AD	)		

GRANT NO: DAMD17-94-J-4376

TITLE: "Positron Emitter I124 Iododeoxyuridine as a Tracer to Follow DNA Metabolism on Scans and in Tumor Samples in Advanced Breast Cancer"

PRINCIPAL INVESTIGATOR(S): Teresa Ann Gilewski, M.D.

CONTRACTING ORGANIZATION: Sloan-Kettering Institute for

Cancer Research

New York, New York 10021

REPORT DATE: September 1995

TYPE OF REPORT: Annual



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.

# REPORT DOCUMENTATION PAGE

Form Approved OMB No. 0704-0188

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 the 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.

Davis Highway, 30tte 1204, Armigton, VA 22202-3308			A CAMPAGE OF THE PROPERTY OF T
1. AGENCY USE OMAY (Leave blank)		3. REPORT TYPE AND	
C. STACTS P. A DACS. #28POTS573 P.	September 1995	Annual I Sep	94 - 31 Aug 95 5. FUNDING NUMBERS
6. THE AND SUBTIFIE Positron Emitter I124 Iod	odeoxvuridine as a	Tracer to Follow	
DNA Metabolism on Scans a Breast Cancer	nd in Tumor Samples	in Advanced	DAMD17-94-J-4376
6. AUTHOR(S)			
Teresa Ann Gilewski, M.D.			
7. PERFORMING ORGANIZATION NAME Sloan-Kettering Institute New York, New York 10021	for Cancer Researc	h	8. PERFORMING ORGANIZATION REPORT NURBER
9. SPOUSORING/MONITORING AGENC	v name(s) and addiess(es	,	10. SPONSORING / MONITORING
U.S. Army Medical Rese Fort Detrick, Maryland	earch and Materiel (		AGENCY REPORT NUMBER
11. Supplementary notes			
12a. DISTRIBUTION / AVAILABILITY STA	TEMENT		126. DISTRIBUTION CODE
A	alagas digtribution	n unlimited	
Approved for public re	erease; distribution	i uniimiteu	
13. ABSTRACT (Meximum 200 words)			
,			
			0.
14. Subject Tennas Positron Emission T	ine, 15. NUMBER OF PAGES		
Advanced Breast Can Humans, Clinical Tr	cer, DNA Metabol	ism, Response	16. PRICE CODE
17. SECURITY CLASSIFICATION 18. OF REPORT	SECURITY CLASSIFICATION OF THIS PAGE	19. SECURITY CLASSIFI OF ADSTRACT	CATION 20. LIMITATION OF ABSTRA
Unclassified	Unclassified	Unclassified	Unlimited

## GENERAL INSTRUCTIONS FOR COMPLETING SF 298

The Report Perturbantation Page (RDP) is used in announcing and cataloging reports. It is important that this information be consistent with the rest of the report, particularly the cover and title page. Instructions for filling in each block of the form follow. It is important to stay within the lines to meet optical spanning requirements.

Elack 1. Agency Use Only (Leave blank).

Elected. Respot Date. Full publication date including day, mouth, and year, if available (e.g. 1 Jan 80). Mant cito at least the year.

Moch 3. Type of Report and Dates Covered. State whether report is interim, final, etc. If applicable, enter inclusive report dates (e.g. 10 Jun 87 - 36 Jun 88).

Block 4. <u>Title and Subtitle</u>. A title is taken from the part of the report that provides the most meeningful and complete information. When a report is proposed in more than one volume, report the primary title, add volume number, and include subtitle for the specific volume. On classified decuments enter the title classification in parantheres.

Electric funding Numbers. To include contract and great numbers; may include program element number(s), project number(s), task number(s), and work unit number(s). Use the following labels:

C - Contract

PR - Project

6 - Grant P5 - Brogram TA - Task WU - Work Unit

VVU - Work Unit Accession No.

Electic. <u>Authoris</u>. Name(s) of person(s) responsible for writing the report, performing the research, or credited with the content of the report. If editor or compiler, this should follow the name(s).

Black 7. <u>Performing Organization Name(s) and Addressfeet</u>. Salf-captanatory.

Black E. <u>Furiorating Organization Report</u>
<u>Kumbar</u>, Enter the unique alphanumeric report number(s) estimated by the organization performing the report.

Float C. Supposition (Monitoring Agency Name(s) and Address(ed). Self-explanatory.

