

FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

OMB APPROVAL

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| hours per response: | 0.5 |

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 * | 2. Issuer Name and Ticker or Trading Symbol | 5. Relationship of Reporting Person(s) to Issuer (Check all applicable) |
| <u>Primus Guy Warren</u> | <u>LIXTE BIOTECHNOLOGY HOLDINGS, INC.</u> | <input checked="" type="checkbox"/> Director 10% Owner |
| (Last) (First) (Middle) | [LIXT] | Officer (give title below) Other (specify below) |
| <u>680 E. COLORADO BOULEVARD,</u> | 3. Date of Earliest Transaction (Month/Day/Year) | |
| <u>SUITE 180</u> | <u>12/24/2025</u> | |
| (Street) | 4. If Amendment, Date of Original Filed (Month/Day/Year) | 6. Individual or Joint/Group Filing (Check Applicable Line) |
| <u>PASADENA CA 91101</u> | | <input checked="" type="checkbox"/> Form filed by One Reporting Person |
| (City) (State) (Zip) | | 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 | | | |

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 Amount or Number of Shares | | | | |
| Stock Option (right to buy) | \$4.05 | 12/24/2025 | | A | 25,000 | 12/24/2025 ⁽¹⁾ 12/24/2030 | Common Stock 25,000 | \$0 | 25,000 | D | |

Explanation of Responses:

1. Fifty percent (50%) of the option vested and became exercisable on the grant date. The remaining fifty percent (50%) vests in equal installments of twelve and one-half percent (12.5%) on March 31, 2026 and on the last date of each subsequent calendar quarter thereafter until fully vested, subject to the reporting person's continuous service through each applicable vesting date.

/s/ Guy Primus 12/29/2025
** 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.