UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, DC 20549

FORM 8-K

CURRENT REPORT

Pursuant to Section 13 or 15(d) of The Securities Exchange Act of 1934

Date of Report (Date of earliest event reported)

December 12, 2005

CYTOKINETICS, INCORPORATED

(Exact name of registrant as specified in its charter)

DELAWARE 000-50633

(State or other jurisdiction of

incorporation)

(Commission File Number)

94-3291317 (IRS Employer Identification No.)

280 East Grand Avenue South San Francisco, California 94080

(Address of principal executive offices, including zip code)

650-624-3000

(Registrant's telephone number, including area code)

(Former name or former address, if changed since last report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions (see General Instruction A.2. below):

| Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425) |
|--|
| Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12) |
| Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b)) |
| Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c)) |

Item 1.01. ENTRY INTO A MATERIAL DEFINITIVE AGREEMENT.

On August 24, 2005, Cytokinetics, Incorporated (the "Company") filed a Current Report on Form 8-K (the "Original Report") with respect to the execution on August 22, 2005 of an Executive Employment Agreement ("Employment Agreement") with Andrew Wolff, M.D., F.A.C.C., the Company's Senior Vice President, Clinical Research and Development and Chief Medical Officer. A copy of the Employment Agreement is attached to this Current Report on Form 8-K ("Current Report") as Exhibit 10.57, and is incorporated herein by reference. A description of the material terms of the Employment Agreement is set forth in the Original Report.

Item 8.01. OTHER EVENTS.

On November 17, 2005, the Company issued a press release related to the announcement of certain clinical and non-clinical results with respect to the Company's drug candidate, ispinesib. A copy of this press release is attached to this Current Report as Exhibit 99.1.

On December 7, 2005, the Company issued a press release related to the announcement of the selection of a development candidate, CK-1827452, for the treatment of chronic heart failure. A copy of this press release is attached to this Current Report as Exhibit 99.2.

On December 8, 2005, the Company issued a press release related to the announcement of certain clinical data with respect to a Phase II clinical trial of the Company's drug candidate, ispinesib, in patients with locally advanced or metastatic breast cancer. A copy of this press release is attached to this Current Report as Exhibit 99.3.

Item 9.01. FINANCIAL STATEMENTS AND EXHIBITS.

(c) Exhibits.

The following Exhibits are filed as part of this Current Report:

| Exhibit No. | Description |
|-------------|--|
| 10.57 | Executive Employment Agreement, dated as of August 22, 2005, by and between the Registrant and Andrew Wolff. |
| 99.1 | Press release dated November 17, 2005, announcing certain clinical and non-clinical results with respect to ispinesib. |
| 99.2 | Press release dated December 7, 2005, announcing the selection of CK-1827452 as a development candidate for the treatment of chronic heart failure. |
| 99.3 | Press release dated December 8, 2005, announcing certain clinical data with respect to the Phase II clinical trial of ispinesib in patients with locally advanced or metastatic breast cancer. |

SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

CYTOKINETICS, INCORPORATED

By: /s/ James H. Sabry

James H. Sabry

President and Chief Executive Officer

Date: December 12, 2005

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EXHIBIT INDEX

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CYTOKINETICS, INCORPORATED

EXECUTIVE EMPLOYMENT AGREEMENT

This Executive Employment Agreement (the "Agreement") is made and entered into by and between Andrew Wolff (the "Executive") and Cytokinetics, Incorporated, a Delaware Corporation (the "Company"), effective as of August 22, 2005 (the "Effective Date").

RECITALS

WHEREAS: It is expected that the Company from time to time will consider the possibility of an acquisition by another company or other change of control. The Board of Directors of the Company (the "Board") recognizes that such consideration can be a distraction to Executive and can cause Executive to consider alternative employment opportunities. The Board has determined that it is in the best interests of the Company and its stockholders to assure that the Company will have the continued dedication and objectivity of Executive, notwithstanding the possibility, threat or occurrence of a Change of Control of the Company.

WHEREAS: The Board believes that it is in the best interests of the Company and its stockholders to provide Executive with an incentive to continue his or her employment and to motivate Executive to maximize the value of the Company upon a Change of Control for the benefit of its stockholders.

WHEREAS: The Board believes that it is imperative to provide Executive with certain severance benefits upon Executive's termination of employment following a Change of Control. These benefits will provide Executive with enhanced financial security and incentive and encouragement to remain with the Company notwithstanding the possibility of a Change of Control.

WHEREAS: Certain capitalized terms used in the Agreement are defined in Section 11 below.

AGREEMENT

NOW, THEREFORE, in consideration of the mutual covenants contained herein, the parties hereto agree as follows:

- 1. <u>Term of Agreement</u>. This Agreement shall terminate upon the date that all of the obligations of the parties hereto with respect to this Agreement have been satisfied.
- 2. <u>At-Will Employment</u>. The Company and Executive acknowledge that Executive's employment is and shall continue to be at-will, as defined under applicable law. If Executive's employment terminates for any reason, including (without limitation) any termination prior to a Change of Control, Executive shall not be entitled to any payments, benefits, damages, awards or compensation other than as provided by this Agreement or by law.
 - 3. Duties and Scope of Employment.
- (a) <u>Positions and Duties</u>. As of the Effective Date, Executive will serve as the Senior Vice President of Clinical Research and Development and Chief Medical Officer. Executive will

render such business and professional services in the performance of his duties, consistent with Executive's position within the Company, as will reasonably be assigned to him by the Company's Board of Directors.

(b) <u>Obligations</u>. During such time as the Executive is employed by the Company, Executive will perform his duties faithfully and to the best of his ability and will devote his full business efforts and time to the Company. During such time as the Executive is employed by the Company, Executive agrees not to actively engage in any other employment, occupation or consulting activity for any material direct or indirect remuneration without the prior approval of the Board.

