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

SCHEDULE 13G

Under the Securities Exchange Act of 1934

(Amendment No.)*

| Homology Medicines, Inc. | | | | | | |
|---------------------------------------------------------|--|--|--|--|--|--|
| (Name of Issuer) | | | | | | |
| | | | | | | |
| | | | | | | |
| Common Stock par value \$0.0001 per share | | | | | | |
| (Title of Class of Securities) | | | | | | |
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| | | | | | | |
| 438083 10 7 | | | | | | |
| (CUSIP Number) | | | | | | |
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| | | | | | | |
| December 31, 2018 | | | | | | |
| (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 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 ("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).

| CUSIP No. 438083 10 7 | | | | 13G | Page 2 of 14 Pages |
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| 1 | I.R.S. | IDENTI | EPORTING PI FICATION NO Fund VIII, L.P. | ERSONS OS. OF ABOVE PERSONS (ENTITIES ONLY) | |
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| 9 | AGGREGATE AMOUNT BENEFICIALLY OWNED BY EACH REPORTING PERSON 6,068,695 | | | | PERSON |
| 10 | CHECK IF THE AGGREGATE AMOUNT IN ROW (9) EXCLUDES CERTAIN SHARES* | | | | |
| 11 | PERCENT OF CLASS REPRESENTED BY AMOUNT IN ROW (9) 16.2% | | | | |
| 12 | TYPE PN | OF REI | PORTING PER | SON* | |

| CUSIP No. 438083 10 7 | | | | 13G | Page 3 of 14 Pages | |
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| CUSIP No. 438083 10 7 | | | | 13G | Page 7 of 14 Pages | |
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| CUSIP No. 438083 10 7 | | | | 13G | Page 8 of 14 Pages | |
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| CUSIP No. 438083 10 7 | 13G | Page 9 of 14 Pages |
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Item 1(a). Name of Issuer

Homology Medicines, Inc. (the "Issuer").

Item 1(b). Address of Issuer's Principal Executive Offices

1 Patriots Park, Bedford, MA 01730

Item 2(a). Name of Person Filing

ARCH Venture Fund VIII, L.P. ("ARCH Venture Fund VIII"); ARCH Venture Fund VIII Overage, L.P. ("AVF VIII Overage LP"); ARCH Venture Partners VIII, L.P. ("AVP VIII LP"); ARCH Venture Partners VIII, LLC ("AVP VIII LLC") (collectively, the "Reporting Entities" and individually, each a "Reporting Entity"); and Keith Crandell ("Crandell"), Robert Nelsen ("Nelsen") and Clinton Bybee ("Bybee") (collectively, the "Managing Directors" and individually, each a "Managing Director"). The Reporting Entities and the Managing Directors collectively are referred to as the "Reporting Persons".

Item 2(b). Address of Principal Business Office or, if none, Residence

8755 W. Higgins Avenue, Suite 1025, Chicago, IL 60631

Item 2(c). Citizenship

ARCH Venture Fund VIII, AVF VIII Overage LP and AVP VIII LP are limited partnerships organized under the laws of the State of Delaware. AVP VIII LLC is a limited liability company organized under the laws of the State of Delaware. Each Managing Director is a US citizen.

Item 2(d). Title of Class of Securities

Common stock, par value \$0.0001 per share.

Item 2(e). CUSIP Number

438083107

Item 3. If this statement is filed pursuant to Rule 13d-1(b), or 13d-2(b) or (c), Check Whether the Person Filing is a:

Not Applicable.

Item 4. Ownership

(a) Amount beneficially owned:

ARCH Venture Fund VIII is the record owner of 4,871,868 shares of Common Stock (the "ARCH VIII Shares") as of December 31, 2018. AVP VIII LP, as the sole general partner of ARCH Venture Fund VIII, may be deemed to beneficially own the ARCH VIII Shares. AVF VIII Overage LP is the record owner of 1,196,827 shares of Common Stock (the "Overage Shares"; combined with ARCH VIII Shares, the "Record Shares") as of December 31, 2018. AVP VIII LLC, as the sole general partner of AVP VIII LP and AVF VIII Overage LP, may be deemed to beneficially own the Record Shares. As managing directors of AVP VIII LLC, each Managing Director may also be deemed to share the power to direct the disposition and vote of the Record Shares.

