UNITED STATES SECURITIES AND EXCHANGE COMMISSION Washington, D.C. 20549

FORM S-3 REGISTRATION STATEMENT

Under The Securities Act of 1933

	Skye Bios	cience, Inc.	
	(Exact name of Registrar	nt as specified in its charter)	
Nevada		45	5-0692882
(State or other jurisdic			S. Employer
incorporation or organ	ization)	Identific	cation Number)
(Address, including zip code, an	San Diege (858) 4	Boulevard, Suite 320 b, CA 92121 i10-0266 uding area code, of Registrant's	principal executive offices)
	Punit Chief Exec Skye Bios 5910 Pacific Center San Dieg (858) 4	Dhillon utive Officer science, Inc. Boulevard, Suite 320 b, CA 92121 i10-0266 e number, including area code, o	
	Сор	ies to:	
	Greenberg 1201 K Stro Sacrament	Lee, Esq. Traurig, LLP eet, Suite 1100 o, CA 95814 868-0630	
Approximate date of proposed sale to the publ	ic: From time to time after	this Registration Statement become	nes effective.
If the only securities being registered on this For \Box	m are being offered pursua	ant to dividend or interest reinvest	ment plans, please check the following box.
If any of the securities being registered on this F 1933, other than securities offered only in connec			
If this Form is filed to register additional securiti list the Securities Act registration statement number			
If this Form is a post-effective amendment filed Act registration statement number of the earlier e			eck the following box and list the Securities
If this Form is a registration statement pursuant t with the Commission pursuant to Rule 462(e) und		*	ereto that shall become effective upon filing
If this Form is a post-effective amendment to a additional classes of securities pursuant to Rule 4			I.D. filed to register additional securities or
Indicate by check mark whether the registrant is an emerging growth company. See definitions company" in Rule 12b-2 of the Exchange Act.			
Large accelerated filer Non-accelerated filer		Accelerated filer Smaller reporting company Emerging growth company	
If an emerging growth company, indicate by checor revised financial accounting standards provided			ansition period for complying with any new

CALCULATION OF REGISTRATION FEE

Title of Each Class of Securities to be Registered ⁽¹⁾	Amount to be Registered ⁽¹⁾ (2)(3)	Proposed Maximum Offering Price Per Unit ⁽²⁾⁽³⁾	Proposed Maximum Aggregate Offering Price (1)(4)	Amount of Registration Fee ⁽⁵⁾
Common Stock, \$0.001 par value	_	_	_	_
Preferred Stock, \$0.001 par value				
Warrants (6)	_	_	_	_
Units (7)				
Total	_	_	\$ 75,000,000	\$ 8,182.50

- (1) There are being registered hereunder such indeterminate number of shares of common stock and preferred stock, such indeterminate number of warrants to purchase common stock and/or preferred stock, and such indeterminate number of units, as may be offered and sold by the registrant from time to time and that together shall have an aggregate initial offering price not to exceed \$75,000,000. Any securities registered hereunder may be sold separately or as units with other securities registered hereunder. The securities registered also include indeterminate amounts of such securities as may be issued upon exercise, conversion or exchange of, or pursuant to anti-dilution adjustments with respect to, securities that provide for such issuance, exercise, conversion, exchange or adjustment. Separate consideration may or may not be received for any of these securities. In addition, pursuant to Rule 416 under the Securities Act of 1933, as amended (the "Securities Act"), the securities being registered hereunder include an indeterminate number of shares of common stock that may be issuable with respect to the shares being registered hereunder as a result of stock splits, stock dividends, or similar transactions.
- (2) Not required to be included in accordance with General Instruction II.D. of Form S-3 and Rule 457(o) under the Securities Act.
- (3) The proposed maximum offering price per unit and the proposed maximum aggregate offering price per class of security will be determined from time to time by the registrant in connection with the issuance by the registrant of the securities registered hereunder.
- (4) Estimated solely to calculate the registration fee in accordance with Rule 457(o) under the Securities Act. The aggregate maximum offering price of all securities issued pursuant to this registration statement will not exceed \$75,000,000.
- (5) Calculated pursuant to Rule 457(o) under the Securities Act based on the proposed maximum aggregate offering price.
- (6) The warrants covered by this registration statement may be warrants to purchase common stock and/or preferred stock.
- (7) Any securities registered hereunder may be sold separately or as units with other securities registered hereunder.

The registrant hereby amends this registration statement on such date or dates as may be necessary to delay its effective date until the registrant shall file a further amendment which specifically states that this registration statement shall thereafter become effective in accordance with section 8(a) of the Securities Act of 1933 or until the registration statement shall become effective on such date as the Commission, acting pursuant to said section 8(a), may determine.

The information in this prospectus is not complete and may be changed. We may not sell these securities until the registration statement filed with the Securities and Exchange Commission is effective. This prospectus is not an offer to sell these securities, and the selling stockholders are not soliciting offers to buy these securities, in any state where the offer or sale of these securities is not permitted.

SUBJECT TO COMPLETION, DATED July 28, 2021

PROSPECTUS



SKYE BIOSCIENCE, INC.

\$75,000,000

Common Stock Preferred Stock Warrants Units

We may offer and sell from time to time, in one or more series or issuances and on terms that we will determine at the time of the offering, any combination of the securities described in this prospectus, up to an aggregate amount of \$75,000,000.

This prospectus provides a general description of the securities we may offer. Each time we sell securities, we will provide specific terms of any offering in a supplement to this prospectus. We may also authorize one or more free writing prospectuses to be provided to you in connection with these offerings. Any prospectus supplement and any related free writing prospectus may also add, update, or change information contained in this prospectus. You should carefully read this prospectus, the applicable prospectus supplement and any related free writing prospectus, as well as the documents incorporated or deemed to be incorporated by reference in this prospectus before you purchase any of the securities offered hereby. This prospectus may not be used to consummate a sale of securities by us unless accompanied by the applicable prospectus supplement.

These securities may be offered and sold in the same offering or in separate offerings; to or through underwriters, dealers, and agents; or directly to institutional purchasers. The names of any underwriters, dealers, or agents involved in the sale of our securities, their compensation and any overallotment options held by them will be described in the applicable prospectus supplement. For a more complete description of the plan of distribution of these securities, see the section entitled "Plan of Distribution" beginning on page 19 of this prospectus.

Our common stock, par value \$0.001 per share ("Common Stock"), is currently quoted on the OTCQB Marketplace operated by OTC Markets Group Inc. (the "OTCQB") under the trading symbol "SKYE". On July 27, 2021, the last reported sale price of our Common Stock on the OTCQB Market was \$0.1495 per share. We will provide information in any applicable prospectus supplement regarding any listing of securities other than shares of our Common Stock on any securities exchange.

The aggregate market value of our outstanding Common Stock held by non-affiliates was \$75,661,160, which was calculated based on 303,860,082 shares of outstanding Common Stock held by non-affiliates as of July 27, 2021, and a price per share of \$0.249, the last reported sale price of our Common Stock on June 2, 2021.

Investing in our securities involves a high degree of risk. You should carefully read and consider the risk factors described in, and incorporated by reference under, "Risk Factors" beginning on page 8 of this prospectus and in the applicable prospectus supplement before investing in any securities.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or passed upon the adequacy or accuracy of this prospectus. Any representation to the contrary is a criminal offense.

This prospectus is dated July 28, 2021

TABLE OF CONTENTS

	Page
ABOUT THIS PROSPECTUS	6
PROSPECTUS SUMMARY	7
RISK FACTORS	8
FORWARD-LOOKING STATEMENTS	9
<u>USE OF PROCEEDS</u>	10
DESCRIPTION OF CAPITAL STOCK	11
DESCRIPTION OF WARRANTS	15
DESCRIPTION OF UNITS	18
PLAN OF DISTRIBUTION	19
<u>LEGAL MATTERS</u>	21
EXPERTS	21
WHERE YOU CAN FIND MORE INFORMATION	21
INCORPORATION OF CERTAIN DOCUMENTS BY REFERENCE	22

ABOUT THIS PROSPECTUS

This prospectus is part of a registration statement we filed with the Securities and Exchange Commission, or the "Commission" or the "SEC," using the "shelf" registration process. Under the shelf registration process, using this prospectus, together with a prospectus supplement, we may sell from time to time any combination of the securities described in this prospectus in one or more offerings. This prospectus provides you with a general description of the securities that may be offered. Each time we sell securities pursuant to this prospectus, we will provide a prospectus supplement that will contain specific information about the terms of the securities being offered. A prospectus supplement may include a discussion of any risk factors or other special considerations applicable to those securities or to us. The prospectus supplement may also add to, update or change information contained in this prospectus and, accordingly, to the extent inconsistent, the information in this prospectus will be superseded by the information in the prospectus supplement. You should read this prospectus, any applicable prospectus supplement and any related issuer free writing prospectus, as well as the additional information incorporated by reference in this prospectus described below under "Where You Can Find More Information" and "Information Incorporated by Reference" or in any applicable prospectus supplement and any related issuer free writing prospectus, before making an investment in our securities.