4. Compensation.

- (a) <u>Base Salary</u>. During such time as the Executive is employed by the Company, the Company will pay Executive an annual salary as determined in the discretion of the Board of Directors or any committee thereof. The base salary will be paid periodically in accordance with the Company's normal payroll practices and will be subject to the usual, required withholding. Executive's salary will be subject to review and adjustments will be made based upon the Company's normal performance review practices.
- (b) <u>Performance Bonus</u>. Executive will be eligible to receive an annual bonus and other bonuses, less applicable withholding taxes, as determined by the Board of Directors or any committee thereof in the Board's or such committee's sole discretion.
- (c) <u>Equity Compensation</u>. Executive will be eligible to receive stock and option grants, and other equity compensation awards, as determined by the Board of Directors or any committee thereof in the Board's or such committee's sole discretion.
- 5. Employee Benefits. During the time that Executive is an employee of the Company, Executive will be entitled to participate in the Benefit Plans currently and hereafter maintained by the Company of general applicability to other senior executives of the Company. The Company reserves the right to cancel or change the Benefit Plans it offers to its employees at any time.
- 6. <u>Vacation</u>. Executive will be entitled to vacation in accordance with the Company's vacation policy, with the timing and duration of specific vacations mutually and reasonably agreed to by the parties hereto.
- 7. Expenses. The Company will reimburse Executive for reasonable travel, entertainment or other expenses incurred by Executive in the furtherance of or in connection with the performance of Executive's duties as an employee of the Company, in accordance with the Company's expense reimbursement policy as in effect from time to time.

8. Severance Benefits.

(a) <u>Involuntary Termination Following a Change of Control</u>. If within eighteen (18) months following a Change of Control (X)(i) Executive terminates his or her employment with the Company (or any parent or subsidiary of the Company) for Good Reason or (ii) the Company (or any parent or subsidiary of the Company) terminates Executive's employment for other than Cause,

and (Y) Executive signs and does not revoke a standard release of claims with the Company in a form reasonably acceptable to the Company, then Executive shall receive the following severance from the Company:

- (i) <u>Severance Payment</u>. Executive will be entitled to (i) receive continuing payments of severance pay (less applicable withholding taxes) at a rate equal to his base salary rate, as then in effect, for a period of eighteen (18) months from the date of such termination, to be paid periodically in accordance with the Company's normal payroll policies; and (B) a lump-sum payment equal to 100% of Executive's target annual bonus as of the date of such termination.
- (ii) Options: Restricted Stock. All of Executive's then outstanding options to purchase shares of the Company's Common Stock (the "Options") shall immediately vest and become exercisable (that is, in addition to the shares subject to the Options which have vested and become exercisable as of the date of such termination), but in no event shall the number of shares subject to such Options which so vest exceed the total number of shares subject to such Options. Additionally, all of the shares of the Company's Common Stock then held by Executive subject to a Company right of repurchase (the "Restricted Stock") shall immediately vest and have such Company right of repurchase with respect to such shares of Restricted Stock lapse (that is, in addition to the shares of Restricted Stock which have vested as of the date of such termination), but in no event shall the number of shares which so vest exceed the number of shares of Restricted Stock outstanding immediately prior to such termination.
- (iii) <u>Continued Employee Benefits</u>. Executive shall receive Company-paid coverage for Executive and Executive's eligible dependents under the Company's Benefit Plans for a period equal to the shorter of (i) eighteen (18) months or (ii) such time as Executive secures employment with benefits generally similar to those provided in the Company's Benefit Plans.
- (b) <u>Timing of Severance Payments</u>. Any lump-sum severance payment to which Executive is entitled shall be paid by the Company to Executive in cash and in full, not later than ten (10) calendar days after the date of the termination of Executive's employment as provided in Section 8(a), and any other severance payments shall be paid in accordance with normal payroll policies as provided in Section 8(a). If Executive should die before all amounts have been paid, such unpaid amounts shall be paid in a lump-sum payment to Executive's designated beneficiary, if living, or otherwise to the personal representative of Executive's estate.
- (c) <u>Voluntary Resignation</u>; <u>Termination for Cause</u>. If Executive's employment with the Company terminates (i) voluntarily by Executive other than for Good Reason or (ii) for Cause by the Company, then Executive shall not be entitled to receive severance or other benefits except for those as may then be established under the Company's then existing severance and Benefits Plans or pursuant to other written agreements with the Company.
- (d) <u>Disability: Death.</u> If the Company terminates Executive's employment as a result of Executive's Disability, or Executive's employment terminates due to his or her death, then Executive shall not be entitled to receive severance or other benefits except for those as may then be established under the Company's then existing written severance and Benefits Plans or pursuant to other written agreements with the Company.

- (e) <u>Termination Apart from Change of Control</u>. In the event Executive's employment is terminated for any reason, either prior to the occurrence of a Change of Control or after the eighteen (18) month period following a Change of Control, then Executive shall be entitled to receive severance and any other benefits only as may then be established under the Company's existing written severance and Benefits Plans, if any, or pursuant to any other written agreements with the Company.
- (f) Exclusive Remedy. In the event of a termination of Executive's employment within eighteen (18) months following a Change of Control, the provisions of this Section 8 are intended to be and are exclusive and in lieu of any other rights or remedies to which Executive or the Company may otherwise be entitled, whether at law, tort or contract, in equity, or under this Agreement. Executive shall be entitled to no benefits, compensation or other payments or rights upon termination of employment following a Change in Control other than those benefits expressly set forth in this Section 8.

