UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

SCHEDULE 13G

Under the Securities Exchange Act of 1934 (Amendment No. 2)*

Lyra Therapeutics, Inc.

(Name of Issuer)

Common Stock, \$0.001 par value per share (Title of Class of Securities)

> 55234L105 (CUSIP Number)

December 31, 2021 (Date of Event which Requires Filing of this Statement)

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

⊠ Rule 13d-1(b)

□ Rule 13d-1(c)

□ Rule 13d-1(d)

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

The information required in the remainder of this cover page shall not be deemed to be "filed" for the purpose of Section 18 of the Securities Exchange Act of 1934, as amended ("*Act*") or otherwise subject to the liabilities of that section of the Act but shall be subject to all other provisions of the Act (however, see the *Notes*).

Name, Address and Telephone Number of Person Authorized to Receive Notices and Communications:

Evan K. Hall, Esq. Haynes and Boone, LLP 2323 Victory Avenue, Suite 700 Dallas, Texas 75219-7673 (214) 651-5000

| CUSIP No. 55234L105 | | | | | | | |
|---|---|-------|--------------------------------------|--|--|--|--|
| 1 | 1 Names of Reporting Persons | | | | | | |
| | Ikarian Capital, LLC | | | | | | |
| 2 Check the appropriate box if a member of a Group (see instructions) (a) □ (b) ⊠ | | | | | | | |
| | | | | | | | |
| 3 Sec Use Only | | | y | | | | |
| | | | | | | | |
| 4 Citizenship or Place of Organization | | ip (| or Place of Organization | | | | |
| | Delawa | | | | | | |
| | | 5 | Sole Voting Power | | | | |
| N | umber of | | 0 | | | | |
| | Shares neficially | 6 | Shared Voting Power | | | | |
| | wned by | | 394,536 (1) | | | | |
| R | Each eporting | 7 | Sole Dispositive Power | | | | |
| | Person With: | | 0 | | | | |
| | with: | 8 | Shared Dispositive Power | | | | |
| | | | 394,536 (1) | | | | |
| 9 | Aggregate Amount Beneficially Owned by Each Reporting Person | | | | | | |
| | 394,536 (1) | | | | | | |
| 10 | 0 Check box if the aggregate amount in row (9) excludes certain shares (See Instructions) | | | | | | |
| | | | | | | | |
| 11 | Percent | of cl | ass represented by amount in row (9) | | | | |
| | 3.0% (1)(2) | | | | | | |
| 12 | 12 Type of Reporting Person (See Instructions) | | | | | | |
| | IA | | | | | | |

(1) Represents shares held by Ikarian Healthcare Master Fund, L.P., a Cayman Islands exempted limited partnership, and certain separately managed accounts. *See* Item 2 for more information.

(2) Based upon 13,004,578 shares of Common Stock, \$0.001 par value per share, outstanding as of November 1, 2021, as disclosed in the Issuer's quarterly report on Form 10-Q for the quarterly period ended September 30, 2021, filed by the Issuer with the U.S. Securities and Exchange Commission on November 9, 2021.

| CUSIP No. 55234L105 | | | | | | | | |
|---|--|------|---|--|--|--|--|--|
| 1 | 1 Names of Reporting Persons | | | | | | | |
| Chart Westcott | | | | | | | | |
| 2 Check the appropriate box if a member of a Group (see instructions) (a) □ (b) ⊠ | | | | | | | | |
| | | | | | | | | |
| 3 | 3 Sec Use Only | | | | | | | |
| 4 Citizenship or Place of Organization | | | | | | | | |
| | United | | tates | | | | | |
| | | 5 | Sole Voting Power | | | | | |
| N | umber of | | 0 | | | | | |
| | Shares neficially | 6 | Shared Voting Power | | | | | |
| | wned by | | 394,536 (1) | | | | | |
| R | Each eporting | 7 | Sole Dispositive Power | | | | | |
| | Person With: | | 0 | | | | | |
| | with: | 8 | Shared Dispositive Power | | | | | |
| | | | 394,536 (1) | | | | | |
| 9 | Aggrega | te A | mount Beneficially Owned by Each Reporting Person | | | | | |
| | 394,536 (1) | | | | | | | |
| 10 | 10 Check box if the aggregate amount in row (9) excludes certain shares (See Instructions) | | | | | | | |
| | | | | | | | | |
| 11 | 11 Percent of class represented by amount in row (9) | | | | | | | |
| 3.0% (1)(2) | | | | | | | | |
| 12 | 12 Type of Reporting Person (See Instructions) | | | | | | | |
| | IN; HC | | | | | | | |

