

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

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

Check this box to indicate that a
transaction was made pursuant to a
contract, instruction or written plan
for the purchase or sale of equity
securities of the issuer that is
intended to satisfy the affirmative
defense conditions of Rule 10b5-
1(c). See Instruction 10.

1. Name and Address of Reporting Person* <u>Third Rock Ventures V, L.P.</u> (Last) (First) (Middle) <u>C/O THIRD ROCK VENTURES, LLC</u> <u>201 BROOKLINE AVE, SUITE 1401</u> (Street) <u>BOSTON</u> <u>MA</u> <u>02215</u> (City) (State) (Zip)	2. Issuer Name and Ticker or Trading Symbol <u>Septerna, Inc. [SEPN]</u> Foreign Trading Symbol 3. Date of Earliest Transaction (Month/Day/Year) <u>08/12/2026</u> 4. If Amendment, Date of Original Filed (Month/Day/Year)	5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director ☒ 10% Owner Officer (give title below) Other (specify below) Form filed by One Reporting Person ☒ Form filed by More than One Reporting Person
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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	08/12/2026		S ⁽¹⁾		147,059	A	\$50.27 ⁽²⁾	6,068,532	D ⁽³⁾	
Common Stock	08/12/2026		S ⁽¹⁾		141	A	\$51.08 ⁽⁴⁾	6,068,391	D ⁽³⁾	
Common Stock								4,211,659	I	By Third Rock Ventures VI, L.P. ⁽⁵⁾

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)
						Date Exercisable	Expiration Date	Title	Amount or Number of Shares			

1. Name and Address of Reporting Person* <u>Third Rock Ventures V, L.P.</u> (Last) (First) (Middle) <u>C/O THIRD ROCK VENTURES, LLC</u> <u>201 BROOKLINE AVE, SUITE 1401</u> (Street) <u>BOSTON</u> <u>MA</u> <u>02215</u> (City) (State) (Zip)	1. Name and Address of Reporting Person* <u>Third Rock Ventures GP V, LP</u> (Last) (First) (Middle) <u>201 BROOKLINE AVE, SUITE 1401</u>
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(Street)	BOSTON	MA	02215
(City)	(State)	(Zip)	
1. Name and Address of Reporting Person*			
TRV GP V, LLC			
(Last)	(First)	(Middle)	
201 BROOKLINE AVE, SUITE 1401			
(Street)	BOSTON	MA	02215
(City)	(State)	(Zip)	
1. Name and Address of Reporting Person*			
Third Rock Ventures VI, L.P.			
(Last)	(First)	(Middle)	
C/O THIRD ROCK VENTURES, LLC 201 BROOKLINE AVE, SUITE 1401			
(Street)	BOSTON	MA	02215
(City)	(State)	(Zip)	
1. Name and Address of Reporting Person*			
Third Rock Ventures GP VI, L.P.			
(Last)	(First)	(Middle)	
201 BROOKLINE AVE, SUITE 1401			
(Street)	BOSTON	MA	02215
(City)	(State)	(Zip)	
1. Name and Address of Reporting Person*			
TRV GP VI, LLC			
(Last)	(First)	(Middle)	
201 BROOKLINE AVE, SUITE 1401			
(Street)	BOSTON	MA	02215
(City)	(State)	(Zip)	

Explanation of Responses:

1. The reported transactions were effected pursuant to a Rule 10b5-1 trading plan dated March 26, 2026, previously adopted by the Reporting Person.
2. The price reported is a weighted average price. These shares were sold in multiple transactions at prices ranging from \$50.05 to \$50.81, inclusive. The reporting person undertakes to provide to the Issuer, any security holder of the Issuer, or the staff of the Securities and Exchange Commission, upon request, full information regarding the number of shares sold at each separate price within the ranges set forth in this footnote (2).
3. These shares are directly held by Third Rock Ventures V, L.P. ("TRV V"). The general partner of TRV V is Third Rock Ventures GP V, L.P. ("TRV GP V"). The general partner of TRV GP V is TRV GP V, LLC ("TRV GP V LLC"). Each of TRV GP V and TRV GP V LLC disclaims beneficial ownership of the shares except to the extent of its pecuniary interest therein, if any, and this report shall not be deemed an admission that it is the beneficial owner of such shares.
4. The price reported is a weighted average price. These shares were sold in multiple transactions at prices ranging from \$51.00 to \$51.10, inclusive. The reporting person undertakes to provide to the Issuer, any security holder of the Issuer, or the staff of the Securities and Exchange Commission, upon request, full information regarding the number of shares sold at each separate price within the ranges set forth in this footnote (4).
5. The shares are held directly by Third Rock Ventures VI, L.P. ("TRV VI"). The general partner of TRV VI is Third Rock Ventures GP VI, L.P. ("TRV GP VI"). The general partner of TRV GP VI is TRV GP VI, LLC ("TRV GP VI LLC"). Each of TRV GP VI and TRV GP VI LLC, disclaims beneficial ownership of these shares except to the extent of its pecuniary interest therein, if any, and this report shall not be deemed an admission that it is the beneficial owner of such shares. Each of Reporting Persons disclaims the existence of a Section 13(d) "group" as between any TRV V related parties and any TRV VI related parties and this report shall not be deemed an admission that any of such parties is or may be part of such a group with any of the other parties.

[/s/ Kevin Gillis, Chief
Operating Officer of TRV GP
V, LLC, general partner of
Third Rock Ventures GP V,
L.P., general partner of Third
Rock Ventures V, L.P.](#) 08/14/2026

[/s/ Kevin Gillis, Chief
Operating Officer of TRV GP
V, LLC, general partner of
Third Rock Ventures GP V,
L.P.](#) 08/14/2026

/s/ Kevin Gillis, Chief
Operating Officer of TRV GP 08/14/2026
V, LLC

/s/ Kevin Gillis, Chief
Operating Officer of TRV GP
VI, LLC, general partner of
Third Rock Ventures GP VI, 08/14/2026
L.P., general partner of Third
Rock Ventures VI, L.P.

/s/ Kevin Gillis, Chief
Operating Officer of TRV GP
VI, LLC, general partner of 08/14/2026
Third Rock Ventures GP VI,
L.P.

/s/ Kevin Gillis, Chief
Operating Officer of TRV GP 08/14/2026
VI, LLC

** Signature of Reporting Person Date

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.