ARQULE INC Form SC 13D March 13, 2008

OMB APPROVAL

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

SCHEDULE 13D Under the Securities Exchange Act of 1934

ArQule, Inc. (Name of Issuer)

Common Stock (Title of Class of Securities)

04269E107 (CUSIP Number)

William Roiniotis BVF Partners L.P. 900 North Michigan Avenue Suite 1100 Chicago, Illinois 60611 (312) 506-6500

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

March 10, 2008
(Date of Event which Requires Filing of this Statement)

If the filing person has previously filed a statement on Schedule 13G to report the acquisition that is the subject of this Schedule 13D, and is filing this schedule because of ss.ss.240.13d-1(e), 240.13d-1(f) or 240.13d-1(g), check the following box $\mid _ \mid$

The information required on 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).

| | Biotech | nnology Value Fund, L.P. |
|---|--------------------------------------|---|
| 2. | Check t Instruc (a) X (b) | |
| 3. SEC Use Only | | |
| 4. Source of Funds (| See Inst | ructions) WC |
| 5. | | Lf Disclosure of Legal Proceedings Is Required at to Items 2(d) or 2(e) |
| | [] | |
| 6. Citizenship or Pl | ace of (| |
| Number of Shares Beneficially Owned by Each Reporting Person With | 7. | Sole Voting Power 0 |
| reison with | 8. | Shared Voting Power |
| | | 2,310,360 |
| | 9. | Sole Dispositive Power |
| | | 0 |
| | 10. | Shared Dispositive Power |
| | | 2,310,360 |
| 11. | Aggrega Person: | ate Amount Beneficially Owned by Each Reporting 2,310,360 |
| 12. | | If the Aggregate Amount in Row (11) Excludes Certain (See Instructions)[] |
| 13. | Percent | of Class Represented by Amount in Row (11): 5.3% |
| 14. | Type of | Reporting Person (See Instructions): PN |
| | | 2 |
| 1. | Names o | of Reporting Persons. |
| | Biotech | nnology Value Fund II, L.P. |
| 2. | Check t Instruc (a) X | |

| 4. Source of Funds | (See Ins | tructions) WC |
|---|------------------|--|
| 5. | | if Disclosure of Legal Proceedings Is Required nt to Items 2(d) or 2(e) |
| | [] | |
| 6. Citizenship or Pi | lace of Delawa | |
| Number of Shares Beneficially Owned by Each Reporting Person With | 7. | Sole Voting Power |
| | 8. | Shared Voting Power |
| | 9. | Sole Dispositive Power |
| | 10. | Shared Dispositive Power |
| 11. | Aggreg Person | ate Amount Beneficially Owned by Each Reporting |
| 12. | | if the Aggregate Amount in Row (11) Excludes Certai (See Instructions)[] |
| 13. | Percen | t of Class Represented by Amount in Row (11): 3.7 |
| 14. | Type o | f Reporting Person (See Instructions): PN |
| | | 3 |
| L. | Names | of Reporting Persons. |
| | BVF In | vestments, L.L.C. |
| 2. | | |
| 3. SEC Use Only | | |
| 4. Source of Funds | (See Ins | tructions) WC |
| 5. | | if Disclosure of Legal Proceedings Is Required nt to Items 2(d) or 2(e) |

| | [] | |
|--|------------------|--|
| 6. | Citize | nship or Place of Organization |
| | Delawa | re |
| Number of Shares | 7. | Sole Voting Power |
| Beneficially Owned by Each Reporting Person With | | 0 |
| | 8. | Shared Voting Power |
| | | 5,543,800 |
| | 9. | Sole Dispositive Power |
| | | 0 |
| | 10. | Shared Dispositive Power |
| | | 5,543,800 |
| 11. | Aggreg Person | ate Amount Beneficially Owned by Each Reporting: 5,543,800 |
| 12. | | if the Aggregate Amount in Row (11) Excludes Certain (See Instructions)[] |
| 13. | Percen | t of Class Represented by Amount in Row (11): 12.7% |
| 14. | Type o | f Reporting Person (See Instructions): 00 |
| | | 4 |
| 1. | Names | of Reporting Persons. |
| | Invest | ment 10, L.L.C. |
| 2. | | |
| 3. SEC Use Only | | |
| 4. Source of Funds | (See Ins | tructions) WC |
| 5. | | if Disclosure of Legal Proceedings Is Required nt to Items 2(d) or 2(e) |
| | [] | |
| 6. Citizenship or P | lace of | Organization |