9. Conditional Nature of Severance Payments.

- (a) <u>Proprietary Information and Invention Assignment Agreement</u>. If Executive is in material breach of the terms of the Proprietary Information and Invention Assignment Agreement, by and between the Company and Executive, dated as of September 20, 2004 (the "Invention Agreement"), including, without limitation, Executive's obligations of confidentiality and of non-solicitation contained in the Invention Agreement, then upon such breach by Executive: (i) Executive shall refund to the Company all cash paid to Executive pursuant to Section 8 of this Agreement; and (ii) all severance benefits pursuant to this Agreement shall immediately cease.
- (b) Non-Competition. Executive acknowledges that the nature of the Company's business is such that if Executive were to become employed by, or substantially involved in, the business of a competitor of the Company during the eighteen (18) months following the termination of Executive's employment with the Company, it would be very difficult for Executive not to rely on or use the Company's trade secrets and confidential information. Thus, to avoid the inevitable disclosure of the Company's trade secrets and confidential information, Executive agrees and acknowledges that Executive's right to receive the severance payments set forth in this Agreement (to the extent Executive is otherwise entitled to such payments) will be conditioned upon Executive not directly or indirectly engaging in (whether as an employee, consultant, agent, proprietor, principal, partner, stockholder, corporate officer, director or otherwise), nor having any ownership interest in or participating in the financing, operation, management or control of, any person, firm, corporation or business that competes with the Company or is a customer of the Company. Notwithstanding the foregoing, Executive may own, directly or indirectly, up to 1% of the capital stock of a company that competes with the Company, provided such capital stock is traded on a national securities exchange or through the automated quotation system of a registered securities association. Upon any breach of this section, all severance payments pursuant to this Agreement will immediately cease.
- (c) <u>Understanding of Obligations</u>. Executive represents that he is fully aware of his obligations under the Invention Agreement and hereunder, including, without limitation, the reasonableness of the length of time, scope and geographic coverage of any such obligations.

10. <u>Limitation on Payments</u>. In the event that the severance and other benefits provided for in this Agreement or otherwise payable to Executive (i) constitute "parachute payments" within the meaning of Section 280G of the Internal Revenue Code of 1986, as amended (the "Code") and (ii) but for this Section 10, would be subject to the excise tax imposed by Section 4999 of the Code, then Executive's severance benefits shall be either:

- (a) delivered in full, or
- (b) delivered as to such lesser extent which would result in no portion of such severance benefits being subject to excise tax under Section 4999 of the Code,

whichever of the foregoing amounts, taking into account the applicable federal, state and local income taxes and the excise tax imposed by Section 4999, results in the receipt by Executive on an after-tax basis, of the greatest amount of severance benefits, notwithstanding that all or some portion of such severance benefits may be taxable under Section 4999 of the Code. Unless the Company and Executive otherwise agree in writing, any determination required under this Section 10 shall be made in writing by the Company's independent public accountants immediately prior to Change of Control (the "Accountants"), whose determination shall be conclusive and binding upon Executive and the Company for all purposes. For purposes of making the calculations required by this Section 10, the Accountants may make reasonable assumptions and approximations concerning applicable taxes and may rely on reasonable, good faith interpretations concerning the application of Sections 280G and 4999 of the Code. The Company and Executive shall furnish to the Accountants such information and documents as the Accountants may reasonably request in order to make a determination under this Section. The Company shall bear all costs the Accountants may reasonably incur in connection with any calculations contemplated by this Section 10. If there is a reduction pursuant to this Section 10 of the severance benefits to be delivered to Executive, such reduction shall first be applied to any cash amounts to be delivered to the Executive under this Agreement and thereafter to any other severance benefits of Executive hereunder.

- 11. <u>Definition of Terms</u>. The following terms referred to in this Agreement shall have the following meanings:
- (a) <u>Benefit Plans</u>. "Benefit Plans" means plans, policies or arrangements that the Company sponsors (or participates in) and that immediately prior to Executive's termination of employment provide Executive and/or Executive's eligible dependents with medical, dental, vision and/or financial counseling benefits. Benefit Plans do not include any other type of benefit (including, but not by way of limitation, disability, life insurance or retirement benefits). A requirement that the Company provide Executive and Executive's eligible dependents with coverage under the Benefit Plans will not be satisfied unless the coverage is no less favorable than that provided to Executive and Executive's eligible dependents immediately prior to Executive's termination of employment. Notwithstanding any contrary provision of this Section 11, but subject to the immediately preceding sentence, the Company may, at its option, satisfy any requirement that the Company provide coverage under any Benefit Plan by instead providing coverage under a separate plan or plans providing coverage that is no less favorable or by paying Executive a lump-sum payment sufficient to provide Executive and Executive's eligible dependents with equivalent coverage under a third party plan that is reasonably available to Executive and Executive's eligible dependents.

- (b) Cause. "Cause" means any of the following: (i) the failure by you to substantially perform your duties with the Company (other than due to your incapacity as a result of physical or mental illness for a period not to exceed 90 days); (ii) the engaging by you in conduct which is materially injurious to the Company, its business or reputation, or which constitutes gross misconduct; (iii) your material breach of the terms of this Agreement, the Invention Agreement or any other agreements between you and the Company; (iv) the material breach or taking of any action in material contravention of the policies of the Company adopted by the Board of Directors or any committee thereof, including, without limitation, the Company's Code of Ethics, Insider Trading Compliance Program, Disclosure Process and Procedures or Corporate Governance Guidelines; (v) your conviction for or admission or plea of no contest with respect to a felony; or (vi) an act of fraud against the Company, the misappropriation of material property belonging to the Company, or an act of violence against an officer, director, employee or consultant of the Company; provided, however, that in the event that any of the foregoing events in (i), (iii) or (iv) is capable of being cured, the Company shall provide written notice to you describing the nature of such event, and you shall thereafter have thirty (30) business days to cure such event.
 - (c) Change of Control. "Change of Control" means the occurrence of any of the following:
- (i) Any "person" (as such term is used in Sections 13(d) and 14(d) of the Securities Exchange Act of 1934, as amended) becomes the "beneficial owner" (as defined in Rule 13d-3 under said Act), directly or indirectly, of securities of the Company representing fifty percent (50%) or more of the total voting power represented by the Company's then outstanding voting securities; or
- (ii) Any action or event occurring within a two-year period, as a result of which fewer than a majority of the directors are Incumbent Directors. "Incumbent Directors" shall mean directors who either (A) are directors of the Company as of the date hereof, or (B) are elected, or nominated for election, to the Board with the affirmative votes of at least a majority of the Incumbent Directors at the time of such election or nomination (but shall not include an individual whose election or nomination is in connection with an actual or threatened proxy contest relating to the election of directors to the Company); or
- (iii) The consummation of a merger or consolidation of the Company with any other corporation, other than a merger or consolidation which would result in the voting securities of the Company outstanding immediately prior thereto continuing to represent (either by remaining outstanding or by being converted into voting securities of the surviving entity) at least fifty percent (50%) of the total voting power represented by the voting securities of the Company or such surviving entity outstanding immediately after such merger or consolidation; or
 - (iv) The consummation of the sale, lease or other disposition by the Company of all or substantially all the Company's assets.
- (d) <u>Disability</u>. "Disability" shall mean that Executive has been unable to perform his Company duties as the result of his incapacity due to physical or mental illness, and such inability, at least twenty-six (26) weeks after its commencement, is determined to be total and permanent by a physician selected by the Company or its insurers and reasonably acceptable to Executive or