This prospectus contains summaries of certain provisions contained in some of the documents described herein, but reference is made to the actual documents for complete information. All of the summaries are qualified in their entirety by the actual documents. Copies of the documents referred to herein have been filed, or will be filed or incorporated by reference as exhibits to the registration statement of which this prospectus is a part, and you may obtain copies of those documents as described below under "Where You Can Find More Information."

Neither the delivery of this prospectus, any accompanying prospectus supplement or any free writing prospectus prepared by us, nor any sale made under this prospectus, any accompanying prospectus supplement or any free writing prospectus prepared by us, implies that there has been no change in our affairs or that the information therein is correct as of any date after the date of this prospectus or of such prospectus supplement or free writing prospectus, as applicable. You should not assume that the information in this prospectus, including any information incorporated in this prospectus by reference, the accompanying prospectus supplement or any free writing prospectus prepared by us, is accurate as of any date other than the date on the front of those documents. Our business, financial condition, results of operations and prospects may have changed since that date.

You may rely only on the information contained or incorporated by reference in this prospectus. Neither we nor any other person has authorized anyone to provide any information other than that contained or incorporated by reference in this prospectus, any accompanying prospectus supplement or in any free writing prospectus prepared by or on behalf of us or to which we have referred you. We take no responsibility for and can provide no assurance as to the reliability of, any other information that others may give you. Neither this prospectus nor any accompanying prospectus supplement nor any free writing prospectus prepared by or on behalf of us or to which we have referred you constitutes an offer to sell nor a solicitation of an offer to buy any securities other than those registered by this prospectus, or an offer to sell or a solicitation of an offer to buy securities where an offer or solicitation would be unlawful. This prospectus does not contain all of the information included in the registration statement. For a more complete understanding of the offering of the securities, you should refer to the registration statement, including its exhibits.

PROSPECTUS SUMMARY

This summary highlights information contained elsewhere in this prospectus and does not contain all of the information that you should consider in making your investment decision. Before deciding to invest in our securities, you should read this entire prospectus carefully, including the sections of this prospectus entitled "Risk Factors" and "Management's Discussion and Analysis of Financial Condition and Results of Operations" and our consolidated financial statements and related notes contained elsewhere in this prospectus. Unless the context otherwise requires, references in this prospectus to the "Company," "Skye Bioscience," "we," "us", and "our" refer to Skye Bioscience, Inc., together with its wholly owned subsidiaries, Nemus, a California corporation, and Skye Bioscience Pty Ltd, an Australian proprietary limited company.

Overview

We are a biopharmaceutical company targeting the discovery, development and commercialization of cannabinoid-based therapeutics through a number of license agreements with the University of Mississippi which has significant expertise in the chemistry and physiology of cannabinoid molecules. We were established as, and continue to be, focused on research and development of proprietary cannabinoid molecules.

Corporate information

We were incorporated in Nevada on March 16, 2011. Our principal executive offices are located at 5910 Pacific Center Boulevard, Suite 320, San Diego, CA 92121, and our telephone number is (858) 410-0266. Our website address is http://www.skyebioscience.com. The information contained on our website is not incorporated by reference into this prospectus, and you should not consider any information contained on, or that can be accessed through, our website as part of this prospectus, or in deciding whether to purchase our securities.

Description of the Securities We May Offer

The descriptions of the securities contained in this prospectus, together with the applicable prospectus supplements, summarize the material terms and provisions of the various types of securities that we may offer. We will describe in the applicable prospectus supplement relating to any securities the particular amounts, prices and other terms of the securities offered by that prospectus supplement. If we so indicate in the applicable prospectus supplement, the terms of the securities may differ from the terms we have summarized below. We will also include in the prospectus supplement information, where applicable, about material U.S. federal income tax considerations relating to the securities, and the securities exchange, if any, on which the securities will be listed.

In this prospectus, we refer to the Common Stock, preferred stock, warrants to purchase Common Stock and/or preferred stock and units registered hereunder collectively as "securities." The total dollar amount of all securities that we may issue under this prospectus will not exceed \$75,000,000.

This prospectus may not be used by us to consummate a sale of securities unless it is accompanied by a prospectus supplement.

RISK FACTORS

An investment in our securities involves a high degree of risk. Certain risks relating to us and our business are described under the headings "Business" and "Risk Factors" in our Annual Report on Form 10-K for the year ended December 31, 2020, filed with the Commission on March 2, 2021, and our Quarterly Report on Form 10-Q for the quarter ended March 31, 2021, filed with the Commission on May 7, 2021, which are incorporated by reference into this prospectus and any accompanying prospectus supplement and which you should carefully review and consider, along with the other information contained in this prospectus and any accompanying prospectus supplement or incorporated by reference herein, as updated by our subsequent filings under the Securities Exchange Act of 1934, as amended, or the "Exchange Act" before making an investment in any of our securities.

Additional risks, as well as updates or changes to the risks described in the documents incorporated by reference herein, may be included in any applicable prospectus supplement. Our business, financial condition or results of operations could be materially adversely affected by any of these risks. The market or trading price of our securities could decline due to any of these risks, and you may lose all or part of your investment. Prior to making a decision to invest in our securities you should consider carefully the specific factors discussed under the caption "Risk Factors" in the applicable prospectus supplement, together with any other information contained in the applicable prospectus supplement or appearing or incorporated by reference in this prospectus.

In addition, please read the section of this prospectus captioned "Special Note Regarding Forward-Looking Statements," in which we describe additional uncertainties associated with our business and the forward-looking statements included or incorporated by reference in this prospectus. Please note that additional risks not presently known to us or that we currently deem immaterial may also impair our business and operations.

Investment in any securities offered pursuant to this prospectus involves risks and uncertainties. If one or more of the events discussed in the risk factors were to occur, our business, financial condition, results of operations or liquidity, as well as the value of an investment in our securities, could be materially adversely affected.

You should carefully consider the risk factors as well as the other information contained and incorporated by reference in this prospectus before deciding to invest.

FORWARD-LOOKING STATEMENTS

Information presented in this prospectus, any accompanying prospectus supplement and any free writing prospectus prepared by or on behalf of us or to which we have referred you, and in other documents which are incorporated by reference in this prospectus under the sections of this prospectus entitled "Where You Can Find More Information" and "Information Incorporated by Reference," that discusses financial projections, information or expectations about our business plans, results of operations, products or markets, or otherwise makes statements about future events, are forward-looking statements. Forward-looking statements can be identified by the use of words such as "intends," "anticipates," "believes," "estimates," "projects," "forecasts," "expects," "plans," and "proposes." Although we believe that the expectations reflected in these forward-looking statements are based on reasonable assumptions, there are a number of risks and uncertainties that could cause actual results to differ materially from these forward-looking statements. These include, among others, the cautionary statements in the "Risk Factors" section of this prospectus set forth above. These cautionary statements identify important factors that could cause actual results to differ materially from those described in the forward-looking statements. When considering forward-looking statements in this prospectus, you should keep in mind the cautionary statements in the "Risk Factors" section and other sections of this prospectus, and other cautionary statements in any accompanying prospectus supplement and any free writing prospectus prepared by or on behalf of us or to which we have referred you and any documents which are incorporated by reference in this prospectus and listed in the "Where You Can Find More Information" and "Information Incorporated by Reference" sections set forth below.

Certain risks, uncertainties, and other factors are incorporated herein by reference to our most recent Annual Report on Form 10-K and our subsequent Quarterly Report on Form 10-Q, along with the other information contained in this prospectus, as updated by our subsequent filings under the Exchange Act. Except as otherwise required by applicable securities laws, we undertake no obligation to publicly update or revise any forward-looking statements, whether as a result of new information, future events, changed circumstances, or any other reason, after the date of this prospectus.

USE OF PROCEEDS

Unless otherwise indicated in the prospectus supplement, we will use the net proceeds from the sale of securities offered by this prospectus for general corporate purposes, which may include working capital, capital expenditures, other corporate expenses, the refinancing or repayment of existing indebtedness and acquisitions of complementary products, product candidates, technologies or businesses. However, we currently have no present agreements or commitments for any such acquisitions. The timing and amount of our actual expenditures will be based on many factors, including cash flows from operations and the anticipated growth of our business. As a result, unless otherwise indicated in the prospectus supplement, our management will have broad discretion to allocate the net proceeds of the offerings.

DESCRIPTION OF CAPITAL STOCK

The following description of our Common Stock and preferred stock, together with the additional information we include in any applicable prospectus supplements, summarizes the material terms and provisions of the Common Stock and/or preferred stock that we may offer under this prospectus. It may not contain all the information that is important to you. For the complete terms of our Common Stock and preferred stock, please refer to our amended and restated articles of incorporation and amended and restated bylaws, as may be amended from time to time, and which are incorporated by reference into the registration statement which includes this prospectus. The Nevada Revised Statutes (as amended from time to time, the "NRS") may also affect the terms of these securities.

