

NYO - 2181 - 4

EPIDEMIOLOGICAL FOLLOW-UP OF THE NEW JERSEY RADIUM CASES

- 1. REPORT OF THE EPIDEMIOLOGICAL AND MEDICAL STUDY GROUP AND THE PROJECT SUBCONTRACTORS
- II. RESUME OF COOPERATIVE LABORATORY STUDIES ON PROJECT CASES AND SPECIMENS
- III. RESUMES OF FINDINGS FROM INDIVIDUAL CASE STUDIES

Progress Report to October 1, 1966

March 1967 [DTIE Issuance Date] DTIC QUALATY INSPECTSO (

20000912 088

Radium Research Project New Jersey State Department of Health West Orange, New Jersey

DUDUEENT LIBRARY

26635

Reproduced From Best Available Copy

Distribution Control Programment Approved for Public Release Approved for Public Release Distribution Unlimited March DTIE Radiu New West

UNITED STATES ATOMIC ENERGY COMMISSION · DIVISION OF TECHNICAL INFORMATION

LEGAL NOTICE

This report was prepared as an account of Government sponsored work. Neither the United States, nor the Commission, nor any person acting on behalf of the Commission:

A. Makes any warranty or representation, expressed or implied, with respect to the accuracy, completeness, or usefulness of the information contained in this report, or that the use of any information, apparatus, method, or process disclosed in this report may not infringe privately owned rights; or

B. Assumes any liabilities with respect to the use of, or for damages resulting from the use of any information, apparatus, method, or process disclosed in this report.

As used in the above, "person acting on behalf of the Commission" includes any employee or contractor of the Commission, or employee of such contractor, to the extent that such employee or contractor of the Commission, or employee of such contractor prepares, disseminates, or provides access to, any information pursuant to his employment or contract with the Commission, or his employment with such contractor.

This report has been reproduced directly from the best available copy.

Printed in USA. Price \$3.00. Available from the Clearinghouse for Federal Scientific and Technical Information, National Bureau of Standards, U. S. Department of Commerce, Springfield, Virginia 22151.

NYO-2181-4 BIOLOGY AND MEDICINE (TID - 4500)

New Jersey State Department of Health Radium Research Project 11 Washington Street West Orange, New Jersey

EPIDEMIOLOGICAL FOLLOW-UP OF THE NEW JERSEY RADIUM CASES

- I. REPORT OF THE EPIDEMIOLOGICAL AND MEDICAL STUDY GROUP AND THE PROJECT SUBCONTRACTORS
- II. RÉSUMÉ OF COOPERATIVE LABORATORY STUDIES ON PROJECT CASES AND SPECIMENS
- III. RÉSUMÉS OF FINDINGS FROM INDIVIDUAL CASE STUDIES

Progress Report to October 1, 1966

Hyman W. Fisher, M.D.Project Medical DirectorCarye-Belle Henle, M.D.Project RadiologistRobert Bonda, D.D.S.Project DentistSamuel C. Ingraham 2nd, M.D.*Project Director

Supported by U. S. Atomic Energy Commission Contract AT(30-1)-2181

*On loan from Division of Radiological Health U. S. Public Health Service

SUBCONTRACTORS AND SPECIAL CONSULTANTS

- General Histopathology Studies-New Jersey College of Medicine and Dentistry Hugh G. Grady, M.D., Professor of Pathology William D. Sharpe, M.D., Assistant Professor Pathology
- Oral Histopathology Studies-Georgetown University School of Dentistry Joseph L. Bernier, D.D.S., Professor & Chairman, Oral Pathology Department Joseph D. Belzile, D.D.S., Consultant, Oral Pathology
- Clinical Laboratory Studies-Newark Beth Israel Hospital Lester M. Goldman, M.D., Director of Laboratories
- Statistical Studies-Rutgers, the State University Ellis R. Ott, Ph.D., Director, Statistics Center Thomas Hayton, Ph.D., Associate Professor, Statistics Center

COOPERATING PROJECTS

Whole Body Counting, Breath Radon Measurements, Radiochemistry Studies

Merril Eisenbud, Sc.D., Professor & Director Henry G. Petrow, Ph.D., Research Scientist and staff of Environmental Radiation Laboratory Institute of Environmental Medicine New York University Medical Center supported by U. S. Atomic Energy Commission Contract AT(30-1)-3086

Breath Radon Measurements

Robley D. Evans, Professor and staff of Department of Physics Massachusetts Institute of Technology supported by U. S. Atomic Energy Commission Contract AT(30-1)-952

ii

PREFACE

This is the final annual report of a series which describes the epidemiological follow-up of a group of dial painters and other radium workers exposed more than 40 years ago to radium-226 and/or radium-228 (mesothorium) used as the activator for radioluminous watch, clock and instrument dial paint. Many of these individuals still carry measureable body burdens of the isotopes.

Summaries of data from fifteen individual cases not previously reported continue the procedure established in prior reports (NYO-2760, NYO-10604, NYO-2181-2, NYO-2181-3). Additional information is known about each case but not all available data are presented, merely the highlights of the results of medical, dental and roentgenographic studies, and estimates of body burden of radium.

Additional data about the entire group of cases may be found in reports previously published, namely:

- NYO-2759: The Epidemiological Follow-Up of the New Jersey Radium Dial Painters, Progress Report to January 31, 1962
- NYO-2760: Résumés of Findings from Individual Case Studies, Epidemiological Follow-Up of Radium Cases, July 1, 1962
- NYO-10604: Epidemiological Follow-Up of the New Jersey Radium Cases, Progress Report to July 1963
- NYO-2761: Atlas of Current Roentgenographic Findings in the New Jersey Radium Cases, October 1963
- NYO-2181-1: Current Data from the Epidemiological Follow-Up of the New Jersey Radium Cases, October 1, 1963
- NYO-2181-2: Epidemiological Follow-Up of the New Jersey Radium Cases, Progress Report to April 1964
- NYO-2181-3: Epidemiological Follow-Up of the New Jersey Radium Cases, Progress Report to September 1, 1965

Further information regarding the New Jersey Radium Research Project cases is contained in the complete file maintained on each case in the Project office. A group of approximately 100 controls has been recruited and is being studied by the same clinical and laboratory examinations as used for cases.

Field study and collection of data on the cases and controls of this Project will be completed about February 28, 1967. Statistical collation and evaluation of the data relating retained body burdens of radium to observed bio-medical effects is concurrently in progress. An evaluative report summarizing information collected over the ten years of the Project is being prepared for publication at conclusion of this U. S. Atomic Energy Commission Research Contract, approximately June 30, 1967.

iii

Recognition is given to the individuals who performed the studies and prepared this report because the New Jersey Radium Research Project is an Epidemiological Field Office coordinating consultant medical, dental, radiographic and medical laboratory clinical studies and subcontracting or cooperating in histopathological, radiochemical and radiophysical laboratory studies of its cases, its controls and clinical and postmortem specimens obtained from them.

> Acknowledgement is made of contribution to the progress of this Project and to the preparation of this report by:

William G. Bernhard, M.D., St. Barnabas Medical Center Gwilym S. Lodwick, M.D., University of Missouri School of Medicine Hubert A. Sissons, M.B., F.C.Path., Royal National Orthopaedic Hospital, London

> by the following members of the staff of the Radium Research Project:

> > Mary Cull Emily Haag, R.N. Beatrice Hait Diane Kopesky Ann Johnson Medora MacLaren Nancy Manner, R.N. Mary Rallo, R.N. Catharine Rowland Elizabeth Stewart

and by the following staff involved in histopathology studies:

Gerome Bell, H.T. (A.S.C.P.) Eugene Zimmerman John J. Raines

ABSTRACT

The 1966 Progress Report of the New Jersey Radium Research Project gives a brief résumé of the goals and organization of the Project; summary of clinical, laboratory and statistical studies conducted during the reporting period; revised schema for coding x-ray bone lesions; and résumés of findings from fifteen cases not previously published.

TABLE OF CONTENTS

SUBCONTRACTORS AN PREFACE	ND SPECIAL	CONSULTANTS,	COOPERATING	PROJECTS	ii iii
ABSTRACT					v
INTRODUCTION					1
Project Organ	ization				2

PART I REPORT OF THE EPIDEMIOLOGICAL AND MEDICAL STUDY GROUP AND THE PROJECT SUBCONTRACTORS

I. CLINICAL STUDIES

II.

III.

Medical Studies	9
Roentgenographic Studies	12
Dental Studies	14
LABORATORY STUDIES	
Clinical Laboratory	15
Histopathology (Soft Tissue and Bone) General Pathology Oral Pathology	16 19
Body Burden Assessment	21
STATISTICAL STUDIES	24

PART II RESUME OF COOPERATIVE LABORATORY STUDIES ON PROJECT CASES AND SPECIMENS

I.	"Estimates of Total Body Radium in the New Jersey Dial Painters"	1-1
II.	"A Study of the Distribution of Ra-226, Ra-228, Pb-210 and Th-228 in Bone and Soft Tissue of Radium Dial Painters"	2-1
II.	"Discussion of Possible Mechanisms Involved in the Apparent	

I Concentration of ^{22b}Ra and ²¹⁰Pb in Aorta and Thyroid" 7-1

vi

Page

PART III RÉSUMÉS OF FINDINGS FROM INDIVIDUAL CASE STUDIES

			Page
Case	No.	5181	1
Case	No.	5208	3
Case	No.	5250	4
Case	No.	5303	5
Case	No.	5368	6
Case	No.	5394	7
Case	No.	5396	8
Case	No.	5519	9
Case	No.	5561	10
Case	No.	5671	11
Case	No.	5674	12
Case	No.	5750	13
Case	No.	5804	14
Case	No.	5831	15
Case	No.	5908	16

APPENDIX

EXHIBIT	I	Case Finding and Follow-Up Progress Report to October 1, 1966
EXHIBIT	2	Table of Organization
EXHIBIT	З	Lodwick Schema for Coding X-Rays (Revised)

vii

INTRODUCTION

This report deals with progress of the Radium Research Project of the New Jersey State Department of Health in epidemiological follow-up and special studies on radium dial painters, other radium workers and selected controls from September 1965 to October 1966.

Previous reports (See Preface) detail information on Project background, methods employed, résumés of cases, observations and impressions based on case finding; occupational and medical histories; medical, dental and roentgenographic examinations; clinical, pathological, radiochemical and radiophysical laboratory studies on cases; and analysis of samples of radioactive paints to which the cases were originally exposed.

As of October 1, 1966, 215 cases and 89 controls have undergone complete or partial study to estimate the long-term effects of a radium body burden in man. In previous reports, data from various phases of the Project have been tabulated and studied. At present, these listings are being brought up to date and additional listings are being prepared. Evaluation of accumulated data is under way. A final set of studies on all cooperating cases and controls will be completed about February 28, 1967. Addition of data from these studies will permit evaluation of most of the information collected by the Project.

Project Organization*

Purpose

The Radium Research Project of the New Jersey State Department of Health, supported by U. S. Atomic Energy Commission Contract AT(30-1)-2181, is reexamining through application of modern epidemiological methodology and reevaluating the low level, long-term, biomedical effects of retained body burdens of radioactive substances (radium-226 and radium-228) in former radium dial painters and other radium workers. The chief interest of the Project lies in refining radiation safety standards, especially for internal body burdens of radium and other bone-seeking isotopes.

This study concentrates on approximately 200 cooperative individuals from the slightly more than 1000 former workers of the radium dial painting industry in Northern New Jersey (See Exhibit 1). Most of these people were employed in the industry between 1913 and 1928. A selected group of presumptively unexposed sibling controls is also under study.

Overt diseased states such as pathological fracture, osteogenic sarcoma, paranasal sinus carcinoma and massive bone necrosis, while dramatic and of commanding interest in understanding biomedical effects of irradiation, are probably results of relatively obscure changes produced in cells or tissues by prolonged or repeated exposures. In the minds of the Project consultants and other Project scientific personnel, both the beginning change and the final disaster to the individual are equally important. If the radiation safety goal of the Project is to be approached, studies must concentrate on relatively low body burden cases with particular attention in them to the unexplained radiolucent areas in mandibles and long bones,

*Revised and reprinted this year to insure availability of this information to individuals not seeing previous Project publications or reports.

uneven thickening of bone cortices, changes in trabecular patterns and calcification, microanatomical changes in soft tissues or individual cells, and subclinical changes in blood cellular or serum components which may be precursors of later, fatal irradiation sequellae. The material appears favorable for this approach. Other studies are left to sister projects, such as the ones under Robley D. Evans, Department of Physics, Massachusetts Institute of Technology (AEC Contract AT(30-1)-952); Merril Eisenbud, Institute of Environmental Medicine, New York University Medical Center (AEC Contract AT(30-1)-3086); Asher J. Finkel, Director of Health Division and L. D. Marinelli, Director of Radiological Physics Division, Argonne National Laboratory; and Robert J. Hasterlik, Department of Medicine, Argonne Cancer Research Hospital.

Goals

Simply stated, the goals of the Project are four-fold:

1. Detection of the earliest possible biomedical changes associated with internal irradiation among radium workers which initiate series of events that culminate in grossly harmful or lethal injuries.

irradiation → biomedical effects + time → deleterious effect
2. Assessment of aggregate internal radiation exposures of tissues,
cells and bodies of radium workers.

3. Collation of aggregate radiation exposures with objective biomedical effects observed in radium workers.

4. Evaluation of radiation doses (body burdens) and biomedical effects observed in radium workers in terms of radiological health or radiation safety standards applicable to radiation workers and to the populace.

Structure (See Exhibit 2)

The New Jersey Radium Research Project, operating under the general direction of the Commissioner, New Jersey State Department of Health, consists of a Project Director, a Project Medical and Dental Staff, a Field and Office Staff, several clinical laboratory consultants, several cooperating independent research groups and several scientific subcontractors.

Epidemiological activities such as case finding, field case follow-up, obtaining of occupational histories, compilation of record folders, storage and distribution of autopsy specimens, etc., are done in the Project office under direction of the Project Director. This immediate Project office group is responsible for general administration of the Project, obtaining vital statistics records on cases and controls, arranging for post-mortem examinations, coding and punching data for statistical processing, etc.

The Project Medical and Dental Staff together with the Project Director comprise the steering group for the Project. They also examine cases and controls, perform the roentgenographic surveys on them, evaluate the clinical findings, evaluate the roentgenograms and help advise on evaluation of data collected.

The clinical laboratory consultants are responsible for performing and reporting clinical laboratory tests on cases and controls.

Special radiation studies such as whole body counting and breath radon measurement for estimating body burdens, and radiochemical analysis of soft tissue and bone specimens for radioisotope content are performed at New York University Medical Center under cooperative arrangement with

Professor Merril Eisenbud and his staff. For interproject comparison purposes, breath radon measurements, and in the past, whole body counts, are done on selected cases under cooperative arrangement at the Massachusetts Institute of Technology Department of Physics with Professor Robley Evans and his staff. Special excretion studies have been done at New York University Medical Center and at Argonne National Laboratory, Radiological Physics Division.

Processing, reading and evaluating of surgical and post-mortem specimens from cases and controls are done under subcontract agreements. General pathological studies are done at the New Jersey College of Medicine and Dentistry where Dr. Hugh Grady and Dr. William Sharpe provide histopathologic interpretation of soft tissues and bone and conduct a special study of early radiation osteitis. Oral pathological studies are done at Georgetown University School of Dentistry where Dr. Joseph Bernier directs histopathologic interpretation of oral specimens with the assistance of Dr. Joseph Belzile.