Illinois

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| Number of Shares Beneficially Owned | 7. | Sole Voting Power |
|--|------------------|---|
| by Each Reporting Person With | | 0 |
| | 8. | Shared Voting Power |
| | | 681,000 |
| | 9. | Sole Dispositive Power |
| | | 0 |
| | 10. | Shared Dispositive Power |
| | | 681,000 |
| 11. | Aggreg Person | ate Amount Beneficially Owned by Each Reporting: 681,000 |
| 12. | | if the Aggregate Amount in Row (11) Excludes Certain (See Instructions)[] |
| 13. | Percen | t of Class Represented by Amount in Row (11): 1.6% |
| 14. | Type o | f Reporting Person (See Instructions): 00 |
| 1 | Namog | of Donorting Dorgons |
| 1. | Names | of Reporting Persons. |
| | BVF Pa | rtners L.P. |
| 2. | | |
| 3. SEC Use Only | | |
| 4. Source of Funds | (See Ins | tructions) 00 |
| 5. | | if Disclosure of Legal Proceedings Is Required nt to Items 2(d) or 2(e) |
| | [] | |
| 6. Citizenship or P | lace of | Organization |
| | Delawa | re |
| | 7. | |
| Beneficially by Owned by Each | | 0 |

Reporting Person With _____ Shared Voting Power 10,162,060 Sole Dispositive Power 10. Shared Dispositive Power 10,162,060 Aggregate Amount Beneficially Owned by Each Reporting Person: 10,162,060 12. Check if the Aggregate Amount in Row (11) Excludes Certain Shares (See Instructions)[] Percent of Class Represented by Amount in Row (11): 23.3% Type of Reporting Person (See Instructions): PN, HC 6 ______ Names of Reporting Persons. BVF Inc. ______ Check the Appropriate Box if a Member of a Group (See Instructions) (a) |X| (b) | | 3. SEC Use Only 4. Source of Funds (See Instructions) 00 Check if Disclosure of Legal Proceedings Is Required 5. Pursuant to Items 2(d) or 2(e) [] _____ 6. Citizenship or Place of Organization Delaware .____ Number of Shares 7. Sole Voting Power Beneficially by Owned by Each 0 Reporting Person With _____

8. Shared Voting Power

| | | 10,162,060 |
|---------|----------|--|
| | 9. | Sole Dispositive Power |
| | | 0 |
| | 10. | Shared Dispositive Power |
| | | 10,162,060 |
| 11. | Person: | te Amount Beneficially Owned by Each Reporting 10,162,060 |
| 12. | Check if | the Aggregate Amount in Row (11) Excludes Certain (See Instructions)[] |
| 13. | | of Class Represented by Amount in Row (11): 23.3% |
| 14. | | (See Instructions): CO, HC |
| | | |

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ITEM 1. SECURITY AND ISSUER

This Schedule 13D relates to the common stock, par value \$0.01 per share (the "Common Stock") of ArQule, Inc., a Delaware corporation ("ArQule"). The principal executive office of ArQule is located at 19 Presidential Way, Woburn, MA 01801-5140.

ITEM 2. IDENTITY AND BACKGROUND

The persons filing this Statement, the persons enumerated in Instruction C of this Statement and, where applicable, their respective places of organization, principal office, general partners, managers, directors, executive officers, controlling persons and certain information regarding each of them, are as follows:

- (a) Biotechnology Value Fund, L.P., a Delaware limited partnership ("BVF"), Biotechnology Value Fund II, L.P., a Delaware limited partnership ("BVF2"), BVF Investments, L.L.C., a Delaware limited liability company ("Investments"), Investment 10, L.L.C., an Illinois limited liability company ("ILL10"), BVF Partners L.P., a Delaware limited partnership ("Partners") and BVF Inc., a Delaware corporation ("BVF Inc.") specialize in holding biotechnology stocks for investment purposes. Together, BVF, BVF2, Investments, ILL10, Partners and BVF Inc. are the "Reporting Persons." Mark N. Lampert, an individual, ("Lampert"), is the sole shareholder, sole director and an officer of BVF Inc.
- (b) The business address of BVF, BVF2, Investments, ILL10 and Partners is 900 North Michigan Avenue, Suite 1100, Chicago, Illinois 60611. The business address of BVF Inc. and Lampert is One Sansome Street, 31st Floor, San Francisco, California 94104.
- (c) Partners is the general partner of BVF and BVF2, which are investment funds. Partners also is the manager of Investments, which is also an investment fund. ILL10 is a managed account, which Partners advises pursuant to an investment management agreement. BVF Inc. is an investment adviser to and

general partner of Partners. For Lampert's occupation, please refer to (a) above.

