

Award Number: DAMD17-00-1-0271

TITLE: A Comprehensive Postdoctoral Training Program in
Breast Cancer

PRINCIPAL INVESTIGATOR: Robert B. Dickson, Ph.D.

CONTRACTING ORGANIZATION: Georgetown University Medical Center
Washington, D.C. 20057

REPORT DATE: July 2003

TYPE OF REPORT: Annual Summary

PREPARED FOR: U.S. Army Medical Research and Materiel Command
Fort Detrick, Maryland 21702-5012

DISTRIBUTION STATEMENT: Approved for Public Release;
Distribution Unlimited

The views, opinions and/or findings contained in this report are those of the author(s) and should not be construed as an official Department of the Army position, policy or decision unless so designated by other documentation.

20031112 193

REPORT DOCUMENTATION PAGEForm Approved
OMB No. 074-0188

Public reporting burden for this collection of information is estimated to average 1 hour per response, including the time for reviewing instructions, searching existing data sources, gathering and maintaining the data needed, and completing and reviewing this collection of information. Send comments regarding this burden estimate or any other aspect of this collection of information, including suggestions for reducing this burden to Washington Headquarters Services, Directorate for Information Operations and Reports, 1215 Jefferson Davis Highway, Suite 1204, Arlington, VA 22202-4302, and to the Office of Management and Budget, Paperwork Reduction Project (0704-0188), Washington, DC 20503

1. AGENCY USE ONLY (Leave blank)		2. REPORT DATE July 2003	3. REPORT TYPE AND DATES COVERED Annual Summary (1 Jul 02 - 30 Jun 03)	
4. TITLE AND SUBTITLE A Comprehensive Postdoctoral Training Program in Breast Cancer			5. FUNDING NUMBERS DAMD17-00-1-0271	
6. AUTHOR(S) Robert B. Dickson, Ph.D.				
7. PERFORMING ORGANIZATION NAME(S) AND ADDRESS(ES) Georgetown University Medical Center Washington, D.C. 20057 E-Mail: dicksonr@georgetown.edu			8. PERFORMING ORGANIZATION REPORT NUMBER	
9. SPONSORING / MONITORING AGENCY NAME(S) AND ADDRESS(ES) U.S. Army Medical Research and Materiel Command Fort Detrick, Maryland 21702-5012			10. SPONSORING / MONITORING AGENCY REPORT NUMBER	
11. SUPPLEMENTARY NOTES				
12a. DISTRIBUTION / AVAILABILITY STATEMENT Approved for Public Release; Distribution Unlimited			12b. DISTRIBUTION CODE	
13. ABSTRACT (Maximum 200 Words) The Comprehensive Postdoctoral Training Program In Breast Cancer Biology has successfully completed its third year. The goal of the program is to significantly extend our existing, highly successful Doctoral Training Program in Tumor Biology and several Cancer Center mechanisms that provide traditional postdoctoral training and junior faculty career development. The education and training of new investigators is essential to our progress in the prevention, detection, diagnosis, and treatment of breast cancer. The Postdoctoral Training Program in Breast Cancer provides comprehensive career development and integrates interactive research in the basic biology of breast cancer, formal and informal course work in key areas, and individualized guidance in career development. The program is enriched with both new and existing courses of interest to postdoctoral fellows covering scientific writing, teaching methodologies, scientific resources and technologies for cancer research, ethics in science, career development seminar series, and provides a firm foundation in the skills needed to succeed in a career science, as well as a focused research experience in basic breast cancer biology in a strong laboratory environment.				
14. SUBJECT TERMS Breast Cancer, translational research, prevention			15. NUMBER OF PAGES 7	
			16. PRICE CODE	
17. SECURITY CLASSIFICATION OF REPORT Unclassified	18. SECURITY CLASSIFICATION OF THIS PAGE Unclassified	19. SECURITY CLASSIFICATION OF ABSTRACT Unclassified	20. LIMITATION OF ABSTRACT Unlimited	

