CUSIP No: 293602207

☐ Rule 13d-1(b)

SECURITIES AND EXCHANGE COMMISSION Washington, D.C. 20549

SCHEDULE 13G

(Rule 13d-102)

INFORMATION TO BE INCLUDED IN STATEMENTS FILED PURSUANT TO \S 240.13d-1(b), (c) AND (d) AND AMENDMENTS THERETO FILED PURSUANT TO \S 240.13d-2.

(Amendment No. ____)*

Ensysce Biosciences, Inc.

(Name of Issuer)

Common Stock, \$0.0001 par value per share (Title of Class of Securities)

293602207 (CUSIP Number)

December 7, 2022 (Date of Event Which Requires Filing of this statement)

Check the appropriate box to designate the rule pursuant to which this Schedule is filed:

| | Rule 13d-1(d) | |
|----|-----------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------|
| | | |
| *Т | The remainder of this cover page shall be filled out for a reporting person's initial filing on this form with respect to the subject class of securities | and for any subsequer |

*The remainder of this cover page shall be filled out for a reporting person's initial filing on this form with respect to the subject class of securities, and for any subsequent amendment containing information which would alter the disclosures provided in a prior cover page.

| | | | the remainder of this cover page shall not be deemed to be "filed" for purposes of Section 18 of the Securities Exchange s of that section of the Act but shall be subject to all other provisions of the Act (however, see the Notes). | Act of 1934 ("Act") or | | |
|--------------------------|-----------------------|--------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------|--|--|
| | | | | | | |
| CUSIP No: 2936 | 02207 | | | | | |
| (1) | NAMES | OF R | EPORTING PERSONS | | | |
| | CVI Investments, Inc. | | | | | |
| (2) | CHECK | THE | APPROPRIATE BOX IF A MEMBER OF A GROUP (SEE INSTRUCTIONS) | | | |
| | | | | (a) □ (b) □ | | |
| (3) | SEC US | E ONI | .Y | | | |
| (4) | CITIZEN | NSHIF | OR PLACE OF ORGANIZATION | | | |
| | Cayman | Island | s | | | |
| NUMBER OF SHARES | | (5) | SOLE VOTING POWER | | | |
| BENEFICIALLY OWNED BY | <i>I</i> | | 0 | | | |
| EACH REPORTING | | (6) | SHARED VOTING POWER ** | | | |
| PERSON WITH | | | 500,000 | | | |
| | | (7) | SOLE DISPOSITIVE POWER | | | |
| | | | 0 | | | |

| | | (8) | SHARED DISPOSITIVE POWER ** | |
|-------------------------------|----------|--------|----------------------------------------------------------------------------------------------------------------------------------|----------------|
| | | | 500,000 | |
| (9) | AGGRE | EGATE | E AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON | |
| | 500,000 | | | |
| (10) | CHECK | BOX | IF THE AGGREGATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHARES (SEE INSTRUCTIONS) | |
| | | | | |
| (11) | PERCE | NT OF | CLASS REPRESENTED BY AMOUNT IN ROW (9) | |
| | 7.8% | | | |
| (12) | TYPE O | F REF | PORTING PERSON (SEE INSTRUCTIONS) | |
| | СО | | | |
| ** Heights Capit | al Manag | gement | Inc. is the investment manager to CVI Investments, Inc. and as such may exercise voting and dispositive power over these shares. | |
| | | | | |
| CUSIP No: 2936 | 02207 | | | |
| (1) | NAMES | S OF R | EPORTING PERSONS | |
| | Heights | Capita | l Management, Inc. | |
| (2) | CHECK | THE | APPROPRIATE BOX IF A MEMBER OF A GROUP (SEE INSTRUCTIONS) | |
| | | | | (a) □ (b) □ |
| (3) | SEC US | E ONI | .Y | (0) |
| (4) | CITIZE | NSHIP | OR PLACE OF ORGANIZATION | |
| | Delawar | re | | |
| NUMBER OF | | (5) | SOLE VOTING POWER | |
| SHARES BENEFICIALLY | Y | | 0 | |
| OWNED BY EACH REPORTING | | (6) | SHARED VOTING POWER ** | |
| PERSON WITH | | | 500,000 | |
| | | (7) | SOLE DISPOSITIVE POWER | |
| | | | 0 | |
| | | (8) | SHARED DISPOSITIVE POWER ** | |
| | | | 500,000 | |
| (9) | AGGRE | EGATE | E AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON | |
| | 500,000 | | | |
| (10) | CHECK | BOX | IF THE AGGREGATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHARES (SEE INSTRUCTIONS) | |
| | | | | |
| (11) | | NT OF | CLASS REPRESENTED BY AMOUNT IN ROW (9) | |
| | 7.8% | | | |
| (12) | TYPE O | F REF | PORTING PERSON (SEE INSTRUCTIONS) | |
| | CO | | | |
| ** Heights Capit | al Manag | gement | Inc. is the investment manager to CVI Investments, Inc. and as such may exercise voting and dispositive power over these shares. | |