Under subcontract at the Rutgers University Statistics Center, Professors Ellis Ott and Thomas Hayton provide consultation and data processing toward statistical evaluation of the data collected and recorded under the other activities of the Project.

Activities

The New Jersey Radium Research Project began in November 1957 as a feasibility study, converted to an ongoing epidemiological project in March 1958. Major effort during 1957 to 1960 went into identifying by name and locating by address the early dial painters and radium workers

of northern New Jersey. During 1959 to 1966 the Project collected medical, dental, x-ray and laboratory data on cases willing to cooperate. Current efforts are directed toward data collation and evaluation but with a final effort being expended in completing the data collection on cases and in developing and examining the selected control group.

Areas of data compilation and of clinical and laboratory study are as follows:

1. Studies performed on cases

a. Occupational history

- b. Medical, family and spouse history
- c. Dental history
- d. Physical examination
- e. Dental examination
- f. X-Ray survey and examination
- g. Clinical laboratory series of tests
- h. Whole body gamma count (NYU)

Two sets of medical, dental, etc. studies plus three additional laboratory sets are planned for each case.

2. Studies performed on controls include one set of medical,

dental, etc. plus four additional laboratory sets.

3. Additional special items as appropriate

- a. Copies of birth and death certificates
- b. Post-mortem examination at which extensive collection of bone specimens is done
- c. Histopathology on surgical, biopsy and post-mortem specimens
- d. A special research study of early radiation osteitis in bones of cases.

4. Development of objective recording systems for the data observed

in (a) histopathology (general and oral) (b) roentgenograms and (c) clinical laboratory.

5. Collation of medical and other findings with gross body burdens.

6. Evaluation of clinical laboratory data for significant variations and comparison with body burden.

7. Radiochemical analysis of selected bone specimens from cases and controls:

- a. A standard six bones per case whenever appropriate specimens are available
- b. Special comparative study of radioisotope content (Ra-226, Ra-228, Pb-210) of multiple bone and soft tissue specimens from several selected cases.

8. With compilation of data nearing completion, attention is now being directed to the points of interest they delineate and to meaningful conclusions that may be drawn therefrom. PART I

REPORT OF THE EPIDEMIOLOGICAL AND MEDICAL STUDY GROUP AND THE PROJECT SUBCONTRACTORS

I. CLINICAL STUDIES

Medical Studies

During this reporting period, three new radium case examinations and seven follow-up examinations on radium cases were performed by the Project Medical Director, as well as examinations of 34 controls cases. Partial medical examinations were performed on three cases.

Summaries of fifteen additional radium cases are presented as Part III of this report, supplementing previously reported case summaries.

Three radium cases who had been under study died during the period covered by this report. Post-mortem examination was performed on one of these. Ten radium cases who had not been under active study also died.

Malignancies Among Cases Under Study

Four malignancies among living radium cases under study have been brought to our attention; these have not been previously reported; and all diagnoses have been verified by histological reports.

Case No. Malignancy

- 5153 Mucoepidermoid carcinoma of right parotid salivary gland (1963) (Bowen's Disease of skin of finger previously reported) Basal cell carcinoma of scrotum (1966)
- 5246 Basal cell carcinoma of forehead and chin (1964) (Squamous cell carcinomas of skin of fingers and basal cell carcinomas of skin of lip, left ear, nose previously reported)
- 5290 Basal cell epithelioma, left upper lip (1955) (Basal cell epithelioma of skin of temple previously reported)

5368 Squamous carcinoma of cervix (1951)

Deaths of Cases Who Died While Under Study

 Case 5155. Case résumé is in NYO-2760. This patient died in December 1965. Post-mortem diagnoses were:

History of automobile accident. Comminuted compound fracture, right tibia and fibula Lacerations, multiple, right and left knees, right temporal area Eccymoses, right ear lobe Multiple rib fractures, status post-surgical cut-down, ankle Acute pulmonary edema, myocardial hypertrophy Acromegaly Hirsutism Large face and hands Coarseness of skin, face Prominent mandible and supra-orbital ridges Generalized organomegaly History of exposure to radium Cavitations of tibial and humeral medulla Osteoporosis Status post hysterectomy

- 2. Case 5136. Case résumé is in NYO-2181-3. This patient died in July 1966 with death certificate diagnoses of coronary thrombosis due to hypertensive arteriosclerotic heart disease. No post-mortem examination was performed.
- Case 5255. Case résumé is in NYO-2760. This patient died in August 1966 with death certificate diagnoses of myocardial infarction due to arteriosclerotic heart disease. No post-mortem examination was performed.

Data from Death Certificates

Five additional malignancies were diagnosed on newly received death

certificates.

Case No.	Year of Birth	Year of Death	Causes of Death
5180	1898	1966	Carcinoma of lung with metastases
5497	1893	1965	Pulmonary embolus Generalized carcinomatosis Primary carcinoma of breast
5954	1892	1965	Post-operative infection and inanition Resection of cancer of sigmoid
5999	1893	1965	Carcinomatosis. Cancer of colon
6008	1908	1966	Multiple myeloma

Control Cases

No controls have died since the last report. In December 1965 a possible basal cell carcinoma, skin of left side of nasal bridge, was diagnosed on Control 6574.

Roentgenographic Studies

During the past year the roentgen examination has proceeded without change in basic policy or equipment. Forty-three complete bone surveys were made, nine of which were on former workers in the luminous dial industry, and the remainder on presumably unexposed controls. Studies were made on osseous material resulting from two autopsies.

As examination of workers and controls nears completion, the evaluation of data accumulated in the past seven years becomes of paramount importance. To be of maximum use, films must be minutely reexamined and carefully classified. The coding forms (See Exhibit 3) devised in 1964 and last revised October 15, 1965, have been used in 25 cases and found suitable. By means of these forms any designated deviations from the normal are classified by location, type, distribution, frequency and size. Record is kept of definite clinical and pathological entities present. The degree of osteoporosis is evaluated by measurement of the combined cortical thickness of the proximal radius and second metacarpal bones as detailed in previous report NYO-2181-3.

A second consultation was held with Dr. Gwilym Lodwick, Professor and Chairman of the Department of Radiology of the University of Missouri School of Medicine, an authority on radiological computer investigation. Agreement was found in the practical aspects of the coding. Dr. Lodwick offered us the facilities of his special computer for analyzing the frequency distributions of our findings. When this point has been reached the importance of the individual findings may be appraised, as well as their relationship to age, sex, exposure, body burden, medical, dental,

pathologic and radiochemical findings. The control material will prove invaluable in this study.

An exhibit was prepared and shown at the Annual Meeting of the Radiological Society of North America, November 28 to December 3, 1965. This exhibit demonstrated the most frequent changes found in the bones of radium workers and compared them with radiographs of the corresponding pathological specimens. It also demonstrated our findings in the measurement of the combined cortical thickness in the radii and hands of radium workers compared with the unexposed population.

While the process of coding appears to be the most important activity at the time of this report, and must be completed if minimal specific changes are to be recognized and evaluated, another important study to be undertaken is correlation of gross and histological pathology with the roentgen image. Seldom, if ever, has such a fund of documented and coordinated material been assembled and it is planned that full use will be made of this unusual opportunity. It is estimated that an additional two years would be required to make full use of the valuable radiological data that has been and is still being accumulated.

Dental Studies

During this reporting period 38 cases were examined clinically and dental radiographs were taken. This total number includes one initial examination of a known exposed case, six follow-up studies of known exposed cases and 30 initial examinations of control cases plus one control follow-up.

No attempt has been made to analyze these data or correlate them with cases previously studied. This will be the subject of our forthcoming final report.

Re Case 5281 (Body Burden $.603\mu$ c Ra²²⁶ and $.0019\mu$ c Ra²²⁸), the x-ray views and histopathology studies of the defect, the bone and soft tissue contained therein, within the body of the mandible were presented and discussed at the April 18, 1966 AEC Group Meeting held in Washington, D.C. The initial impression of the tissue removed is that of a necrotic mass with scavenger cells and remnants of acute inflammatory cells. Additional considerations are some form of degenerated connective tissue, hyalin type of degeneration and plasma protein. As the histopathology studies progress and these findings can be compared to those found throughout the skeletal structure, Case 5281 will be the subject of a separate report.

The oral histopathology findings are summarized elsewhere in this report. While the microscopic slides of the specimens are being read by a pathologist who has no knowledge of retained body burdens of patients from whom the specimens were taken, the processing of the total findings; clinical, roentgenographic, histopathology, radiochemistry, retained body burdens, etc., will hopefully be completed within the coming year and an analysis tying together these independent observations will be contained in our final Project report.

II. LABORATORY STUDIES

Clinical Laboratory

The Clinical Laboratory, under the direction of Dr. Lester M. Goldman, Director of Laboratories, Newark Beth Israel Hospital, continued to perform its preselected battery of tests on cases and controls of the New Jersey Radium Research Project. During the reporting year, 54 sets of examinations were done on radium workers and 164 sets of examinations were done on controls. Collation and statistical testing evaluation of the results of the laboratory examinations are currently in process.

Histopathology (Soft Tissue and Bone) General Pathology

Quantitative Report

Substantial progress was made between 1 September 1965 and 1 September 1966. 403 complete specimens representing 1215 tissue blocks (missing 23 blocks which are at least temporarily "not available") have been reported, and 121 bones from 23 cases have been examined in sufficient detail to begin the definition of changes. About 700 blocks have been described but are not yet assembled into complete specimen reports because of randomization necessary for blind study. No correlation has so far been attempted with radium burdens of any sort.

Limitation of Material

Because of the necessary delay between death and freezing of the tissues, much of the bone is poorly fixed, but the anatomic outline of various structures and lamellar architecture are satisfactory. Because osteocytic criteria for bone ischemia and metabolic injury, insofar as established, are not applicable, microradiographic evidence of occluded vessels and calcified osteocyte lacunae will be necessary to assess the presence and extent of bone infarction and death in borderline areas.

Histologic Abnormalities

Without knowledge of body or local bone burdens of radium at the time of interpretation of histologic preparations, many of the changes noted presumably represent only the usual and expected changes of aging: patchy and irregular loss of osteocytes from their lacunae, calcification of vascular channels and canals, increased porosity of both cortical and

cancellous bone, and structural changes in joints and periosteal surfaces. Changes consistent with varying stages and degrees of Paget's disease (osteitis deformans) have often been noted. Subsequent statistical analysis against a control population may define whether these changes are in fact accelerated in patients with higher than background radiation burdens.

Conferences with Dr. Hubert A. Sissons in London gave independent support to specific changes which have not been encountered in the enormous material seen at the Royal National Orthopaedic Hospital, which have apparently not hitherto been described, and which may tentatively be presumed related in some way to exposure to radiation. Other abnormalities may subsequently be identified, and only analysis of the whole material available will be able to assess possible relationship to radium burdens. Correlation of these histologic changes with whole body and fractional bone radiation burdens will require a larger sampling than has been accomplished to date.

For adequate scientific completion of the bone histopathology component of the Project, financial support is needed for at least two more years, and probably longer.

The time is probably ripe to initiate direct, fractional analyses of radium bearing bones, because the total radioactivity is so small that autoradiographs may prove wholly unreliable except for assessment of "hot spots." Exposure times for autoradiographs may approach two years, and direct analyses seem a more useful approach.

Current Queries

Review of current and past literature suggests that early bone infarction and "radium osteitis" were not distinguished by writers generally before World War II, and that confusion between these processes makes their reports subject to reinterpretation. The fibrillar cortical and granular basophilic material reported in early descriptions of both bone infarction and "radium osteitis" have never been adequately characterized. The special staining program which is under way may yield useful information along these lines, and further review of the literature will continue.

Oral Histopathology

Material studied during the reporting period includes fifteen individual specimens from nine cases. Specimens were from the oral cavity except for two cases where specimens were from the maxillary sinuses.

All tissues were processed in a conventional manner at the Department of Oral Pathology of Georgetown University. All sections were read blind with no knowledge of body burdens.

Serial sections of all specimens were prepared and every tenth section was stained with hematoxylin and eosin. All blocks were paraffin embedded; thus far no special staining has been found necessary. All unstained sections are mounted on glass slides, carefully protected by paraffin sheet film and stored.

The data are presented in condensed form in terms of geographical area. A more detailed report on each specimen has been recorded and is in the Project files. Specimens were examined from the following areas:

- (1) Mucosa and bone of the maxillary sinus
- (2) Mucosa of the oral cavity, including tongue, palate, cheek, alveolar mucosa and gingiva
- (3) Bone of the mandible and maxilla

The histopathologic findings in the above areas were:

(1) Maxillary Sinus: The mucosa presented a non-specific inflammatory proliferative polyposis which, judging from the eosinophilia could be interpreted as allergic.

(2) Mucosa of the Oral Cavity: The findings in the oral cavity were inflammatory or benign, proliferative pseudo-tumors such as papilloma, fibromas, acanthosis, etc. Two specimens did, however, show epithelial dyskeratosis. Serial examination of these two specimens revealed no extension to carcinoma in situ or submucosal invasion. Reactions in mucosal adnexae, such as the minor salivary glands showed non-specific ductal ectasia, but no tumors were encountered.

(3) Bone: Specimens from two cases showed no unusual periosteal or endosteal changes and neither specimen presented any tumors. Trabecular spaces did, however, appear enlarged and Haversian systems in cortical bone were unusually prominent. Large empty spaces were encountered frequently in cortical bone.

As the study continues, soft tissue changes are still non-specific, but many specimens remain for examination and it is impossible at this time to correlate findings with body burdens. Bone lesions examined in two specimens are as previously reported, large empty spaces within cortical bone. It is, however, too soon to determine if this will be a constant finding. Unfortunately, due to the technical difficulties of processing, bone tissue examination does not progress as rapidly as soft tissue. No malignant changes have been uncovered thus far.

Body Burden Assessment

Through interproject cooperation the New York University Medical Center group, working under Merril Eisenbud and supported by U. S. Atomic Energy Commission Contract AT(30-1)-3086, has performed 73 whole body counts and 36 breath radon determinations on cases and controls of the Radium Research Project since September 1, 1965. Additionally, the Massachusetts Institute of Technology group, working under Robley Evans and supported by U. S. Atomic Energy Commission Contract AT(30-1)-952, has performed 13 breath radon determinations on Project cases.

As is reported in NYO-3086-6 (10/1/66), special additional studies on materials from Project cases were done by the New York University Medical Center group:

(1) "Estimates of Total Body Radium in the New Jersey Dial Painters" by H. Peterson, N. Cohen, and H. W. Berk.

(2) "A Study of the Distribution of Ra-226, Ra-228, Pb-210 and Th-228 in Bone and Soft Tissue of Radium Dial Painters" by Henry Petrow.

(3) "Discussion of Possible Mechanisms Involved in the Apparent Concentration of ²²⁶Ra and ²¹⁰Pb in Aorta and Thyroid" by Norman Cohen.

Since the data presented in these three reports constitutes an integral part of the information about radium workers in the case group of the Radium Research Project, by permission of Dr. Eisenbud, Dr. Petrow and Mr. Cohen, the summaries of these three portions of NYO-3086-6 are reproduced here and copies of the full reports are included as Part II of this current progress report of the Project.

Summary

The effective half-life of radium-226, calculated from RaC retention, has been found to be approximately 11 years. It is believed that this value, which is shorter than the half-life proposed by others, might be due to a natural aging effect such as osteoporosis which results in increased bone resorption. No difference is apparent between the exponential and power function retention models.