NSN 7540-01-280-5500

Standard Form 298 (Rev. 2-89)
Prescribed by ANSI Std. Z39-18
298-102

Table of Contents

Cover.....	1
SF 298.....	2
Introduction.....	4
Body.....	4
Key Research Accomplishments.....	5
Reportable Outcomes.....	6
Conclusions.....	7

COMPREHENSIVE POSTDOCTORAL TRAINING PROGRAM IN BREAST CANCER BIOLOGY

INTRODUCTION

The Comprehensive Postdoctoral Training Program in Breast Cancer Biology has successfully completed its third year. The goal of the program is to significantly extend our existing, highly successful Doctoral Training Program in Tumor Biology and several Cancer Center mechanisms that provide traditional postdoctoral training and junior faculty career development. The education and training of new investigators is essential to our progress in the prevention, detection, diagnosis, and treatment of breast cancer. The Postdoctoral Training Program in Breast Cancer provides comprehensive career development and integrates interactive research in the basic biology of breast cancer, formal and informal course work in key areas, and individualized guidance in career development. The program is enriched with both new and existing courses of interest to postdoctoral fellows covering scientific writing, teaching methodologies, scientific resources and technologies for cancer research, ethics in science, career development seminar series, and provides a firm foundation in the skills needed to succeed in a career science, as well as a focused research experience in basic breast cancer biology in a strong laboratory environment. Additional programmatic activities include monthly Oncology Grand Rounds, weekly Tumor Biology Seminar Series, and weekly journal clubs. The program makes use of the existing organizational structure of the Interdisciplinary Doctoral Training Program in Tumor Biology and incorporates a multi-disciplinary faculty who are devoted to research and education in breast cancer.

Postdoctoral fellows are given the opportunity to seek research and career advice from a number of senior faculty. The further development of this structured Tumor Biology Postdoctoral training program will provide excellent preparation for successful careers in cancer research.

BODY

Training and Research Accomplishments

The accomplishments of this program include the recruitment and progress of postdoctoral fellows. In the past 10 years, over 100 postdoctoral fellows have been trained by our Tumor Biology program preceptors. These fellows have an outstanding record of research productivity and peer-reviewed publication, and over 50% of them have obtained academic faculty level positions.

In the first year of the program, two postdoctoral fellows, Fadwa Attiga, Ph.D. and Tushar Deb, Ph.D., were recruited to the program. Dr. Attiga has successfully

completed her postdoctoral fellowship in Dr. Stephen Byer's lab, researching the cross regulation between the IKK and the beta-catenin signaling pathways in breast and colon cancers. Her research topics include a study of how different kinases regulate the level of oncogene beta-catenin in tumor cells and identification of the key players in the signaling cascades that alter the beta-catenin protein level and transcriptional activity in tumors.

Dr. Deb, has completed his postdoctoral fellowship in Dr. Robert Dickson's lab and is now a Junior Faculty member at Georgetown University. The main focus of Dr. Deb's research is to explore the EGF cell-survival signaling in transgenic MMTV-c-Myc mammary epithelial cells. Proto-oncogene c-Myc sensitizes apoptosis in absence of serum in these cells. The specific role of a calmodulin dependent kinase in this survival signaling is also being explored.

Marcia Noble, Ph.D., and Kerrie O'Brien, Ph.D., were recruited into the second year of the program. Dr. Noble began her postdoctoral fellowship in September 2001, under collaborative mentorship of Dr. Michael Johnson and Dr. Robert Dickson. Dr. Noble's research focus is on the mechanism of VEGF to promote breast cancer metastases in transgenic mouse models. Dr. Noble is continuing her postdoctoral fellowship with Dr. Dickson. Dr. O'Brien began her postdoctoral fellowship in March 2002 under mentorship of Dr. Robert Clarke. Dr. O'Brien's research focuses on the resistance to retinoids in breast cancer. Constanze Hample, PhD was also recruited into this class of the postdoctoral fellowship program in September 2002 in the laboratory of Dr. Stephen Byers. Dr. Hample is currently focusing on Cadherin-11 and breast cancer invasion.

