

FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

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).

☒ 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.

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>King-Jones Heidy</u> (Last) (First) (Middle) <u>221 CRESCENT STREET, BUILDING 23, SUITE 105</u> (Street) <u>WALTHAM MA 02453</u> (City) (State) (Zip)	2. Issuer Name and Ticker or Trading Symbol <u>Spyre Therapeutics, Inc. [SYRE]</u> Foreign Trading Symbol	5. Relationship of Reporting Person(s) to Issuer (Check all applicable) Director 10% Owner ☒ Officer (give title below) Other (specify below) <u>See Remarks</u>
	3. Date of Earliest Transaction (Month/Day/Year) <u>09/01/2026</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			
Common Stock	09/01/2026		M ⁽¹⁾		27,999	A	\$14.5	30,844 ⁽²⁾	D	
Common Stock	09/01/2026		S ⁽¹⁾		4,845	D	\$86.4 ⁽³⁾	25,999 ⁽²⁾	D	
Common Stock	09/01/2026		S ⁽¹⁾		16,548	D	\$87.2 ⁽⁴⁾	9,451 ⁽²⁾	D	
Common Stock	09/01/2026		S ⁽¹⁾		6,206	D	\$88.17 ⁽⁵⁾	3,245 ⁽²⁾	D	
Common Stock	09/01/2026		S ⁽¹⁾		400	D	\$89.38	2,845 ⁽²⁾	D	
Common Stock	09/02/2026		M ⁽¹⁾		400	A	\$14.5	3,245 ⁽²⁾	D	
Common Stock	09/02/2026		S ⁽¹⁾		400	D	\$90	2,845 ⁽²⁾	D	

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		Date Exercisable	Expiration Date	Title	Amount or Number of Shares				
Stock Option (Right to Buy)	\$14.5	09/01/2026		M ⁽¹⁾		27,999	(6)	09/01/2033	Common Stock	27,999	\$0	511,811	D	
Stock Option (Right to Buy)	\$14.5	09/02/2026		M ⁽¹⁾		400	(6)	09/01/2033	Common Stock	400	\$0	511,411	D	

Explanation of Responses:

1. This transaction was executed pursuant to a Rule 10b5-1 trading plan adopted on April 16, 2026.
2. Includes 489 shares of common stock acquired by the Reporting Person on August 15, 2024, 495 shares of common stock acquired by the Reporting Person on February 15, 2025, 850 shares of common stock acquired by the Reporting Person on August 15, 2025, 325 shares of common stock acquired by the Reporting Person on December 31, 2025, and 686 shares of common stock acquired by the Reporting Person on June 30, 2026 pursuant to the Issuer's employee stock purchase plan.
3. The price reported above is a weighted average price. The shares were sold in multiple transactions at prices ranging from \$85.71 to \$86.70, 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 (the "SEC"), upon request, full information regarding the number of shares sold at each separate price within the range.
4. The price reported above is a weighted average price. The shares were sold in multiple transactions at prices ranging from \$86.71 to \$87.70, inclusive. The Reporting Person undertakes to provide to the Issuer, any security holder of the Issuer, or the staff of the SEC, upon request, full information regarding the number of shares sold at each separate price within the range.
5. The price reported above is a weighted average price. The shares were sold in multiple transactions at prices ranging from \$87.73 to \$88.69, inclusive. The Reporting Person undertakes to provide to the Issuer, any security holder of the Issuer, or the staff of the SEC, upon request, full information regarding the number of shares sold at each separate price within the range.
6. This option represents the right to purchase 539,810 shares of the Issuer's common stock (which have been adjusted to reflect the Issuer's 1-for-25 reverse stock split on September 8, 2023), one quarter of which vested and became exercisable on September 1, 2024, with the remaining three quarters vesting in monthly installments over the following three years, subject to the Reporting Person's continued employment with the Issuer.

Remarks:

/s/ Heidy King-Jones

09/03/2026

** 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.