FORM 4

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

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| OMB Number: | 3235-0287 |
|--------------------------|-----------|
| Estimated average burden | |
| l | 0.5 |

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

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

| 10b5-1(c). See In | struction 10. | | | |
|--|---|-------|--|---|
| Name and Address of Reporting Person* Grayson Paul A. | | | 2. Issuer Name and Ticker or Trading Symbol Skye Bioscience, Inc. [SKYE] | Relationship of Reporting Person(s) to Issuer (Check all applicable) Provided the second of the second o |
| (Last) | , | | 3. Date of Earliest Transaction (Month/Day/Year) 08/25/2023 | X Director 10% Owner Officer (give title Other (specify below) |
| C/O SKYE BIOSCIENCE, INC. 1250 EL CAMINO REAL SUITE 100 | | | If Amendment, Date of Original Filed (Month/Day/Year) | 6. Individual or Joint/Group Filing (Check Applicable Line) X Form filed by One Reporting Person |
| (Street) SAN DIEGO | CA | 92130 | | Form filed by More than One Reporting Person |
| (City) | (State) | (Zip) | | |

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) | Transaction Code (Instr. | | 4. Securities Acquired (A) or Disposed Of (D) (Instr. 3, 4 and 5) | | | 5. Amount of Securities Beneficially Owned Following Reported | Form: Direct (D) or Indirect (I) (Instr. 4) | Beneficial Ownership |
|---------------------------------|--|---|--------------------------|---|--|---|------------------------------------|--|---|-------------------------|
| | | | Code | v | Amount (A) or (D) Price | | Transaction(s) (Instr. 3 and 4) | | (Instr. 4) | |
| Common Stock | 08/25/2023 | | A | | 18,132,596(1) | A | (2) | 18,132,596 | 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 | 3A. Deemed Execution Date, if any (Month/Day/Year) | 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 | Ownership Form: Direct (D) | 11. Nature of Indirect Beneficial Ownership (Instr. 4) |
|--|---|---|--------------------|---|--|-----|--|--------------------|--|-------------------------------------|---|--|----------------------------------|--|
| | | | Code | v | (A) | (D) | Date Exercisable | Expiration Date | Title | Amount or Number of Shares | | Transaction(s) (Instr. 4) | | |

Explanation of Responses:

1. Represents a restricted stock unit ("RSU") award that vests on the following performance milestones: 25% vests upon achieving a market cap of \$125M, an additional 25% vests upon achieving a market cap of \$400M, an additional 25% vest at an exit of \$500M or greater; provided, however, that no RSUs shall vest until the compensation committee of the Issuer determines that shares can be sold into the market to cover withholding tax obligations associated with the vesting of the RSUs. Notwithstanding anything to the foregoing, all of the RSUS shall vest if an exit greater than \$500M is achieved at an earlier date.

2. Each RSU represents a contingent right to receive one share of the Issuer's common stock.

Remarks:

/s/ Kaitlyn Arsenault, as Attorneyin-Fact 08/29/2023

** 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.