A possible correlation between a respiratory parameter, minute volume (liters/min), and radon retention has been found. It is believed that this variation might explain short-term fluctuations in radon retention measurements. Additional data is needed to confirm this observation.

A study has been made of the distribution of Ra-226, Ra-228, Pb-210 and Th-228 in bone and soft tissue of two radium dial painters. The data are presented in a separate section of this report, available on microfilm from the AEC (NYO-3086-5). The results of this study showed the usual heterogeneous deposition pattern for radium in the skeleton. The average skeletal Pb-210 to Ra-226 ratios demonstrate that lead and radium have equal biological half-lives. The variation in the ratio were site dependent. Shafts of long bone and rib had high ratio values, while the ends of long bones and vertebrae had low ratios. In long bone, low ratios were associated with high Ra-226 concentrations. The cause of the ratio variations cannot be proven unequivocably, but is believed to be due to variations in the radon retention factor.

The Th-228 to Ra-228 ratios were also measured. These ratios vary considerably, but in a random manner. The cause of the variation is believed to be Th-228 translocation. The average skeletal ratio of Th-228 to Ra-228 offers a second method of estimating radium biological half-life over the last 5 - 10 years of life. This value is 10.4 years. The analysis of beagle bone obtained from an animal injected with Ra-228 indicated a similar ratio pattern, with large, random variations being observed. The radium biological half-life, as calculated from the data, is 16 years.

The analysis of tissues from an individual with a 50 year old radium burden indicated that 0.05% of the Ra-226 and 0.65% of the Pb-210 in the body is found in soft tissue. The aorta contained the most Ra-226 and Pb-210 of all soft tissues analyzed with the thyroid being the second highest.

Since, of all the soft tissues obtained from a deceased radium dial painter, the aorta had the highest concentrations of both Ra-226 and Pb-210, a survey of the literature concerned with the uptake of polyvalent cations by the aorta has been made, and consideration given to the radiobiological implications of the known facts, as they pertain to radium cases.

III. STATISTICAL STUDIES

The coding and punching of further data has proceeded through the year. Specifically the whole body counter determinations have been assigned a deck, likewise the breath radon and thoron determinations. A coding scheme has been devised for the revised x-ray scoring system pending the actual release of evaluations by the Project radiologist. For day-to-day working purposes average values have been promulgated of the clinical laboratory parameters.

In the expectation that Project pathologists will require statistical processing of their bone histological findings a preliminary study of these findings has been made to gain insight into the phenomena. On the basis of this, the pathologists' requirements can be rapidly executed. Likewise in the expectation of evaluations by the Project consultant on hematological data, a study of the interrelations of the components of the electrophoretic pattern of controls and subjects with less than detectable body burdens has been mounted.

With a view to final analysis, a review of radium retention data has been made. It has been possible to supplement this by the computations of certain parameters from the New Jersey data.

PART II

RÉSUMÉ OF COOPERATIVE LABORATORY STUDIES ON PROJECT CASES AND SPECIMENS

Reprinted recognizing credit due to and with permission of Merril Eisenbud and Henry Petrow and staff, Environmental Radiation Laboratory, Institute of Environmental Medicine, New York University Medical Center. This work done under U. S. Atomic Energy Commission Contract AT(30-1)-3086 and originally published as part of NYO-3086-6 (October 1, 1966)

PART I

Estimates of Total Body Radium in the New Jersey Dial Painters

H. Peterson, N. Cohen, H.W. Berk

Method of Whole Body Counting

In previous progress reports (1962 and 1965) the technique used to determine the breath radon estimates has been reviewed. This section will describe the procedures employed in the whole body gamma spectrometric analysis.

Figure (la) illustrates the gamma ray spectrum of a normal individual measured with a 8" x 4" sodium iodide (T1) detector and an energy calibration of 12 keV/channel. This represents the energy region 0 - 3 MeV when 255 channels are The peak in channel 120 is due to the 1.46 MeV employed. photon from 40 K which occurs naturally in potassium with a specific activity of 829 pCi 40 K per gram of potassium. The other prominent photopeak is due to 137 Cs (0.661 MeV) which is present due to fallout from nuclear weapons testing. In order to avoid interferences from these nuclides and from variations in their concentration from subject to subject only the energy region above 1.60 MeV is used for study.

Figure (1c) is the gamma spectra of a 1.0 μ Ci source of 226 Ra taken under the same conditions as Figure (1a). The prominent photopeak in the region above 1.60 MeV is due to

1-1

the 1.76 MeV gamma of ²¹⁴ Bi(RaC). Since this nuclide is a daughter of the 3.83 day ²²²Rn it represents only the fraction of the ²²⁶Ra body burden due to retained radon. For most of our cases this fraction is approximately 0.33 (see Section2). The region containing this photopeak extends from channels 140 (1.68 MeV) to 159 (1.91 MeV) and is identified in Figure (1c) as the RaC region. Radium-228 (Mesothorium) is determined by measuring the 2.61 MeV gamma of its daughter ²⁰⁸Tl (ThC"). The intervening radon isotope, 220 En (Thoron), has a half life of 56 seconds and therefore is presumed to decay entirely in the body.

Thallium-208, in equilibrium with Thorium-232, is used as a standard for the mesothorium estimation. Due to its short (5.7 yr) half life 228 Ra is not used as a standard, thus eliminating decay corrections. A correction is made, however, to take into account the equilibrium ratio of 208 Tl to 228 Ra. This is 0.5 for 228 Ra in equilibrium, whereas it is 0.33 for 232 Th . The gamma ray spectrum of 232 Th is shown in Figure (1d); the region encompassing the 2.61 MeV peak of 208 Tl (channels 206-229, 2.47-2.74 MeV) is indicated.

Figure (1b) is the gamma spectrum of case #5025 who has an estimated body burden of 0.085 $\mu{\rm Ci}~^{226}{\rm Ra}.$

The principal decay products of importance in the two chains are: $\begin{array}{c}232\\\text{Th}\end{array}$, $\begin{array}{c}228\\\text{Ra}\end{array}$, $\begin{array}{c}228\\\text{Ra}\end{array}$, $\begin{array}{c}228\\\text{Th}\end{array}$, $\begin{array}{c}1.9\\\text{yr.}$, \\\begin{array}{c}208\\\text{Th}\end{array}, $\begin{array}{c}228\\\text{Ra}\end{array}$, $\begin{array}{c}228\\\text{Th}\end{array}$, $\begin{array}{c}228\\\text{Th}\end{array}$, $\begin{array}{c}208\\\text{Th}\end{array}$, $\begin{array}{c}218\\\text{Th}\end{array}$, $\begin{array}{c}228\\\text{Th}\end{array}$, $\begin{array}{c}228\\\text{Th}\end{array}$, $\begin{array}{c}218\\\text{Th}\end{array}$, $\begin{array}{c}218\\\text{Th}$, $\begin{array}{c}218\\\text{Th}$, $\begin{array}{c}218\\\text{Th}\end{array}$, $\begin{array}{c}218\\\text{Th}$, \begin{array}
If the thorium chain is in equilibrium, the activity of any daughter equals that of the parent (secular equilibrium), and $A(^{232}Th) = A(^{228}Ra) = A(^{228}Th) = 3 A(^{208}T1)$. The factor of 3 is due to a branch in the decay chain such that the 208 Tl yield is 33%.

Under these conditions in the ²²⁸ Ra chain, however, transient equilibrium exists, therefore 3 $A(^{208}TI) =$ $A(^{228}\text{Th}) = \frac{\lambda^{228}\text{Th}}{\lambda^{228}\text{Th}} - \lambda^{228}\text{Ra} = 1.5 A(^{228}\text{Ra}).$

Therefore, $\frac{A({}^{208}\text{Tl}) / A({}^{228}\text{Ra})}{A({}^{208}\text{Tl}) / A({}^{232}\text{Th})} = 1.5$. However, it does

not necessarily follow that the ratio of 228 Th to 228 Ra in the body is equal to 1.5, since this ratio is a function of the pertinent effective half lives. Petrow has measured the ratio of Th to Ra in a skeleton and found it to be 2.08, which reflects the fact that radium has a shorter biological half life than thorium (14). Since this ratio may vary from case to case, this is a possible source of error in any ²²⁸Ra measurement.

The counting rate in the RaC and MsTh regions may be represented by

$$S_{RaC} = {}^{B}_{RaC} + {}^{R}_{RaC} + {}^{T}_{Rac}$$
(1)
$$S_{MsTh} = {}^{B}_{Msth} + {}^{R}_{MsTh} + {}^{T}_{MsTh}$$
(2)

^TMsTh

(2)

and RaC = gross count rate in RaC region.where

 B_{RaC} = background count rate in RaC region.

^R RaC	=	count rate in RaC region due to 226 Ra and its daughters.
T RaC	=	count rate in the RaC region due to 228 Ra and its daughters.
S _{MsTh}	=	gross count rate in Msth region.
B MsTh	=	background count rate in MsTh region.
R _{MsTh}	-	count rate in the MsTh region due to 226 Ra.
T _{MsTh}	=	count rate in the MsTh region due to $\frac{228}{Ra}$ and its daughters.

An estimate of B can be made by determining the counting rate obtained under the same conditions but without the subject present. Because of variations in the cosmic ray intensity and in the radon-220 and radon-222 levels in the counting room and surrounding area as well as changes in the calibration (gain drift, etc.) the background measurement is made either immediately before or after the measurement of the subject.

The estimation of R and T is somewhat more difficult. As can be seen from Figures (lc) and (ld) there are counts contributed to the RaC region from thorium daughters and counts contributed to the MsTh region from radium-226 daughters. The most accurate method of determining these contributions would be to employ phantoms having the same anthropometric characteristics as the subject and containing known amounts of radium and thorium. In practice, however, this would be unsatisfactory for the following reasons:

1.) A large number of different sized phantoms would be required to simulate the variety of body sizes encountered in the subjects.

2.) Homogeneous phantoms generally contain aqueous solutions of the standards. The possibility of contamination from leakage as well as the difficulty of handling large phantoms limits their desirability. Also the deposition of radium in the skeleton is heterogeneous, but not reproducibily so.
3.) Leakage of radon would affect the concentration of ²¹⁴Bi in the phantom and result in the overestimation of the body burden.

The standards are in the form of sealed sources and are counted in the crook of the chair (see Figure 2). From these measurements the contribution to each region is determined. These factors are evaluated for each run and are approximately,

a =
$$\frac{159}{5}$$
 thorium
a = $\frac{140}{229}$ = 0.23
b = $\frac{229}{5}$ thorium
b = $\frac{229}{506}$ = 0.028
159
5 radium
140

Differences between the Compton scattering contributions in the standard and the subject are assumed to be second order

effects.

Using these contribution factors equations 1 and 2 become -B_{RaC} N_{RaC} = R RaC a T MsTh SRAC + (3)т MsTh, MsTh = $-B_{MSTh} = bR_{RaC} +$ S (4)MsTh where ^NRaC and ^NMsTh are the net counting rates in the two regions.

These equations can be solved for ^TMsTh and ^RRaC giving

$$R_{RaC} = \frac{N_{RaC} - aN_{MsTh}}{1 - ab}$$
(5)

$$^{T}MsTh = ^{N}MsTh - b\left(\frac{\overset{N}{RaC} - \overset{aN}{MsTh}}{1 - ab}\right)$$
(6)

If ^NMsTh is negative, it is set equal to zero so as not to overestimate the radium counting rate.

The difference in the counting efficiency (geometry) between a point source and a human subject is compensated for by means of a chair factor. The chair factor is defined as f = Counting rate per unit known activity in a person (7)

f = Counting rate per unit known activity in a person Counting rate per unit activity of a standard source and was found to be 1.2 by measuring 5 individuals having

known Ra burdens.

The body burdens of RaC and Mesothorium are given by

$$\frac{BB_{RaC}}{f N+} = \frac{R_{RaC} A_{Ra}}{f N+}$$
RaC (8)

$${}^{BB}_{MSTh} = \frac{{}^{T}_{MSTh} {}^{A}_{Th}}{\frac{f(1.5)}{f(1.5)} {}^{N+}_{N+}}$$
(9)

Where

and

$${}^{A}Ra = activity of radium standard$$

 ${}^{A}Th = activity of thorium standard$
 $f = chair factor = 1.2$
 $1.5 = equilibrium correction factor (see pp.tl-2; l-3)$
 ${}^{N+}RaC = net RaC$ counting rate of radium standard
 ${}^{N+}MsTh = net MsTh$ counting rate of thorium standard

These calculations have been programmed for a Control Data 160-A computer in the program "BURDEN", which computes inmaddition to the estimates, the variance due to counting statistics and the MSA* (minimum significant amount) for each run.

Appendix II contains a listing of "BURDEN" and a sample output page.

Reproducibility of Whole Body Data

An additional study (6) of the reproducibility of whole body counter measurements was performed using four subjects having body burdens ranging from 0.003 to 0.083 μ Ci RaC. The body burdens were determined by two separate techniques: The "Burden" program with a radium point source standard which

*The MSA is the minimum amount of measured radioactivity of which it can be stated that, at the 0.10 level of significance, this measured activity is above background.

is used routinely for ²²⁶Ra body burden determinations; and the integral stripping technique of Hallden and Harley (5). The variability of these two techniques, expressed as % standard error / estimated body burden of RaC, are plotted in Figure 3. The data reported in the 1965 report on two cases measured in 1962 at the New York City Facility are also plotted. The curve is the estimated percent standard deviation computed from counting statistics for individual measurements versus body burden RaC. The standard deviation used is the value currently reported as the error in the RaC measurements. It has been found that the variation in repeated measurements of an individual is less than the error predicted from counting statistics for most of the cases studied.

Breath Radon Measurement

The breath radon values obtained thus far in 1966 are the second series of measurements made on the group of dial painters studied in 1965.

The procedures for performing the radon measurements have remained essentially the same as previously described in the 1965 Annual Report (3). The only two modifications in procedure involved subject positioning during collection and a change in the gamma count summation region.

Since it is likely that the retained fractional radon is in some way related to the short term variation of the

1=8

respiratory pattern, every effort was taken to remove all emotional and physiological discomforts. Subjects were allowed to recline in a chaise lounge during sample collection, thereby providing a relaxed atmosphere and achieving a more reproducible breathing pattern. In addition, this procedure makes collection of larger samples more feasible, providing for better counting statistics.

In the 1965 annual report (3) two different counting procedures for the estimation of radon were described. One method involves gamma counting of a charcoal sample containing adsorbed radon. As originally described, only events occurring in the ²¹⁴Bi region were measured. In order to increase sensitivity and to minimize errors due to gain shift, summation is now performed over the energy continuum, 0.22-2.56 MeV. This results in a new MSA (minimium significant activity) (1) of 1.30 pCi for radon.

The second method is essentially that developed by Lucas and consists of desorbing the radon and collecting the gas in a scintillation bottle, and alpha counting radon and its other emitting daughters. This technique is considerably more sensitive than gamma counting and the MSA is 0.016 pCi of radon.

Based on a 100 liter breath sample from an individual with an average minute volume of 7.5 L/min, the minimum significant

body burdens corresponding to the above MSA values are 1.18 nCi and 14.5 pCi, assuming the fractional radon retention is 0.33. Since the chance of procedural error is less likely by the direct gamma counting technique, and the method is rapid, it is used for those cases where the body burden is high enough to allow for acceptable counting statistics. (See figure 4)

The results of 27 breath radon measurements made during the past two years are listed in Table 1. If the effective half life of ²²⁶Ra is assumed to be 11 years, a decrease of approximately 6% in the 1965 body burdens would be expected. A change of this magnitude, however, would be masked by differences due to other factors contributing to the overall experimental error.