| CUSIP No: 293602207 | | | | | | | |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--|--|--|--|--|--|--|
| Item 1. | | | | | | | |
| (a) Name of Issuer | | | | | | | |
| ysce Biosciences, Inc. (the "Company") | | | | | | | |
| (b) Address of Issuer's Principal Executive Offices | | | | | | | |
| 7946 Ivanhoe Avenue, Suite 201, La Jolla, CA 92037 | | | | | | | |
| Item 2(a). Name of Person Filing | | | | | | | |
| This statement is filed by the entities listed below, who are collectively referred to herein as "Reporting Persons," with respect to the shares of common stock of the Company, \$0.0001 par value per share (the "Shares"). | | | | | | | |
| (i) CVI Investments, Inc. | | | | | | | |
| (ii) Heights Capital Management, Inc. | | | | | | | |
| Item 2(b). Address of Principal Business Office or, if none, Residence | | | | | | | |
| The address of the principal business office of CVI Investments, Inc. is: | | | | | | | |
| P.O. Box 309GT Ugland House South Church Street George Town Grand Cayman KY1-1104 Cayman Islands | | | | | | | |
| The address of the principal business office of Heights Capital Management, Inc. is: | | | | | | | |
| 101 California Street, Suite 3250 San Francisco, California 94111 | | | | | | | |
| Item 2(c). Citizenship | | | | | | | |
| Citizenship is set forth in Row 4 of the cover page for each Reporting Person hereto and is incorporated herein by reference for each such Reporting Person. | | | | | | | |
| Item 2(d) Title of Class of Securities | | | | | | | |
| Common stock, \$0.0001 par value per share | | | | | | | |
| Item 2(e) CUSIP Number | | | | | | | |
| 293602207 | | | | | | | |
| Item 3. If this statement is filed pursuant to §§240.13d-1(b) or 240.13d-2(b) or (c), check whether the person filing is a: | | | | | | | |
| (a) ☐ Broker or dealer registered under Section 15 of the Act (15 U.S.C. 78o). | | | | | | | |
| CUSIP No: 293602207 | | | | | | | |
| (b) ☐ Bank as defined in Section 3(a)(6) of the Act (15 U.S.C. 78c). | | | | | | | |
| (c) ☐ Insurance company as defined in Section 3(a)(19) of the Act (15 U.S.C. 78c). | | | | | | | |
| (d) ☐ Investment company registered under Section 8 of the Investment Company Act of 1940 (15 U.S.C. 80a-8). | | | | | | | |
| (e) ☐ An investment adviser in accordance with §240.13d-1(b)(1)(ii)(E); | | | | | | | |
| (f) \square An employee benefit plan or endowment fund in accordance with §240.13d-1(b)(1)(ii)(F); | | | | | | | |
| (g) ☐ A parent holding company or control person in accordance with §240.13d-1(b)(1)(ii)(G); | | | | | | | |
| (h) ☐ A savings association as defined in Section 3(b) of the Federal Deposit Insurance Act (12 U.S.C. 1813); | | | | | | | |
| (i) A church plan that is excluded from the definition of an investment company under Section 3(c)(14) of the Investment Company Act of 1940 (15 U.S.C. 80a-3); | | | | | | | |
| (j) \square A non-U.S. institution in accordance with §240.13d-1(b)(1)(ii)(J); | | | | | | | |

(k) \square Group, in accordance with Rule 13d-1(b)(1)(ii)(K).