General

Our amended and restated articles of incorporation authorize us to issue 5,000,000,000 shares of common stock, par value \$0.001 per share, of which 417,247,333 shares are issued and outstanding as of July 27, 2021, and 50,000,000 shares of preferred stock, par value \$0.001 per share, none of which are outstanding.

Common Stock

Holders of our Common Stock are entitled to one vote for each share held of record on all matters on which stockholders are entitled to vote generally, including the election or removal of directors. Holders of our Common Stock do not have cumulative voting rights in the election of directors. Holders of Common Stock are not subject to further calls or assessments as a result of their holding shares of Common Stock.

Holders of shares of our Common Stock are entitled to receive dividends when, as and if declared by our board of directors out of funds legally available therefor, subject to any statutory or contractual restrictions on the payment of dividends and to any restrictions on the payment of dividends imposed by the terms of any outstanding preferred stock.

Upon our liquidation, dissolution or winding up and after payment in full of all amounts required to be paid to creditors and to the holders of preferred stock having liquidation preferences, if any, the holders of shares of our Common Stock will be entitled to share ratably our remaining assets available for distribution.

Holders of shares of our Common Stock do not have preemptive, subscription, redemption or conversion rights. There are no redemption or sinking fund provisions applicable to the Common Stock.

Our Common Stock is currently quoted on the OTCQB Marketplace operated by OTC Markets Group Inc. under the trading symbol "SKYE". The transfer agent and registrar for our Common Stock is ClearTrust LLC, 16540 Pointe Village Drive, Suite 210, Lutz, Florida 33558.

Preferred Stock

Our amended and restated articles of incorporation authorize us to issue up to 50,000,000 shares of preferred stock, par value \$0.001 per share, in one or more series and with rights and preferences that may be fixed or designated by our board of directors without any further action by our stockholders. Each series of preferred stock will have the number of shares, designations, preferences, voting powers, qualifications and special or relative rights or privileges as shall be determined by our board of directors, which may include, among others, dividend rights, voting rights, liquidation preferences, conversion rights and preemptive rights.

Our board of directors may issue authorized shares of preferred stock, as well as authorized but unissued shares of Common Stock, without further stockholder action, unless stockholder action is required by applicable law or by the rules of a stock exchange or quotation system on which any series of our stock may be listed or quoted.

When we offer to sell a particular series of preferred stock, we will describe the specific terms of the securities in a supplement to this prospectus. The prospectus supplement will also indicate whether the general terms and provisions described in this prospectus apply to the particular series of preferred stock. The preferred stock will be issued under a certificate of designations relating to each series of preferred stock and is also subject to our amended and restated articles of incorporation. The certificate of designations will be filed with the SEC in connection with an offering of preferred stock.

The prospectus supplement relating to any preferred stock that we may offer will describe the terms of any preferred stock being offered, including:

- the number of shares and designation or title of the shares;
- any liquidation preference per share;
- any date of maturity;
- any redemption, repayment or sinking fund provisions;
- any dividend rate or rates and the dates of payment (or the method for determining the dividend rates or dates of payment);
- any voting rights;
- if other than the currency of the United States, the currency or currencies including composite currencies in which the preferred stock is denominated and/or in which payments will or may be payable;
- the method by which amounts in respect of the preferred stock may be calculated and any commodities, currencies or indices, or value, rate or price, relevant to such calculation;
- whether the preferred stock is convertible or exchangeable and, if so, the securities or rights into which the preferred stock is convertible or exchangeable, and the terms and conditions of conversion or exchange;
- the place or places where dividends and other payments on the preferred stock will be payable; and
- any additional voting, dividend, liquidation, redemption, preemption, transfer restrictions, and other rights, preferences, privileges, limitations and restrictions.

All shares of preferred stock offered will be fully paid and non-assessable. Any shares of preferred stock that are issued will have priority over the Common Stock with respect to dividend or liquidation rights or both.

Our board of directors could create and issue a series of preferred stock with rights, privileges or restrictions which effectively discriminate against an existing or prospective holder of preferred stock as a result of the holder beneficially owning or commencing a tender offer for a substantial amount of Common Stock. One of the effects of authorized but unissued and unreserved shares of capital stock may be to make it more difficult or discourage an attempt by a potential acquirer to obtain control of our company by means of a merger, tender offer, proxy contest or otherwise. This protects the continuity of our management. The issuance of these shares of capital stock may defer or prevent a change in control of our company without any further stockholder action.

The transfer agent for each series of preferred stock will be described in the prospectus supplement.

Limitations on Liability and Indemnification of Officers and Directors

Our amended and restated articles of incorporation provide that the liability of our directors and officers shall be eliminated or limited to the fullest extent permitted by the NRS. NRS 78.138(7) provides that, subject to very limited statutory exceptions and unless the articles of incorporation or an amendment thereto (in each case filed on or after October 1, 2003) provide for greater individual liability, a director or officer is not individually liable to a corporation or its stockholders or creditors for any damages as a result of any act or failure to act in his or her capacity as a director or officer unless it is proven that: (i) the act or failure to act constituted a breach of his or her fiduciary duties as a director or officer and (ii) the breach of those duties involved intentional misconduct, fraud or a knowing violation of law.

Our amended and restated articles of incorporation and amended and restated bylaws provide that we must indemnify and advance expenses to our directors and officers to the fullest extent permitted under the NRS. We also are expressly authorized to carry directors' and officers' liability insurance providing indemnification for our directors, officers and certain employees for some liabilities.

Anti-Takeover Effects of Our Amended and Restated Articles of Incorporation and Amended and Restated Bylaws and Certain Provisions of Nevada Law

The provisions of our amended and restated articles of incorporation and amended and restated bylaws and of the NRS summarized below may have an anti-takeover effect and may delay, defer or prevent a tender offer or takeover attempt that a stockholder might consider in the stockholder's best interest, including an attempt that might result in the stockholder's receipt of a premium over the market price for the stockholder's shares. These provisions are also designed, in part, to encourage persons seeking to acquire control of us to first negotiate with our board of directors, which could result in an improvement of their terms.

Blank Check Preferred Stock

Our amended and restated articles of incorporation authorize the issuance of up to 50,000,000 shares of preferred stock in one or more series with such designations, voting powers, if any, preferences and relative, participating, optional or other special rights, and such qualifications, limitations and restrictions, as are determined by our board of directors in accordance with Nevada law.

Board of Directors

Our amended and restated articles of incorporation and amended and restated bylaws provide that the number of directors will be fixed from time to time exclusively pursuant to a resolution adopted by the board of directors.

Business Combinations and Acquisition of Control Shares

Pursuant to provisions in our amended and restated articles of incorporation, we have elected not to be governed by certain Nevada statutes that may have the effect of discouraging corporate takeovers.

Nevada's "combinations with interested stockholders" statutes (NRS 78.411 through 78.444, inclusive) prohibit specified types of business "combinations" between certain Nevada corporations and any person deemed to be an "interested stockholder" for two years after such person first becomes an "interested stockholder" unless the corporation's board of directors approves the combination (or the transaction by which such person becomes an "interested stockholder") in advance, or unless the combination is approved by the board of directors and sixty percent of the corporation's voting power not beneficially owned by the interested stockholder, its affiliates and associates. Furthermore, in the absence of prior approval certain restrictions may apply even after such two-year period. For purposes of these statutes, an "interested stockholder" is any person who is (1) the beneficial owner, directly or indirectly, of ten percent or more of the voting power of the corporation and at any time within the two previous years was the beneficial owner, directly or indirectly, of ten percent or more of the voting power of the then-outstanding shares of the corporation. The definition of the term "combination" is sufficiently broad to cover most significant transactions between a corporation and an "interested stockholder." Our amended and restated articles of incorporation provide that these statutes do not apply to us.

Nevada's "acquisition of controlling interest" statutes (NRS 78.378 through 78.3793, inclusive) contain provisions governing the acquisition of a controlling interest in certain Nevada corporations. These "control share" laws provide generally that any person that acquires a "controlling interest" in certain Nevada corporations may be denied voting rights, unless a majority of the disinterested stockholders of the corporation elects to restore such voting rights. Our amended and restated articles of incorporation provide that these statutes will not apply to us and such provision remains in effect. Absent such a provision in our articles of incorporation or bylaws, these laws would apply to us if we were to have 200 or more stockholders of record (at least 100 of whom have addresses in Nevada appearing on our stock ledger) and do business in the State of Nevada directly or through an affiliated corporation, unless our articles of incorporation or bylaws in effect on the tenth day after the acquisition of a controlling interest provide otherwise. These laws provide that a person acquires a "controlling interest" whenever a person acquires shares of a subject corporation that, but for the application of these provisions of the NRS, would enable that person to exercise (1) one-fifth or more, but less than one-third, (2) one-third or more, but less than a majority or (3) a majority or more, of all of the voting power of the corporation in the election of directors. Once an acquirer crosses one of these thresholds, shares which it acquired in the transaction taking it over the threshold and within the 90 days immediately preceding the date when the acquiring person acquired or offered to acquire a controlling interest become "control shares" to which the voting restrictions described above apply.

