

Clinical Use of



Radioisotopes

WILLIAM H. BEIERWALTES, M.D.

Associate Professor of Internal Medicine and Coordinator, Clinical Radioisotope Unit, University Hospital, Ann Arbor

PHILIP C. JOHNSON, M.D.

Assistant Professor of Internal Medicine and Chief, Radioisotope Unit, Veterans Administration Hospital, University of Oklahoma Medical School, Oklahoma City

ARTHUR J. SOLARI, B.S., M.S. (Physics)

Instructor in Radiation Physics, Department of Radiology, Radiation Physicist for Clinical Radioisotope Unit and Kresge Research Isotope Unit, University Hospital, Ann Arbor

Iilustrated

W. B. SAUNDERS COMPANY

Philadelphia

1957

London

© 1957, by W. B. Saunders Company

Copyright under the International Copyright Union

All Rights Reserved. This book is protected by copyright. No part of it may be duplicated or reproduced in any manner without written permission from the publisher. Made in the United States of America. Press of W. B. Saunders Company. Library of Congress Catalog Card Number: 56-10933

Preface



THIS TEXT for the clinician was originally conceived as a means of reducing the load of repetitive teaching that we have carried in the Clinical Radioisotope Unit, University Hospital, since 1947. Our first students were technicians. Then the departments of radiology, medicine and surgery made the Clinical Radioisotope Unit a regular rotation for every resident in these departments. In addition, many residents in pediatrics, obstetrics and gynecology, ophthalmology and other specialties sought training to use radioisotopes for specialized techniques, in clinical tests or for clinical research projects.

Since the Atomic Energy Commission agreed, in 1952, to allow certain physicians to use radioactive isotopes in private practice, we have given a preceptorship type of training to many men in practice so that they might become licensed by the Atomic Energy Commission to use radioactive isotopes, or so that they might learn how to use existing radioisotope centers most intelligently for the diagnosis and treatment of their patients.

Recently, every medical student at the University of Michigan Medical School has been assigned to the Clinical Radioisotope Unit as a part of his regular rotation during some part of his junior year.

This text was written primarily to help instruct such persons in the most common clinical uses of radioactive isotopes. The space devoted to each subject is weighted largely according to the popularity of each use. Chapter 10, covering the less commonly used radioisotopes, was added because these substances have been used to some extent in human beings or because it seemed likely to the authors that they might well be adapted to routine use in the future.

Physicians, medical students and technicians have shown us what we must teach and how it must be taught. Dean Ralph Sawyer and Dr. Henry Gomberg of the Michigan Memorial Phoenix Project and Dr. Albert Kerlikowske, Director of University Hospital, made our start possible. Dr. C. C. Sturgis, Chairman of the Department of Medicine, and Dr. Fred Hodges, Chairman of the Department of Radiology, promoted the start of the Isotope Unit. Drs. Isadore Lampe and Henry

Gomberg taught the senior author the approach to radioisotope work and a healthy respect for radiation. Dr. F. A. Coller, Chairman of the Department of Surgery, has contributed almost daily in innumerable ways to insure the best possible operation of the unit. Ardath Emmons, Associate Radiation Safety Officer of the University, has been indispensable in his help and instruction in radiation health physics. Dean A. C. Furstenberg has never failed us in our endless requests.

WILLIAM H. BEIERWALTES PHILIP C. JOHNSON ARTHUR J. SOLARI

January, 1957

Contents

	Chapter 1	
CE	RTAIN PRELIMINARIES	
5	Who Should Train?	2
	The Atom Isotopes Radioactivity Measurement of Radioactivity, 31.	2 2
	Half Life	3
	The Energy Unit—The Mev The Alpha Particle The Beta Particle Positrons and Annihilation Radiation Gamma Rays Shielding of Gamma Rays, 40; Inverse Square Law, 41. The Roentgen The Rep, RBE, Rem and Rad The Detection of Radiation Gas-Filled Detectors Ionization Chamber, 43; Proportional Counters, 44; Geiger Counters, 44; Self-Quenching Counters (Organic), 46; Bismuth Cathode Geiger-Mueller Tubes, 46.	3 3 3 4 4 4 4 4
	Scintillation Counters	5
	Chapter 2	
ТН	IYROID PHYSIOLOGY AND RADIOACTIVE IODINE, I ¹³¹	5
	Definitions	5' 5'