Executive's legal representative. Termination resulting from Disability may only be effected after at least thirty (30) days' written notice by the Company of its intention to terminate Executive's employment. In the event that Executive resumes the performance of substantially all of his or her duties hereunder before the termination of his or her employment becomes effective, the notice of intent to terminate shall automatically be deemed to have been revoked.

(e) Good Reason. "Good Reason" means any of the following unless such event is agreed to, in writing or as set forth below, by you: (i) a material reduction in your salary or benefits (excluding the substitution of substantially equivalent compensation and benefits), other than as a result of a reduction in compensation affecting employees of the Company, or its successor entity, generally; (ii) a material diminution of your duties or responsibilities relative to your duties and responsibilities in effect immediately prior to the Change of Control, provided however, that, in the case of the Company being acquired and made part of a larger organization, a change in your title or reporting requirements where your duties, responsibilities and authority after the Change of Control are functionally similar to your duties, responsibilities and authority prior to the Change of Control (as, for example, when the Vice-President, Sales of the Company remains responsible for sales of the Company's products following a Change of Control but is not made the Vice President, Sales of the acquiring corporation) shall not constitute "Good Reason;" (iii) relocation of your place of employment to a location more than 50 miles from the Company's office location at the time of the Change of Control; and (iv) failure of a successor entity in any Change of Control to assume and perform under this Agreement. If any of the events set forth above shall occur, you shall give prompt written notice of such event to the Company, or its successor entity, and if such event is not cured within thirty (30) days from such notice you may exercise your rights to resign for Good Reason, provided that if you have not exercised such right within 45 days of the date of such notice you shall be deemed to have agreed to the occurrence of such event.

12. Arbitration.

(a) General. In consideration of Executive's service to the Company, its promise to arbitrate all employment related disputes and Executive's receipt of the compensation, pay raises and other benefits paid to Executive by the Company, at present and in the future, Executive agrees that any and all controversies, claims, or disputes with anyone (including the Company and any employee, officer, director, shareholder or benefit plan of the Company in their capacity as such or otherwise) arising out of, relating to, or resulting from Executive's service to the Company under this Agreement or otherwise or the termination of Executive's service with the Company, including any breach of this Agreement, will be subject to binding arbitration under the Arbitration Rules set forth in California Code of Civil Procedure Section 1280 through 1294.2, including Section 1283.05 (the "Rules") and pursuant to California law. Disputes which Executive agrees to arbitrate, and thereby agrees to waive any right to a trial by jury, include any statutory claims under state or federal law, including, but not limited to, claims under Title VII of the Civil Rights Act of 1964, the Americans with Disabilities Act of 1990, the Age Discrimination in Employment Act of 1967, the Older Workers Benefit Protection Act, the California Fair Employment and Housing Act, the California Labor Code, claims of harassment, discrimination or wrongful termination and any statutory claims. Executive further understands that this Agreement to arbitrate also applies to any disputes that the Company may have with Executive.

- (b) <u>Procedure</u>. Executive agrees that any arbitration will be administered by the American Arbitration Association ("AAA") and that a neutral arbitrator will be selected in a manner consistent with its National Rules for the Resolution of Employment Disputes. The arbitration proceedings will allow for discovery according to the rules set forth in the *National Rules for the Resolution of Employment Disputes or California Code of Civil Procedure*. Executive agrees that the arbitrator will have the power to decide any motions brought by any party to the arbitration, including motions for summary judgment and/or adjudication and motions to dismiss and demurrers, prior to any arbitration hearing. Executive agrees that the arbitrator will issue a written decision on the merits. Executive also agrees that the arbitrator will have the power to award any remedies, including attorneys' fees and costs, available under applicable law. Executive understands the Company will pay for any administrative or hearing fees charged by the arbitrator or AAA except that Executive will pay the first \$125.00 of any filing fees associated with any arbitration Executive initiates. Executive agrees that the arbitrator will administer and conduct any arbitration in a manner consistent with the Rules and that to the extent that the AAA's National Rules for the Resolution of Employment Disputes conflict with the Rules, the Rules will take precedence.
- (c) Remedy. Except as provided by the Rules, arbitration will be the sole, exclusive and final remedy for any dispute between Executive and the Company. Accordingly, except as provided for by the Rules, neither Executive nor the Company will be permitted to pursue court action regarding claims that are subject to arbitration. Notwithstanding, the arbitrator will not have the authority to disregard or refuse to enforce any lawful Company policy, and the arbitrator will not order or require the Company to adopt a policy not otherwise required by law which the Company has not adopted.
- (d) <u>Availability of Injunctive Relief.</u> In addition to the right under the Rules to petition the court for provisional relief, Executive agrees that any party may also petition the court for injunctive relief where either party alleges or claims a violation of this Agreement or the Confidentiality Agreement or any other agreement regarding trade secrets, confidential information, nonsolicitation or Labor Code §2870. In the event either party seeks injunctive relief, the prevailing party will be entitled to recover reasonable costs and attorneys fees.
- (e) <u>Administrative Relief.</u> Executive understands that this Agreement does not prohibit Executive from pursuing an administrative claim with a local, state or federal administrative body such as the Department of Fair Employment and Housing, the Equal Employment Opportunity Commission or the workers' compensation board. This Agreement does, however, preclude Executive from pursuing court action regarding any such claim.
- (f) Voluntary Nature of Agreement. Executive acknowledges and agrees that Executive is executing this Agreement voluntarily and without any duress or undue influence by the Company or anyone else. Executive further acknowledges and agrees that Executive has carefully read this Agreement and that Executive has asked any questions needed for Executive to understand the terms, consequences and binding effect of this Agreement and fully understand it, including that Executive is waiving Executive's right to a jury trial. Finally, Executive agrees that Executive has been provided an opportunity to seek the advice of an attorney of Executive's choice before signing this Agreement.