In addition, NRS 78.139 also provides that directors may resist a change or potential change in control if the directors, by majority vote of a quorum, determine that the change is opposed to, or not in, the best interests of the corporation.

Removal of Directors; Vacancies

Under NRS 78.335, one or more of the incumbent directors may be removed from office by the vote of stockholders representing two-thirds or more of the voting power of the issued and outstanding stock entitled to vote. Our amended and restated articles of incorporation provide that any newly created position on the board of directors that results from an increase in the total number of directors and any vacancies on the board of directors will be filled only by the affirmative vote of a majority of the remaining directors, even if less than a quorum, by a sole remaining director.

No Cumulative Voting

The NRS does not permit stockholders to cumulate their votes other than in the election of directors, and then only if expressly authorized by the corporation's articles of incorporation. Our amended and restated articles of incorporation do not permit cumulative voting.

Special Stockholder Meetings

Our amended and restated articles of incorporation provide that special meetings of our stockholders may be called at any time only by or at the direction of (i) the board of directors, (ii) the chairman of the board of directors or (iii) two or more of the members of our board of directors. Our amended and restated bylaws prohibit the conduct of any business at a special meeting other than as specified in the notice for such meeting.

Requirements for Advance Notification of Director Nominations and Stockholder Proposals

Our amended and restated bylaws established advance notice procedures with respect to stockholder proposals and the nomination of candidates for election as directors, other than nominations made by or at the direction of the board of directors or a committee of the board of directors. In order for any matter to be properly brought before a meeting of our stockholders, the stockholder submitting the proposal or nomination will have to comply with advance notice requirements and provide us with certain information.

Exclusive Forum

Our amended and restated articles of incorporation provide that unless we consent to the selection of an alternative forum that the Eighth Judicial District Court of Clark County, Nevada shall be the sole and exclusive forum for any (i) derivative action or proceeding brought in the name or right of the corporation or on its behalf, (ii) action asserting a claim of breach of a fiduciary duty owed by any of our directors, officers, employees or agents to the corporation or any of our stockholders, creditors or other constituents or stakeholders, (iii) action asserting a claim arising pursuant to any provision of Chapters 78 or 92A of the NRS or any provision of our articles of incorporation or bylaws or (iv) any action asserting a claim governed by the internal affairs doctrine.

DESCRIPTION OF WARRANTS

General

We may issue warrants for the purchase of our Common Stock and/or preferred stock. Warrants may be issued independently or together with our Common Stock and/or preferred stock and may be attached to or separate from any offered securities. The warrants may be attached to or separate from those offered securities. We may issue the warrants under warrant agreements to be entered into between us and a bank or trust company to be named in the applicable prospectus supplement, as warrant agent, all as described in the applicable prospectus supplement. The warrant agent will act solely as our agent in connection with the warrants and will not assume any obligation or relationship of agency or trust for or with any holders or beneficial owners of warrants. If we offer warrants, we will file the warrant agreement relating to the offered warrants as an exhibit to, or incorporate it by reference in, the registration statement of which this prospectus is a part.

Warrants

The prospectus supplement relating to any warrants that we may offer will contain the specific terms of the warrants. These terms may include the following:

- · the title of the warrants;
- · the offering price for the warrants, if any;
- · the aggregate number of warrants;
- the designation and terms of the Common Stock and/or preferred stock that may be purchased upon exercise of the warrants;
- · if applicable, the designation and terms of the securities with which the warrants are issued and the number of warrants issued with each security;
- · if applicable, the date from and after which the warrants and any securities issued with the warrants will be separately transferable;
- the number of shares of Common Stock and/or preferred stock that may be purchased upon exercise of a warrant and the exercise price for the warrants:
- · whether the exercise price may be paid in cash, by the exchange of warrants or other securities or both, and the method of exercising the warrants;
- whether the warrants will be settled by delivery of the underlying securities or other property or in cash;
- the dates on which the right to exercise the warrants shall commence and expire;
- · whether and under what circumstances we may cancel the warrants prior to their expiration date, in which case the holders will be entitled to receive only the applicable cancellation amount, which may be either a fixed amount or an amount that varies during the term of the warrants in accordance with a schedule or formula;
- · if applicable, the minimum or maximum amount of the warrants that may be exercised at any one time;
- the currency or currency units in which the offering price, if any, and the exercise price are payable;
- the identities of the warrant agent, any depositaries and any paying, transfer, calculation or other agents for the warrants;
- · any securities exchange or quotation system on which the warrants or any securities deliverable upon exercise of the warrants may be listed;

- whether the warrants are to be sold separately or with other securities, and if the warrants are to be sold with the securities of another company or other companies, certain information regarding such company or companies;
- · if applicable, a discussion of material U.S. federal income tax considerations;
- the anti-dilution provisions of the warrants, if any;
- · the redemption or call provisions, if any, applicable to the warrants;
- any provisions with respect to the holder's right to require us to repurchase the warrants upon a change in control or similar event; and
- · any additional terms of the warrants, including procedures, and limitations relating to the exchange, exercise and settlement of the warrants.

Holders of equity warrants will not be entitled:

- to vote, consent or receive dividends;
- · receive notice as stockholders with respect to any meeting of stockholders for the election of our directors or any other matter; or
- · exercise any rights as stockholders of us.

Exercise of Warrants

Each warrant will entitle the holder to purchase the securities that we specify in the applicable prospectus supplement at the exercise price that we describe in the applicable prospectus supplement. Unless we otherwise specify in the applicable prospectus supplement, holders of the warrants may exercise the warrants at any time up to the specified time on the expiration date that we set forth in the applicable prospectus supplement. After the close of business on the expiration date, unexercised warrants will become void.

Unless we otherwise specify in the applicable prospectus supplement, holders of the warrants may exercise the warrants by delivering the warrant certificate representing the warrants to be exercised together with specified information, and paying the required amount to the warrant agent in immediately available funds, as provided in the applicable prospectus supplement. We will set forth on the reverse side of the warrant certificate and in the applicable prospectus supplement the information that the holder of the warrant will be required to deliver to the warrant agent in connection with the exercise of the warrant.

Upon receipt of the required payment and the warrant certificate properly completed and duly executed at the corporate trust office of the warrant agent or any other office indicated in the applicable prospectus supplement, we will issue and deliver the securities purchasable upon such exercise. If fewer than all of the warrants represented by the warrant certificate are exercised, then we will issue a new warrant certificate for the remaining amount of warrants. If we so indicate in the applicable prospectus supplement, holders of the warrants may surrender securities as all or part of the exercise price for the warrants.

Outstanding Warrants

As of July 27, 2021, we had warrants issued and outstanding for the purchase of up to 51,695,001 shares of our Common Stock, at exercise prices ranging from \$0.08 to \$5.00. Outstanding warrants to purchase our Common Stock are as follows:

Source	1	Exercise Price	Term (Years)	Number of Warrants Vested and Outstanding
Pre 2015 Common Stock Warrants	\$	1.00	6-10	1,110,000
2015 Common Stock Warrants	\$	5.00	10	100,000
2016 Common Stock Warrants to Service Providers	\$	1.15	10	40,000
2016 Series C Common Stock Warrants to Placement Agent	\$	0.40	5	125,000
2017 Series D Common Stock Warrants to Placement Agent	\$	0.25	5	480,000
2017 Common Stock Warrants to Service Provider	\$	0.41	5	125,000
2018 Emerald Financing Warrants	\$	0.10	5	3,400,000
Emerald Multi-Draw Credit Agreement Warrants	\$	0.50	5	7,500,000
2019 Common Stock Warrants	\$	0.35	5	8,000,000
2020 Common Stock Warrants to Placement Agent	\$	0.08	4.99	8,166,667
2021 Common Stock Warrants	\$	0.15	5	21,166,667
2021 Common Stock Warrants to Placement Agent	\$	0.19	5	1,481,667
Total warrants vested and outstanding as of July 27, 2021				51,695,001

DESCRIPTION OF UNITS

General

We may, from time to time, issue units comprised of one or more of the other securities described in this prospectus in any combination. A prospectus supplement will describe the specific terms of the units offered under that prospectus supplement, and any special considerations applicable to investing in those units. You must look at the applicable prospectus supplement and any applicable unit agreement for a full understanding of the specific terms of any units. We will incorporate by reference into the registration statement of which this prospectus is a part the form of unit agreement, including a form of unit certificate, if any, that describes the terms of the series of units we are offering before the issuance of the related series of units. While the terms we have summarized below will generally apply to any future units that we may offer under this prospectus, we will describe the particular terms of any series of units that we may offer in more detail in the applicable prospectus supplement and incorporated documents. The terms of any units offered under a prospectus supplement may differ from the terms described below.

Units

We may issue units consisting of Common Stock, preferred stock, warrants or any combination thereof. Each unit will be issued so that the holder of the unit is also the holder of each security included in the unit. Thus, the holder of a unit will have the rights and obligations of a holder of each included security. The unit agreement under which a unit is issued may provide that the securities included in the unit may not be held or transferred separately, at any time, or at any time before a specified date.