(b) Percent of class:

See line 2 of the cover sheets. The percentages set forth on the cover sheet for each Reporting Person is based upon 37,495,773 shares of common stock outstanding as of November 5, 2018 as reported on the Issuer's Form 10-Q as filed with the Securities and Exchange Commission on November 13, 2018.

| CUSIP | No. 43 | 8083 10 7 | 13G | Page 10 of 14 Pages | | | | | | | |
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| | | | | | | | | | | | |
| | (c) | Number of shares as to which such | ch person has: | | | | | | | | |
| | | (i) Sole power to vote or to dire | ect the vote: | | | | | | | | |
| | | See line 5 of the cover sheets | 5. | | | | | | | | |
| | | (ii) Shared power to vote or to d | lirect the vote: | | | | | | | | |
| | | See line 6 of the cover sheets | 5. | | | | | | | | |
| | | (iii) Sole power to dispose or to | direct the disposition: | | | | | | | | |
| | | See line 7 of the cover sheet | S. | | | | | | | | |
| | | (iv) Shared power to dispose or t | to direct the disposition: | | | | | | | | |
| | | See line 8 of the cover sheet: | S. | | | | | | | | |
| Each Repo | rting Per | rson disclaims beneficial ownership | of such shares of Common Stock except for the share | res, if any, such Reporting Person holds of record. | | | | | | | |
| Item 5. | Owne | ership of Five Percent or Less of a | Class | | | | | | | | |
| | Not A | pplicable. | | | | | | | | | |
| Item 6. | Owne | ershin of More Than Five Percent | on Behalf of Another Person | | | | | | | | |
| rem o. | Ownership of More Than Five Percent on Behalf of Another Person Not Applicable. | | | | | | | | | | |
| | | | | | | | | | | | |
| Item 7. | Identi | Identification and Classification of the Subsidiary Which Acquired the Security Being Reported on By the Parent Holding Company | | | | | | | | | |
| | Not A | pplicable. | | | | | | | | | |
| Item 8. | Identi | ification and Classification of Men | nbers of the Group | | | | | | | | |
| | Not A | pplicable. | | | | | | | | | |
| Item 9. | Notice | e of Dissolution of Group | | | | | | | | | |
| Tem 5. | | pplicable. | | | | | | | | | |
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| | Certif | fication | | | | | | | | | |
| Item 10. | | | | | | | | | | | |

| CUSIP No. 438083 10 7 | 13G |
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| | |
| | <u>SIGNATURE</u> |
| After reasonable inquiry and to the best of my knowle | edge and belief, I certify that the information set forth |
| Dated: February 14, 2019 | |
| | ARCH VENTURE FUND VIII, L.P. |
| | By: ARCH Venture Partners VIII, L.P. its General Partner |
| | By: ARCH Venture Partners VIII, LLC its General Partner |
| | By:* |
| | Keith Crandell Managing Director |
| | ARCH VENTURE PARTNERS VIII, L.P. |
| | By: ARCH Venture Partners VIII, LLC its General Partner |
| | By:* |
| | Keith Crandell Managing Director |
| | ARCH VENTURE PARTNERS VIII, LLC |
| | By:* |
| | Keith Crandell Managing Director |
| | * |
| | Keith Crandell |
| | * |
| | Robert Nelsen |
| | * |
| | Clinton Bybee |
| | ARCH VENTURE FUND VIII OVERAGE, L.P. |
| | By: ARCH Venture Partners VIII, LLC |

* By: /s/ Mark McDonnell

Mark McDonnell as
Attorney-in-Fact

This Schedule 13G was executed by Mark McDonnell pursuant to Powers of Attorney attached hereto as Exhibit 2.0 and Exhibit 2.1 and incorporated herein by reference.

its General Partner

Keith Crandell Managing Director

By:

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in this statement is true, complete and correct.

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Exhibit 1

AGREEMENT

Pursuant to Rule 13d-1-(k)(1) under the Securities Exchange Act of 1934, the undersigned hereby agree that only one statement containing the information required by Schedule 13G need be filed with respect to the ownership by each of the undersigned of shares of stock of Homology Medicines, Inc.

This Agreement may be executed in any number of counterparts, each of which shall be deemed an original.

| Dated: | February 14, 2019 | ARCH VENTURE FUND VIII, L.P. |
|--------|-------------------|----------------------------------------------------------|
| | | By: ARCH Venture Partners VIII, L.P. its General Partner |
| | | By: ARCH Venture Partners VIII, LLC its General Partner |
| | | By:* |
| | | Keith Crandell Managing Director |
| | | ARCH VENTURE PARTNERS VIII, L.P. |
| | | By: ARCH Venture Partners VIII, LLC its General Partner |
| | | By:* |
| | | Keith Crandell |
| | | Managing Director |
| | | ARCH VENTURE PARTNERS VIII, LLC |
| | | By:* |
| | | Keith Crandell |
| | | Managing Director |
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| | | Keith Crandell |
| | | * |
| | | Robert Nelsen |
| | | * |
| | | Clinton Bybee |
| | | ARCH VENTURE FUND VIII OVERAGE, L.P. |
| | | By: ARCH Venture Partners VIII, LLC |
| | | its General Partner |
| | | By:* |
| | | Keith Crandell |
| | | Managing Director |

* By: /s/ Mark McDonnell

Mark McDonnell as

Attorney-in-Fact

This Agreement was executed by Mark McDonnell pursuant to Powers of Attorney attached hereto as Exhibit 2.0 and Exhibit 2.1 and incorporated herein by reference.