Elith 18. Spanishing/Manitoring Agency Resert Number. (M. Imourn)

Each St. Supplementary Notes. Enter information and included elsewhere such as: Prepared in cooperation with...; Trans. of...; To be published in.... When a report is revised, include a distance whather the new report supersedes on rupplement who the older report.

Block 12a. <u>Distribution/Availability Statement.</u>
Denotes public availability or limitations. Cite any availability to the public. Enter additional limitations or special markings in all capitals (e.g. NOFORN, REL, ITAR).

DOD - See DoDD 5230.24, "Distribution Statements on Technical Documents."

DOE - See authorities.

NASA - See Handbook NHB 2200.2.

NTIS - Leave blank.

Block 12b. <u>Distribution Code</u>.

DOD - Leave blank.

DOE - Enter DOE distribution categories from the Standard Distribution for Unclassified Scientific and Technical Reports.

NASA - Leave blank.

NTIS - Leave blank.

Block 13. <u>Abstract</u>. Include a brief (*Maximum* 200 words) factual summary of the most significant information contained in the report.

**Block 14.** Subject Terms. Keywords or phrases identifying major subjects in the report.

**Block 15.** <u>Number of Pages</u>. Enter the total number of pages.

Block 16. <u>Price Code</u>. Enter appropriate price code (NTIS only).

Blocks 17. - 19. <u>Security Classifications</u>. Self-explanatory. Enter U.S. Security Classification in accordance with U.S. Security Regulations (i.e., UNCLASSIFIED). If form contains classified information, stamp classification on the top and bottom of the page.

Block 20. <u>Limitation of Abstract</u>. This block must be completed to assign a limitation to the abstract. Enter either UL (unlimited) or SAR (same as report). An entry in this block is necessary if the abstract is to be limited. If blank, the abstract is assumed to be unlimited.

## FOREWORD

Opinions, interpretations, conclusions and recommendations are those of the author and are not necessarily endorsed by the US Army.

Where copyrighted material is quoted, permission has been obtained to use such material.

Where material from documents designated for limited distribution is quoted, permission has been obtained to use the material.

Citations of commercial organizations and trade names in this report do not constitute an official Department of Army endorsement or approval of the products or services of these organizations.

In conducting research using animals, the investigator(s)

adhered to the "Guide for the Care and Use of Laboratory
Animals," prepared by the Committee on Care and Use of Laboratory
Animals of the Institute of Laboratory Resources, National
Research Council (NIH Publication No. 86-23, Revised 1985).

For the protection of human subjects, the investigator(s) adhered to policies of applicable Federal Law 45 CFR 46.

In conducting research utilizing recombinant DNA technology, the investigator(s) adhered to current guidelines promulgated by the National Institutes of Health.

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.

#### SUMMARY

#### Grant # DMAD 17-94-J-4376

"Positron emitter  $I^{124}$  iododeoxyuridine as a tracer to follow DNA metabolism on scans and in tumor samples in advanced breast cancer."

The objectives of this study are: (1) to determine whether the biologic activity of locally advanced Stage III breast cancer as measured by <sup>124</sup>I-iododeoxyuridine (IUdR) uptake on positron emission tomography (PET) scans pre and post chemotherapy can be correlated with the clinical response as determined by physical examination and conventional radiographic studies, (2) to demonstrate that incorporation of IUdR is into the DNA contained within the tumor and that it correlates with the subsequent tumor response and proliferative activity of the tumor, (3) to further assess the biologic activity of tumor sites and clinical response by using a program which fuses PET scan images on computed tomography (CT) scans, magnetic resonance imaging (MRI) or SPECT bone scans.

Patients with Stage III breast cancer will have a complete extent of disease evaluation including routine radiographic studies. A PET scan with IUdR will be obtained within 2 weeks prior to therapy and after 4 cycles of chemotherapy. Whole body emission scans will be performed 24 hours after intravenous injection of 8 mCi of IUdR. Tumor biopsies will be obtained on the day of the PET scan and assessed for incorporation of IRdR into DNA. Flow cytometry and Ki-67 stains will also be obtained. Fusion imaging will generate resliced PET images that correspond to appropriate original CT, MRI or SPECT bone scan images.

There has been an unforseen delay in the construction of the site for the positron emission tomography (PET) scan. It is expected that the PET scan will be operational in November 1995.

SecoA	sion For				
NTIS	GRA&I				
DTIC	TAB				
Unannousced					
Justi	fication_				
By Distr	1but boo				
Avai	lability*	Coces			
	Avall and	/or			
<b>Bist</b>	Spec 11	,			
9.1	THE CANADA TO COLOR OF TELESCOPE				