13. Successors.

- (a) The Company's Successors. Any successor to the Company (whether direct or indirect and whether by purchase, merger, consolidation, liquidation or otherwise) to all or substantially all of the Company's business and/or assets shall assume the obligations under this Agreement and agree expressly to perform the obligations under this Agreement in the same manner and to the same extent as the Company would be required to perform such obligations in the absence of a succession. For all purposes under this Agreement, the term "Company" shall include any successor to the Company's business and/or assets which executes and delivers the assumption agreement described in this Section 13(a) or which becomes bound by the terms of this Agreement by operation of law.
- (b) <u>The Executive's Successors</u>. The terms of this Agreement and all rights of Executive hereunder shall inure to the benefit of, and be enforceable by, Executive's personal or legal representatives, executors, administrators, successors, heirs, distributees, devisees and legatees.

14. Notice.

- (a) <u>General</u>. Notices and all other communications contemplated by this Agreement shall be in writing and shall be deemed to have been duly given when personally delivered or when mailed by U.S. registered or certified mail, return receipt requested and postage prepaid. In the case of Executive, mailed notices shall be addressed to him or her at the home address which he or she most recently communicated to the Company in writing. In the case of the Company, mailed notices shall be addressed to its corporate headquarters, and all notices shall be directed to the attention of its Chief Financial Officer.
- (b) Notice of Termination. Any termination by the Company for Cause or by Executive for Good Reason or as a result of a voluntary resignation shall be communicated by a notice of termination to the other party hereto given in accordance with Section 14(a) of this Agreement. Such notice shall indicate the specific termination provision in this Agreement relied upon, shall set forth in reasonable detail the facts and circumstances claimed to provide a basis for termination under the provision so indicated, and shall specify the termination date (which shall be not more than thirty (30) days after the giving of such notice).

15. Miscellaneous Provisions.

- (a) No Duty to Mitigate. Executive shall not be required to mitigate the amount of any payment contemplated by this Agreement, nor, except as otherwise contemplated in this Agreement, shall any such payment be reduced by any earnings that Executive may receive from any other source.
- (b) <u>Waiver</u>. No provision of this Agreement shall be modified, waived or discharged unless the modification, waiver or discharge is agreed to in writing and signed by Executive and by an authorized officer of the Company (other than Executive). No waiver by either party of any breach of, or of compliance with, any condition or provision of this Agreement by the other party

shall be considered a waiver of any other condition or provision or of the same condition or provision at another time.

- (c) Headings. All captions and section headings used in this Agreement are for convenient reference only and do not form a part of this Agreement.
- (d) Entire Agreement. This Agreement and the Invention Agreement constitute the entire agreement of the parties hereto and supersedes in their entirety all prior representations, understandings, understandings or agreements (whether oral or written and whether expressed or implied) of the parties with respect to the subject matter hereof. No future agreements between the Company and Executive may supersede this Agreement, unless they are in writing and specifically mentioned this Agreement.
- (e) <u>Choice of Law</u>. The laws of the State of California (without reference to its choice of laws provisions) shall govern the validity, interpretation, construction and performance of this Agreement.
- (f) <u>Severability</u>. The invalidity or unenforceability of any provision or provisions of this Agreement shall not affect the validity or enforceability of any other provision hereof, which shall remain in full force and effect.
 - (g) Withholding. All payments made pursuant to this Agreement will be subject to withholding of applicable income and employment taxes.
- (h) Counterparts. This Agreement may be executed in counterparts, each of which shall be deemed an original, but all of which together will constitute one and the same instrument.

[Remainder of Page Intentionally Left Blank]

IN WITNESS WHEREOF, each of the parties has executed this Agreement, in the case of the Company by its duly authorized officer, as of the day and year set forth below.

CYTOKINETICS, INCORPORATED COMPANY

/s/ James H. Sabry

Title: CEO/President

EXECUTIVE By: /s/ Andrew Wolff

Andrew Wolff, Senior Vice President of Clinical Research and Development and Chief Medical Officer

Contacts:
Cytokinetics, Incorporated
Robert I. Blum
EVP, Corporate Development and Commercial Operations & CBO
(650) 624-3000

Burns McClellan, Inc. Clay Kramer (investors) Justin Jackson (media) (212) 213-0006

For immediate release

CYTOKINETICS ANNOUNCES CLINICAL AND NON-CLINICAL RESULTS FOR ISPINESIB AT THE 2005 ANNUAL AACR-NCI-EORTC MEETING

Data from Two Phase Ib Combination Trials Presented

South San Francisco, CA, November 17, 2005 – Cytokinetics, Incorporated (Nasdaq: CYTK) announced today that data from Phase Ib combination clinical trials and non-clinical studies of ispinesib (SB-715992) were presented at the 2005 AACR-NCI-EORTC International Meeting in Philadelphia, Pennsylvania. The two clinical poster presentations highlighted data from two Phase Ib combination trials, one evaluating ispinesib in combination with capecitabine and the other evaluating ispinesib in combination with docetaxel. The Phase Ib combination clinical trials were designed to assess the safety, tolerability and pharmacokinetics of ispinesib in combination with the standard chemotherapeutics, and to determine the optimally tolerated regimen of ispinesib in combination with each of the two drugs. In addition, four poster presentations describing the further non-clinical characterization of ispinesib, including data relating to resistance mechanisms, pharmacodynamic markers, potential therapeutic index and suitability for combination with cisplatin, will also be presented.