Fractional expired radon vs body burden for both 1965 and 1966 measurements is plotted in Figure 5. No significant correlation between body burden and the radon loss rate is evident. If the standard deviation of the outgassed radon values due to counting statistics is plotted against body burden, (Figure 6), it is obvious that the distribution in Figure 5 is a function of the associated counting error. The plot of percent counting error vs body burden (Figure 7), illustrates the statistical degree of uncertainty in the outgassed radon value as a function of body burden.

There appears, in the data accumulated to date, to be

a possible correlation ($\rho = 0.76$, significant at the 95% confidence level) between minute volume (1/m) and outgassed radon (Figure 8). It has been assumed in the past that radon retention may be affected by the respiratory pattern but until now there has been no evidence of correlation between expired radon and any respiratory parameter.

The rate of radon elimination from bone is considered to be the controlling factor in radon retention, (15) and while this process may be time dependent it can probably be considered constant for periods as short as one year for old radium burdens. Percent retention as a function of time may be possibly follow the slope illustrated in Figure 9 which was plotted from the data of several investigators (4,7,8,11, 13,15-17) of the retention factors for man, dogs and rats with body burdens of 11 days to 40 years duration. A comparison of several models of radon retention is presented in Table II.

Short term variation of the expired radon fraction on the other hand is much greater than that predicted from these models and may be a function of more immediate physiological parameters. Further experimentation is necessary to confirm the observed correlation between percent radon expired and minute volume.

Radium Retention

In the previous report (3) the effectivehalf-life of

radium was computed based on whole body measurements of the RaC daughter made in this laboratory from 1960 to 1965.

Re-analysis of 11 cases selected in 1965 with the addition of data obtained in 1966 gave the ²²⁶Ra effective half-life estimates shown in Table III. The statistical techniques used for obtaining the pooled regression estimate are described in appendix I. The value thus obtained of the effective half-life of radium for these 11 cases was $15.7 \stackrel{+}{-} 2.4$ years in 1965 and 14.73 $\stackrel{+}{-} 2.31$ years when the 1966 data were included. If the data are normalized to percentages of the initial measurement (taken as 100%), and are combined and fitted by least squares technique, the estimate of the effective half-life of radium is $14.2 \pm$ 3.08 years when the 1966 data are added. Based on pooled regression analysis of 21 cases having body burdens greater than MSA (0.002 $_{\rm LL}$ Ci RaC) an effective half-life of 11.1 $^+$ 1.9 years was reported in 1965.

If more stringent conditions are imposed upon the selection of data by choosing only cases where the observations have been made over a period greater than three years, having more than two measurements, and having body burdens greater than 0.01 μ Ci RaC; only six cases would be included (5014, 5025, 5215, 5281, 5284, 5917) and a pooled half-life of 11.32 \pm 0.94 years is obtained. These six cases represent 42 points.

On the other hand, if all cases having a body burden greater than MSA (0.002 μ Ci RaC) (1) are used, a value of 10.71 $\stackrel{+}{-}$ 2.58 years results. This study includes 124 points (30 cases). These data are given in Table IV.

The variation of the half-life estimate with RaC body burden is shown in Figure 10. No general trend is evident but it should be noted that the negative values are all for cases having a 1960 estimate body burden of less than 0.01 μ Ci RaC. Most of the half-life estimates in the region above 0.01 μ Ci that are less than 5 years are from cases represented by only a few points.

:

The pooled half-life estimate of 10.7 ± 2.58 years, although considerably lower than the 45 years presently assumed for radium, is in excellant agreement with a value of 10.4 years calculated by Petrow (14) for 228 Ra in bone (case 5281) from the 228 Th/ 228 Ra ratio. The reason for this rapid elimination is not known, but it should be noted that our population is an aged one (mean age 66.6 years) and all measurements made were made in the past 6 years. Hence, this relative short half-life may be due to an aging effect such as osteoporosis.

Due to an error in the 1965 report the power function model for radium elimination waser**r**oneously thought to be inapplicable to our data. We now find that this conclusion

was based on a numerical miscalculation and that the power function does in fact yield estimates of elimination rates that are consistant with our data.

The power function retention model is

$$R(t) = at$$

where a = R(1); then at some time later, x, let $R(t + x) = \frac{1}{2} R(t)$, therefore $\frac{1}{2} a t^{-b} = a (t + x)^{-b}$, and x should then be the apparent half-life. then $-bln(t) = 1\frac{n}{2}$ -bln (t + x) b (ln (t + x) - ln (t)) = ln 2 $b (ln \frac{t + x}{t}) = ln 2$ $ln (1 + \frac{x}{t}) = \frac{1}{b} ln 2$ $1 + \frac{x}{t} = 2^{1/b}$

therefore $x = t (2^{1/b} - 1)$ (For the derivation used last year the 1 was assumed to be small compared with $2^{1/b}$.)

The principal error in the previous analysis was that the value of b (0.1518) was determined by employing the data of initial measurement as the initial date of exposure. Although this assumption is inconsequential when employing an exponential model it does lead to serious error when applied to the power function. The mean time of exposure is now used and is taken as the midpoint of the period of active exposure to radium. These data were obtained from the Radium Research Project files and are listed in Table V. When the analysis is redone using

this initial point a value of b of 2.78 results. Then

x = 43.1 (1.274 - 1) = 11.8 years considerably more in agreement with the exponential values of 10.7 and 11.2 years than the value of 4140 years that would be obtained if b = 0.1518. Norris (12) has reported a more rigorous derivation of an apparent half-life based on setting the slopes of the two equations obtained after logarithmic transform equal to one another:

 $\frac{1}{R} \begin{pmatrix} t \end{pmatrix} = -\frac{b}{t} \text{ from the power function and}$ $\frac{1}{R} \begin{pmatrix} t \end{pmatrix} = -\lambda = -\frac{0.693}{T_{1/2}} \text{ if the exponential is used;}$

then $T_{1/2}$ apparent = $\frac{0.693 \text{ t}}{\text{b}}$.

With t = 43.1 years and b = 2.78 this gives an apparent half-life of 10.78 years, in excellent agreement with the exponential values. Although radium retention in aged subjects can be described by a power function this function is quite different than that reported by Norris (11).

The fact that RaC and not 226 Ra is being measured produces a bias in the effective half-life estimate only if the fraction of retained radon is changing with time. Since the RaC produced in the body is directly proportional to this quantity:

RaC = (fractional Rn retention) (^{226}Ra) , then if the radon retention is changing with time the true biological half-life of ^{226}Ra will be different than that inferred

from the RaC measurements.

If the equation describing increasing radon retention reported by Miller and Finkel (10) is used with the value we report for the power function exponent, the effective halflife of radium would be 10.1 years, in good agreement with the reported values.

One might expect the short term changes in radon retention described in the second section of this report to have an effect upon the measured half-life. However, if the changes in radon retention are related to the minute volume then one might assume that they are either random or produce a bias in one direction. In either of these cases the effects should be negligible when a large number of cases are used to compute the pooled slope, however they might contribute to the variability of the estimates obtained from individual cases.

Appendix I - Statistical Techniques

In testing the applicability of the exponential function the regression model is $B(t) = B_0 e^{-\lambda_E t}$ where T_E , the (1) effective half-life is $T_E = \frac{\ln 2}{R_E}$. This can be transformed into the linear equation Y = a + bx (2) letting $Y = \ln A(t)$

etting
$$Y = \ln A(t)$$

 $a = \ln A_0$
 $x = t$
and $b = -\lambda_F$

For the power function the regression model is

$$A(t) = qt^{-p}$$
(3)

which can also be transformed into equation (2) with the following change of parameters.

$$Y = \ln A(t)$$

a = ln q
x = ln t
b = -p

The parameters obtained are therefore those which give the best fit to equation 2 in the log - log plane and not to equation 3. It has been shown (9) that these parameters can be different than those obtained from iterative calculations based on equation 3 directly. However, most of the studies in the literature use the log - log transformation.

Keeping in mind the different transformations employed to obtain the linear equation the remaining techniques are the same for both cases.

If there are k cases having n_i points each then the estimate of the individual slope for each case is given by

$$b_{i} = \frac{S_{xy}^{i}}{Xx} \qquad (4)$$

$$\sum_{i=1}^{n_{i}} x_{xy} \sum_{j=1}^{n_{i}} x_{ij} \sum_{j=1}^{n_{i}} y_{ij}$$

where $S_{xy}^{i} = \sum_{j=1}^{n_{i}} x_{ij}y_{ij} - \frac{\sum_{j=1}^{n_{i}} j_{j} - y_{ij}}{n_{i}}$ (5)

and
$$S_{XX}^{i} = \begin{cases} n_{i} & 2 \\ j = 1 & x_{ij} & -\begin{pmatrix} n_{i} & \\ y = 1 & x_{ij} \end{pmatrix}^{2} \\ \frac{j=1}{n_{i}} \end{cases}$$
 (6)

The standard error of this estimate is

$$S_{.}E_{.}(b_{i}) = s_{i} / \sqrt{S_{XX}^{i}}$$
 (7)

where S_i^2 , is the mean square due to error,

$$s_{i}^{2} = \frac{1}{n_{i} - 2} \begin{bmatrix} s_{yy}^{i} - b_{i} & \frac{s_{xy}^{i}}{s_{xx}^{i}} \end{bmatrix}$$
 (8)

and
$$S_{yy}^{i} = \sum_{j=1}^{n_{i}} y_{i_{j}}^{2} - \left(\sum_{\substack{j=1\\n_{i}}}^{n_{i}} y_{i_{j}}\right)^{2}$$
 (9)

The pooled slope, b, is estimated assuming parallel lines and is given by reference 2;

$$\overline{b} = \frac{\sum_{i}^{k} S_{xy}^{i}}{\sum_{i}^{k} S_{xx}^{i}}$$
(10)

The standard error of this estimate is given by

$$S.E.(b) = S^{+} / \sqrt{\sum_{i=1}^{k} S_{xx}^{i}}$$
 (11)

where \textbf{S}^{+2} is given by

$$\frac{\underset{i=1}{\overset{k}{\sum}} s_{yy}^{i} - \underset{i=1}{\overset{k}{\sum}} \left(\frac{s_{xy}^{i}}{s_{xx}^{i}} \right)^{2}}{\underset{i=1}{\overset{k}{\sum}} (12)}$$

ø

	APPENDIX II COMPUTER PROGRAM FOR RaC & MsTh DETERMINATIONS
С	BURDEN 11 - PROGRAM FOR RADIUM AND THORIUM BODY BURDEN CALCULATIONS
С	DEVELOPED BY H. PETERSON AND H. BERK, NYU MEDICAL CENTER,
С	INSTITUTE OF ENVIRONMENTAL MEDICINE.
C	MODIFIED OCTOBER 18+1965 BY H.PETERSON AND G.LAURER TO INCLUDE
C	MSA DETERMINATIONS AND CORRECTION OF PROCEDURAL ERROR
G	KA = 1 READ NEW SERIES STARTING WITH BACKGROUND
C	KA # 2 KLAU NEW SAMPLE CARD
-	DIMENSION BRODU 250 JAPADSU 250 JATUDDSU 250 JACAMU 250 JATUTU CULUA
	DIMENSION ANAMEAR)
· 2	READ 10+TMB+(TITIE(J)+J=1+11)
10	FORMAT(F6.1)11A6)
	PRINT +TMB+(TITLE(J)+J=++++)
11	FORMAT(1H1)F6.1.11A6)
	READ 12+(BKGD(1)+1=1+255)
12	FORMAT(IOF6.0)
	READ IO, TMR, (TITLE(J), J=1,11)
~	PRINI [0] [MR)(I] TLE(J))J=[]]
u	$\frac{REAU}{129} \frac{129}{RAU5} \frac{1391 = 19255}{120}$
7	
	READ 12+(THORS(1)+1=1+255)
200	RARRA * 0.0
	RARTH =0.0
	RARBK = 0.0
	THRRA = 0.0
	THRBK = 0.0
	THRTH = 0.0
	THURIUM REGIUN AT 12 REV/CHANNEL = CHANNELS 205-22912.59 MEV TO
· · · · · · · · · · · · · · · · · · ·	2.75 HEV) ENCOMPASSING THE 2.61 HEV PEAK OF THURIUM CFF TE-208).
	THRRA = THRRA + RADS(1)
	THRTH = THRTH + THORS(1)
20	THRBK = THRBK + RKGD(1)
C	RADIUM REGION AT 12 KEV/CHANNEL = CHANNELS 140-159(1.68 MEV TO
<u>C</u>	1.91 MEV) ENCOMPASSING THE 1.76 MEV PEAK OF RAC(B1-214).
	D0 25 I=140,159
	RARIH = RARIH + IHURS(I)
25	PARK = PARK + RAUS(1)
C 23	MSA(SIGMIN) PROCEDURE FROM B. ALTSCHILLER. B. PASTERNACK . HEALTH
C	PHYSICS,9,293,1963.
	CALL SIGMIN (RARBK)
	RMIN = CMIN
	CALL SIGMIN (THRBK)
· -	THMIN = CMIN
	PRINT 150
	FORMATCHD + 25X + 1 HBACKGROUND + 15X + 16HRADIUM STANDARD + 15X + 17HTHORI
151	FORMAT(140+13H SUM 140+159 +12X+F8-0+21X+F8-0+21X+F8-0-1
	RARBK = RARBK/TMB
•	RARRA = RARRA/TMR
	RARTH = RARTH/TMT
	PRINT 140+RARBK+RARRA+RARTH
140	FORMAT(1H0+15H COUNTS/MINUTE +12X+F9.3+20X+F9.3+20X+F9.3)
	ERARBK = RARBK/TMB
	ERARRA = SUKIF(ADSF((MAKKA/INK) + ERARBK))
	DADDA - DADDA-DADRK
•••••	RARTH = RARTH-RARBK
····· ·	PRINT 152+RARBK+RARRA+ERARRA+RARTH+FRARTH

