

☐ 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

OMB APPROVAL	
OMB Number:	3235-0287
Estimated average burden hours per response:	0.5

1. Name and Address of Reporting Person* <u>Burrows Scott L</u> (Last) (First) (Middle) <u>221 CRESCENT STREET, BUILDING 23,</u> <u>SUITE 105</u> (Street) <u>WALTHAM</u> <u>MA</u> <u>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>Chief Financial Officer</u>
	3. Date of Earliest Transaction (Month/Day/Year) <u>09/01/2026</u>	
	4. If Amendment, Date of Original Filed (Month/Day/Year)	6. Individual or Joint/Group Filing (Check Applicable Line) ☒ Form filed by One Reporting Person Form filed by More than One Reporting Person

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⁾		7,500	A	\$14.5	105,494	D	
Common Stock	09/01/2026		S ⁽¹⁾		900	D	\$86.25 ⁽²⁾	104,594	D	
Common Stock	09/01/2026		S ⁽¹⁾		4,384	D	\$87.11 ⁽³⁾	100,210	D	
Common Stock	09/01/2026		S ⁽¹⁾		2,016	D	\$88.03 ⁽⁴⁾	98,194	D	
Common Stock	09/01/2026		S ⁽¹⁾		200	D	\$89.04 ⁽⁵⁾	97,994	D	
Common Stock	09/02/2026		S		18,232 ⁽⁶⁾	D	\$89.35	79,762 ⁽⁷⁾	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					
Stock Option (Right to Buy)	\$14.5	09/01/2026		M ⁽¹⁾		7,500	(8)	09/01/2033	Common Stock	7,500	\$0	337,357	D

Explanation of Responses:

1. This transaction was executed pursuant to a Rule 10b5-1 trading plan adopted on November 10, 2025.
2. The price reported above is a weighted average price. The shares were sold in multiple transactions at prices ranging from \$85.60 to \$86.53, 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.
3. The price reported above is a weighted average price. The shares were sold in multiple transactions at prices ranging from \$86.60 to \$87.57, 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.
4. The price reported above is a weighted average price. The shares were sold in multiple transactions at prices ranging from \$87.64 to \$88.59, 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 \$88.69 to \$89.38, 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. The transaction reported reflects the sale of shares of the Issuer's common stock in satisfaction of the Reporting Person's tax liability in connection with the settlement of 33,738 restricted stock units ("RSUs"). This sale was made to satisfy tax withholding obligations through a "sell to cover" transaction and does not represent a discretionary trade made by the Reporting Person.
7. Includes 33,738 RSUs. Each RSU represents a contingent right to receive, upon vesting, one share of the Issuer's common stock. The RSUs vest on September 1, 2027, subject to the Reporting Person's continued employment with the Issuer.
8. This option represents a right to purchase 404,857 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, as
Attorney-in-Fact

09/03/2026

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.