We will describe in the applicable prospectus supplement and any incorporated documents the terms of the series of units, including the following:

- the designation and terms of the units and of the securities comprising the units, including whether and under what circumstances those securities may be held or transferred separately;
- · any unit agreement under which the units will be issued; and
- any provisions for the issuance, payment, settlement, transfer, or exchange of the units or of the securities comprising the units.

The provisions described in this section, as well as those described under "Description of Capital Stock" and "Description of Warrants" will apply to each unit and to any Common Stock, preferred stock or warrant included in each unit, respectively.

Issuance in Series

We may issue units in such amounts and in such numerous distinct series as we determine.

Enforceability of Rights by Holders of Units

Each unit agent will act solely as our agent under the applicable unit agreement and will not assume any obligation or relationship of agency or trust with any holder of any unit. A single bank or trust company may act as unit agent for more than one series of units. A unit agent will have no duty or responsibility in case of any default by us under the applicable unit agreement or unit, including any duty or responsibility to initiate any proceedings at law or otherwise, or to make any demand upon us. Any holder of a unit, without the consent of the related unit agent or the holder of any other unit, may enforce by appropriate legal action its rights as holder under any security included in the unit.

Title

We, the unit agent, and any of their agents may treat the registered holder of any unit certificate as an absolute owner of the units evidenced by that certificate for any purposes and as the person entitled to exercise the rights attaching to the units so requested, despite any notice to the contrary.

PLAN OF DISTRIBUTION

We may sell the securities offered by this prospectus from time to time in one or more transactions,

- directly to purchasers;
- through agents;
- · to or through underwriters or dealers; or
- through a combination of these methods.

A distribution of the securities offered by this prospectus may also be effected through the issuance of derivative securities, including warrants.

In addition, the manner in which we may sell some or all of the securities covered by this prospectus includes, without limitation, through:

- a block trade in which a broker-dealer will attempt to sell as agent, but may position or resell a portion of the block, as principal, in order to facilitate the transaction;
- purchases by a broker-dealer, as principal, and resale by the broker-dealer for its account; or
- ordinary brokerage transactions and transactions in which a broker solicits purchasers.

In addition, we may enter into derivative or hedging transactions with third parties, or sell securities not covered by this prospectus to third parties in privately negotiated transactions. In connection with such a transaction, the third parties may sell securities covered by and pursuant to this prospectus and an applicable prospectus supplement or other offering materials, as the case may be. If so, the third party may use securities borrowed from us or others to settle such sales and may use securities received from us to close out any related short positions. We may also loan or pledge securities covered by this prospectus and an applicable prospectus supplement to third parties, who may sell the loaned securities or, in an event of default in the case of a pledge, sell the pledged securities pursuant to this prospectus and the applicable prospectus supplement or other offering materials, as the case may be.

A prospectus supplement with respect to each series of securities will state the terms of the offering of the securities, including:

- the terms of the offering;
- the name or names of any underwriters or agents and the amounts of securities underwritten or purchased by each of them, if any;
- the public offering price or purchase price of the securities and the net proceeds to be received by us from the sale;
- any delayed delivery arrangements;
- any initial public offering price;
- any underwriting discounts or agency fees and other items constituting underwriters' or agents' compensation;
- any discounts or concessions allowed or reallowed or paid to dealers; and
- any securities exchange on which the securities may be listed.

The offer and sale of the securities described in this prospectus by us, the underwriters or the third parties described above may be effected from time to time in one or more transactions, including privately negotiated transactions, either:

- at a fixed price or prices, which may be changed;
- in an "at the market" offering within the meaning of Rule 415(a)(4) of the Securities Act;
- at prices related to the prevailing market prices; or
- at negotiated prices.

General

Underwriters, dealers, agents and remarketing firms that participate in the distribution of the offered securities may be "underwriters" as defined in the Securities Act. Any discounts or commissions they receive from us and any profits they receive on the resale of the offered securities may be treated as underwriting discounts and commissions under the Securities Act. We will identify any underwriters, agents or dealers and describe their commissions, fees or discounts in the applicable prospectus supplement, as the case may be.

Underwriters and Agents

If underwriters are used in a sale, they will acquire the offered securities for their own account. The underwriters may resell the offered securities in one or more transactions, including negotiated transactions. These sales will be made at a fixed public offering price or at varying prices determined at the time of the sale. We may offer the securities to the public through an underwriting syndicate or through a single underwriter. The underwriters in any particular offering will be named in the applicable prospectus supplement or other offering materials, as the case may be.

Unless the applicable prospectus supplement states otherwise, the obligations of the underwriters to purchase the offered securities will be subject to certain conditions contained in an underwriting agreement that we will enter into with the underwriters at the time of the sale to them. The underwriters will be obligated to purchase all of the securities of the series offered if any of the securities are purchased, unless the applicable prospectus supplement says otherwise. Any initial public offering price and any discounts or concessions allowed, reallowed or paid to dealers may be changed from time to time.

We may designate agents to sell the offered securities. Unless the applicable prospectus supplement states otherwise, the agents will agree to use their best efforts to solicit purchases for the period of their appointment. We may also sell the offered securities to one or more remarketing firms, acting as principals for their own accounts or as agents for us. These firms will remarket the offered securities upon purchasing them in accordance with a redemption or repayment pursuant to the terms of the offered securities. A prospectus supplement or other offering materials, as the case may be, will identify any remarketing firm and will describe the terms of its agreement, if any, with us and its compensation.

In connection with offerings made through underwriters or agents, we may enter into agreements with such underwriters or agents pursuant to which we receive our outstanding securities in consideration for the securities being offered to the public for cash. In connection with these arrangements, the underwriters or agents may also sell securities covered by this prospectus to hedge their positions in these outstanding securities, including in short sale transactions. If so, the underwriters or agents may use the securities received from us under these arrangements to close out any related open borrowings of securities.

Dealers

We may sell the offered securities to dealers as principals. The dealer may then resell such securities to the public either at varying prices to be determined by the dealer or at a fixed offering price agreed to with us at the time of resale.

Direct Sales

We may choose to sell the offered securities directly. In this case, no underwriters or agents would be involved.

Institutional Purchasers

We may authorize agents, dealers or underwriters to solicit certain institutional investors to purchase offered securities on a delayed delivery basis pursuant to delayed delivery contracts providing for payment and delivery on a specified future date. The applicable prospectus supplement or other offering materials, as the case may be, will provide the details of any such arrangement, including the offering price and commissions payable on the solicitations.

We may enter into such delayed contracts only with institutional purchasers that we approve. These institutions may include commercial and savings banks, insurance companies, pension funds, investment companies and educational and charitable institutions.

Indemnification; Other Relationships

We may have agreements with agents, underwriters, dealers and remarketing firms to indemnify them against certain civil liabilities, including liabilities under the Securities Act. Agents, underwriters, dealers and remarketing firms, and their affiliates, may engage in transactions with, or perform services for, us in the ordinary course of business. This includes commercial banking and investment banking transactions.

Market-Making, Stabilization and Other Transactions

There is currently no market for any of the offered securities, other than our common stock which is traded on OTCQB. If the offered securities are traded after their initial issuance, they may trade at a discount from their initial offering price, depending upon prevailing interest rates, the market for similar securities and other factors. While it is possible that an underwriter could inform us that it intends to make a market in the offered securities, any such underwriter would not be obligated to do so, and any such market-making could be discontinued at any time without notice. Therefore, no assurance can be given as to whether an active trading market will develop for the offered securities. We have no current plans for listing of the warrants on any securities exchange or quotation system. Any such listing with respect to any particular warrants will be described in the applicable prospectus supplement or other offering materials, as the case may be.

Any underwriter may engage in stabilizing transactions, syndicate covering transactions and penalty bids in accordance with Rule 104 under the Exchange Act. Stabilizing transactions involve bids to purchase the underlying security in the open market for the purpose of pegging, fixing or maintaining the price of the securities. Syndicate covering transactions involve purchases of the securities in the open market after the distribution has been completed in order to cover syndicate short positions.

Penalty bids permit the underwriters to reclaim a selling concession from a syndicate member when the securities originally sold by the syndicate member are purchased in a syndicate covering transaction to cover syndicate short positions. Stabilizing transactions, syndicate covering transactions and penalty bids may cause the price of the securities to be higher than it would be in the absence of these transactions. The underwriters may, if they commence these transactions, discontinue them at any time.

LEGAL MATTERS

Unless otherwise specified in the applicable prospectus supplement, the validity of the securities offered by this prospectus will be passed upon for us by Greenberg Traurig, LLP, Sacramento, California. If legal matters in connection with offerings made by this prospectus are passed on by counsel for the underwriters, dealers or agents, if any, that counsel will be named in the applicable prospectus supplement.

EXPERTS

The consolidated financial statements incorporated in this Prospectus by reference to the Annual Report on Form 10-K for the year ended December 31, 2020, have been so incorporated in reliance on the report of Mayer Hoffman McCann P.C., an independent registered public accounting firm, given on the authority of said firm as experts in auditing and accounting.