- (d) During the last five years, none of the Reporting Persons or Lampert has been convicted in a criminal proceeding (excluding traffic violations and similar misdemeanors).
- (e) During the last five years, none of the Reporting Persons or Lampert has been a party to a civil proceeding of a judicial or administrative body of competent jurisdiction nor, as a result of such proceeding, was or is subject to a judgment, decree or final order enjoining future violations of, or prohibiting or mandating activities subject to, federal or state securities laws or finding any violation with respect to such laws.
- (f) Lampert is a citizen of the United States of America. For the citizenship of each of BVF, BVF2, Investments, ILL10, Partners, and BVF Inc., please refer to (a) above

ITEM 3. SOURCE AND AMOUNT OF FUNDS OR OTHER CONSIDERATION

Partners, in its capacity as (i) general partner of BVF, has purchased on behalf of such limited partnership an aggregate number of 2,310,360 shares of the Common Stock for an aggregate consideration of \$12,417,882.25, (ii) general partner of BVF2, has purchased on behalf of such limited partnership an aggregate number of 1,626,900 shares of Common Stock for an aggregate consideration of \$8,602,421.60, (iii) manager of Investments, has purchased on

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behalf of such limited liability company an aggregate number of 5,543,800 shares of Common Stock for an aggregate consideration of \$30,452,981.50, and (iv) investment adviser to ILL10, purchased on behalf of such limited liability company an aggregate of 681,000 shares of Common Stock for an aggregate consideration of \$3,643,426.10. Each of BVF, BVF2, Investments and ILL10 purchased the shares held by it using its own working capital. No borrowed funds were used to purchase the Common Stock other than any borrowed funds used for working capital purposes in the ordinary course of business.

ITEM 4. PURPOSE OF TRANSACTION

The Reporting Persons are engaged in the investment business. In pursuing this business, the Reporting Persons analyze the operations, capital structure and markets of companies, including the Issuer, on a continuous basis through analysis of documentation and discussions with knowledgeable industry and market observers and with representatives of such companies (often at the invitation of management). From time to time, one or more of the Reporting Persons may hold discussions with third parties or with management of such companies in which the Reporting Persons may suggest or take a position with respect to potential changes in the operations, management or capital structure of such companies as a means of enhancing shareholder value. Such suggestions or positions may relate to one or more of the transactions specified in clauses (a) through (j) of Item 4 of Schedule 13D of the Exchange Act, including, without limitation, such matters as disposing of or selling all or a portion of the company or acquiring another company or business, changing operating or marketing strategies, adopting or not adopting certain types of anti-takeover measures and restructuring the Issuer's capitalization or dividend policy.

Except as set forth above, the Reporting Persons do not have any present plans or proposals that relate to or would result in any of the actions required to be described in Item 4 of Schedule 13D. Each of the Reporting Persons may, at any time, review or reconsider its position with respect to the Issuer and

formulate plans or proposals with respect to any of such matters, but has no present intention of doing so.

ITEM 5. INTEREST IN SECURITIES OF THE ISSUER

The Reporting Persons' percentage ownership of Common Stock is based on 43,710,379 shares being outstanding.

- (a) As of March 10, 2008, BVF beneficially owns 2,310,360 shares of Common Stock, BVF2 beneficially owns 1,626,900 shares of Common Stock, Investments beneficially owns 5,543,800 shares of Common Stock, ILL10 beneficially owns 681,000 shares of Common Stock and each of Partners and BVF Inc. may be deemed to beneficially own 10,162,060 shares of Common Stock, representing percentage ownership of approximately 5.3%, 3.7%, 12.7%, 1.6% and 23.3%, respectively.
- (b) Each of BVF, BVF2, Investments and ILL10 shares with Partners voting and dispositive power over the Common Stock each such entity beneficially owns. Partners and BVF Inc. share voting and dispositive power over the 10,162,060 shares of Common Stock they may be deemed beneficially to own with BVF, BVF2 and Investments, and ILL10.

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(c) The following purchases of Common Stock have been made by the following Reporting Persons within the last 60 days. All such transactions were made for cash in open market, over-the-counter transactions,