I ¹⁸¹ Absorption, Transport and Excretion Functions of the Thyroid Gland 1. Uptake of Iodine and Tyrosine I ¹⁸¹ Practical Considerations, 62. 2. Protein Binding of Iodine	59 60 60
Mechanism of Action of Antithyroid Drugs, 67; I ¹³¹ Practical Considerations, 68.	
 Storage of Thyroid Hormone	70 71
I ¹⁸¹ Practical Considerations, 72. Transport, Action and Metabolism of Thyroid Hormone	72
Action of Thyroxine, 72.	
Chapter 3	
I ¹⁸¹ TESTS OF THYROID FUNCTION	78
Thyroid Function versus Patient Function	78 80
The Mechanics of Three Thyroid Function Tests The Two Hour I ¹³¹ Uptake Test Technique, 85; Reporting the Impression, 90.	85 85
The 24 Hour Uptake Test	91
The One Hour Uptake Test	96
I ¹⁸¹ THYROIDECTOM¥	99
Treatment of Hyperthyroidism with I ¹⁸¹	99 99
Indications for Radioiodine Treatment in Hyperthyroidism Previous Thyroidectomy for Hyperthyroidism, 103; Exophthalmic Goiter in Persons over Forty Years of Age, 104; Impractically Long Preoperative Preparation, 104; Hyperthyroidism in the Presence of Well Accepted Contraindications for Surgery, 104; Hyperthyroidism with "Malignant" Exophthalmos at Any Age, 104.	ř
Principal Reason for I ¹⁸¹ Treatment Dosage of I ¹⁸¹ One Dose Method, 106; Fractionation of Dosage, 110; Prediction Tests, 110.	106
Unpleasant Effects from I ¹⁸¹	111
Persistent or Recurrent Hyperthyroidism and Ilai Freatment	112

Contents	V 11
Goiter in Relation to I ¹³¹ Additional Aspects of I ¹⁸¹ Thyroidectomy	
Nodules in Recurrent Exophthalmic Goiter, 115; Thyroid- ectomies after I ¹³¹ , 116; Pathology, 116; Auricular Fibrillation, 117; Exophthalmos, 118.	
Results to be Expected and Follow-up Care	118
Summary: Use of I ¹³¹ in Treatment of Hyperthyroidism	
I ¹³¹ Therapy for Angina Pectoris or Intractable Congestive Failure Rationale, 121; Indications, 122; Contraindications, 122; Preparation and Dosage, 123; Results, 124.	
I ¹³¹ Therapy for Intractable Symptoms of Pulmonary Insufficiency Rationale, 125; Indications, 125; Contraindications, 125; Preparation and Dosage, 125; Results, 125.	
I ¹³ Therapy for Intractable Parkinsonism	126
I ¹³¹ Therapy for Intermittent Claudication	127
•	
Chapter 5	
I ¹⁸¹ IN THE DIAGNOSIS AND TREATMENT OF THYROID NEO- PLASMS	121
Diagnosis and Treatment of Goiters Not Yet Proved to Harbor Carcinoma Non-nodular Toxic Goiter	131
Nodular Toxic Goiter	132
Diagnostic Use of I ¹⁸¹ When Surgery Is Contraindicated, 132; Recurrence after Previous Thyroidectomy, 132.	
Nodular Non-toxic Goiters Solitary Nodules, 132; Non-toxic Multinodular Goiter, 138.	
Recurrent Goiter with Hyperthyroidism	
I ¹³¹ in the Differential Diagnosis of Carcinoma	142
Radioiodine Treatment of Proved Carcinoma of the Thyroid Gland	147
Surgical Treatment of Thyroid Carcinoma	147
Roentgen Ray Therapy	149
Indications for I ¹⁸¹	149
I ¹³¹ Concentration in Relation to Type of Carcinoma Follicular Carcinoma, 150; Papillary Carcinoma, 150; Papillary	150
and Follicular Carcinoma, 152; More Undifferentiated Types,	
155; Primary and Metastatic Types, 156.	156
Methods of Determining I ¹³¹ Concentration by Tissue	130
Urinary Excretion, 167; Blood I ¹³¹ Level, 168; Indices of Thy-	
roid Function, 168. Enhancing Uptake of I ¹⁸¹ by Thyroid Carcinoma Metastases	168
General Program of Treatment and Follow-up	170