WHERE YOU CAN FIND MORE INFORMATION

We file annual, quarterly current and other reports, proxy statements and other information with the SEC. Our SEC filings are available to the public over the Internet at the SEC's website at http://www.sec.gov. Our Annual Report on Form 10-K, Quarterly Reports on Form 10-Q, and Current Reports on Form 8-K, including any amendments to those reports, and other information that we file with or furnish to the SEC pursuant to Section 13(a) or 15(d) of the Exchange Act can also be accessed free of charge through the Internet. These filings will be available as soon as reasonably practicable after we electronically file such material with, or furnish it to, the SEC.

We have filed with the SEC a registration statement under the Securities Act relating to the offering of these securities. The registration statement, including the attached exhibits, contains additional relevant information about us and the securities. This prospectus does not contain all of the information set forth in the registration statement. You can obtain a copy of the registration statement, at prescribed rates, from the SEC at the address listed above. The registration statement and the documents referred to below under "Incorporation of Certain Documents by Reference" are also available on our website, http://www.skyebioscience.com. We have not incorporated by reference into this prospectus the information on our website, and you should not consider it to be a part of this prospectus.

INCORPORATION OF CERTAIN DOCUMENTS BY REFERENCE

The SEC allows us to "incorporate by reference" much of the information we file with it, which means that we can disclose important information to you by referring you to other documents we have filed or will file with the SEC. The information incorporated by reference is deemed to be part of this prospectus, and any of our subsequent filings with the SEC will automatically update and supersede this information. This means that you must look at all of the SEC filings that we incorporate by reference to determine if any statements in the prospectus or any document previously incorporated by reference have been modified or superseded. This prospectus incorporates by reference the documents set forth below (excluding any portions of any Current Report on Form 8-K that are not deemed "filed" pursuant to the General Instructions of Form 8-K):

- our Annual Report on Form 10-K for the year ended December 31, 2020, filed with the SEC on March 2, 2021;
- our Quarterly Report on Form 10-Q for the quarter ended March 31, 2021, filed with the SEC on May 7, 2021; and
- · our description of our Common Stock contained in the Registration Statement on Form 8-A12G filed with the SEC on January 28, 2014, including any subsequent amendments or reports filed for the purpose of updating such description.

We also incorporate by reference all documents we file in the future pursuant to Section 13(a), 13(c), 14 or 15(d) of the Exchange Act after the date of this prospectus and prior to the completion or termination of the offering (including all such documents filed with the SEC after the date of the initial filing of the registration statement that contains this prospectus and prior to effectiveness of the registration statement or after such effectiveness), except in each case the information contained in such document to the extent "furnished" and not "filed."

We will provide, upon written or oral request, to each person to whom a prospectus is delivered, including any beneficial owner, a copy of any or all of the information that has been incorporated by reference in the prospectus but not delivered with the prospectus. You may request a copy of these filings, at no cost, by writing or calling us at:

Skye Bioscience, Inc.
5910 Pacific Center Boulevard, Suite 320
San Diego, CA 92121
Attn: Chief Executive Officer
(858) 410-0266

You may also access the documents incorporated by reference in this prospectus through our website at http://www.skyebioscience.com. Except for the specific incorporated documents listed above, no information available on or through our website shall be deemed to be incorporated in this prospectus or the registration statement of which it forms a part.

You should rely only on the information incorporated by reference or provided in this prospectus or any supplement. We have not authorized anyone else to provide you with different information. You should not assume that information in this prospectus or any supplement is accurate as of any date other than the date on the front of these documents.



SKYE BIOSCIENCE, INC.

\$75,000,000

Common Stock Preferred Stock Warrants Units

PROSPECTUS

23

Part II

Information Not Required in the Prospectus

Item 14. Other Expenses of Issuance and Distribution

The following table sets forth an estimate of the costs and expenses, other than underwriting discounts and commissions, payable by the registrant in connection with the sale of the securities being registered. All of the amounts shown are estimates except the Securities and Exchange Commission registration fee and the FINRA filing fee.

SEC registration fee	\$ 8,182.50
FINRA filing fee	*
Accounting fees and expenses	*
Legal fees and expenses	*
Printing and engraving	*
Fees and expenses of the transfer agent or registrar	*
Miscellaneous	*
Total	\$ *

These fees and expenses depend on the securities offered and the number of issuances, and accordingly cannot be estimated at this time.

Item 15. Indemnification of Directors and Officers

Nevada Revised Statutes 78.138(7) provides that, subject to limited statutory exceptions and unless the articles of incorporation or an amendment thereto (in each case filed on or after October 1, 2003) provide for greater individual liability, a director or officer is not individually liable to a corporation or its stockholders or creditors for any damages as a result of any act or failure to act in his or her capacity as a director or officer unless it is proven that: (i) the act or failure to act constituted a breach of his or her fiduciary duties as a director or officer and (ii) the breach of those duties involved intentional misconduct, fraud or a knowing violation of law.

Our amended and restated articles of incorporation and amended and restated bylaws provide that we must indemnify and advance expenses to our directors and officers to the fullest extent permitted under the Nevada Revised Statutes.

We have entered into indemnification agreements with each of our directors and executive officers that require us to indemnify these persons against expenses, judgments, fines, settlements and other amounts actually and reasonably incurred (including expenses of a derivative action) in connection with any proceeding, whether actual or threatened, to which any such person may be made a party by reason of the fact that the person is or was a director or officer of our Company or any of our affiliated enterprises.

We also maintain any policy of directors' and officers' liability insurance that insures its directors and officers against the cost of defense, settlement or payment of a judgment under any circumstances.

Insofar as indemnification for liabilities arising under the Securities Act may be permitted to our directors, officers and controlling persons pursuant to the foregoing provisions, or otherwise, we have been advised that in the opinion of the SEC such indemnification is against public policy as expressed in the Act and is, therefore, unenforceable.

Item 16. Exhibits

A list of exhibits filed herewith is contained in the exhibit index that immediately precedes such exhibits, which exhibit index is incorporated herein by reference.

Item 17. Undertakings

- (a) The undersigned registrant hereby undertakes:
 - (1) To file, during any period in which offers or sales are being made, a post-effective amendment to this registration statement:
 - (i) To include any prospectus required by Section 10(a)(3) of the Securities Act of 1933;
 - (ii) To reflect in the prospectus any facts or events arising after the effective date of the registration statement (or the most recent post-effective amendment thereof) which, individually or in the aggregate, represent a fundamental change in the information set forth in the registration statement. Notwithstanding the foregoing, any increase or decrease in volume of securities offered (if the total dollar value of securities offered would not exceed that which was registered) and any deviation from the low or high end of the estimated maximum offering range may be reflected in the form of prospectus filed with the SEC pursuant to Rule 424(b) if, in the aggregate, the changes in volume and price represent no more than a 20 percent change in the maximum aggregate offering price set forth in the "Calculation of Registration Fee" table in the effective registration statement; and
 - (iii) To include any material information with respect to the plan of distribution not previously disclosed in the registration statement or any material change to such information in the registration statement;

provided, however; that paragraphs (a)(1)(i), (a)(1)(ii) and (a)(1)(iii) of this section do not apply if the information required to be included in a post-effective amendment by those paragraphs is contained in reports filed with or furnished to the SEC by the registrant pursuant to Section 13 or Section 15(d) of the Securities Exchange Act of 1934 that are incorporated by reference in the registration statement, or is contained in a form of prospectus filed pursuant to Rule 424(b) that is part of the registration statement

- (2) That, for the purpose of determining any liability under the Securities Act of 1933, each such post-effective amendment shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial bona fide offering thereof.
- (3) To remove from registration by means of a post-effective amendment any of the securities being registered which remain unsold at the termination of the offering.
- (4) That, for the purpose of determining liability under the Securities Act of 1933 to any purchaser:
 - (i) Each prospectus filed by the registrant pursuant to Rule 424(b)(3) shall be deemed to be part of the registration statement as of the date the filed prospectus was deemed part of and included in the registration statement; and