PURCHASE OF SHARES

| REPORTING PERSON | DATE | SHARES PURCHASED | PRICE PER SHARE |
|--------------------------|------------|------------------|-----------------|
| BVF | | | |
| | 01/10/2008 | 3,200 | \$4.26 |
| | 01/10/2008 | 77,027 | \$4.37 |
| | 01/14/2008 | 5,800 | \$4.50 |
| | 01/15/2008 | 26,300 | \$4.65 |
| | 01/23/2008 | 38,600 | \$4.47 |
| | 01/31/2008 | 126,000 | \$4.71 |
| | 02/01/2008 | 41,000 | \$5.00 |
| | 02/04/2008 | 52,100 | \$5.08 |
| | 02/07/2008 | 91,300 | \$4.90 |
| | 02/08/2008 | 63,000 | \$4.98 |
| | 02/11/2008 | 20,700 | \$4.96 |
| | 02/15/2008 | 12,600 | \$5.08 |
| | 02/19/2008 | 8,300 | \$5.10 |
| | 02/20/2008 | 7,100 | \$5.05 |
| | 02/21/2008 | 45,000 | \$4.95 |
| | 02/22/2008 | 2,200 | \$4.75 |
| | 03/10/2008 | 337,283 | \$3.90 |
| | 5.1 | | |
| Reporting Person BVF2 | Date | Shares Purchased | Sales Price |
| | 01/10/2008 | 3,000 | \$4.26 |
| | 01/10/2008 | 52,000 | \$4.37 |
| | 01/14/2008 | 4,000 | \$4.50 |
| | 01/15/2008 | 18,000 | \$4.65 |
| | 01/23/2008 | 26,000 | \$4.47 |
| | 01/31/2008 | 86,000 | \$4.71 |

| 02/01/2008 | 28,000 | \$5.00 |
|------------|---------|--------|
| 02/04/2008 | 35,000 | \$5.08 |
| 02/07/2008 | 62,000 | \$4.90 |
| 02/08/2008 | 44,000 | \$4.98 |
| 02/11/2008 | 14,000 | \$4.96 |
| 02/15/2008 | 8,000 | \$5.08 |
| 02/19/2008 | 5,000 | \$5.10 |
| 02/20/2008 | 5,000 | \$5.05 |
| 02/21/2008 | 32,000 | \$4.95 |
| 02/22/2008 | 1,000 | \$4.75 |
| 03/10/2008 | 231,000 | \$3.90 |
| | | |

| REPORTING PERSON | DATE | SHARES PURCHASED | SALES PRICE |
|------------------|--|---|--|
| BVFLLC | | | |
| | 01/10/2008 | 9,000 | \$4.26 |
| | 01/10/2008 | 180,000 | \$4.37 |
| | 01/14/2008 | 15,000 | \$4.50 |
| | 01/15/2008 | 63,000 | \$4.65 |
| | 01/23/2008 | 90,000 | \$4.47 |
| | 01/31/2008 | 298,000 | \$4.71 |
| | 02/01/2008 | 96,000 | \$5.00 |
| | 02/04/2008 | 123,000 | \$5.08 |
| | 02/07/2008 | 215,000 | \$4.90 |
| | 02/08/2008 | 152,000 | \$4.98 |
| | 02/11/2008 | 49,000 | \$4.96 |
| | 02/15/2008 | 29,000 | \$5.08 |
| | 02/19/2008 | 19,000 | \$5.10 |
| | 02/20/2008 | 16,000 | \$5.05 |
| | 02/21/2008 | 109,000 | \$4.95 |
| | 02/22/2008 | 5,000 | \$4.75 |
| | 03/10/2008 | 795,000 | \$3.90 |
| | 03/10/2000 | 730,000 | , , , , , |
| REPORTING PERSON | DATE | SHARES PURCHASED | SALES PRICE |
| | | | · |
| | | | · |
| | DATE | SHARES PURCHASED | SALES PRICE |
| | DATE 01/10/2008 | SHARES PURCHASED | SALES PRICE |
| | DATE 01/10/2008 01/10/2008 | SHARES PURCHASED | \$4.26 \$4.37 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 | SHARES PURCHASED 1,000 22,000 2,000 | \$4.26 \$4.37 \$4.50 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 | 1,000 22,000 2,000 8,000 11,000 | \$4.26 \$4.37 \$4.50 \$4.65 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 | 1,000 22,000 2,000 8,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 | SHARES PURCHASED 1,000 22,000 2,000 8,000 11,000 37,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 | SHARES PURCHASED 1,000 22,000 2,000 8,000 11,000 37,000 12,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/04/2008 | SHARES PURCHASED 1,000 22,000 2,000 8,000 11,000 37,000 12,000 15,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 \$5.08 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/04/2008 02/07/2008 | 1,000 22,000 2,000 8,000 11,000 37,000 12,000 15,000 27,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 \$5.08 \$4.90 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/04/2008 02/07/2008 02/08/2008 | SHARES PURCHASED 1,000 22,000 2,000 8,000 11,000 37,000 12,000 15,000 27,000 19,000 | \$4.26 \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 \$5.08 \$4.90 \$4.98 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/04/2008 02/07/2008 02/08/2008 02/11/2008 | 1,000 22,000 22,000 8,000 11,000 37,000 12,000 15,000 27,000 19,000 6,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 \$5.08 \$4.90 \$4.98 \$4.96 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/04/2008 02/07/2008 02/08/2008 02/11/2008 02/11/2008 | 1,000 22,000 2,000 8,000 11,000 37,000 12,000 15,000 27,000 19,000 6,000 4,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 \$5.08 \$4.90 \$4.96 \$5.08 |
| REPORTING PERSON | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/04/2008 02/07/2008 02/07/2008 02/08/2008 02/11/2008 02/15/2008 02/19/2008 | 1,000 22,000 22,000 2,000 8,000 11,000 37,000 12,000 15,000 27,000 19,000 6,000 4,000 2,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$4.71 \$5.00 \$5.08 \$4.90 \$4.98 \$4.96 \$5.08 \$5.10 |
| | DATE 01/10/2008 01/10/2008 01/14/2008 01/15/2008 01/23/2008 01/31/2008 02/01/2008 02/01/2008 02/07/2008 02/07/2008 02/08/2008 02/11/2008 02/15/2008 02/19/2008 | 1,000 22,000 22,000 2,000 8,000 11,000 37,000 12,000 15,000 27,000 19,000 6,000 4,000 2,000 2,000 | \$4.26 \$4.37 \$4.50 \$4.65 \$4.47 \$5.00 \$5.08 \$4.90 \$4.98 \$4.96 \$5.08 \$5.10 \$5.05 |