Contents

Some Clinical Aspects of Thyroid Cancer Seen in a Radioisotope Unit Age of Patient and Colloid Production of Carcinoma, 170, Presenting Symptom, 172; Patterns Exhibited by Metastases of Thyroid Carcinoma in Lungs, 172. Treatment Dosage of I ¹³¹ Toxic Effects of Treatment Dose Local Swelling, 175; Irradiation Sickness, 179; Radiation Sialadenitis, 179; Urinary Tract Effects, 180; Depression of Bone Marrow Function, 180; Ovarian Function, 182. Urinary Excretion of I ¹³¹ after Treatment Dose Post-treatment Hypothyroidism Treatment of the Thyroid Carcinoma Patient and Concomitant Desiccated Thyroid Administration, 184. Results Pathology of Lesions and Survival after Radioiodine, 185; Deaths, 188; Patients Apparently Well, 188.	174 175 182 183
Chapter 6	
OTHER CLINICAL USES OF I131	192
Human Serum Albumin (IHSA or RISA) Labeled with Radioactive Iodine Chemical and Physical Properties Pharmacology Mixing with Blood, 192; Mixing with Extravascular Pools, 193; The Metabolic Degradation of the Isotopically Labeled Protein, 193.	192
Localization of IHSA	-
Radiation Hazard from IHSA 1131 HSA Blood Volume Determination Other Physiologic Considerations, 195; Technique of I ¹³¹ HSA Blood Volume, 198; Variations in Blood Volume, 203.	
Other Uses of I ¹³¹ -Labeled Human Serum Albumin (IHSA or RISA) Serian Tumor Localization, 203; Circulation Time, Dilution Curve and Cardiac Output, 210; Radiocardiography, 217; Saphenous Circulation Time, 220; Peripheral Circulation, 220; Myeloscintigram, 223; Diagnosis of Carcinoma Metastatic to Liver, 226; Diagnosis of Pancreatic Insufficiency, 228.	203
Other Uses of I ¹³¹	231
Rationale, 231; Technique, 234; Interpretation, 234.	
Chapter 7	040
RADIOACTIVE PHOSPHORUS (P32)	240 240
Pharmacology	470

	~				
- (in	n	te	n	ts

Contents	IA
Diagnostic Uses of P ³² Brain Tumor Localization Localization, 241: Technique, 241; Uses, 241; Tumor Type, 242; Time Interval, 242; Age of Patient, 243.	241 241
Eye Tumor Localization	243
Determination of Viability of Tissue	216
of Circulation in Tubed Pedicle Flaps, 246. Blood Volume Determination	247
Utility, 248. Therapeutic Use of Radioactive Phosphorus	
Duration of Remissions Produced and Comparison with Other Forms of Therapy, 251.	051
Radioactive Phosphorus in Leukemia	
Radioactive Chromic Phosphate	
Method of Preparation Technique	
Chapter 8	
RADIOACTIVE GOLD	257
Pharmacology	257
Intracavitary Use	258
Indications Method of Administration	
Equipment, 259; Technique, 259. Dosage	261
Mode of Action, 261.	
Reactions to Au ¹⁹⁸ in Therapy	261
Therapeutic Results	
Direct Tumor Injection	
Injection into Prostatic Carcinoma	
Injection for Other Pelvic Cancer	266
Rationale, 266; Technique, 266; Reactions, 267; Results, 267. Treatment of Carcinoma of Lung	

