

OMB APPROVAL	
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**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>Klein Uwe</u> <hr/> (Last) (First) (Middle) <u>C/O SEPTERNA, INC.</u> <u>250 EAST GRAND AVENUE</u> <hr/> (Street) <u>SOUTH SAN FRANCISCO CA 94080</u> <hr/> (City) (State) (Zip)	2. Issuer Name and Ticker or Trading Symbol <u>Septerna, Inc. [ SEPN ]</u>	5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director 10% Owner <input checked="" type="checkbox"/> Officer (give title below) Other (specify below) <u>SVP, Biological Sciences</u>
	3. Date of Earliest Transaction (Month/Day/Year) <u>02/06/2026</u>	
	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			

**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						Title
Stock Option (Right to Buy)	\$25.41	02/06/2026		A		80,000		(1)	02/05/2036	Common Stock	80,000	\$0	80,000	D	

**Explanation of Responses:**

1. 2/48th of the shares subject to such option shall vest and become exercisable on March 1, 2026 and the remaining shares subject to the option shall vest in 46 substantially equal monthly installments on each monthly anniversary thereafter, subject to the Reporting Person's continuous service to the Issuer on each such date.

**Remarks:**

Exhibit 24.2 - Substitute Power of Attorney

/s/ Mark A. Wilson, Attorney-in-Fact 02/10/2026

\*\* 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.

## SUBSTITUTE POWER OF ATTORNEY

Under the terms of powers of attorney (each, a "Power of Attorney") previously filed with the U.S. Securities and Exchange Commission, the undersigned was appointed an attorney-in-fact for the below listed individuals to, among other things, execute for and on behalf of the following individuals any Form 3, Form 4, Form 5, Schedule 13D, or Schedule 13G, or any amendment(s) thereto, in accordance with Section 13 or Section 16 of the Securities Exchange Act of 1934, as amended, and the rules and regulations promulgated thereunder:

Abraham Bassan  
Liz Bhatt  
Bernard Coulie  
Alan Ezekowitz  
Jeffrey Finer  
Jae B. Kim  
Uwe Klein  
Daniel D. Long  
Shalini Sharp  
Jake Simson  
Samira Shaikhly  
Jeffrey K. Tong

In accordance with the authority granted under each Power of Attorney, including the full power of substitution, the undersigned hereby appoints each of Gil M. Labrucherie and Mark A. Wilson, as substitute attorneys-in-fact, on behalf of the individuals listed above, with the power to exercise and execute all of the powers granted or conferred in the original Power of Attorney. For the avoidance of doubt, the foregoing appointment shall not serve as a revocation of the powers granted to the undersigned herself in each Power of Attorney.

This substitute Power of Attorney shall remain in full force and effect with respect to each Power of Attorney until such underlying Power of Attorney is revoked or terminated, unless earlier revoked by the undersigned in a signed writing.

Date: February 10, 2026

By: /s/ Ran Xiao  
Name: Ran Xiao  
Title: Attorney-in-Fact

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