| If filing | as a non-U.S. institution in accordance with §240.13d-1(b)(1)(ii)(J), please specify | the type of institution: | | | | | |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--|--|--|--|--|
| Item 4. | Ownership | | | | | | |
| | Provide the following information regarding the aggregate number and percent | age of the class of securities of the issuer identified in Item 1. | | | | | |
| | The information required by Items $4(a) - (c)$ is set forth in Rows $5 - 11$ of the coveach such Reporting Person. | ver page for each Reporting Person hereto and is incorporated herein by reference for | | | | | |
| | The Company's Prospectus (Registration No. 333-268038), filed on December 9, warrants issued at the same time) as of the completion of the offering of the Share | 2022, indicates there were 6,413,324 Shares outstanding (excluding Shares underlying es referred to therein. | | | | | |
| Heights Capital Management, Inc., which serves as the investment manager to CVI Investments, Inc., may be deemed to be the beneficial owner of all Shar CVI Investments, Inc. Each of the Reporting Persons hereby disclaims any beneficial ownership of any such Shares, except for their pecuniary interest there | | | | | | | |
| Item 5. | Ownership of Five Percent or Less of a Class | | | | | | |
| class of | If this statement is being filed to report the fact that as of the date hereof the resecurities, check the following: [] | porting person has ceased to be the beneficial owner of more than five percent of the | | | | | |
| | | | | | | | |
| | | | | | | | |
| CUSIP 1 | No: 293602207 | | | | | | |
| Item 6. | Ownership of More than Five Percent on Behalf of Another Person | | | | | | |
| | Not applicable. | | | | | | |
| Item 7. | Identification and Classification of the Subsidiary Which Acquired the Security Being Reported on by the Parent Holding Company or Control Person | | | | | | |
| | Not applicable. | | | | | | |
| Item 8. | Identification and Classification of Members of the Group | | | | | | |
| | Not applicable. | | | | | | |
| Item 9. | Notice of Dissolution of Group | | | | | | |
| | Not applicable. | | | | | | |
| Item 10. | Certification | | | | | | |
| | , e e | ledge and belief, the securities referred to above were not acquired and are not held for securities and were not acquired and are not held in connection with or as a participant | | | | | |
| | | | | | | | |
| CUSIP 1 | No: 293602207 | | | | | | |
| SIGNAT | TURES | | | | | | |
| is true, c | After reasonable inquiry and to the best of its knowledge and belief, each of the omplete, and correct. | e undersigned certifies that the information with respect to it set forth in this statement | | | | | |
| Dated: I | December 16, 2022 | | | | | | |
| CVI INV | VESTMENTS, INC. | HEIGHTS CAPITAL MANAGEMENT, INC. | | | | | |
| | ghts Capital Management, Inc. to a Limited Power of Attorney, a copy of which is attached as Exhibit I hereto | By: /s/ Brian Sopinsky Name: Brian Sopinsky Title: Secretary | | | | | |
| Name: | Brian Sopinsky Brian Sopinsky Secretary | | | | | | |

CUSIP No: 293602207

EXHIBIT DESCRIPTION

I Limited Power of Attorney
II Joint Filing Agreement

CUSIP No: 293602207

Exhibit I

LIMITED POWER OF ATTORNEY

THIS LIMITED POWER OF ATTORNEY given on the 16th day of July, 2015 by CVI Investments, Inc. (hereinafter called "the Company"), whose Registered Office is situated at PO Box 309GT, Ugland House, South Church Street, George Town, Grand Cayman, Cayman Islands.

WHEREAS, by agreement dated July 16, 2015, by and between the Company and Heights Capital Management, Inc., the Company expressly authorized Heights Capital Management, Inc. to enter into transactions in certain designated areas as defined in the Discretionary Investment Management Agreement attached hereto marked "Appendix 1"

NOW THIS DEED WITNESSETH that William Walmsley, Director of the Company, hereby appoints on behalf of the Company the firm of HEIGHTS CAPITAL MANAGEMENT, INC., which through its officers, directors and employees is hereby formally granted limited power of attorney for the purpose of entering into transactions on behalf and for the account of the Company; and to take all actions on behalf of the Company as may be necessary to consummate such transactions, including but not limited to making, negotiating; signing, endorsing, executing, acknowledging and delivering in the name of the Company all applications, contracts, agreements, notes, statements, certificates, proxies and any other instruments of whatever kind and nature as may be necessary or proper in connection with the entering into of such transactions, instructing the transfer of funds where necessary with respect to such transactions, and performing all of the services specified under the Discretionary Investment Management Agreement with respect to such transactions.

IN WITNESS WHEREOF, the Company has caused this Limited Power of Attorney to take effect on the day and year above written.

CVI Investments Inc.

By: /s/ William Walmsley

William Walmsley, Director

CUSIP No: 293602207

EXHIBIT II

JOINT FILING AGREEMENT

This will confirm the agreement by and among the undersigned that the Schedule 13G filed with the Securities and Exchange Commission on or about the date hereof with respect to the beneficial ownership by the undersigned of the shares of common stock of Ensysce Biosciences, Inc., \$0.0001 par value per share, is being filed, and all amendments thereto will be filed, on behalf of each of the persons and entities named below in accordance with Rule 13d-1(k) under the Securities Exchange Act of 1934, as amended. This Agreement may be executed in two or more counterparts, each of which shall be deemed an original, but all of which together shall constitute one and the same instrument

Dated as of December 16, 2022

CVI INVESTMENTS, INC.

HEIGHTS CAPITAL MANAGEMENT, INC.

By: Heights Capital Management, Inc. pursuant to a Limited Power of Attorney

By: /s/ Brian Sopinsky
Name: Brian Sopinsky
Title: Secretary

By: /s/ Brian Sopinsky

Name: Brian Sopinsky Title: Secretary