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| CUSIP No. 55234L105 | | | | | | | |
|---|--|---|--|--|--|--|--|
| 1 Names of Reporting Persons | | | | | | | |
| Neil Shahrestani | | | | | | | |
| 2 Check the appropriate box if a member of a Group (see instructions) (a) □ (b) ⊠ | | | | | | | |
| | | | | | | | |
| 3 Sec Use Only | | | | | | | |
| 4 Citizensl | nip (| or Place of Organization | | | | | |
| United | ted States | | | | | | |
| | 5 | Sole Voting Power | | | | | |
| Number of | | 0 | | | | | |
| Shares Beneficially | 6 | Shared Voting Power | | | | | |
| Owned by | | 394,536 (1) | | | | | |
| Each Reporting | 7 | Sole Dispositive Power | | | | | |
| Person With: | | 0 | | | | | |
| , , , , , , , , , , , , , , , , , , , | 8 | Shared Dispositive Power | | | | | |
| | | 394,536 (1) | | | | | |
| 9 Aggrega | te A | mount Beneficially Owned by Each Reporting Person | | | | | |
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| 10 Check b | 10 Check box if the aggregate amount in row (9) excludes certain shares (See Instructions) | | | | | | |
| | | | | | | | |
| 11 Percent | ot cl | ass represented by amount in row (9) | | | | | |
| 3.0% (1)(2) | | | | | | | |
| 12 Type of Reporting Person (See Instructions) | | | | | | | |
| IN; HC | | | | | | | |

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Item 1.

(a) Name of Issuer:

Lyra Therapeutics, Inc.

(b) Address of Issuer's Principal Executive Offices:

480 Arsenal Way, Watertown, MA 02472

Item 2.

(a) Name of Person Filing:

This statement is filed jointly by and on behalf of each of Ikarian Capital, LLC, a Delaware limited liability company ("*Ikarian Capital*"), Chart Westcott and Neil Shahrestani (collectively referred herein as the "*Reporting Persons*"). Ikarian Healthcare Master Fund, L.P., a Cayman Islands exempted limited partnership (the "*Fund*"), and certain separately managed accounts managed by Ikarian Capital (collectively, the "*Managed Accounts*") are the record owners of the securities covered by this statement. Ikarian Capital is an investment adviser registered under the Investment Advisers Act of 1940, as amended, and serves as investment manager to the Fund and as sub-adviser to the Managed Accounts, and may be deemed to have beneficial ownership of the securities covered by this statement through the investment discretion it has over the Fund and the Managed Accounts. Ikarian Capital is ultimately controlled, indirectly, by Mr. Westcott, and Mr. Shahrestani. Accordingly, each of Mr. Westcott and Mr. Shahrestani may be deemed to indirectly beneficially own securities beneficially owned by Ikarian Capital. The Fund disclaims beneficial ownership of the shares held by the Managed Accounts. The Managed Accounts disclaim beneficial ownership of the shares held by the Fund.

Each Reporting Person declares that neither the filing of this statement nor anything herein shall be construed as an admission that such person is, for the purposes of Section 13(d) or 13(g) of the Act or any other purpose, the beneficial owner of any securities covered by this statement.

Each Reporting Person may be deemed to be a member of a group with respect to the issuer or securities of the issuer for the purpose of Section 13(d) or 13(g) of the Act. Each of the Reporting Persons declares that neither the filing of this statement nor anything herein shall be construed as an admission that such person is, for the purpose of Section 13(d) or 13(g) of the Act or any other purpose, (i) acting (or has agreed or is agreeing to act together with any other person) as a partnership, limited partnership, syndicate, or other group for the purpose of acquiring, holding, or disposing of securities of the issuer or otherwise with respect to the issuer or any securities of the issuer or (ii) a member of any group with respect to the issuer or any securities of the issuer.

(b) Address of Principal Business Office or, if None, Residence:

The address of the principal business office of each of the Reporting Persons is c/o Ikarian Capital, LLC, 100 Crescent Court, Suite 1620, Dallas, Texas 75201.