- (ii) Each prospectus required to be filed pursuant to Rule 424(b)(2), (b)(5) or (b)(7) as part of a registration statement in reliance on Rule 430B relating to an offering made pursuant to Rule 415(a)(1)(i), (vii) or (x) for the purpose of providing the information required by Section 10(a) of the Securities Act of 1933 shall be deemed to be part of and included in the registration statement as of the earlier of the date such form of prospectus is first used after effectiveness or the date of the first contract of sale of securities in the offering described in the prospectus. As provided in Rule 430B, for liability purposes of the issuer and any person that is at that date an underwriter, such date shall be deemed to be a new effective date of the registration statement relating to the securities in the registration statement to which that prospectus relates, and the offering of such securities at that time shall be deemed to be the initial bona fide offering thereof. Provided, however, that no statement made in a registration statement or prospectus that is part of the registration statement or made in a document incorporated or deemed incorporated by reference into the registration statement or prospectus that is part of the registration statement will, as to a purchaser with a time of contract of sale prior to such effective date, supersede or modify any statement that was made in the registration statement or prospectus that was part of the registration statement or made in any such document immediately prior to such effective date.
- (5) That, for the purpose of determining liability of the registrant under the Securities Act of 1933 to any purchaser in the initial distribution of the securities, the undersigned registrant undertakes that in a primary offering of securities of the undersigned registrant pursuant to this registration statement, regardless of the underwriting method used to sell the securities to the purchaser, if the securities are offered or sold to such purchaser by means of any of the following communications, the undersigned registrant will be a seller to the purchaser and will be considered to offer or sell such securities to such purchaser:
 - (i) Any preliminary prospectus or prospectus of the undersigned registrant relating to the offering required to be filed pursuant to Rule 424:
 - (ii) Any free writing prospectus relating to the offering prepared by or on behalf of the undersigned registrant or used or referred to by the undersigned registrant;
 - (iii) The portion of any other free writing prospectus relating to the offering containing material information about the undersigned registrant or its securities provided by or on behalf of the undersigned registrant; and
 - (iv) Any other communication that is an offer in the offering made by the undersigned registrant to the purchaser.
- (b) The undersigned registrant hereby undertakes that, for purposes of determining any liability under the Securities Act of 1933, each filing of the registrant's annual report pursuant to Section 13(a) or Section 15(d) of the Securities Exchange Act of 1934 (and, where applicable, each filing of an employee benefit plan's annual report pursuant to Section 15(d) of the Securities Exchange Act of 1934) that is incorporated by reference in the registration statement shall be deemed to be a new registration statement relating to the securities offered therein, and the offering of such securities at that time shall be deemed to be the initial *bona fide* offering thereof.
- (c) Insofar as indemnification for liabilities arising under the Securities Act of 1933 may be permitted to directors, officers and controlling persons of the registrant pursuant to the foregoing provisions, or otherwise, the registrant has been advised that in the opinion of the Securities and Exchange Commission such indemnification is against public policy as expressed in the Act and is, therefore, unenforceable. In the event that a claim for indemnification against such liabilities (other than the payment by the registrant of expenses incurred or paid by a director, officer or controlling person of the registrant in the successful defense of any action, suit or proceeding), is asserted by such director, officer or controlling person in connection with the securities being registered, the registrant will, unless in the opinion of its counsel the matter has been settled by controlling precedent, submit to a court of appropriate jurisdiction the question whether such indemnification by it is against public policy as expressed in the Act and will be governed by the final adjudication of such issue.

EXHIBIT INDEX

Exhibit Number	Description of Exhibit ¹
1.1 **	Form of Underwriting Agreement
<u>3.1</u>	Amended and Restated Articles of Incorporation (1)
<u>3.2</u>	Amended and Restated Bylaws (1)
4.1	Form of Warrants issued by Nemus to certain security holders to purchase an aggregate of 3,000,000 shares of commons stock (2)
4.2	Form of Warrants issued by Nemus to certain security holders to purchase an aggregate of 3,000,000 shares of commons stock ⁽²⁾
4.3	Form of Common Stock Purchase Warrant to certain security holders to purchase shares of common stock (3)
4.4	Form of Warrant dated April 24, 2015 issued by Nemus Bioscience, Inc. to holder to purchase 100,000 shares of common stock 4)
4.5	Form of Warrant dated April 28, 2015 issued by Nemus Bioscience, Inc. to holder to purchase 90,000 shares of common stock
4.6	Form of Warrant dated April 28, 2015 issued by Nemus Bioscience, Inc. to holder to purchase 6,000 shares of common stock
<u>4.7</u>	Form of Warrant dated June 8, 2015 issued by Nemus Bioscience, Inc. to holder to purchase 10,000 shares of common stock (6)
4.8	Form of Warrant to certain security holders to purchase shares of common stock (4)
4.9	Registration Rights Agreement, dated January 7, 2015, by and between Nemus Bioscience, Inc. and certain investors (7)
4.10	Form of Warrant ⁽⁸⁾
4.11	Form of Warrant (9)
4.12	Form of Common Warrant (11)
4.13	Form of Common Stock Purchase Warrant (11)
4.14 **	Form of Warrant Certificate
4.15**	Form of Warrant Agreement
4.16 **	Form of Certificate of Designation with respect to the Preferred Stock
4.10	Form of Certificate of Designation with respect to the Freience Stock
4.17**	Form of Unit Agreement (including form of Unit Certificate)
5.14	
5.1*	Opinion of Greenberg Traurig, LLP
23.1*	Consent of Mayer Hoffman McCann P.C., Independent Registered Public Accounting Firm
23.2*	Consent of Greenberg Traurig, LLP (included in Exhibit 5.1)
24.1*	D CAM ('.1.1.1') (1
24.1*	Power of Attorney (included on signature page to this Registration Statement on Form S-3)
· /	led as exhibit to our Annual Report on Form 10-K filed on March 2, 2021.

- (2) Included as exhibit to our Current Report on Form 8-K filed on November 3, 2014.
- (3) Included as exhibit to our Current Report on Form 8-K filed April 7, 2015.
- (4) Included as exhibit to our Current Report on Form 8-K filed August 20, 2015.
- (5) Included as exhibit to our Quarterly Report on Form 10-Q filed May 13, 2015
- (6) Included as exhibit to our Quarterly Report on Form 10-Q filed August 14, 2015
- (7) Included as exhibit to our Current Report on Form 8-K filed on January 9, 2015.
- (8) Included as exhibit to our Annual Report on Form 10-K filed on March 14, 2019.
- (9) Included as exhibit to our Quarterly Report on Form 10-Q filed on September 30, 2019.
- (10) Included as exhibit to our Current Report on Form 8-K filed on August 5, 2020.
- (11) Included as exhibit to our Current Report on Form 8-K filed on July 22, 2021.
- * Filed herewith
- ** To be filed by amendment or as an exhibit to a document incorporated by reference or deemed to be incorporated by reference in this registration statement, including a current report on Form 8-K, in connection with the offering of any securities, as appropriate.

SIGNATURES

Pursuant to the requirements of the Securities Act of 1933, the registrant has duly caused this registration statement to be signed on its behalf by the undersigned, thereunto duly authorized in San Diego, California, on this 28th day of July, 2021.

SKYE BIOSCIENCE, INC.

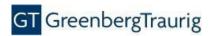
By: /s/ Punit Dhillon Punit Dhillon Chief Executive Officer, Chairman

POWER OF ATTORNEY

KNOW ALL MEN BY THESE PRESENTS, that each person whose signature appears below, hereby constitutes and appoints Punit Dhillon and Richard Janney, and each of them, his or her true and lawful attorneys-in-fact and agents, with full power of substitution and resubstitution, for him or her and in his or her name, place and stead, in any and all capacities, to sign any and all amendments to the registration statement, including post-effective amendments, and registration statements filed pursuant to Rule 462 under the Securities Act of 1933, and to file the same, with all exhibits thereto, and other documents in connection therewith, with the Securities and Exchange Commission, and does hereby grant unto said attorneys-in-fact and agents, and each of them, full power and authority to do and perform each and every act and thing requisite and necessary to be done in and about the foregoing, as fully to all intents and purposes as he or she might or could do in person, hereby ratifying and confirming all that said attorneys-in-fact and agents, or any of them, or their or his or her substitutes, may lawfully do or cause to be done by virtue hereof.

Pursuant to the requirements of the Securities Act, this Registration Statement has been signed by the following persons in the capacities and on the dates indicated:

By:	/s/ Punit Dhillon	July 28, 2021
	Punit Dhillon	
Its:	Chief Executive Officer, Chairman, Director	
	(Principal Executive Officer)	
D.,,	/a/ Dialand I ama	Index 29, 2021
By:	/s/ Richard Janney	July 28, 2021
Its:	Richard Janney	
	Interim Principal Accounting Officer	
	(Principal Financial and Accounting Officer)	
By:	/s/ Margaret Dalesandro	July 28, 2021
	Margaret Dalesandro	,,
Its:	Director	
113.	Director	
By:	/s/ Jim Heppell	July 28, 2021
Its:	Jim Heppell	
	Director	
D.	/a/ Puguaga Tula	July 29, 2021
By:	/s/ Praveen Tyle	July 28, 2021
Its:	Praveen Tyle	
	Director	
		28



July 28, 2021

Skye Bioscience, Inc. 5910 Pacific Center Boulevard Suite 320 San Diego, California 92121

Ladies and Gentlemen:

We have acted as legal counsel to Skye Bioscience, Inc., a Nevada corporation (the "Company"), in connection with the proposed issuance and sale from time to time by the Company of (i) shares of its common stock, \$0.001 par value per share (the "Common Stock"), (ii) shares of its preferred stock, \$0.001 par value per share (the "Preferred Stock"), (iii) warrants to purchase shares of Common Stock or Preferred Stock (the "Warrants"), and (iv) units consisting of any combination of the Common Stock, Preferred Stock, and Warrants (the "Units") with an aggregate offering price of up to \$75,000,000 (the Common Stock, Preferred Stock, Warrants and Units are each referred to herein as a "Security," and collectively as the "Securities"). The Securities may be issued and sold by the Company pursuant to a registration statement on Form S-3 (the "Registration Statement") being filed by the Company with the Securities and Exchange Commission (the "Commission") under the Securities Act of 1933, as amended (the "Act"), on the date hereof. The Registration Statement includes a base prospectus (the "Base Prospectus"), which provides that it will be supplemented in the future by one or more supplements to the Base Prospectus (each, a "Prospectus Supplement", and together with the Base Prospectus, a "Prospectus").