152	FURMATCH J20H NET COUNTS/MINUTE 1/X3F9.399X3F9.293H+- 3F9.2914X3F
	19.2,3H+- ,F9.2)
	PRINT 153, THRBK, THRRA, THRTH
153	FORMAT(1H0+13H SUM 206-229 +12X+F8+0+21X+F8+0+21X+F8+0)
	THRBK = THRBK/TMB
	ETHRBK = THRBK/TMB
	THRTH = THRTH/TMT
	THRRA = THRRA/TMR
	ETHRRA = SQRTF(ABSF((THRRA/TMR) + ETHRBK))
	ETHRTH = SQRTF(ABSF((THRTH/TMT) + ETHRBK))
	PRINT 140+THRBK+THRRA+THRTH
	THRTH = THRTH - THRok
c	A - CONTOINING THE THOUSING OWNERS CANNA PUOTONS TO PARTIM DECION
C	A - CONTRIBUTION OF THORIDA CONFION GAPHA PHOTONS TO RADION REGION
•	EA = AOSURIFICERARTH/RARTH/OO2 + EITHRIM/INRIH/OO2 + O
C	BE CONTRIBUTION OF HIGHER RADIUM PEAKS TO THURIUM REGION.
	B = THRRA/RARRA
	EB = B&SQRTF((ETHRRA/THRRA) + (ERARRA/RARRA) +)
	Q=1.0-(A+B)
·	PRINT 154+A+EA+B+EB+Q
154	FORMAT(1H0,5H A = +F9.4+3H+- +F9.4+ 10X+5H B = +F9.4+3H+- +F9.4+10
	$X_{2}OH I - A + B = 2F_{2} + 1//$
	PRINT 155
- · ·	PRINT 143, RMIN, THMIN
с	MSC = MINIMIUM SIGNIFICANT COUNT
	EORMAT(1H0+7H MSC +15X+FA-3+44X+FA-3)
. 145	RMIN = RMIN/TMR
c	MSCD - MINIMUM SIGNIFICANT COUNTING DATE
	$ \begin{array}{c} FOR(PAIL(ID)) FOR(F) $
•	
C	MSA = HINIHIUH SIGNIFICANI AHUUNI
	PRINT 1457AMIN BMIN
145	FORMATCIHOF7H MSA FI3XFF8.5F3H UCF40XFF8.5F3H UC ///)
	READ 6+KA
6	FORMAT(IG)
5	READ 14+TMS+CASE+(ANAME(J)+J=1+8)+IM+ID+IY+RUN
14	FORMAT(F6,1,9A6,312,A6)
	IF(TMB-TMS) 174,301,174
174	PRINT 175
175	FORMAT(1H0 + 75H MSA NOT VALID FOR RUN. DIFFERENT COUNTING TIME USED
	FOR PATIENT AND BKGD. ///)
	AMIN = 0.0
	BMIN = 0.0
301	PRINT IGITMSICASEI(ANAME(J))J=118)IMIIDIIYIRUN
16	FORMATCIHOFES, 1, 12H NJRRP CASE +AS+5X+8AS+8H RUN ON +12+3H / +12+3
1	H / 12,4X,8H RON (A6) -
	RFAD 12+(SAM(1)+1+1+255)
201	
201	
	D0 30 1 = 140+150
20	
30	NANJAH - NANJAH JAH
	TUDCAM - TUDCAM - CAM(1)
35	100000 - 100000 + 300117
	PKINI 155
155	FURMATCIHU/20X/ISH RADIUM REGION /30X/I6H THORIUM REGION /25X/ 9H
1	140-159 35X310H 206-229)
	PRINI ISSIKARSAMITHRSAM

```
156 FORMAT(1H0+14H TOTAL COUNTS + 6x+F9+0+39x+F9+0 )
     RARSAM = RARSAM/TMS
     THRSAM = THRSAM/THS
                                         PRINT 146, RARSAM, THRSAM
 146 FORMAT( 1H +12HCOUNTS/MIN +11X+F9.3+39X+F9.3)
     ERASAM = SORTF(ABSF((RARSAM/TMS)+ERARBK))
     ETHSAM = SQRTF(ABSF((THRSAM/TMS) + ETHRBK))
     RARSAM = RARSAM-PARBK
                                       .....
     THRSAM = THRSAM-THRBK
                                      the second method of the second
     PRINT 157+RARSAM+ERASAM+THRSAM+ETHSAM
 157 FORMAT(1H +13H NET COUNTS +4X+F8.3+3H+- +F8.3+27X+F8.3+3H+- +F8.3
    1)
    )
EAB = (A \diamond B) \diamond SQRTF((EA/A) \diamond 2 + (EB/B) \diamond 2)
     IF(THRSAM) 170+170+171
 170 TIRA = RARSAM
     E2RA = ERASAM
     GO TO 172
 171 TIRA = RARSAM-(A+THRSAM)
     EIRA = (A+THRSAM)+SORTF((EA/A)++2 + (ETHSAM /THRSAM)++2)
     E2RA = SQRTF((ERASAM )++ (E1RA)++2)
 172 T2RA = T1RA/(1.2-(A+B))
                                           TRA = T2RA/(1.2*RARRA)
     ACTIVITY OF RADIUM STANDARD = 1.0 UC
С
     1.2 = CHAIR FACTOR = RATIO OF ACTIVITY IN BODY TO SAME ACTIVITY A
С
Ĉ
     POINT SOURCE DUE TO BETTER BODY CRYSTAL GEOMETRIC RELATION.
    ERA=ABSF(TRA)+SQRTF((E2RA/TIRA)++(EAB/(A+B))++2+(ERARRA/RARRA)++
  TITH = THRSAM-(BOT2RA)
                                          TTH = (T1TH+0.72)/(1.8+THRTH)
     0.72 = ACTIVITY IN UC OF THORIUM STANDARD.
С
     1.8 = 1.2 CHAIR FACTOR # 1.5(RATIO OF TL-208 YIELD FROM TH-232
C
    STANDARD TO THE TL-208 YIELD FROM RA-228 IN PATIENT
C
    EOTH = ABSF(B+T2RA)+SQRTF((EB/B)++ (E2RA/T2RA)++2
    EITH = SQRTF((ETHSAM) + 2 + (EOTH) + 2)
     ETH = ABSF(TTH)+SQRTF((EITH/TITH)++2 + (ETHRTH/THRTH)++2)
    PRINT 100
                                MICROCURIES (UC) )
 100 FORMAT( 1H0+19X+33H BODY BURDEN
     PRINT 50, TRA, ERA
  50 FORMAT( 1H0+28H RADIUM C BODY BURDEN
                                      = +F12.6+4H +- +F12.6)
     PRINT SIJTHJETH
  51 FORMAT(1H0,28H MESO THORIUM BODY BURDEN = +F12.6,44H +- +F12.6)
     IF(TRA-AMIN) 160+160+162
 160 PRINT 161
 161 FORMATCIHO,22H RADIUM C BODY BURDEN ,5X,3H = ,14X,3HMSA )
 162 IF(TTH-BMIN) 163,163,165
 164 FORMAT(1H0+28H MESO THORIUM BODY BURDEN = +14X+3HMSA )
 165 GO TO (2+1)+KA
                   END
     SUBROUTINE SIGMIN(S)
                       COMMON CMIN
     SQ = SQRTF(2.0+S)
                    SA = 8.0*S
                    ......
     SB = SORTF(SA)
     SC = 4.0*S
     ALPHA = 1.2816
     CMIN = ALPHA+(SQ+(SQRTF(1+0 +(ALPHA+2)/SA) + (ALPHA/SB)))
     RETURN
     END
```

1 - 21

30.0 BACKGROUND RUN 1443 10/7/65 15.0 RADIUM 226 1.0 UC RUN 1445 10/M/65 15.0 THOPIUM 232 0.72 UC RUN 1444 10/7/65

SAMPLE OUTPUT PAGE FROM BURDEN

.

4.08 7.72 ì THORIUM STANDARD 882.866 860.19+-234,533 203.23+-3518. 13243. -995 Ħ |-A≎B RADIUM STANDARD 8.42 16.1 1049.200 43.600 654. 15738. • 0018 20.93+--+06-1101 • 0205+-H æ 31.300 22.666 22.666 BACKGROUND 680. 939. • 005 . COUNTS/MINUTE Net Counts/MINUTE NET COUNTS/MINUTE .2362+-COUNTS/MINUTE SUM 140-159 SUM_206-229 ∎ ∀

RUN **6**5 ~ ř RUN ON 10 / 1.603 .00074 UC 1.210 48.091 THORIUM REGION THORIUM REGION 21.300 639. 206-229 206-229 -1.366+-•002789 .0005c3 MICROCURIES (UC) .025146 +--+ 000929 +-1.762 RADIUM REGION RADIUM REGION 140-159 140-159 61.866 BODY BURDEN 1.878 .00154 UC 56.366 1856. R 30.566+-MESO THOKIUM BODY RURDEN 5001 RADIUM C BODY BURDEN ł 30.0 NJRRP CASE TOTAL COUNTS COUNTS/MIN NET COUNTS MSCR HSC MSA

MSA

MESO THORIUM BODY BURDEN =

TABLE I

RADIUM DIAL PAINTER BODY BURDEN MEASUREMENTS 1965 - 1966

NIMBER	DATE	na Fvni nod	ם (ה) Phil	WHOLF BA		BLADEN	(¹)	RADON F	XPTRED	VOLUME
		nattdva		COUNTING	IUU		$\hat{\mathbf{b}}$			
5021	3/15/65	•0134+-	•0026	• 0036+-	•0000	-+0110+	•0027	•7882+-	1991 •	10+35
5284	3/10/65	.1472+-	• 00 3 I	•0886+-	• 00 1 8	.2358+-	• 0035	.6242+-	.0162	6.44
5014	4 14/65	•0895+-	•0034	•0569+-	•0034	-++911-	•0039	•6113+-	.0765	8.05
5551	4/28/65	-+2+10.	• 006 1	-+6200.	1100.	•0226+-	.0061	• 6 5 0 4 + -	.3235	7.26
5818	6/02 65	•0132+-	.0037	-+1010.	.0012	•0233+-	• 0038	• 5665+-	.1848	7.15
5413	5/11 65	•0108+-	•0080	•0036+-	•0007	-+ ** 10 •	•0080	•7500+-	•6954	8.55
5321	5 19/65	-+2#10+	• 0026	•0089+-	• 0008	•0236+-	.0027	.6228+-	•1314	5.37
5040	7 13/65	•0077+-	•0012	•0056+-	.0012	•0133+-	• 0016	•5789+-	• 1 1 6 6	6.54
5121	5 18/65	•0066+-	•0027	•0035+-	.0007	-+1010-	• 0027	•6534+-	.3225	7.06
5358	5 04/65	•2513+-	•0175	•0868+-	• 0074	•3381+-	0610.	• 74 32+-	• 0665	01-11
5251	9 29/65	•0315+-	•0028	•0156+-	• 00 1 4	-+12+0•	•0031	•6687+-	.0742	8.25
5001	10/07/65	-+06 #0	•0024	•0251+-	0100.	•0681+-	•0025	• e314+-	.0426	7.10
5025	11/09 65	-+26 h0-	•0003	•0355+-	•0034	•0847+-	•0035	-5808+-	•0263	6.50
5358	6 23/66	-2040+-	•0028	•0772+-	•0047	.2812+-	•0054	•7254+-	.0172	9.10
5121	6 15/66	•0085+-	• 000 •	-+1+00*	• 0018	.0126+-	• 0 0 I 8	•6746+-	•0968	7.24
5040	6 10/66	•0080+-	• 0002	•0053+-	6100°	•0133+-	• 00 I G	• 6015+-	.0877	6.86
5321	6 07/66	•0183+-	1100.	•0131+-	.0020	•0314+-	• 0022	.5828+-	• 0549	6.12
5025	6/01 66	•0203+-	• 00 15	•0330+-	•0029	•0833+-	•0032	• 6038+-	.0297	7.00
5413	5/26 66	-+0600*	1100.	•0048+-	•0018	•0138+-	.0021	.6521+-	.1276	7.30
5818	5/11 66	~+C610*	• 0012	-+1010.	.0020	-+#£20•	•0023	• 6564+-	.0661	7.70
5215	4/06 66	1.1209+-	•0069	•4280+-	•0289	1.5489_{+-}	.0297	.7236+-	•0145	7.78
5551	4 01/66	•0175+-	• 0017	•0067+-	.0016	• 0242+-	• 0023	-1231+-	0660.	11.40
5014	3 31/66	•0813+-	• 0012	•0459+-	•0036	.1278+-	•0037	• 6408+	.0212	6.26
5021	3/25/66	-+1810.	• 00 1 4	•0022+-	.0015	•0159+-	•0020	- 8616+-	.1418	8.95
5284	3/23/66	• I 5 I 0 + -	• 0017	•0862+-	•0060	.2372+-	•0062	•6365+-	.0182	5.47
5119	6 28/66	•0277+-	•0026	•0051+-	• 00 1 8	•0328+-	•0031	8445+-	•1136	8.39
2917	8/10/66	•0416+-	•0001	•0412+-	•0035	.0828+-	•0035	• 5024+-	.0219	3.82
AVERAGE	0.6.F. =	.662967	+- 0.08	2656						

TABLE II

COMPARISON OF MODELS FOR RADON RETENTION

Test	% Retention	Predict	ed Radon	Retention	Reference
Subject	(t in days)	29 yrs	35 yrs	46 yrs.	
Mice	5.8 t ^{0.1804}	30.9	31.8	33.2	10
Dogs	5.7 $t^{0.20}$	36.2	37.8	40.0	8
Mice	5.16 t ^{0.208}	36.1	37.9	39.5	7
Mice	4.95 t ^{0.224}	39.3	41.0	43.9	4
Dogs(RaC) Humans(Ra)	$3.89 t^{0.24}$	35.9	37.9	40.8	11,16
Humans	Measured(8 cases)	31.2			10
Humans	Measured(6 cases)	-	34.4		10
Humans	Measured(2 cases)	26.5			This report
Humans	Measured(28 cases)			33.7	This report

•

TABLE III

COMPARISON OF 1965 AND 1966 EFFECTIVE HALF LIFE ESTIMATES

	19	965		1966
Case #	No.Points	Estimated T ₁ (years) ²	No,Points	Estimated T <u>1</u> (years)
5001	4	-37.73 <u>+</u> 74.57	6	+ 19.91 <u>+</u> 22.14 ^{**}
5014	5	25.19 <u>+</u> 11.92	7	13.60 <u>+</u> 3.78
5025	7	18.88 <u>+</u> 5.97	11	12.50 ± 1.92
5096	2	$14.03 \pm 0.31^*$	2	$14.03 \pm 0.31^*$
5251	2	$-5.55 \pm 0.046^*$	4	- 9.12 <u>+</u> 6.17
5278	2	$10.16 \pm 0.052^*$	2	$10.16 \pm 0.052^*$
5281	4	18.71 <u>+</u> 15.73	4	18.71 <u>+</u> 15.73
5284	6	6.80 <u>+</u> 1.57	9	8.61 ± 1.40
5818	2	$-4.15 \pm 0.011^*$	4	- 40.0 <u>+</u> 120 ***
5917	4	10.74 ± 2.39	5	9.04 ± 0.89
5962	2	$7.46 \pm 0.020^*$	2	$7.46 \pm 0.020^*$
Pooled M	lean 38	15.7 ± 2.4	56	14.73 ± 2.31

Notes:

	*	Underestimate	of	standard	error	when	only	two	points	are	used.
--	---	---------------	----	----------	-------	------	------	-----	--------	-----	-------

- ** Negative half life reported in 1965 is mainly due to 1 point which is out of line.
- *** Large value due to very small change in body burden from 1965 to 1966.

TABLE IV

COMPARISON OF HALF LIVES RESULTING FROM AN EXPONENTIAL MODEL AND THE APPARENT HALF LIFE PREDICTED BY FITTING A POWER FUNCTION TO

THE DATA

Case #	Number of Points	T ₁ Exponential ² (years)	T _{app} Power Funct. (years)
5001	6	19.91 ± 22.12	20.32 <u>+</u> 23.03
5014	7	13.60 ± 3.78	13.64 <u>+</u> 3.88
5021	6	3.86 <u>+</u> 1.19	3.91 <u>+</u> 1.24
5025	11	12.50 ± 1.92	12.55 <u>+</u> 1.96
5040	4	-81.35 <u>+</u> 748	-97.06 ± 1042
5089	2	-8.36 ± 0.54	-8.34 <u>+</u> 0.61
5096	2	14.03 <u>+</u> 0.17	14.05 ± 0.51
5102	2	5.95 ± 0.12	5.93 <u>+</u> 0.38
5118	2	1.42 ± 0.02	1.41 <u>+</u> 0.07
5119	4	13.58 ± 17.97	13.50 <u>+</u> 17.64
5121	5	43.69 <u>+</u> 178	39.82 <u>+</u> 147
5153	6	4.91 <u>+</u> 3.09	4.93 <u>+</u> 3.15
5184	3	4.10 <u>+</u> 0.62	4.15 <u>+</u> 0.58
5215	5	15.37 <u>+</u> 0.68	15.15 <u>+</u> 0.61
5251	4	-9.12 ± 6.17	-9.38 <u>+</u> 6.56
5257	2	0.77 <u>+</u> 0.011	0.77 <u>+</u> 0.016
5278	2	10.16 <u>+</u> 0.052	10.17 <u>+</u> 0.15
5281	4	18.71 <u>+</u> 15.73	18.54 + 15.32
5284	9	8.61 + 1.40	8.61 + 1.38

Continued

TABLE IV (Contd.)