(c) Citizenship:

See Item 4 on the cover page(s) hereto.

| (d) | Title and Class of Securities: | | | | | |
|---------|--|----------|--|--|--|--|
| | Common Stock, \$0.001 par value per share. | | | | | |
| (e) | CUSIP No.: 55234L105 | | | | | |
| Item 3. | ent is filed pursuant to §§ 240.13d-1(b) or 240.13d-2(b) or (c), check whether the person filing is a: | | | | | |
| | (a) | | Broker or dealer registered under Section 15 of the Act; | | | |
| | (b) | | Bank as defined in Section 3(a)(6) of the Act; | | | |
| | (c) | | Insurance company as defined in Section 3(a)(19) of the Act; | | | |
| | (d) | | Investment company registered under Section 8 of the Investment Company Act of 1940; | | | |
| | (e) | \times | An investment adviser in accordance with Rule 13d-1(b)(1)(ii)(E); | | | |
| | (f) | | An employee benefit plan or endowment fund in accordance with Rule 13d-1(b)(1)(ii)(F); | | | |
| | (g) | \times | A parent holding company or control person in accordance with Rule 13d-1(b)(1)(ii)(G); | | | |
| | (h) | | A savings associations as defined in Section 3(b) of the Federal Deposit Insurance Act (12 U.S.C. 1813); | | | |
| | (i) | | A church plan that is excluded from the definition of an investment company under section 3(c)(14) of the Investment Company Act of 1940; | | | |
| | (j) | | A non-U.S. institution in accordance with Rule 240.13d-1(b)(1)(ii)(J); | | | |
| | (k) | | Group, in accordance with Rule 240.13d-1(b)(1)(ii)(K). If filing as a non-U.S. institution in accordance with Rule 240.13d-1(b)(1)(ii)(J), please specify the type of institution: | | | |
| Item 4. | Ownership | | | | | |

The information set forth in Item 2(a) is incorporated by reference herein.

- (a) Amount Beneficially Owned as of December 31, 2021: See Item 9 on the cover pages(s) hereto.
- (b) **Percent of Class as of December 31, 2021:** See Item 11 on the cover page(s) hereto.
- (c) As of December 31, 2021, number of shares as to which such person has:
 - (i) Sole power to vote or to direct the vote: See Item 5 on the cover pages hereto.
 - (ii) Shared power to vote or to direct the vote: See Item 6 on the cover pages hereto.
 - (iii) Sole power to dispose or to direct the disposition of: See Item 7 on the cover pages hereto.
 - (iv) Shared power to dispose or to direct the disposition of: See Item 8 on the cover pages hereto.

Item 5. Ownership of Five Percent or Less of a Class.

If this statement is being filed to report the fact that as of the date hereof the Reporting Persons have ceased to be the beneficial owners of more than five percent of the class of securities, check the following 🗵.

Item 6. Ownership of more than Five Percent on Behalf of Another Person.

The information set forth in Item 2(a) is incorporated by reference herein.

Item 7. Identification and classification of the subsidiary which acquired the security being reported on by the parent holding company or control person.

Not applicable.

Item 8. Identification and classification of members of the group.

Not applicable.

Item 9. Notice of Dissolution of Group.

Not applicable.

Item 10. Certifications.

By signing below I certify that, to the best of my knowledge and belief, the securities referred to above were not acquired and are not held for the purpose of or with the effect of changing or influencing the control of the issuer of the securities and were not acquired and are not held in connection with or as a participant in any transaction having that purpose or effect other than activities solely in connection with a nomination under § 240.14a-11.

SIGNATURE

After reasonable inquiry and to the best of my knowledge and belief, I certify that the information set forth in this statement is true, complete and correct.

Dated: February 14, 2022

IKARIAN CAPITAL, LLC

By: Chart Westcott Living Trust Its: Manager

By: /s/ Chart Westcott

Chart Westcott, Trustee

CHART WESTCOTT

/s/ Chart Westcott

NEIL SHAHRESTANI

/s/ Neil Shahrestani

EXHIBIT INDEX

Exhibit Description of Exhibit

99.1 Joint Filing Agreement (filed herewith).

JOINT FILING AGREEMENT

February 14, 2022

Pursuant to and in accordance with the Securities Exchange Act of 1934, as amended (the "*Act*"), and the rules and regulations thereunder, each party hereto hereby agrees to the joint filing, on behalf of each of them, of any filing required by such party under Section 13 or Section 16 of the Act or any rule or regulation thereunder (including any amendment, restatement, supplement, and/or exhibit thereto) with the Securities and Exchange Commission (and, if such security is registered on a national securities exchange, also with the exchange), and further agrees to the filing, furnishing, and/or incorporation by reference of this Joint Filing Agreement as an exhibit thereto. This Joint Filing Agreement shall remain in full force and effect until revoked by any party hereto in a signed writing provided to each other party hereto, and then only with respect to such revoking party.

IN WITNESS WHEREOF, each party hereto, being duly authorized, has caused this Joint Filing Agreement to be executed and effective as of the date first written above.

IKARIAN CAPITAL, LLC

By: Chart Westcott Living Trust Its: Manager

By: /s/ Chart Westcott Chart Westcott, Trustee

CHART WESTCOTT

/s/ Chart Westcott

NEIL SHAHRESTANI

/s/ Neil Shahrestani