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GRANT NUMBER DAMD17-96-1-6170

TITLE: Use of p53 Mutation Analysis for Staging Breast Cancer

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Fort Detrick, Maryland 21702-5012

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FOREWORD

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_____ In the conduct of research utilizing recombinant DNA, the investigator(s) adhered to the NIH Guidelines for Research Involving Recombinant DNA Molecules.

_____ In the conduct of research involving hazardous organisms, the investigator(s) adhered to the CDC-NIH Guide for Biosafety in Microbiological and Biomedical Laboratories.

Mary Belle White 8/31/98
PI - Signature Date

TABLE OF CONTENTS

PART 1.

SF 298 Report Documentation Page **Pg. 2**

PART 2.

Foreword **Pg. 3**

PART 3.

Table of Contents **Pg. 4**

PART 4.

Introduction, Body and Conclusions **Pg. 5**

PART 5.

Summary of Work on Samples **Pg. 6**

Attachments

Personnel/Voucher History

**“Use of p53 Mutation Analysis For Staging Breast Cancer”
Final Report**

Introduction, Body and Conclusion

This grant began with a collaboration between Oncormed and Dr. Timothy O’Leary from the Department of the Army. Dr. O’Leary was to provide the samples for the evaluation of the Molecular Staging Assay and its applications for detection of micrometastasis in breast cancer. The samples Dr. O’Leary provided were from a retrospective study. When these samples were collected, the protocols in use at the time were not compatible with retaining the genetic material intact. As a result, the DNA that was isolated to perform the Molecular Staging Assay was quite degraded. Oncormed extended a great deal of energy in attempts to salvage DNA of sufficient quality and quantity from these samples to do the study without success (see report from ?).

After it was decided that Dr. O’Leary’s samples could not be used to complete this study, another collaborator was identified. Dr. Bruce Ponder from ? had 100 samples from breast cancer patients that could be used to complete this study. An initial evaluation of the samples indicated they were of sufficient quality and quantity. The work was initiated on these samples and a summary of the progress is attached.

Over the past several months, the focus of Oncormed has been changed. Our Director of Scientific Affairs, Dr. Christine Carter, left the company in the summer of 1997 and our Grants Department was eliminated. I then took over as Principal Investigator. However, over the past year I and my department have been forced to place the majority of the resources on basic business considerations related to the survival of the company. This has now culminated in the acquisition of Oncormed by another biotechnology company, Gene Logic. As a result of this purchase all current on-going projects at Oncormed have been evaluated for their compatibility with the goals of the new combined entity. It has been decided that this project does not fit into the goals of Gene Logic.

It is a great disappointment to myself and the Oncormed staff that we will be unable to complete this study. We hope that the unused portion of the funding can be redirected to other worthy studies.

**“Use of p53 Mutation Analysis For Staging Breast Cancer”
Final Report**

Summary of Work on Army Grant Samples

The following is a brief summary of the various stages of sample processing the Army Grant samples have been through as of April, 1998:

Stage I. DNA
Extraction from Paraffin Embedded Samples (slides).

50

paraffin samples (5 slides / samples or 250 slides) were scrapped from slides for DNA extraction.

Stage II. PCR
Amplification using p53 primers for exons 5, - 9.

266

PCR reactions were conducted on the above mentioned DNA samples.

Samples were analyzed on three, large 1.2% agarose gels.

Stage III PCR
Purification of amplified samples.

Samples are currently being purified for dye attachment -sequencing reaction to verify the presence or absence of p53 mutations.

Stage IV Dye -
Attachment / Sequencing Reaction. *(present status)

Samples that have been successfully amplified by PCR and purified will undergo dye - attachment.

The project was canceled and at the time of cancellation, samples were being prepared for dye attachment. Samples that had been dye-attached but had unsuccessful runs were slated for re-attachment however due to the cancellation of the project these samples were never re-dye attached. The remaining paraffin samples were sent back to Cambridge along with the DNA that was isolated from these paraffin samples. PCR fragments were not mailed because of quality

concerns due to freeze/thaw procedures involving samples diluted in water as opposed to TE buffer.

**PUBLIC VOUCHER FOR PURCHASES AND
SERVICES OTHER THAN PERSONAL**

Continuation Sheet

Voucher # 1
Sheet No. 1

U.S. Department, Bureau, or Establishment:
U.S. Army Medical Research Acquisition Activity

Oncormed, Inc.
Contract No. DAMD17-96-1-6170

Reimbursable costs from 8/1/97 thru 4/30/98

<u>Direct Costs</u>	<u>Amount billed for current period</u>	<u>Cumulative amount from inception to date of this billing</u>
Salaries	\$3,840	\$10,405
Fringe Benefits	\$511	\$1,384
Travel	\$0	\$1,042
Supplies	\$692	\$5,675
Subcontract	\$0	\$120
Other	\$0	\$0
Equipment	\$0	\$0
Total Direct Costs	\$5,043	\$18,626
 <u>Indirect Costs</u>		
Indirect Cost (60% of Labor + Fringe)	\$2,611	\$7,074
G&A Cost (10% of All Cost)	\$765	\$2,570
Total Indirect Costs	\$3,376	\$9,644
Total Amount Claimed	\$8,419	\$28,270

**PUBLIC VOUCHER FOR PURCHASES AND
SERVICES OTHER THAN PERSONAL**

Continuation Sheet

Voucher # 1
Sheet No. 2

U.S. Department, Bureau, or Establishment:
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Oncormed, Inc.
Contract No. DAMD17-96-1-6170

Reimbursable costs from 8/1/97 thru 4/30/98

<u>Direct Costs</u>	<u>Amount billed for current period</u>	<u>Cumulative amount from inception to date of this billing</u>
<u>Salaries</u>	\$3,840	\$10,405
<u>Travel</u>		
Transporation	\$0	\$286
Loding	\$0	\$626
Meals	\$0	\$130
Total Travel Costs	\$0	\$1,042
<u>Supplies</u>		
Paraffin/DNA preparation reagents	\$250	\$850
PCR amplification reagents	\$442	\$4,042
Barrier pipet tips	\$0	\$300
Camera film	\$0	\$30
Laboratory disposables	\$0	\$453
Total Supplies Costs	\$692	\$5675

**PUBLIC VOUCHER FOR PURCHASES AND
SERVICES OTHER THAN PERSONAL**

Continuation Sheet

Voucher # 1
Sheet No. 3

U.S. Department, Bureau, or Establishment:
U.S. Army Medical Research Acquisition Activity

Oncormed, Inc.
Contract No. DAMD17-96-1-6170

Reimbursable costs from 8/1/97 thru 4/30/98

<u>Direct Costs</u>	<u>Amount billed for current period</u>	<u>Cumulative amount from inception to date of this billing</u>
<u>Subcontract</u>		
Consultant came to Oncormed to demonstrate testing procedure in the laboratory	\$0	\$120
Total Subcontract Costs	\$0	\$120
<u>Fringe Benefits</u>		
13.30% of labor	\$511	\$1,384
<u>Indirect Costs</u>	\$2,611	\$7,074
60% of labor + fringe		
<u>G&A Cost</u>	\$765	\$2,570
10% of all costs		
Total Amount Claimed	\$8,419	\$28,270