Intravenous Administration of Au ¹⁹⁸ Distribution Particle Size, 268; Tissues and Organs, 268. Toxicity	268
For Treatment of Chronic Granulocytic Leukemia For Treatment of Carcinoma in Liver Intrathecal Injection Rationale, 270; Toxicity, 270; Distribution, 270; Results, 270.	269 27(
Chapter 9	
RADIOACTIVE CHROMIUM, CR ⁵¹ , COBALT, CO ⁶⁰ , AND IRON, FE ⁵⁵ AND FE ⁵⁹	273
Radioactive Chromium	273 273
Blood Volume Determinations	275
Use of Cr ⁵¹ RBC Labeling for Study of RBC Survival in Normal	280
Rationale, 280; Technique, 281; Results and Interpretation, 281. Cr ⁵¹ Red Blood Cell Half Life	282
Rationale, 283. Use of Other Radioisotopes in the Differential Diagnosis of Anemia Use of Radioactive Cobalt (Co60) Labeled Vitamin B12 to Diagnose	284
Prenicious Anemia	284
Radioactive Iron, Fe ⁵⁹	
Chapter 10	
OTHER LESS COMMONLY USED RADIOACTIVE ISOTOPES	296
Radioactive Arsenic, As ⁷⁴	•
Ast atine, At ²¹¹	298
Bationale, 299; Technique, 300; Toxicity, 300; Results, 300	
Radioactive Bromine, Br ⁸² (and Chlorine, Cl ⁸⁶)	. 301
Estimation of Total Body Chloride in Man by Radiobromide	. 30 1
Technique, 301; Results, 302; Radiation Hazard, 302. Radioactive Calcium and Strontium	. 302
Pharmacology, 302; Clinical Studies, 303. 此为试读,需要完整PDF请访问: www.ertongboo	ok.

完整PDF请访问: www.ertongbook.com

Contents.	хi
Radioactive Carbon, C ¹⁴ Radioactive Cerium Radioactive Cesium Radioactive Chlorine Radioactive Cobalt	304 304 304
For Teletherapy, 304. Radioactive Copper, Cu ⁸⁴ Radioactive Fluorine, F ¹⁸ Radioactive Gallium, Ga ⁷² Radioactive Germanium, Ge ⁷¹ Hydrogen ³ (Tritium) Method, 307; Toxicity, 308; Results, 308.	306 306 307
Radioactive Iridium, Ir ¹⁹²	309
Radioactive Lead, Pb ²¹⁰ Radioactive Manganese, Mn ⁵² Radiomercury, Hg ²⁰⁸ Radioactive Potassium, K ⁴² (and Sodium, Na ^{22, 24}) Method of K ⁴² Determination of Exchangeable Potassium, 312.	310 311 311
Radioactive Sodium	
Exchangeable Body Sodium Determination with Sodium Rubidium, Rb ⁸⁶ Radioactive Silver, Ag ¹¹⁰ , ¹¹¹ Use in Liver Function Test, 320. Radioactive Sodium, Na ²² , ²⁴	319 320
Radioactive Strontium, Sr ⁸⁹ , ⁹⁰	320 320
Radioactive Thallium, Tl ²⁰⁴ Uranium ²⁸⁸ Yttrium ^{90, 91} Distribution, 323.	322 323 323
Radioactive Zinc, Zn ⁶⁵	
BIOLOGIC EFFECTS OF RADIATION	329
Acute Penetrating Radiation Syndrome in Man	
The Chronic Radiation Syndrome	335

Contents

Radiosensitivity of Cells	336
Chapter 12	
HEALTH PHYSICS IN THE CLINICAL USE OF RADIOISOTOPES	345
Types of Radiation Hazards External Hazards Tolerance Dose, 346; Protection from External Radiation, 346. Internal Radiation Hazards Comparison of External and Internal Hazards, 359; Calculation of Radiation Exposure from Internal Emitters, 359; Tolerance Levels, 361; Protection from Internal Hazards, 361; Hazards to Experiments, 362.	346 358
Radioactive Waste Disposal	365
Twelve Unsafe Practices in Handling Radioisotopes	·369
Instrumentation in the clinical radioisotope unit \dots	371
Scalers	
Count Rate Meters External Recorder, 376; Special Rate Meters, 376.	376
Gas-Filled Detectors	
Solid Counters	380
Linear Amplifiers	385
Coincidence and Anticoincidence Circuits Ionization Chambers Determination of Operating Voltage Calibration of Counters Calibration of Scintillation Well-Type Counter with I ¹³¹ , 392 Calibration of a Thin-Window Geiger Tube for P ³² or I ¹³¹ , 393 Calibration with a Radium D and E Source, 393: Calibration by Coincidence Counting, 395.	388 390 392
Calibration of the Electroscope	. 396