As noted earlier, our Postdoctoral Program incorporates elements of our existing Tumor Biology Ph.D. program, as well as new, Postdoctoral Training-specific elements as a part of a Breast Cancer Prevention Track. In addition to the existing core courses of the Interdisciplinary Doctoral Training Program in Tumor Biology, new course components have been incorporated into the Breast Cancer Prevention track in Spring 2002. These include a course in Biostatistics, *Applied Biostatistics*, that has been refocused on statistical design and methodology for research rather than biostatistics theory, and a Cancer Genetics course, *Genetics, Health, and Society in the 21st Century*, which focuses on practical and ethical questions raised by genetic information and technology. Both courses had a very successful first year and will continue to be offered. A new course in Genetics, *Human and Microbial Genetics*, has been developed and will be offered in the Fall.

KEY ACCOMPLISHMENTS

- *Recruitment of Trainees and Progress of Trainees:*
 - Two postdoctoral fellows have been recruited into the fourth class of the Comprehensive Postdoctoral Training Program in Breast Cancer Biology: **Rebecca Riggins**, start date 6/23/03 in Dr. Robert Clarke's lab, and **Fransiscus Utama** (start date 01/02/04).

- One postdoctoral fellow, Dr. Constanze Hampel, was recruited into the third class of the program. Dr. Hampel began her fellowship in September 2003 in the laboratory of Dr. Stephen Byers.
- Two postdoctoral fellows were recruited into the second class of the Comprehensive Postdoctoral Training Program In Breast Cancer Biology: Marcia Noble, Ph.D., and Kerrie O'Brien, Ph.D. Dr. Noble began the program in September, 2001 in Dr. Michael Johnson's and Dr. Robert Dickson's laboratories. Dr. O'Brien began March, 2002 in Dr. Robert Clarke's laboratory.
- Dr. Deb, from the first class, has successfully completed his postdoctoral fellowship and is currently a Junior Faculty member at Georgetown University.
- Dr. Attiga, from the first class, completed her first year with Dr. Stephen Byers and is currently on a maternity leave of absence from the program.
- *New Courses:*
 - *Human and Microbial Genetics* had a very successful Fall semester and will be offered again next fall.
 - Two courses, *Applied Biostatistics* and *Genetics, Health and Society in the 21st Century* both will continue to be offered.

REPORTABLE OUTCOMES

- *Publications:*
 - Pei XF, Noble M, Johnson MD, Rosfjord E, and Dickson RB. Expant-Cell Culture of Primary Mammary Tumors from MMTV-c-Myc Transgenic Mice. In Vitro Cell Biology 2003 (In Press).
 - Clarke R, Liu MC, Bouker KB, Gu Z, Lee RY, Zhu Y, Skaar TC, Gomez B, O'Brien K, Wang Y, and Hilakivi-Clarke LA. Antiestrogen Resistance in Breast Cancer and the Role of Estrogen Receptor Signaling. Oncogene 2003 (In Press).
- *Abstracts:*
 - Deb, T, and Dickson, R. Calmodulin as a Positive Modulator of EGF Survival Signaling in MMTV-C-MYC Mouse Mammary Epithelial Cells. DOD ERA of Hope Meeting, Orlando, FL, 2002.
 - Noble M, Rosfjord EC, Sharp R, Merlino G, and Dickson RB. Ectopic VEGF Expression Promotes Metastasis in Bitransgenic Breast Cancer Model. DOD ERA of Hope Meeting, Orlando, FL, 2002.

- Teo M, Attiga F, Jarret C, and Byers S. Cytokine Regulation of β -catenin Signaling. Fourth Annual Lombardi Research Fair. Georgetown University Medical Center, Washington, D.C. 2002.

CONCLUSIONS

The goal of the program is to significantly extend our existing, highly successful Postdoctoral Training Program in Cancer with a new specialization in Breast Cancer Biology. We have successfully completed our first year of the Comprehensive Postdoctoral Training Program in Breast Cancer Biology and have three postdoctoral fellows who are now entering their second year of research. We have recruited two new postdoctoral fellows to the program.