ITEM 6. CONTRACTS, ARRANGEMENTS, UNDERSTANDINGS OR RELATIONSHIPS WITH RESPECT TO SECURITIES OF THE ISSUER

Partners is the general partner of each of BVF and BVF2 pursuant to limited partnership agreements which authorize Partners, among other things, to invest the funds of BVF and BVF2 in the shares of Common Stock and other securities and to vote, exercise or convert and dispose of such securities. Pursuant to such limited partnership agreements, Partners is entitled to receive fees based on assets under management and allocations based on realized and unrealized gains on such assets. Pursuant to the operating agreement of Investments, Partners is authorized, among other things, to invest the funds of Ziff Asset Management, L.P., the majority member of Investments, in the shares of Common Stock and other securities and to vote, exercise or convert and dispose of such securities and is entitled to receive fees based on assets under management and allocations based on realized and unrealized gains on such assets. Pursuant to an investment management agreement with ILL10, Partners and BVF Inc. have authority, among other things, to invest funds of ILL10 in the shares of Common Stock and other securities and to vote, exercise or convert and dispose of such securities. Pursuant to such investment management agreement, Partners and BVF Inc. receive fees based on assets under management and realized and unrealized gains thereon. BVF Inc. is the general partner of Partners and may be deemed to own beneficially securities over which Partners exercises voting and dispositive power.

ITEM 7. MATERIAL TO BE FILED AS EXHIBITS

A Joint filing Agreement on behalf of the Reporting Persons is filed herewith as Exhibit A.

Signature

After reasonable inquiry and to the best of my knowledge and belief, ${\tt I}$ certify that the information set forth in this Statement is true, complete and correct.

Dated: March 13, 2008

BIOTECHNOLOGY VALUE FUND, L.P.*

By: BVF Partners L.P., its general partner

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

BIOTECHNOLOGY VALUE FUND II, L.P.*

By: BVF Partners L.P., its general partner

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

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BVF INVESTMENTS, L.L.C.*

By: BVF Partners L.P., its manager

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

INVESTMENT 10, L.L.C.*

By: BVF Partners L.P., as investment manager

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

BVF PARTNERS L.P. *

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

BVF INC.*

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

 ${}^{\star}\mathrm{The}$ Reporting Person disclaims beneficial ownership except to the extent of its pecuniary interest therein.

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Exhibit A

JOINT FILING AGREEMENT

The undersigned agree that this schedule 13D dated March 13, 2008, relating to the common Stock of ArQule, Inc. shall be filed on behalf of the undersigned.

BIOTECHNOLOGY VALUE FUND, L.P.

By: BVF Partners L.P., its general partner

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

BIOTECHNOLOGY VALUE FUND II, L.P.

By: BVF Partners L.P., its general partner

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

BVF INVESTMENTS, L.L.C.

By: BVF Partners L.P., its manager

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

INVESTMENT 10, L.L.C.

By: BVF Partners L.P., as investment manager

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert President

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BVF PARTNERS L.P.

By: BVF Inc., its general partner

By: /s/ MARK N. LAMPERT

Mark N. Lampert

President

BVF INC.

By: /s/ MARK N. LAMPERT

Mark N. Lampert

President