One clinical trial presentation entitled, "Phase I Study of *Ispinesib* in Combination with *Docetaxel* in Patients with Advanced Solid Tumors," contained data from the clinical trial that demonstrated that the combination of *ispinesib* with *docetaxel* has an acceptable tolerability profile on a once every 21 day schedule. The dose limiting toxicity in this combination regimen was prolonged ≥ 5 days) Grade 4 neutropenia. Other common toxicities were observed including neutropenia, infection, anemia, pain, nausea, fatigue, diarrhea and alopecia. The optimally tolerated regimen (OTR) was defined as 10 mg/m^2 of *ispinesib* and 60 mg/m^2 of *docetaxel*, each administered once every 21 days. In addition, plasma concentrations of both *ispinesib* and *docetaxel* were consistent with those previously reported when each drug was given as a monotherapy, suggesting no pharmacokinetic interaction between the two agents. In this trial, a total of 13 patients (out of 24) had a response of stable disease by RECIST criteria (duration 2.25-7.5 months). Two prostate cancer patients had a best response of confirmed PSA partial response (>50% reduction in baseline PSA at least four weeks later) as determined by the investigator.

The other clinical trial presentation entitled, "Phase I Study of *Ispinesib* in Combination with *Capecitabine* in Patients with Advanced Solid Tumors," contained data arising from the ongoing clinical trial that demonstrated that the combination of *ispinesib* and *capecitabine* appears to have an acceptable tolerability profile on the study's treatment schedule. One case of dose limiting toxicity of prolonged (≥ 5 days) Grade 4 neutropenia has been identified to date. Other common toxicities that have been observed include leukopenia, infections, anorexia and fatigue. The OTR in this study has yet to be defined, however, in the cohorts evaluated thus far, *ispinesib* plasma concentrations do not appear to be effected by the presence of *capecitabine*. In this clinical trial, a total of 8 patients out of 16 have had a best response of stable disease by RECIST criteria (duration 2-6.5 months).

"We believe these combination trials suggest that *ispinesib* has an acceptable tolerability and no pharmacokinetic interaction when used with either of these two common chemotherapeutic agents in patients suffering from advanced solid tumors," stated Dr. Andrew A. Wolff, Cytokinetics' Senior Vice President of Clinical Research and Development and Chief Medical Officer. "These data are encouraging about possibilities for *ispinesib* in combination with other commonly used anti-cancer drugs and may guide future clinical trial design for *ispinesib*."

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In addition, results from non-clinical studies involving ispinesib were presented as follows:

- "A Resistance Mechanism for the KSP Inhibitor *Ispinesib* Implicates Point Mutations in the Compound Binding Site." This presentation described a specific genetic mutation in the development of an acquired cellular resistance to the inhibition of Kinesin Spindle Protein (KSP). The implication of this finding in humans is uncertain as initial investigations in human subjects suggest that such germline polymorphisms may not be a prevalent resistance mechanism in humans.
- "Mitotic Arrest in Tumors as a Pharmacodynamic Marker for Inhibition of the Mitotic Kinesin KSP by *Ispinesib*, a Novel KSP Inhibitor." This presentation revealed that *ispinesib* induced dose-dependent accumulation of mitotic cells at doses sufficient to suppress tumor growth in mice. These findings suggest that analysis of pharmacodynamic markers of mitotic arrest in patient biopsies by flow cytometry may prove useful as a rapid and quantitative method of demonstrating the biological activity of KSP inhibitors in clinical studies.
- "Relationship Between the Antitumor Activity of *Ispinesib*, a Novel KSP Inhibitor, and Neutropenia in a Human Xenograft Model." This presentation characterized the induction of neutropenia with *ispinesib* in correlation with the observed anti-tumor activity in mice revealing that tumor regression and increases in life span were achieved at doses of *ispinesib* that did not induce severe neutropenia. These findings suggest that a more clinically relevant endpoint of toxicity in mouse efficacy models may improve their utility in preclinical evaluation.
- "Cisplatin Enhances the Activity of Ispinesib, a Novel KSP Inhibitor, Against Murine P388 Lymphocytic Leukemia." This presentation highlighted enhanced activity of ispinesib in combination with cisplatin in mouse tumor models and demonstrated that the order of administration influenced effectiveness. The data suggest that inducing DNA damage in vivo makes tumor cells more sensitive to the effects of ispinesib.