Case #	Number of Points	T <u>1</u> Exponential 2 (years)	T _{app} .Power Funct. (years)
5307	2	6.52 <u>+</u> 0.04	6.55 <u>+</u> 0.03
5321	5	-12.91 <u>+</u> 15.39	-13.49 + 16.75
5358	2	6.66 ± 0.07	6.66 <u>+</u> 0.16
5369	3	7.28 <u>+</u> 19.24	7.51 <u>+</u> 20.38
5413	4	3.84 <u>+</u> 2.22	3.88 <u>+</u> 2.24
5541	3	0.46 + 0.47	0.47 <u>+</u> 0.48
5551	4	23.82 ± 17.0	24.32 <u>+</u> 17.63
5631	4	31.72 <u>+</u> 125	30.95 <u>+</u> 119
5818	4	-40.0 <u>+</u> 120	-39.13 <u>+</u> 114
5917	5	9.04 <u>+</u> 0.89	8.96 <u>+</u> 0.93
5962	2	7.46 <u>+</u> 0.02	7.46 <u>+</u> 0.10
Pooled	Mean 124	10.71 <u>+</u> 2.58	10.40 ± 2.32

TABLE V

.

EXPOSURE HISTORY OF CASES STUDIED

Case #	Initial Exposure Year	Duration (years)	Zero Time for Calculation	Years Elapsed to 1963
5001	1917	2.92	1918.5	45.97
5014	1916	3.00	1917.5	46.00
5021	1917	0.33	1917.2	47.34
502 5	1917	1.83	1917.92	46.12
5040	1917	1.00	1917.50	46.80
5089	1917	0.500	1917.25	46.55
5096	1918	0.50	1918.25	43.49
5102	1916	6.00	1919.60	42.24
5118	1917	1.333	1917.67	44.18
5119	1924	3.17	1925.58	38.10
512 1	1921	0.67	1921.33	42.49
5153	1923	2.50	1924.25	39.29
5184	1922	3.00	1923.50	39.74
5215	1917	1.50	1917.75	45.13
52 51	1917	8.25	1921.12	42.40
5 2 57	1932	23.33	1943.67	17.23
5278	1917	1.00	1917.50	45.62
5 2 81	1916	2.92	1917.46	44.80
5 2 84	1917	1.50	1917.75	45.95
5307	1944	1.33	1944.67	18.30
5321	1916	0.25	1916.12	48.13
5358	1923	28,33	1937.16	28.75
5369	1919	0.50	1919.25	44.24

TABLE V cont

5413	1916	1.00	1916.50	48.05
5541	1937	7.00	1940.50	24.40
5551	1918	0.75	1918.38	46.42
5631	1917	0.42	1917.21	46.12
5818	1916	2,00	1917.00	48.00
5917	1918	0.75	1918.38	45.52
5962	1932	1.83	1932.92	31,02
				49 19
			mean	43,13





FIGURE 2









FIG. 5





% ERROR ASSOCIATED WITH OUTGASSED RADON






(SAAAY) AT

References

- Altschuler, B., B. Pasternack, <u>Health Physics</u>, 9, 293-296 (1963).
- Brownlee, H.A., <u>Statistical Theory and Methodology</u>
 <u>in Science and Engineering</u>, New York ; V. Wiley and Son (1963) Section 11. 19.
- (3) Eisenbud, M.E., H. Petrow, N.Y.U. Annual Report USAEL
 Contract At (30-1) 3086. NYO 3086-4. (1965).
- (4) Finkel, A.J., C.E. Miller, <u>Radiation Research</u>, 14.
 466 (1961).
- (5) Hallden, N.A., J.H. Harley, <u>Health Physics</u>, 10, 265-9
 (1964).
- (6) Harley, N.A., Masters Thesis New York University, 1966.
- (7) Mays, C.W., T.H. Cochran, W.S.S. Jee <u>Health Physics</u>, <u>9</u> 615-619 (1963).
- Mays, C.W., M.A. Van Dilla, R.L. Floyd, J.S. Arnold
 Radiation Research, 8 480-489 1 (1958).
- (9) Mc Williams, D.C., J.E. Furchner, C.R. Richmond, <u>Health</u> Physics, 10, 817-822 (1964).
- (10) Miller, C.E., A.J. Finkel, <u>Radiation Research</u>, 26, 269-286 (1965).
- Norris, W.P., T.W. Speckman, P.E. Gustafson, <u>Am. J.</u>
 Roentgenology Radium Therapy Nuc. Med. 73, 785-802 (1955).
- (12) Norris. W.P., S.A. Tyler, and A.M. Brues, <u>Science</u>, <u>128</u> 456-462 (1958).

- Norris, W.P., T. Speckman, R.A. Craig, C. Armstrong,
 Argonne National Laboratory Report ANL-5288 pp. 79-82 (1954).
- Petrow, H. Ph. D. Thesis New York University, U.S.A.E.C.Report NYO-3086-5 In Press.
- (15) Rowland, R.E., J. Jowsey, J.H. Marshall, <u>Radiation</u> Research, 8, 298-306 (1958).
- (16) Schlundt, H., J.T. Nerancy, J.P. Morris <u>Am. J.</u> <u>Roentgenology Radium Therapy Nuc. Med.</u>, <u>30</u>, 515-522 (1933).
- (17) Van Dilla, M.A., B.J. Stover, R.L. Floyd, D.R. Atherton,
 D.H. Taysum, Radiation Research, 8, 417-437 (1958).

PART II

A Study of the Distribution of Ra-226, Ra-228 Pb-210 and Th-228 in Bone and Soft Tissue of Radium Dial Painters

Henry G. Petrow

This section is reported here is summary only. The full report (147 pages), which constitutes the doctoral disertation submitted by the author in partial fulfillment of the requirements for the degree of Doctor of Philosophy, is on file with the Division of Biology and Medicine, and is available from the AEC in microfilm form (NYO-3086-5).

Summary

Bone and soft tissue samples from two radium dial painters, who died approximately 50 years after being exposed to Ra-226 and Ra-228, were analyzed for Pb-210 and Ra-226, and in the case of bone tissue, for Ra-228 and Th-228 as well. In addition bone obtained from a beagle injected with pure Ra-228 was analyzed for Ra-228 and Th-228. The purpose of the study was to determine what fraction of an old Ra-226 burden resided in soft tissue, and through measurement of Pb-210 to Ra-226 and Th-228 to Ra-228 ratios, to determine the metabolic behavior of Pb-210 and Th-228 relative to their radium parents; to determine if possible, the relationships of Rn-222 retention factors to the locus of Ra-226 deposition;

to determine the maximum amount of radium, lead, and thorium translocation in the skeleton; to estimate the biological half-life of radium through measurement of the Th-228 to Ra-228 ratio; and to assess the dosimetric significance of the findings.

The soft tissue study for one of the two dial painters indicated that 0.05% of the Ra-226 and 0.65% of the Pb-210 present at death were in soft tissue. The largest concentrations of both of these radionuclides were in the aorta and the thyroid. The dose to both of these organs was low provided their radioactivity content maintained a constant relationship to the skeletal burden throughout exposure and subsequent life. The possibility exists, however, that during exposure, where radium blood levels were undoubtedly high, the dose to both the thyroid and the aorta was high.

The average skeletal ratio of Pb-210 to Ra-226 for both dial painters indicates that these two radionuclides are eliminated from the skeleton with the same biological halflife. The analysis of excreta obtained from one of the dial painters supports this hypothesis.

The variability of the Pb-210 to Ra-226 ratio from site to site in the skeleton follows a distinct, reproducible pattern. For a given bone, the magnitude of the ratio is correlated with the Ra-226 concentration, with low ratios

being found in the region of high radium concentrations. For long bones, the identical correlation was found with respect to ratio value and locus of radium deposition. Low ratios were found in the end pieces of long bone, and high ratios in the shaft. Low ratios were found in vertebra and high ratios in rib.

Two mechanisms have been suggested to explain the variability in the Pb-210 to Ra-226 ratios. Translocation, as the result of radium and lead resorption followed by partial redeposition of the resorbed material is one possibility. If translocation is the sole cause of ratio variability, then the pattern of redeposition must be selective. That is, the redeposition of Pb-210 must proceed preferentially, relative to radium, in the shafts of long bones and in rib. The maximum amount of resorbed radium or lead redeposition is 4% of the terminal body burden. If both radionuclides are redeposited to the same extent, then 2% of the terminal body burden of Ra-226 and Pb-210 was redeposited.

An alternative hypothesis that could explain ratio variability is differences in Rn-222 retention factors. For long bones, this hypothesis is supported by our current knowledge of long bone structure and the mechanism of Rn-222 escape from bone, in that one could expect greater Rn-222

retention in the shaft, relative to the end pieces, and hence, ratios should be higher in the shaft than in the ends. This is the pattern of ratio variability found in long bone. Animal data, to some extent, corrobrate these findings. The reasons for high ratios in rib and low ratios in vertebra are not clear, but may result from differences in bone crystal size, or amorphous mineral content. If Rn-222 retention is the cause of ratio differences, as a consequence of the fact that Pb-210 and Ra-226 have equal biological half-lives, Rn-222 retention factors can be calculated for each ratio. The range of ratios was such for the two cases, that Rn-222 retention varied from 20% to 57%, if ratio differences are ascribed solely to differences in Rn-222 retention.

The study of Th-228 to Ra-228 ratios in the bone of one of the dial painters and in the beagle's bone indicated that this ratio was variable as was that of the Pb-210 to Ra-226 ratio, without, however, showing any discernible pattern. It is suggested that Th-228 translocation was the cause of the variability, and that 27% of the Th-228 translocated for the dial painter and 10% for the beagle. The average skeletal ratio of Th-228 to Ra-228 allows the calculation of the Ra-228 biological half-life, assuming that the effective half-life of Th-228 is equal to its physical half-life. The calculated Ra-228 biological half-life was 10.4 years for the dial painter and 16 years for the beagle.

Using the findings of the study, it is estimated that both dial painters had equal burdens of Ra-226 and Ra-228 after exposure. The alpha dose received from each chain was nearly identical after 50 years. About 5% of the alpha dose from the Ra-226 chain was received from Po-210, the alpha emitting daughter of Pb-210.

PART VII

Discussion of Possible Mechanisms Involved in the Apparent Concentration of ²²⁶Ra and ²¹⁰Pb in Aorta and Thyroid

Norman Cohen

Introduction

The ²²⁶Ra and ²¹⁰Pb content of various soft tissues from two former radium dial painters has been measured to determine what portion of the total body burden is found in soft tissues of individuals with old burdens, what organs concentrate radium and, finally, whether the soft tissue dose is significant relative to the skeletal dose (1). The dosimetric significance of these findings is discussed in Chapter II. In addition the biochemical mechanism responsible for concentration of these nuclides and the associated radiobiological implications will be discussed.

Experimental Results

In Table I are presented the data for 33 soft tissue samples from N.J. Case No. 5281 (estimated body burden at death = 0.6 μ c) and 5 soft tissue samples from N.J. Case No. 5278 (estimated body burden at death = 0.05 μ c). The important organs, as regards both ²²⁶Ra and ²¹⁰Pb concentrations are the aorta and thyroid.

In Case No. 5281, the 226 Ra concentration of the aorta was at least 8 times as great as the next highest value obtained, excluding the thyroid. The aortic tissue is more than 43 times the average of the soft tissues other than thyroid.

In Case No. 5278, the 226 Ra in the aorta was more than 18 times the average 226 Ra content of the soft tissues and at least 9 times as

Table I

The analysis of Soft Tissues From Case 5281 and Case 5278 for

Pb-210 and Ra-226

Organ

Ra-226 pc/wet gram

Pb-210

Case 5281 Lung, Left	0.037 ± 0.004	0.044 ± 0.004
Right Kidney	0.011 ± 0.001	0.031 <u>+</u> 0.003
Brain	0.027 ± 0.002	0.030 ± 0.001
Liver	0.002 ± 0.001	0.076 <u>+</u> 0.010
Lung, Right	0.008 ± 0.003	0.056 <u>+</u> 0.020
Colon	0.003 ± 0.007	0.008 ± 0.004
Left Adrenal	0.053 ± 0.004	0.050 ± 0.010
Breast, Left & Rt.	Not Detectable	0.011 ± 0.005
Right Adrenal	0.017 ± 0.003	0.046 ± 0.015
Stomach	0.006 ± 0.002	0.029 ± 0.015
Duodenum	0.010 ± 0.006	0.035 <u>+</u> 0.035
Brain	0.009 ± 0.001	0.022 ± 0.007
Urinary Bladder	0.003 ± 0.001	0.009 ± 0.005
Uterus and Adnexa	0.004 + 0.001	0.020 + 0.003
Finger Tissue, left	—	_
ring and middle	0.030 + 0.002	0.050 + 0.005
Femoral Lymph Nodes	Not Detectable	Not Detectable
Pancreas	0.009 + 0.001	0.010 + 0.010
Spleen	0.044 + 0.003	0.024 + 0.012
Heart	0.011 + 0.001	0.073 + 0.003
Pituitary	Not Detectable	Not Detectable
Rector Sigmoid	0.018 + 0.004	0.003 + 0.003
Vagina	Not Detectable	Not Detectable
Illeum	Not Detectable	Not Detectable
Esophagus	Not Detectable	Not Detectable
*Thyroid	0.110 + 0.005	0.180 + 0.020
Diaphragm	Not Detectable	0.600 + 0.100
Larvnx	Not Detectable	Not Detectable
Gall Bladder	Not Detectable	Not Detectable
Appendix	Not Detectable	Not Detectable
Jejunum	Not Detectable	Not Detectable
*Aorta	$0 430 \pm 0 020$	1.010 + 0.050
Tracheal Lymph Nodes	Not Detectable	0.070 ± 0.020
Liver	Not Detectable	0.080 ± 0.010
milei	Not Detectable	
Case 5278 Uterus	0.003 ± 0.000	0.004 ± 0.000
Left Kidnev	0.004 + 0.000	0.012 + 0.001
Right Kidnev	0.010 + 0.000	0.009 + 0.000
Heart	0.003 + 0.000	0.012 + 0.001
*Aorta	0.089 + 0.008	1.01 + 0.050
	- ·····	

great as the next highest soft tissue measured.

The concentration of 226 Ra obtained for thyroid tissue in Case No. 5281 was 11 times greater than the average but still 4 times smaller than the concentration in the aorta. Thyroid tissue for Case No. 5278 was not available for analysis.

The concentration of 210 Pb in the aorta of Case No. 5281 was about 39 times as great as the average 210 Pb concentration excluding values for the thyroid, (approximately 7 times greater than the average) and the diaphragm, (approximately 23 times greater than the average).