This opinion letter is being delivered in accordance with the requirements of Item 601(b)(5) of Regulation S-K under the Act, and no opinion is expressed herein as to any matter pertaining to the contents of the Registration Statement or such related applicable Prospectus, other than as expressly stated herein with respect to the sale and issuance of such respective Securities under such related applicable Prospectus.

We have made such legal and factual examinations and inquiries, including an examination of originals or copies certified or otherwise identified to our satisfaction of such documents, corporate records and instruments, as we have deemed necessary or appropriate for purposes of this opinion. We have also examined the Registration Statement as filed with the Commission in accordance with the provisions of the Act, and the rules and regulations of the Commission thereunder. In our examination, we have assumed the genuineness of all signatures, the authenticity of all documents submitted to us as originals and the conformity to authentic original documents of all documents submitted to us as copies. As to facts material to the opinions, statements and assumptions expressed herein, we have, with your consent, relied upon oral or written statements and representations of officers and other representatives of the Company and others. We have not independently verified such factual matters.

In expressing our opinions below, we have assumed, with your consent, that:

- (a) the Registration Statement (including any and all required post-effective amendments thereto) will have become effective under the Act and will comply with all applicable laws;
- (b) the Registration Statement (including any and all required post-effective amendments thereto) will be effective under the Act and will comply with all applicable laws at the time the Securities are offered or sold as contemplated by the Registration Statement (including any and all required post-effective amendments thereto), the Base Prospectus, and the applicable Prospectus Supplement(s);

- (c) no stop order suspending the effectiveness of the Registration Statement (including any and all required post-effective amendments thereto) will have been issued and remain in effect;
- a Prospectus Supplement describing the Securities offered thereby and the offering thereof and complying with all applicable laws will have been prepared and filed with the Commission;
- (e) the Securities will be offered and sold in the form and with the terms set forth in the Registration Statement (including any and all required post-effective amendments thereto), the Base Prospectus, and the applicable Prospectus Supplement(s) and the organizational documents of the Company, as applicable;
- (f) the Securities will be offered and sold in compliance with all applicable federal and state securities laws and in the manner stated in the Registration Statement (including any and all required post-effective amendments thereto), the Base Prospectus, and the applicable Prospectus Supplement(s);
- (g) the Company will have obtained any and all legally required consents, approvals, authorizations and other orders of the Commission and any and all other regulatory authorities and other third parties necessary to offer and sell the Securities being offered;
- (h) a definitive purchase, underwriting, sales or similar agreement (each a "Purchase Agreement") with respect to any Securities offered and sold will have been duly authorized and validly executed and delivered by the Company and the other parties thereto; and
- (i) any Securities or other securities issuable upon conversion, exchange or exercise of any Security being offered and sold will be duly authorized, created and, if appropriate, reserved for issuance upon such conversion, exchange or exercise.

Our opinions expressed in paragraphs 3 and 4 below are subject to the qualifications that we express no opinion as to the applicability of, compliance with or effect of: (i) any bankruptcy, insolvency, reorganization, preference, fraudulent conveyance, fraudulent transfer, moratorium or other similar laws relating to or affecting the rights and remedies of creditors generally; (ii) general principles of equity, whether considered in a proceeding in equity or at law (including the possible unavailability of specific performance or injunctive relief), concepts of materiality, reasonableness, good faith and fair dealing, and the discretion of the court before which a proceeding is brought; or (iii) public policy considerations that may limit the rights of parties to obtain certain remedies.

We express no opinion as to (i) any provision providing for the indemnification of or contribution to a party with respect to a liability where such indemnification or contribution is contrary to public policy, (ii) consents to, or restrictions upon, governing law, jurisdiction, venue, arbitration, remedies or judicial relief, (iii) the securities or "Blue Sky" laws of any state to the offer or sale of the Securities and (iv) the antifraud provisions of the securities or other laws of any jurisdiction.

Subject to the foregoing and the other matters set forth herein, it is our opinion that, as of the date hereof:

- 1. When an issuance of Common Stock has been duly authorized by all necessary corporate action of the Company, upon issuance, delivery and payment therefor in an amount not less than the par value thereof in the manner contemplated by the applicable Prospectus and by such corporate action, and in total amounts and numbers of shares that do not exceed the respective total amounts and numbers of shares (a) available under the Company's articles of incorporation (as amended, the "Articles of Incorporation") and (b) authorized by the board of directors of the Company in connection with the offering contemplated by the applicable Prospectus, such shares of Common Stock will be validly issued, fully paid and nonassessable.
- 2. When a series of Preferred Stock has been duly established in accordance with the terms of the Articles of Incorporation and authorized by all necessary corporate action of the Company, upon issuance, delivery and payment therefor in an amount not less than the par value thereof in the manner contemplated by the applicable Prospectus and by such corporate action, and in total amounts and numbers of shares that do not exceed the respective total amounts and numbers of shares (a) available under the Articles of Incorporation and (b) authorized by the board of directors of the Company in connection with the offering contemplated by the applicable Prospectus, such shares of such series of Preferred Stock will be validly issued, fully paid and nonassessable.

- 3. When (a) a warrant agreement, if any, has been duly authorized by all necessary corporate action of the Company (including, without limitation, the adoption by the board of directors of the Company of a resolution in form and content as required by applicable law duly authorizing the execution and delivery of such warrant agreement) and duly executed and delivered by the Company, (b) the specific terms of a particular issuance of Warrants have been duly established in accordance with such warrant agreement, if any, and all applicable law and authorized by all necessary corporate action of the Company (including, without limitation, the adoption by the board of directors of the Company of a resolution in form and content as required by applicable law duly authorizing the issuance and delivery of the Warrants), and (c) the Warrants have been duly executed, issued and delivered against payment therefor in accordance with such warrant agreement, if any, the applicable Purchase Agreement and all applicable law (and assuming the satisfaction of the conditions described in the applicable numbered paragraphs of this opinion letter with respect to our opinion regarding any Securities issuable upon exercise of the Warrants), such Warrants will be legally valid and binding obligations of the Company, enforceable against the Company in accordance with their terms.
- 4. When (a) a unit agreement has been duly authorized by all necessary corporate action of the Company (including, without limitation, the adoption by the board of directors of the Company of a resolution duly authorizing the execution and delivery of such unit agreement) and duly executed and delivered by the Company, (b) the specific terms of a particular issuance of the related Units have been duly established in accordance with such unit agreement and all applicable law and authorized by all necessary corporate action of the Company (including, without limitation, the adoption by the board of directors of the Company of a resolution in form and content as required by applicable law duly authorizing the issuance and delivery of the Units), and such (c) Units have been duly executed, issued and delivered against payment therefor in accordance with such unit agreement, the applicable Purchase Agreement and all applicable law (and assuming the satisfaction of the conditions described in the applicable numbered paragraphs of this opinion letter with respect to our opinion regarding the Securities underlying the Units), such Units will be legally valid and binding obligations of the Company, enforceable against the Company in accordance with their terms.

This opinion is opining upon and is limited to the current federal laws of the United States and the Nevada Revised Statutes as such laws presently exist and to facts as they presently exist. We express no opinion with respect to the effect or applicability of the laws of any other jurisdiction.

This opinion is being delivered solely for the benefit of the Company and such other persons as are entitled to rely upon it pursuant to applicable provisions of the Act. This opinion may not be used, quoted, relied upon or referred to for any other purpose nor may this opinion be used, quoted, relied upon or referred to by any other person, for any purpose, without our prior written consent.

We consent to your filing this opinion as an exhibit to the Registration Statement and to the reference to our firm in the applicable Prospectus under the heading "Legal Matters." In giving such consent, we do not thereby admit that we are in the category of persons whose consent is required under Section 7 of the Act or the rules and regulations of the Commission thereunder.

Very truly yours,

/s/ Greenberg Traurig, LLP

Greenberg Traurig, LLP

CONSENT OF INDEPENDENT REGISTERED PUBLIC ACCOUNTING FIRM

As independent registered public accountants, we consent to the incorporation by reference in this Registration Statement on Form S-3 of our report dated March 1, 2021, relating to the consolidated financial statements of Skye Bioscience, Inc., formerly known as Emerald Bioscience, Inc., and Subsidiaries as of and for the years ended December 31, 2020 and 2019 (which report includes an explanatory paragraph relating to the uncertainty of the Company's ability to continue as a going concern).

We also consent to the reference to our Firm under the caption "Experts" in the Prospectus, which is part of said Registration Statement.

/s/ Mayer Hoffman McCann P.C.

Irvine, California July 28, 2021