About Ispinesib

Ispinesib is a novel small molecule inhibitor of Kinesin Spindle Protein (KSP), a mitotic kinesin protein essential for proper cell division. Ispinesib is the first drug candidate in clinical development that has arisen from a broad strategic collaboration between Cytokinetics and GlaxoSmithKline (GSK) to discover, develop and commercialize novel small molecule therapeutics targeting human mitotic kinesins for applications in the treatment of cancer and other diseases. GSK is conducting a broad clinical trials program for ispinesib designed to study the drug candidate in multiple tumor types, combination regimens and dosing schedules. GSK is currently evaluating ispinesib in three Phase II studies being conducted in patients with each of non-small cell lung, ovarian and breast cancers and three Phase Ib clinical trials designed to evaluate ispinesib in combination with each of docetaxel, carboplatin and capecitabine. Interim data from the ongoing breast cancer clinical trial and the platinum-refractory treatment arm of the non-small cell lung cancer clinical trial were announced recently. In the Phase II clinical trial enrolling patients with advanced breast cancer, the best overall responses observed with ispinesib administered as monotherapy have been partial responses in three of thirty-three evaluable patients to date. Interim data from this ongoing breast cancer trial will be presented on December 8, 2005 at the San Antonio Breast Cancer Symposium in San Antonio, Texas. In the platinum-refractory treatment arm of a Phase II clinical trial enrolling patients with non-small cell lung cancer, the best overall response observed with ispinesib administered as monotherapy has been disease stabilization in 25% (N=20) of patients with a median time to progression (TTP) of 12 weeks (overall median TTP was six weeks). In addition to the ongoing studies being conducted by GSK, the National Cancer Institute (NCI) continues to enroll patients in five other Phase II clinical trials evaluating ispinesib in other tumor types, including melanoma, head and neck, hepatocellular, colorectal and prostate cancers. In addition, the NCI plans to conduct one additional Phase II clinical trial in patients with renal cell carcinoma. The NCI is also conducting two other Phase I clinical trials evaluating an alternative schedule of ispinesib in leukemia and advanced solid tumors.

About Cytokinetics

Cytokinetics is a leading biopharmaceutical company focused on the discovery, development and commercialization of novel small molecule drugs that specifically target the cytoskeleton. The cytoskeleton is a complex biological infrastructure that plays a fundamental role within every human cell. Cytokinetics' focus on the cytoskeleton enables it to develop novel and potentially safer and more effective classes of drugs directed at treatments for cancer,

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cardiovascular disease and other diseases. Cytokinetics has developed a cell biology driven approach and proprietary technologies to evaluate the function of many interacting proteins in the complex environment of the intact human cell. Cytokinetics employs the PUMATM system and CytometrixTM technologies to enable early identification and automated prioritization of compounds that are highly selective for their intended protein targets without other cellular effects, and may therefore be less likely to give rise to clinical side effects. Cytokinetics and GlaxoSmithKline have entered into a strategic alliance to discover, develop and commercialize small molecule therapeutics targeting human mitotic kinesins for applications in the treatment of cancer and other diseases. GlaxoSmithKline is conducting Phase II and Phase Ib clinical trials for *ispinesib* (SB-715992) and a Phase I clinical trial for SB-743921, each a drug candidate that has emerged from the strategic alliance. Cytokinetics' heart failure program is the second program to leverage the company's expertise in cytoskeletal pharmacology. Cytokinetics recently initiated a Phase I human clinical trial with CK-1827452, a novel small molecule cardiac myosin activator, for the treatment of heart failure. Additional information about Cytokinetics can be obtained at www.cytokinetics.com.

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CYTOKINETICS ANNOUNCES SELECTION OF CK-1827452 AS DEVELOPMENT CANDIDATE FOR CHRONIC HEART FAILURE

Attractive Oral Bioavailability Profile Provides Opportunity for Development Across Continuum of Care

South San Francisco, CA, December 7, 2005 – Cytokinetics, Incorporated (Nasdaq: CYTK) announced today that it has recently selected CK-1827452 as a development candidate for the potential treatment of patients with chronic heart failure treated in an outpatient setting. An intravenous formulation of CK-1827452 is currently in Phase I clinical development as a potential treatment for patients with acute heart failure. Pharmacokinetic data arising from that clinical trial confirms that CK-1827452 has a sufficiently long half-life to support development of a chronic oral dosing formulation. Additional preclinical studies to support oral dosing in humans with CK-1827452 are currently underway. Following successful completion of the enabling preclinical studies, Cytokinetics intends to submit a regulatory filing for the initiation of a Phase I clinical trial meant to confirm in humans the bioavailability seen in preclinical species with an orally administered formulation of CK-1827452.

Cytokinetics' heart failure program focuses on the discovery and development of small molecules that directly activate cardiac myosin, a cytoskeletal protein that drives cardiac muscle contractility. This mechanism of action results in increased cardiac contractility without increasing stimulating beta-andrenergic receptors or inhibiting phosphodiesterase activity to increase intracellular calcium, that may be associated with adverse clinical effects in heart failure patients. Cytokinetics identified CK-1827452 at the beginning of 2005 as a drug candidate arising from this program taking into consideration certain properties including potency, tolerability, pharmacokinetics and pharmaceutics. CK-1827452 was determined to be suitable for development as a novel, next-generation pharmaceutical for the treatment of acute heart failure, but also provided an opportunity for this compound to be used in patients with chronic heart failure.

"We are excited about the potential for this compound," stated James H. Sabry, M.D., Ph.D., Cytokinetics' President and Chief Executive Officer.

"Cytokinetics' expertise in cytoskeletal pharmacology has now generated a development compound with potential application as a next-generation treatment for both acute and chronic heart failure patients. This compound may be able to address heart failure patients' needs across the continuum of care."

"The selection of this compound represents an important step forward for the development of cardiac myosin activators as novel therapeutics in the treatment of heart failure," added Andrew A. Wolff, M.D., F.A.C.C., Cytokinetics' Senior Vice President of Clinical Research and Development and Chief Medical Officer. "We are excited to now explore the potential for this class of compounds in the treatment of chronic heart failure."

Development Status of CK-1827452

CK-1827452, a novel, small-molecule, direct activator of cardiac myosin, is currently in a Phase I, first-in-humans clinical trial for treatment of acute heart failure. The clinical trial is a double-blind, randomized, placebo-controlled, dose-escalation study being conducted to investigate the safety, tolerability, pharmacokinetic, and pharmacodynamic profile of CK-1827452 in normal healthy volunteers. The clinical trial is designed to identify the maximum tolerated dose of a 6-hour intravenous infusion of CK-1827452. The effect of CK-1827452 on the left ventricular function of these healthy volunteers will be evaluated using serial echocardiograms. The cross-over design of this clinical trial ensures that each volunteer serves as his own control to compare the effects of escalating doses of CK-1827452 to those of placebo. The clinical trial is being conducted under a Clinical Trial Application at a clinical investigative center in the United Kingdom.