Van Middlesworth (11) reported that some cattle thyroids contained sufficiently high concentrations of 226 Ra and 228 Ra to be detectable by gamma spectrometry. Workers at MIT have confirmed this estimate that as much as 50pc each of 226 Ra and 228 Ra are present in some cattle thyroids. Tipton (6) has shown concentrations of lead, calcium, barium, and strontium in the thyroid and aorta which are higher than most other soft tissues in the body.

In the case of the aorta, the anomanously high lead and radium values may be explained by cationic binding by mucopolysaccharides. Bunting and Bunting (1953) (2) showed that arterial connective tissue consists chiefly of collagen, elastin and mucopolysaccharides (chondroitin sulfate and hyaluronic acid). Both elastin and mucopolysaccharides might be suspected as binding agents on the basis of histologic distribution: however, biological considerations (3) favor the mucopolysaccharides as the potential cation binding agent. The mucopolysaccharide molecule is a polyanion containing multiple carboxyl and sulfate radicals. Rothstein in 1957 (4) was the first to propose that the mucopolysaccharides in vivo behave as ion exchanges, that is, the binding is mainly electro-

static, of a type not depending on the specific properties of the anionic groups. In the chondroitin polysulfates the binding is shared equally by weakly acidic carboxyl groups and strongly acidic o-sulfate ester groups. It has also been shown (5) that different cations exhibit varying degrees of affinity for the acid mucopolysaccharides. This further supports the idea that, in general the reactions are of the ion exchange type with nonspecific electrostatic binding, rather than of the more specific chelate type.

The common order of affinity for some of the alkaline earths has been given (5) as $Mg^{+2} \lt Ca^{+2} \lt Sr^{+2} \lt Ba^{+2}$. It has been demonstrated that calcium ion has an essential role in the concentration of arterial smooth muscle. The studies of both Waugh (7) and Bohr (8) indicate that calcium is a vital link in the coupling of excitation and contraction in vascular muscle. It also appears to have an effect on membrane excitability in arterial muscle (8).

Lamberts and Van Andel (9) have demonstrated the high bivalent cation affinity by the inner arterial intima membrane using barium and strontium. One hundred rats, each subcutaneously injected with 30 μ c of $^{140}BaSO_4$ and 30 μ c $^{85}SrSO_4$ were measured for bone, aortic wall, skin and lung activity at various times up to 15 weeks after injection. Their data clearly indicate specific soft tissue deposition for the aortic wall. After 14 weeks this tissue on a per gram basis represented 43% of the barium activity present per gram of bone tissue. For strontium the maximum occurred after 7 weeks when the aortic wall represented 1.25% of the activity present per gram of bone. The difference in aortic concentration of barium and strontium may be due to the ion exchange

order of affinity previously noted. The barium deposition displayed periodicity which will be discussed in the next section.

From the foregoing evidence it appears quite reasonable that radium, a bivalent element of the alkaline earth series, should likewise deposit on the aortic wall. In fact, considering the common cation exchange affinity for the alkaline earths, the chondroitin sulfates should have even greater affinity for radium than for barium.

Lead has also been shown to be present in the normal aorta in concentrations exceeding that for most other soft tissues in the body. It is probable that the same biochemical ion exchange mechanism is responsible for this specific site deposition. In fact Schroeder (10) et. al. has demonstrated a rather large increase in lead concentration with age in the normal human aorta. This is not surprising in the light of the fact that the mucopolysaccharide concentration of the human aorta also increases with age.

There is less information presently available to explain the rather high 226 Ra and 210 Pb concentrations found in the thyroid gland. At present it is not known whether or not mucopolysaccharides are present in the thyroid.

Radiobiological Implications

The high sensitivity of mucopolysaccharides to X-rays has been demonstrated by both Lamberts and deBoer (12) and Brinkman et.al. (13) in several ways; the immediate drop of the injection pressure in mucoid and connective tissue, viscosity changes in fresh synovial fluid and the increased permeability of the connective membranes.

Viscosity changes in synovial fluid, (which contains a large proportion of mucopolysaccharides), is explained by an irradiation depolymerization mechanism of the chondroitin sulfates. The hypothesis thus emerges that the periodic type of ¹⁴⁰Ba noted previously is a consequence of the radiation reaching a value sufficient for destruction of deposition factors and its elimination from the wall tissue, (9). The effect of the increased permeability as a result of depolymerization may result in degenerative changes in the mucopolysaccharide matrix leading to infiltration of plasma lipoprotein complexes into the membrane wall. Infiltration of fat in the intimal and medial layers of the arterial wall causes deposition of fat charged macrophages which may lead to the formation of atheromatous plaques (12). Lindsay, et. al. (1962) (14) irradiated the abdominal part of the aorta in dogs and found atherosclerosis in the irradiated vessel in 2 to 48 weeks which could not be distinguished from the "natural" process. Likewise Lamberts and deBoer (12) have described the development of atheromatosis in the carotid artery of hypercholeserolaemic rabbits within a few weeks after X-irradiation of 500-3000r.

From these observations it is evident that elastic arteries do show extensive changes a short time after irradiation with doses of X-rays on the order of 500r. Under hypercholesterolaemic conditions these changes are easily demonstrable by the penetration of fat into the wall and the deposition of lipophages on the intima. The mechanism by which these lesions develop might be explained as follows: the formation and destruction of elastic and reticular fibers, a continuous process, is closely related to an intact mucopolysaccharide matrix.

Changes in this matrix e.g. depolymerization by X-irradiation, will cause changes in the formation of these fibers, leading up to an impairment of the filter barrier function of the arterial wall. This becomes evident by the changes in elastic fibers and fat infiltration into the wall (12).

It is interesting to note that a very large number of the radium dial painters examined at autopsy showed evidence of sclerotic lesions in the arteries (15). Since arterial sclerosis is so common among the aged as almost to be considered "normal", there can be little said concerning this observation. Further experimental and epidemiologic evidence is necessary, including comparison with a control group, before any statement can be made concerning causal effects.

The dose received by the aortic linings are currently being considered. It is clear that most of the dose received by these cases was early in life when the radium was freshly injected and relatively labile. It may be necessary to resort to animal experiments to obtain information about the dose to the aorta from freshly ingested radium.

Acknowledgement:

I would like to thank Dr. Henry G. Petrow for the initiation and supervision of this preliminary study of the radium and lead distributions in the soft tissues of exposed individuals. In addition, Dr. Petrow performed all of the 210 Pb determinations reported here.

References

- 1. Petrow, H.G.: "A Study of the Distribution of Ra^{226} , Ra^{228} , Ra^{210} and Th^{228} in Bone and Soft Tissue of Radium Dial Painters", Ph.D. Thesis, New York University, April 1966.
- 2. Bunting, C. and H. Bunting : Arch. Path 55:257, 1953.
- 3. Headings, V.E., P.A. Rondell and D.F. Bohr : <u>Am. J. Physiol</u> 199:5, pp 783-787, 1960.
- 4. Rothstein, A.L.: In: Ion Exchanges in Organic and Biochemistry, Interscience, New York, pp 213-232, 1957.
- Dunstone, J.R.: Ion Exchange Reactions between Acid Mucopolysaccharides and Various Cations, J. <u>Biochem</u>. 85, pp 336-351, 1962.
- 6. Tipton, I.H. and M J. Cook: Trace Elements in Human Tissue Part II Adult subjects from the United States, <u>Health</u> Physics, 9:2, pp 103-145, 1963.
- 7. Wangh, W.H.: Circulation Research, 11:927, 1962.
- 8. Bohr, D.F.: J. <u>Canad. Med. Assn</u>. 90:174, 1964.
- 9. Lamberts, H.B. and J.G. Van Andel: The Deposition of Ra Ba and Sr in the Aortic Wall, <u>Koninklijke Nederlandse</u> Akademic Van Proceedings Series C Biological and Medical Sciences Vol. LXVIII, #4, North Holland Pub. Co., Amsterdam, 1965.
- 10. Schroeder, H.A. and J.J. Balassa: Adnormal Trace Metals in Man: Lead, J. Chron. Dis., 14:4, 1961.

- 11. Van Middlesworth, L., P. Correa, R.W. Perkins and A.T. Keane: Ra Isotopic Accumulation in Animal Thyroids, Science, 155, pp 453-4, 1966.
- 12. Lamberts, H.B. and W.G.R.M. De Boer: X-ray Induced Atheromatous Lesions in the Arterial Wall of Hypercholesterolaemic Rabbits, <u>Int. J. Rad. Biol.</u>, 6:4, pp 343-350, 1963.
- 13. Brinkman, R., H.B. Lamberts, J. Wadel and J. Zuideveld: Further Analysis of the Immediate Drop in Injection Pressure on Moderate Irradiation, <u>Int. J. Rad. Biol.</u>, 3:2 pp 205-210, 1961.
- 14. Lindsay, S., H.J. Kohn, R.L. Darkin, and J. Jew Circulation Research, 10:51, 1962.
- 15. N.J. Radium Reports NYO-2181-2-3 Epidemiological followup of the New Jersey Radium Cases: Progress Reports 1964 and 1965.

PART III

RÉSUMÉS OF FINDINGS FROM INDIVIDUAL CASE STUDIES

Previously published résumés appear in Reports NYO-2760, NYO-10604, NYO-2181-2 and NYO-2181-3

ţ

VITAL STATISTICS

Living female Born 28 September 1901; Orange, New Jersey

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

First exposure 1917 at age 18, New Jersey Radium Luminous Material Corporation as packer

ESTIMATED BODY BURDEN

MSA radium-226, NYU Whole Body Counter, November 23, 1965 MSA radium-228, NYU Whole Body Counter, November 23, 1965

FAMILY HISTORY

Mother: Father: Brother:	Died age 78 following gall bladder surgery Died age 83, automobile accident (1) Died age 66, heart failure (2) Died age 69, heart failure
	(3) Died age 33, World War I
	(4) Living and well, age 62 (RRP Control 6581)
	(5) Living and well, age 60 (RRP Control 6563)
	(6) Living and well, age 55 (RRP Control 6548)
Sister:	(1) Died age 34, radium poisoning (RRP Case 5056)
	(2) Living and well, age 58
	(3) Living and well, age 53 (RRP Control 6568)
Son:	Living and well, born 1922
Daughter:	(1) Living and well, born 1923
	(2) Living and well, born 1924

MEDICAL HISTORY

Obstetrical: Para 3 Gravida 3 Scarlet fever, childhood (050) Appendectomy, 1929 (551) Pneumonia (490) and bronchitis (501), 1946 Bladder surgery, 1948 Phlebitis, 1956 (464) Arthritis (725)

DIAGNOSES AND IMPRESSIONS FROM MEDICAL, CLINICAL LABORATORY, X-RAY AND DENTAL EXAMINATIONS AS OF 11/22/65

Apparent health: Good Fair Poor

Edentulous (533.7) Slight osteoarthritis of lower cervical spine (723.1)

CASE 5181 continued

Moderate osteoarthritis of mid-dorsal spine (723.1)
Fairly advanced osteoarthritis of lumbo-sacral spine with thinning of 4th and 5th intervertebral discs (723.1)
Degenerative changes of greater tuberosities of humeri at tendon attachments (733)
Osteoarthritis of right acromio-clavicular joint (723)
Fairly advanced osteoarthritis of both knees (723)
Symmetrical deformities with chronic dislocation of 1st and 2nd metatarso-phalangeal joints of both feet
Obesity (287)
Bilateral arcus senilis (388)
Hemangioma right hand, congenital (228)
Mild retinal and peripheral arteriosclerosis (450.0)

VITAL STATISTICS

Male (last known to be alive 1962) Born 7 November 1891

MEANS OF DISCOVERY

Referred by another case

OCCUPATIONAL HISTORY

No information on date of first exposure, age at first exposure or job description Radium Luminous Material Corporation, New Jersey

ESTIMATED BODY BURDEN

1.73 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, April 27, 1958

FAMILY HISTORY

Mother:	No information
Father:	No information
Brother:	Last known to be alive in 1960 (RRP Case 5108)

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Dead Female Born 9 October 1898; Steinbach, Germany Died 15 November 1945; Los Angeles, California

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

First exposure 1918 at age 21, New Jersey Radium Luminous Material Corporation as dial painter

ESTIMATED BODY BURDEN

3.4 micrograms radium-226, Geiger-Müeller Counter, H. S. Martland, M.D., August 24, 1940

FAMILY HISTORY

Died age 72, heart disease
Died age 79, stroke
(1) Died age 54, coronary
(2) Living and well, age 66 (RRP Case 5322)
(3) Living and well, age 56 (RRP Control 6510)
(4) Died age 19, accidental fall in home (RRP Case 5182)
(1) Died age 2, burns
(2) Living and well, age 60 (RRP Control 6511)
(3) Living and well, age 59 (RRP Control 6509)
(1) Living, age and status of health unknown
(2) Living, age and status of health unknown

MEDICAL HISTORY

No information available

DIAGNOSES AND IMPRESSIONS

No studies performed

DATA FROM DEATH CERTIFICATE 11/15/45:

Ulcerative osteogenic sarcoma of left leg (196.9) Metastasis to the lungs (165) due to radium osteitis (E961.3)

VITAL STATISTICS

Female (last known to be alive in 1961) No birthdate available

MEANS OF DISCOVERY

Referred by another case

OCCUPATIONAL HISTORY

No information on date of first exposure or age at first exposure United States Radium Corporation, New York as dial painter

ESTIMATED BODY BURDEN

0.564 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, October 26, 1959

FAMILY HISTORY

Mother:	No information
Father:	No information
Sister:	(1) No information
	(2) No information

MEDICAL HISTORY

Obstetrical: Para 0 Gravida 0 Hypertensive cardiovascular disease, 1961 (443) Herpes Zoster, left chest, 1961 (088) Enlarged heart, 1961 (434.4) Mild myocardial damage, 1961 Infected teeth, 1961 (532.3)

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Living female Born 30 March 1901; Montclair, New Jersey

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

First exposure 1917 at age 16, New Jersey Radium Luminous Material Corporation as clerk

ESTIMATED BODY BURDEN

MSA radium-226, NYU Whole Body Counter, October 25, 1965 MSA radium-228, NYU Whole Body Counter, October 25, 1965

FAMILY HISTORY

MEDICAL HISTORY

Obstetrical: Para 3 Gravida 3 Arthritis (725) Squamous cell carcinoma of the uterus, 1951 (171)

DIAGNOSES AND IMPRESSIONS FROM MEDICAL, CLINICAL LABORATORY, X-RAY AND DENTAL EXAMINATIONS AS OF 10/25/65

Apparent health: Good Fair Poor

Edentulous maxilla (533.7) 6 remaining mandibular teeth Generalized osteopprosis (733) Degenerative disc disease (735) Degenerative joint disease (723.1) Peritendinitis calcarea (741) Bilateral Morton's toes (733) Tumor right lower eyelid, type undetermined (238) Mild varicose vein, right greater saphenous (460) Probable HCVD (443)

VITAL STATISTICS

Male (last known to be alive in 1950) No birthdate available

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

No information on date of first exposure or age at first exposure United States Radium Corporation, New York as purchasing agent

ESTIMATED BODY BURDEN

0.337 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, February 27, 1950

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Female (last known to be alive in 1961) No birthdate available

MEANS OF DISCOVERY

Referred by another case

OCCUPATIONAL HISTORY

No information on date of first exposure or age at first exposure Ansonia Clock Company as dial painter

ESTIMATED BODY BURDEN

0.62 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, October 7, 1954

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Living female Born 16 March 1887, Orange, New Jersey