	Contents	xiii
3 0	Equipment for an Isotope Unit	
Costs and	Servicing	399
Chapter		
STARTING A	AND MANAGING A CLINICAL RADIOISOTOPE UNIT	. 401
The Hosp	ician	. 101
Requ Space and Space I	esting Tests or Treatment Procedures Equipment Basic Requirements, 408; Where to Locate, 408; Delivery, Vent	. 408
Small Large Expenses Interested The Futu	ng and Plumbing, 409. I Hospitals and Private Practitioners Er Clinical Radioisotope Unit Facilities and Income Colleagues re or Physicist Training	. 413 . 415 . 416 . 419
Appendi		
APPENDIX A.	STATISTICS OF COUNTING	. 421
APPENDIX B.	DECAY TABLES OF RADIOISOTOPES	. 427
APPENDIX C.	RADIOAUTOGRAPHY OF THYROID SECTIONS	
APPENDIX D.	SHIELD THICKNESS FOR LABORATORY SOURCES .	. 432
APPENDIX E.	COMPILATIONS OF NUCLEAR DATA	. 434
Appendix F.	FORMAL CLINICAL RADIOISQTOPES TRAINING COURSES	435
Appendix G.	COMMERCIAL SUPPLIERS OF ISOTOPE-LABELE	D 43'

COMPOUNDS 437

Certain Preliminaries

WHO SHOULD TRAIN?

Who should receive training to diagnose and treat patients with radioactive isotopes? When in the physician's career should this training be

acquired? How should one acquire this training?

On February 7 and 8, 1954, a Conference on Training in the Use of Radioactive Isotopes, sponsored by Northwestern University Medical School and Argonne National Laboratory, was held in Chicago to urge certain interested persons to crystallize their thinking on these problems. Our opinions coincide with the conclusions derived from this conference.

1. The Medical Student. The medical student is required to take chemistry and physics in undergraduate school. When he enters medical school, a separate course on radiation biology or clinical techniques should not be taught, both because of curriculum difficulties and because this material can be pertinently presented in relation to material included in existing courses. On the other hand, coverage of certain fundamental topics in this field should not be left to chance. Inclusion of such topics in specific courses should be recommended by the curriculum committees.

Instruction should be given in the freshman year on properties of radiation, principles of radiation detection and measurement, and interaction of radiation and matter. This material, presented in lecture and demonstration, should be included in either 'biological chemistry or physiology, depending upon which subject is presented first in the curriculum. As students are now taught to think in chemical terms of the structure and interrelationships of matter, they should also be encouraged to regard structure and function from a physical point of view. The effect of radiation on tissue might be taught in the second year in pharmacology or pathology. Instruction on radiation hazards and protection should be included in the curriculum, possibly in the junior year, as a part of the course work either in radiology or in public health and preventive medicine. The application of radioisotopes in diagnosis and therapy would be left entirely to the clinical departments as to what and how much to include.

2. The Intern and Resident. Current opinion agrees that radioactive isotopes should be a tool for the use of the skilled clinician in diag-

nosis and treatment. It is suggested, therefore, that the period of radioisotope training might best be presented at the end of two or three
years of specialty training. The major quantity of radioisotopes used in
clinical medicine today are administered by internists, radiologists and
surgeons. Pathology may soon include training in radioisotopes as part
of clinical laboratory pathology. We have made rotation through the
Clinical Radioisotope Unit available to all residents in University Hospital to give them a broader knowledge of modern medicine, to prepare
them to be relatively well informed members of their community in
respect to questions of atomic warfare, civil defense programs and hazards which might arise or be alleged to arise from industrial use of
artificial radioactivity, and, most important, to equip them for the possibility that they may be needed to head a clinical radioisotope laboratory in a university hospital or a leading hospital in a large community.