Background on the Heart Failure Market

Heart failure is a widespread and debilitating syndrome affecting approximately five million people in the United States alone. The high and rapidly growing prevalence of heart failure translates into significant hospitalization rates and associated societal costs. The number of hospital discharges in the United States identified with a primary diagnosis of heart failure rose

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from 550,000 in 1989 to 970,000 in 2002. Heart failure is one of the most common primary discharge diagnoses identified in hospitalized patients over the age 65 in the United States. The annual costs of heart failure in the United States are estimated to be \$27.9 billion, including \$18.3 billion for inpatient care. The market for heart failure drugs was approximately \$2.7 billion in 2001, according to industry reports. Despite currently available therapies, readmission rates for patients over the age of 65 remain high at 30 to 40 percent within six months of hospital discharge and mortality rates over an eight year period range from 70% to 80% for patients under the age of 65. The limited effectiveness of current therapies points to the need for next-generation agents with improved efficacy without increased adverse events.

Background on Cardiac Myosin Activators and Cardiac Contractility

Cardiac myosin is the cytoskeletal motor protein in the cardiac muscle cell that is directly responsible for converting chemical energy into the mechanical force resulting in cardiac contraction. Cardiac contractility is driven by the cardiac sarcomere, the fundamental unit of muscle contraction in the heart that is a highly ordered cytoskeletal structure composed of cardiac myosin, actin and a set of regulatory proteins. The sarcomere represents one of the most thoroughly characterized protein machines in human biology.

Cytokinetics' heart failure program is focused towards the discovery and development of small molecule cardiac myosin activators in order to create next-generation treatments to manage acute and chronic heart failure. Cytokinetics' program is based on the hypothesis that activators of cardiac myosin may address certain mechanistic liabilities of existing positive inotropic agents by increasing cardiac contractility without stimulating beta-adrenergic receptors inhibiting phosphodiesterase activity to increase intracellular calcium, each of which may be associated with adverse clinical effects in heart failure patients. Existing drugs that seek to improve cardiac cell contractility increase the concentration of intracellular calcium, which indirectly activates cardiac myosin, but this effect on calcium levels also has been linked to potentially life-threatening side effects. In contrast, cardiac myosin activators have been shown to work by a novel mechanism that directly stimulates the activity of the cardiac myosin motor protein by accelerating the rate-limiting step of the myosin enzymatic cycle and thereby shifting the enzymatic cycle in favor of the force producing state.

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CYTOKINETICS ANNOUNCES PRESENTATION OF ISPINESIB DATA AT SAN ANTONIO BREAST CANCER SYMPOSIUM

Anti-cancer Activity Supportive of Continuation of Phase II Clinical Trial

South San Francisco, CA, December 8, 2005 – Cytokinetics, Incorporated (Nasdaq: CYTK) announced today that an interim analysis from an ongoing multicenter Phase II clinical trial of ispinesib (SB-715992) in subjects with locally advanced or metastatic breast cancer was presented at the 2005 San Antonio Breast Cancer Symposium (SABCS) at the Henry B. Gonzalez Convention Center in San Antonio, Texas. The clinical poster presentation elaborated on previously announced data from a Phase II clinical trial, which was designed to assess the safety, tolerability and efficacy of ispinesib in these patients with breast cancer.

The poster entitled, "Phase II, Open Label Study of *Ispinesib* in Patients with Locally Advanced or Metastatic Breast Cancer," presented data from the ongoing clinical trial that demonstrated that *ispinesib* has clinical activity in patients with metastatic breast cancer. In this study, patients received *ispinesib* as monotherapy at 18 mg/m² IV as a 1 hour infusion every 21 days. At the time of this interim analysis, the best overall responses observed with *ispinesib* have been partial responses in 3 of 33 evaluable patients, measured by the RECIST criteria. These three patients had maximum decrease in tumor size ranging from 46% to 68% with the duration of response ranging from 7.1 weeks to 13.4 weeks. The overall response rate for all 33 evaluable patients was 9% with a median time to progression of 5.7 weeks. The adverse events were manageable, predictable, and consistent with the Phase I experience. The most common adverse event was Grade 4 neutropenia. *Ispinesib* plasma concentrations were comparable to those observed in Phase I clinical trials.

This ongoing Phase II clinical trial was designed to evaluate the safety and efficacy of *ispinesib* in the second- or third-line treatment of patients with locally advanced or metastatic breast cancer whose disease had recurred or progressed despite treatment with anthracyclines and taxanes. This clinical trial employs a conventional Green-Dahlberg design which specifies that the advancement to the second stage requires the satisfaction of pre-defined efficacy criteria. As a result of the aforementioned data, the predetermined response criteria to progress from Stage 1 to Stage 2 of the clinical trial have been achieved, and patients are currently enrolling in Stage 2 in which an additional 25 patients are planned to be enrolled and evaluated.

"We are encouraged to see this level of clinical activity in these chemorefractory patients," stated Dr. Andrew A. Wolff, Cytokinetics' Senior Vice President of Clinical Research and Development and Chief Medical Officer. "These patients had previously been treated with a range of different approved chemotherapeutics, so to see such decreases in tumor size for certain of these patients, confirmed by an independent review of the radiographic data, is an encouraging finding, and represents the first objective demonstration of the anti-cancer activity of *ispinesib* in patients with malignant disease."

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clinical trials evaluating *ispinesib* in other tumor types, including melanoma, head and neck, hepatocellular, colorectal and prostate cancers. In addition, the NCI plans to conduct one additional Phase II clinical trial in patients with renal cell carcinoma. The NCI is also conducting two other Phase I clinical trials evaluating an alternative schedule of *ispinesib* in leukemia and advanced solid tumors.

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