MEANS OF DISCOVERY

Referred by her physician

OCCUPATIONAL HISTORY

First exposure 1918 at age 31, New Jersey Radium Luminous Material Corporation as dial painter

ESTIMATED BODY BURDEN

1.25 microcurie radium-226, MIT Breath Radon, December 8, 1965

FAMILY HISTORY

Mother: Dead, age and cause unknown Father: Dead, age and cause unknown Sister: Dead, age and cause unknown No further information

MEDICAL HISTORY

Obstetrical: Para O Gravida O Pathological fracture, left femur, 1946 (N821) Pathological fracture, right femur, 1951 (N821) Pathological fracture, left elbow, 1964 (N821)

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Male (last known to be alive in 1957) No birthdate available

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

No information on date of first exposure, age at first exposure or job description United States Radium Corporation, New York

ESTIMATED BODY BURDEN

1.54 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, October 7, 1957

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Male (last known to be alive in 1951) No birthdate available

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

No information on date of first exposure, age at first exposure or job description United States Radium Corporation, New York

ESTIMATED BODY BURDEN

0.49 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, October 14, 1951

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Living male Born 8 September 1922; Reading, Pennsylvania

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

First exposure 1946 at age 25, New York United States Radium Corporation as chemist

ESTIMATED BODY BURDEN

0.0045 microcurie radium-226, NYU Whole Body Counter, September 16, 1965 0.0008 microcurie radium-228, NYU Whole Body Counter, September 16, 1965

FAMILY HISTORY

Mother:	Living	and	well,	age	64
Father:	Living	and	well,	age	65
Sister:	Living	and	well,	age	31
Son:	Living	and	well,	age	19

MEDICAL HISTORY

Pertussis, childhood (056.0) Diphtheria, childhood (055) Tonsillectomy and adenoidectomy, childhood (510.1) Submucous resection of nose, 1942 (514) Fracture, nose, traumatic, 1964 (N802)

DIAGNOSES AND IMPRESSIONS FROM MEDICAL, CLINICAL LABORATORY, X-RAY AND DENTAL EXAMINATIONS AS OF 10/28/65

Apparent health: Good Fair Poor

29 remaining teeth Chronic dental pathology, periapical Functional gastric disturbance (544.2) Borderline hypertension (BP 156/96) (444) One centimeter mass above left testicle, probable hydrocele (613) Deviated nasal septum (514) Degenerative disc disease (735)

VITAL STATISTICS

Living male No birthdate available

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

No information on date of first exposure, age at first exposure or job description United States Radium Corporation, New York

ESTIMATED BODY BURDEN

0.738 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, September 23, 1963

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Male (last known to be alive in 1958) No birthdate available

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

No information on date of first exposure, age at first exposure or job description United States Radium Corporation, New York

ESTIMATED BODY BURDEN

0.28 x 10⁻¹² curies per litre, Breath Radon by Dr. Hess, March 31, 1958

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Living male No birthdate available

MEANS OF DISCOVERY

Search by case finding group

OCCUPATIONAL HISTORY

No information on date of first exposure, age at first exposure or job description United States Radium Corporation, New York

ESTIMATED BODY BURDEN

4.55 x 10^{-12} curies per litre, Breath Radon by Dr. Hess, May 1, 1951

FAMILY HISTORY

No information

MEDICAL HISTORY

No information

DIAGNOSES AND IMPRESSIONS

VITAL STATISTICS

Female (last known to be alive in 1962) Born 24 August 1922; Kulpmont, Pennsylvania

MEANS OF DISCOVERY

Referred by another case

OCCUPATIONAL HISTORY

First exposure 1941 at age 19, United States Radium Corporation, New York No information on job description

ESTIMATED BODY BURDEN

"Inactive" Breath radon by Dr. Hess, February 14, 1950

FAMILY HISTORY

Mother:	Dead, age and cause unknown
Father:	Dead, age and cause unknown
Brother:	Last known to be alive in 1962
Son:	(1) Born 1950, last known to be alive in 1962
	(2) Born 1954, last known to be alive in 1962
	(3) Born 1957, last known to be alive in 1962

MEDICAL HISTORY

Obstetrical: Para 3 Gravida 3 No further information

DIAGNOSES AND IMPRESSIONS

APPENDIX

EXHIBIT 1

Case Finding and Follow-Up Progress Report to October 1, 1966

- I. Estimated potential number of radium cases who were exposed in New Jersey or the metropolitan areas of New York or Philadelphia....1000
- II. Number of names of all radium cases identified who were exposed in the State of New Jersey or the metropolitan areas of New York A. Number of radium cases living..... 332 1. Within New Jersey or the metropolitan areas of New York or Philadelphia..... 300 B. Number of radium cases known to have deceased..... 259 C. Number of radium cases whose current status has not yet been determined..... 387 III. Number of radium cases contacted..... 362 A. Basic medical and radiation evaluation completed..... 150 1. Number of cases currently living..... 114 B. Number of cases currently undergoing studies..... 50 C. Number of cases not studied.....161 2. Dead..... 22
- IV. To date 53 cases have been completed and are on "standby" (with the exception of a yearly whole body count in some instances).
- V. Controls

Α.	Initial studies completed82
B.	Completed all studies (including four
	additional laboratory examinations)
с.	Undergoing initial study 7
D.	Numbered but not vet started 5
U. S. Atomic Energy Commission N. J. State Depart Division of Biology & Medicine Dr. Roscoe P. Kandl	cment of Health Scientific Advisory Le, Commissioner Committee
---	---
Radium Resear Dr. Samuel C. Ingraham 2	ch Project 2nd, Project Director
Consultants Dr. Lester Goldman, Clinical Laboratory Dr. William Bernhard, Special Laboratory	Project Medical & Dental Staff Dr. Hyman W. Fisher, Project Medical Director Dr. Carye-Belle Henle, Project Radiologist Dr. Pohent Bonds Duriest Dentist
Cooperating Agencies for special radiation studies WBC, Br Ra, Radiochemistry New York University Medical Center Drs. Merril Eisenbud & Henry Petrow	Case Follow-Up Nancy Manner, Nurse Supervisor Mary Pailo Field Nurse
Breath Radon Massachusetts Institute of Technology Dr. Robley Evans	Data Collection & Processing
Subcontractors	Catharine Rowland
General Pathology & Special Radio-osteitis Study New Jersey College of Medicine & Dentistry Drs. Hugh Grady & William Sharpe	Storage & Processing of Specimens Ann Johnson
Oral Pathology Georgetown University Dental School Drs. Joseph Bernier & Joseph Belzile	Fiscal & Clerical Medora MacLaren, Administrative Assistant
Statistical Consultation & Data Processing Rutgers University Statistics Center Drs. Ellis Ott & Thomas Hayton	Mary CULL, MEGICAL SECRETARY
EXHIBI	Г 2
Table of Org	ganization



EXHIBIT 3 - Lodwick Schema for Coding X-Rays

*For femur the capital epiphysis is considered to be the "proximal epiphysis" and the two trochanters are included in the "metaphysis"



A-2









 Radius (in AP film): Number of defects in 1 Cm² at distal end of shaft and distal metaphysis at point of greatest slope.



2. Combined thickness (medial and lateral) of cortex of radius at point below radial tuberosity where cortices of radius and ulna become parallel (measured to nearest 0.1 mm).



3. Total width (thickness) of radius at point below radial tuberosity where cortices of radius and ulna become parallel (measured to nearest 0.1 mm).



4. Combined cortical thickness (medial and lateral) of second metacarpal bone(in AP film) through thickest area of cortex (measured to nearest 0.1 mm).





If present, specify: USE SEPARATE SHEET FOR EACH DIAGNOSIS PERTINENT TO BONE

WHO Code No.

INSTRUCTIONS AND DEFINITIONS FOR USE WITH X-RAY CODING FORMS

FORM 1

Form 1 (consisting of two pages) is used to code all bones and portions of bones containing changes in categories listed on Forms 2 and 3. Each set of x-rays on a case will have one Form 1.

Basic information

- 1. Enter month, day and year of x-ray film set being interpreted.
- 2. Enter examination number, also checking block 10 if x-rays are of specimens rather than of case.
- Enter assigned number of radiologist doing the interpretation (Dr. Henle #1, Dr. Bonda #2, etc.).
- 4. Enter source of x-rays (2 Taken by RRP, 3 Taken for RRP, 3 Other).
- 5. Enter case number.

Interpretation

- 1. Check appropriate space for each bone interpreted.
- 2. Check area or areas in which changes of categories listed on Forms 2 (Destruction) and 3 (Proliferation) are located or check appropriate "no changes" block if there are no changes of categories listed on Forms 2 and 3.
- n.b. The blocks marked "DO NOT USE" are for coding at the Project office preparatory to punching IBM cards for this deck.

FORM 2

Form 2 is for coding destructive lesions in bones or portions of bones. There will be a Form 2 or Form 3 or both for each entry in the "changes" columns of Form 1. Normally there will be no Forms 2 or 3 to match entries in the "no changes" columns of Form 1.

Basic Information

- 1. Enter month, day and year of x-ray film set being interpreted.
- 2. Enter examination number.
- 3. Enter case number.
- 4. Enter Area Code number which is comprised of ...

On page 1 of Form 1: the two digits to the left of the name of the bone being interpreted plus the digit above the portion of the bone being interpreted ie. the Area Code for a change in the shaft of the Left Clavicle would be "013".

On page 2 of Form 1: the two digits to the left of the name of the bone being interpreted ie. the Area Code for a change in the Skull (vault) would be "29". Since there is no third digit, box 15 of the Area Code is left blank.

When the interpretation is "no changes" in categories listed, no Form 2 or Form 3 will be prepared.

FORM 2 (CONTINUED)

Interpretation

1. DIFFUSE CORTICAL POROSIS (See 1, 2, 3)

(16) Check one only.

(17)-(18) "Greatest number of defects per Cm^2 " - to quantitate greatest occurrence of cortical defects, select most involved area of bone or bone segment covered by Form 2 and count defects in 1 Cm^2 .

(19)-(21) "% of area involved" is to be a visual estimate of % of bone or bone segment covered by Form 2 that the defects constitute.

2. CORTICAL DEFECTS seen in profile (See 4, 5, 7, 15)

(22)-(24) "No. of defects in area" is to be an actual count or, if defects are very numerous, a best estimate of number of defects in the bone or bone segment described by the Form 2.

(25)-(27) "% of area involved" is to be a visual estimate of % of bone or bone segment covered by Form 2 that the defects constitute.

(28)-(35) "Range of size of defects" is to be actual measurements (rounded off to the nearest 0.1 mm) of the dimensions of the defects visually selected as being the largest and smallest ones present.

- 3. SUBCORTICAL (ENDOSTEAL) DEFECTS seen in profile (See 4, 5, 6) as for CORTICAL DEFECTS.
- 4. MEDULLARY DEFECTS (See 8)

(50)-(55) Length (longest dimension of defect), width (measure at right angle to length of defect) and depth (seen on lateral film, if available) are the three dimensions of a defect mutually at right angles to each other but not necessarily in line with any axis of the bone or bone segment being described. Normally only two of these dimensions are recordable from the available films. N.A. means "Not Applicable".

(56) "Margin": Distinct - margin of the lesion is clearly and easily determined. Less Distinct - margin of the lesion is <u>reasonably</u> clearly and easily determined. Indistinct - margin of the lesion is quite vague.

n.b. These terms refer to determinability of the margin, not its contour.

(57)-(59) "% of area involved" is to be a visual estimate of the % of the study area represented by the Form 2 that appears to be involved.

FORM 3

Form 3 (consisting of two pages) is for coding proliferative lesions in bones or portions of bones. There will be a Form 3 or Form 2 or both for each entry in the "changes columns of Form 1. Normally there will be no Forms 3 or 2 to match entries in the "no changes" columns of Form 1.

Basic Information as for Form 2.

Interpretation

1. DIFFUSE MEDULLARY SCLEROSIS (See 10, 17)

(16)-(18) "% of area involved" is to be a visual estimate of degree of involvement of bone or bone segment covered by the Form 3.

(19)-(22) "Size of lesion" is to be the best reasonable estimate of the size in square centimeters of the area involved.

(23)-(25) "Degree of solidity" is to be a visual estimate of the % of the coded area represented by the Form 3 that appears to be sclerosed.

n.b. These terms refer to determinability of the margin, not its contour

2. FOCAL TRABECULAR SCLEROSIS (See 11, 12, 13, 14, 16)

(27)-(29) "Number of foci in area" is to be an actual count or, if foci are very numerous, a best estimate of number of defects in the bone or bone segment described by the Form 3.

(30)-(41) "Range of size of foci" is to be actual measurements (rounded off to the nearest mm) of the dimensions of the foci visually selected as being the largest and smallest ones present.

(42)-(44) "Degree of solidity" - See (23)-(25) above.

(45) "Margin" - See (26) above.

3. PERIOSTEAL THICKENING (See 15)

(46)-(49) "Degree of thickening" is to be a measure (rounded off to the nearest mm) of the maximum thickness of major area of periosteal thickening.

(50)-(52) "% of length of area" is to be the best visual estimate.

(53) "Regular" and "Irregular" refer to contour of periosteal thickening.

4. CORTICAL THICKENING as for PERIOSTEAL THICKENING.

5. ENDOSTEAL THICKENING (See 5) as for PERIOSTEAL THICKENING.

FORM 3 (CONTINUED)

- 6. SEQUESTRAE (See 9) Check as indicated.
- 7. CORTEX INSIDE OF CORTEX (See 4, 6, 15) Check as indicated.
- 8. TRANSVERSE LINEAR DENSITIES Check as indicated.
- 9. OBLIQUE LINEAR DENSITIES Check as indicated.

FORM 4

Form 4 is for coding special standard measurements made on x-rays of designated bones. There will be a Form 4 for each Form 1.

Basic Information as for Form 2 (Omit Area Code)

Interpretation

- 1. Record number of defects in 1 Cm² in AP film at junction of distal end of shaft of radius and distal metaphysis at point of greatest angle. Right radius is preferred. If both right and left AP views are available, both may be recorded.
- Record combined thickness (medial and lateral) of cortex of radius at point below radial tuberosity where cortices of radius and ulna become parallel. Left radius is preferred. Both AP and lateral views should be measured and recorded if films are available. All measurements are rounded off to the nearest 0.1 mm.
- 3. Record total width (thickness) of radius at point below radial tuberosity where cortices of radius and ulna become parallel. Left radius is preferred. Both AP and lateral views should be measured and recorded if films are available. All measurements are rounded off to the nearest 0.1 mm.
- 4. Record combined thickness of medial and lateral cortex of second metacarpal bone through thickest area. Both measurements are to be made at the same level in the bone. Left second metacarpal is preferred. All measurements are to be rounded off to the nearest 0.1 mm. If both right and left AP views are available, both may be recorded.

FORM 5

Form 5 is for recording each clinical entity affecting bone that is diagnosed for a set of x-rays. A separate Form 5 is used for each diagnosis.

Basic Information as for Form 2 (Omit Area Code)

Interpretation

(16) Check as appropriate. If there is no Definite Clinical Entity Affecting Bone, there will be only one Form 5 with block (16) checked "Absent".

(17) "Degree of certainty" - Check appropriate block.

(18)-(21) "Method(s) of Establishing Diagnosis" - Check appropriate block or blocks.

"Diagnosis" - Enter name of each diagnosis pertinent to bone. The International Code Number for the diagnosis will be entered prior to the punching of the appropriate IBM card.