3. The Postgraduate in Private Practice. Physicians in private practice generally request training in a clinical isotope unit for one of two reasons. The specialist usually wants to learn how to handle a certain radioactive isotope in his practice as a specialized tool in his daily work. The general practitioner, on the other hand, desires information on the indications for the use of all clinical radioisotopes in his patients, how to prepare his patients for these isotopes, and how to handle his patients after the isotopes have been administered to his patients. This latter type of instruction should be available to any physician to keep him informed on the utility of this new development in modern medicine.

RECOMMENDATIONS AND REQUIREMENTS BY THE ATOMIC ENERGY COMMISSION

The Atomic Energy Commission has made several specific recommendations and requirements to be met by the person who wishes to use radioisotopes in clinical medicine. These statements are reproduced in toto as follows.

THE MEDICAL USE OF RADIOISOTOPES

Recommendations and Requirements by the Atomic Energy Commission

I. INTRODUCTION

The present procedures of the Atomic Energy Commission for the allocation of radioisotopes for medical research, diagnosis and therapy are set forth in this announcement. The recommendations for minimum clinical radioisotope training and experience for use of radioisotopes in human subjects have been established in advisement with the Subcommittee on Human Applications of the Atomic Energy Commission's General Advisory Committee on Isotope Distribution.

These recommendations are designed to provide guidelines for physicians and typify the nature rather than the precise extent of the desirable clinical radioisotope experience. For special situa-

tions, other experience may serve in lieu of the particular recommendations set forth in this announcement.

II. ADMINISTRATIVE PROCEDURE FOR RADIOISOTOPE PROCUREMENT

A. APPLICATION

A medical institution or a physician in an individual medical practice desiring to obtain radioisotopes, forwards to the Isotopes Extension*, Form AEC-313, "Application for Byproduct Material Licenses," and Supplement A, (Form AEC-313-a). If the radioisotopes are to be obtained as sealed sources (such as Cobalt 60 for teletherapy units or Strontium 90 for medical eye applicator), the applicant should complete the basic form (Form AEC-313) and Supplement B (Form AEC-313-b). These forms should be completed in accordance with the instructions attached thereto. Applications for use of radioisotopes in human subjects in an INSTITUTIONAL MEDICAL PROGRAM should be supported by the special information described under Section III, Page 4, Section IV, Page 6 and Section V, Page 7 of this announcement. For such use by a physician in his INDIVIDUAL MEDICAL PRACTICE the application should be supported by the special information described under Section III, Page 4, Section VI, Page 11 and Section VII, Page 13.

In considering such applications, the Atomic Energy Commission is concerned primarily with matters of radiological health safety. Toward this end the Commission seeks to determine if the applicant has equipment and facilities appropriate to the proposed use and whether the physician is trained in basic principles of radioactivity and has specific experience in the use of radioisotopes in the clinical situations being proposed. The information is indicated by the applicant on his application form and the supple-

mentary sheets attached thereto.

B. LICENSE

Upon favorable review of the application (See NOTE below), Form AEC-374, "Byproduct Material License," is issued. This license permits the holder to procure radioisotopes in accordance with the conditions stated on the application and license forms and in Title 10 of the Code of Federal Regulations.

NOTE. The Isotopes Extension normally reviews applications proposing new or nonroutine medical uses of radioisotopes in collaboration with the Advisory Subcommittee on Human Applications.

This review usually requires four weeks for completion.

C. TYPES OF CLINICAL RADIOISOTOPE PROGRAMS

The recommendations and requirements established by the Atomic Energy Commission for the clinical use of radioisotopes are designed to provide for two types of medical radioisotope programs. These are defined as follows:

1. <u>Institutional Medical Radioisotope Program</u>

Clinical radioisotope programs established by a medical institution and carried out under the guidance of a medical isotopes committee (See recommendations for membership and duties of a medical isotopes committee in Section IV, A. Page 6) are designated as "Institutional use." Licenses for institutional use require that the physician(s) named on the license form supervise the conduct of

* Allocations Branch, Isotopes Extension, Division of Civilian Application, U.S.

Atomic Energy Commission, P. O. Box E, Oak Ridge, Tennessee.