CUSIP No. 438083 10 7 13G Page 13 of 14 Pages

Exhibit 2.0

POWERS OF ATTORNEY

KNOW ALL MEN BY THESE PRESENTS, that each person whose signature appears below hereby constitutes and appoints Mark McDonnell his true and lawful attorney-in-fact, with full power of substitution, to sign any and all instruments, certificates and documents that may be necessary, desirable or appropriate to be executed on behalf of himself as an individual or in his capacity as a direct or indirect general partner, member, director, officer or manager of any partnership, corporation or limited liability company, pursuant to section 13 or 16 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), and any and all regulations promulgated thereunder, and to file the same, with all exhibits thereto, and any other documents in connection therewith, with the Securities and Exchange Commission, and with any other entity when and if such is mandated by the Exchange Act or by the Financial Industry Regulatory Authority, granting unto said attorney-in-fact full power and authority to do and perform each and every act and thing necessary, desirable or appropriate, fully to all intents and purposes as he might or could do in person, thereby ratifying and confirming all that said attorney-in-fact, or his substitutes, may lawfully do or cause to be done by virtue hereof. This Power of Attorney shall remain in full force and effect with respect to each undersigned person unless and until six months after such person is both no longer a Managing Director of ARCH Venture Partners and no longer serving on the board of directors of any portfolio company of any ARCH Venture Partners fund.

IN WITNESS WHEREOF, this Power of Attorney has been signed as of the 9th day of March, 2015.

ARCH VENTURE FUND VIII, L.P.

By: ARCH Venture Partners VIII, L.P. its General Partner

By: ARCH Venture Partners VIII, LLC its General Partner

By: <u>/s/ Keith Crandell</u>
Managing Director

ARCH VENTURE PARTNERS VIII, L.P.

By: ARCH Venture Partners VIII, LLC its General Partner

By: /s/ Keith Crandell

Managing Director

ARCH VENTURE PARTNERS VIII, LLC

By: <u>/s/ Keith Crandell</u>
Managing Director

/s/ Keith Crandell Keith Crandell

/s/ Robert Nelsen

/s/ Clinton Bybee

Clinton Bybee

/s/ Kristina Burow Kristina Burow

/s/ Paul Thurk

Paul Thurk

CUSIP No. 438083 10 7 13G Page 14 of 14 Pages

Exhibit 2.1

POWERS OF ATTORNEY

KNOW ALL MEN BY THESE PRESENTS, that each person whose signature appears below hereby constitutes and appoints Mark McDonnell his true and lawful attorney-in-fact, with full power of substitution, to sign any and all instruments, certificates and documents that may be necessary, desirable or appropriate to be executed on behalf of himself as an individual or in his capacity as a direct or indirect general partner, member, director, officer or manager of any partnership, corporation or limited liability company, pursuant to section 13 or 16 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), and any and all regulations promulgated thereunder, and to file the same, with all exhibits thereto, and any other documents in connection therewith, with the Securities and Exchange Commission, and with any other entity when and if such is mandated by the Exchange Act or by the Financial Industry Regulatory Authority, granting unto said attorney-in-fact full power and authority to do and perform each and every act and thing necessary, desirable or appropriate, fully to all intents and purposes as he might or could do in person, thereby ratifying and confirming all that said attorney-in-fact, or his substitutes, may lawfully do or cause to be done by virtue hereof. This Power of Attorney shall remain in full force and effect with respect to each undersigned person unless and until six months after such person is both no longer a Managing Director of ARCH Venture Partners and no longer serving on the board of directors of any portfolio company of any ARCH Venture Partners fund.

IN WITNESS WHEREOF, this Power of Attorney has been signed as of the 26th day of July, 2017.

ARCH VENTURE FUND VIII OVERAGE, L.P.

By: ARCH Venture Partners VIII, LLC its General Partner

By: /s/ Keith Crandell

Managing Director

ARCH VENTURE PARTNERS VIII, L.P.

By: ARCH Venture Partners VIII, LLC its General Partner

By: <u>/s/ Keith Crandell</u>
Managing Director