Director</u> <u>of ARCH Venture Partners</u> <u>VIII, LLC, General Partner of</u> <u>ARCH Venture Partners VIII,</u> <u>L.P.</u></p>	<p><u>02/26/2019</u></p>
<p><u>/s/ Mark McDonnell,</u> <u>Attorney-in-Fact for Keith</u> <u>Crandell</u></p>	<p><u>02/26/2019</u></p>
<p><u>/s/ Mark McDonnell,</u> <u>Attorney-in-Fact for Robert</u> <u>Nelsen</u></p>	<p><u>02/26/2019</u></p>
<p><u>/s/ Mark McDonnell,</u> <u>Attorney-in-Fact for Steve</u> <u>Gillis</u></p>	<p><u>02/26/2019</u></p>
<p>** Signature of Reporting Person</p>	<p>Date</p>

Reminder: Report on a separate line for each class of securities beneficially owned directly or indirectly.

* If the form is filed by more than one reporting person, see Instruction 4 (b)(v).

** Intentional misstatements or omissions of facts constitute Federal Criminal Violations See 18 U.S.C. 1001 and 15 U.S.C. 78ff(a).

Note: File three copies of this Form, one of which must be manually signed. If space is insufficient, see Instruction 6 for procedure.

Persons who respond to the collection of information contained in this form are not required to respond unless the form displays a currently valid OMB Number.

POWERS OF ATTORNEY

KNOW ALL MEN BY THESE PRESENTS, that each person whose signature appears below hereby constitutes and appoints Mark McDonnell his or its true and lawful attorney-in-fact, with full power of substitution, to sign any and all instruments, certificates and documents that may be necessary, desirable or appropriate to be executed on behalf of himself as an individual or in his capacity as a direct or indirect general partner, member, director, officer or manager of any partnership, corporation or limited liability company, pursuant to section 13 or 16 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), and any and all regulations promulgated thereunder, and to file the same, with all exhibits thereto, and any other documents in connection therewith, with the Securities and Exchange Commission, and with any other entity when and if such is mandated by the Exchange Act or by the Financial Industry Regulatory Authority, granting unto said attorney-in-fact full power and authority to do and perform each and every act and thing necessary, desirable or appropriate, fully to all intents and purposes as he might or could do in person, thereby ratifying and confirming all that said attorney-in-fact, or his substitutes, may lawfully do or cause to be done by virtue hereof. This Power of Attorney shall remain in full force and effect with respect to each undersigned person unless and until six months after such person is both no longer a Managing Director of ARCH Venture Partners and no longer serving on the board of directors of any portfolio company of any ARCH Venture Partners fund.

IN WITNESS WHEREOF, this Power of Attorney has been signed as of the 21th day of February, 2019.

ARCH VENTURE FUND X, L.P.

By: ARCH Venture Partners X, L.P.

its General Partner

By: ARCH Venture Partners X, LLC

its General Partner

By: /s/ Keith Crandell

Managing Director

ARCH VENTURE PARTNERS X, L.P.

By: ARCH Venture Partners X, LLC

its General Partner

By: /s/ Keith Crandell

Managing Director

ARCH VENTURE FUND X OVERAGE, L.P.

By: ARCH Venture Partners X Overage, L.P.

its General Partner

By: ARCH Venture Partners X, LLC

its General Partner

By: /s/ Keith Crandell

Managing Director

ARCH VENTURE PARTNERS X OVERAGE, L.P.

By: ARCH Venture Partners X, LLC

its General Partner

By: /s/ Keith Crandell

Managing Director

ARCH VENTURE PARTNERS X, LLC

By: /s/Keith Crandell

Managing Director

/s/ Keith Crandell

Keith Crandell

/s/ Kristina Burow

Kristina Burow

/s/ Robert Nelsen

Robert Nelsen

/s/ Steven Gillis

