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TECHNICAL PROGRESS REVIEW

ISOTOPES AND RADIATION TECHNOLOGY

VOLUME 1, NO. 1

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ISOTOPES DEVELOPMENT —8

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TECHNICAL PROGRESS REVIEW
ISOTOPES AND RADIATION TECHNOLOGY
VOLUME 1, NO. 1

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Oak Ridge, Tennessee
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A new Quarterly Technical Progress Review, "Isotopes and Radiation Technology," was approved by the U. S. Atomic Energy Commission in the summer of 1962 and its preparation was assigned to ORNL as a part of the general program of the Isotopes Development Center. The assembly of a staff began almost immediately. By early fall, under the auspices of the AEC's Divisions of Technical Information and Isotopes Development, the preliminary drafts which are now the first issue of this periodical were being developed. "Isotopes and Radiation Technology" is the fifth of a series of Commission-sponsored quarterly technical progress reviews which cover broadly the field of nuclear science and technology. The others are Nuclear Safety, Power Reactor Technology, Reactor Materials, and Reactor Fuel Processing.

This new review will emphasize the reporting of DID-sponsored research and development. By publicizing present applications, it can help to expand the utilization of isotopes; and by helping the Commission and contractors to keep abreast of related activities within the AEC complex it can help prevent overlap of research effort and can aid in predicting isotope usage so as to point the direction for future isotope production efforts.

"Isotopes and Radiation Technology" is a technical progress <u>review</u>. Even guest articles will be of a review nature.\* The editors and the AEC have no intention of duplicating the present efforts of commercial publishers of abstract journals and periodicals covering original research, and overlapping will be avoided. Books, monographs, and bibliographies that are considered pertinent to the areas routinely covered will be noted and occasionally reviewed.

<sup>\*</sup>e.g., see report by G. H. Cartledge on technetium in this issue, p. 3.

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# TECHNICAL PROGRESS REVIEW Isotopes and Radiation Technology

Edit	orial	iii
Feat	ure Article:	
	INHIBITION OF CORROSION BY THE PERTECHNETATE ION	1
	References	12
I.	PRINCIPLES OF ISOTOPE TECHNOLOGY	16
	Introduction	17
	Isotope Production	19
	Isotope Utilization	20
	History and Chronology	22
	References	28
	STATUS OF THE UNITED STATES PROGRAM ON ISOTOPES AND RADIATION DEVELOPMENT	39
II.	ISOTOPE PRODUCTION AND DEVELOPMENT	115
	Fission Products	116
	Stable Isotopes	140
	References	174
III.	APPLICATIONS TECHNOLOGY	
	Analysis	176
	Radiometric	177
	References	206
	Activation Analysis	209
	References	240
	Radioisotope Sources for Radiography	245
	References	253

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vii

	References	366
IX.	AVAILABLE MATERIALS AND SERVICES	368
х.	BOOK REVIEWS	372

VII. APPLICATIONS IN MEDICINE

The literature on medical uses of radioisotopes is briefly reviewed here.\* Since developments in this rapidly expanding field have been summarized in various well-documented publications, learlier original literature was not extensively consulted. Only a few of the many published papers are listed here, and emphasis is on recent, rather than older, work. Since many reports are preliminary and opinions on the effectiveness of some of the isotopes differ widely, in future issues of this quarterly various viewpoints on the medical use of radioisotopes will be presented, and some of the less well-known isotopes, both old and new, will be reviewed. Readers with publications in this field are invited to send in reprints to ensure coverage.

The medical use of artificial radioactive isotopes began in the late thirties. The increasing interest in them is indicated by the numerous conferences, journals, bibliographies, and reviews partly or entirely devoted to this field. Radioisotopes are used in both diagnosis and therapy (Table VII-17), in the former in extremely small amounts; in the latter, chiefly in malignant or malignancy-related conditions. Safety in their use is ensured (1) by requiring the licensing of physicians who use them -- and then only after special training; (2) by standardization of handling techniques and adoption of special safety regulations by hospitals. Early fears of long-term adverse \*The editors of this review gratefully acknowledge the many helpful suggestions made by W. G. Myers, M.D., and M. Brucer, M.D., who reviewed a rough draft of the manuscript. Some of their suggestions that could

not be readily incorporated will be used in future issues.

Table VII-1. Radioisotopes Used in Diagnosis and Therapy\*

	Diagnosis	
Test	Measurement	Usefulness
1 <sup>151</sup> studies of thyroid function; NaI, usually oral	Percentage of dose concentrated in thyroid, percentage of dose excreted in urine; plasma free and bound $1^151$	Valuable index of thyroid function; drugs cause inter- ference
1131 for diagnosis of thyroid carcinoma; oral	Scanning and external procedures	Relatively little value while normal thyroid gland present; highly useful in locating well differentiated metastatic lesions after thyroidectomy
Fe <sup>59</sup> iron metabolism; usually intravenous iron citrate; oral doses used to study absorption	Serial plasma assays for several hours; red cell assays for about 10 days; external counts over spleen, liver, sacrum; blood and fecal assays after oral dose	Helps explain mechanism of some anemias, importance of impaired erythropoiesis versus hemolysis, oral test indicates absorption defects.
Cr51 red cell survival; red cells labeled with Cr <sup>51</sup> in vitro and injected intravenously	Serial assays of blood (red cells) for 2 or 3 weeks; external counts over spleen, liver, heart; fecal assays for gastrointestinal bleeding	Indicates rate of hemolysis and to some extent the role of the spleen in hemolysis
Co <sup>60</sup> -vitamin B <sub>12</sub> ; oral	Fecal and urinary excretion; sometimes external counts over liver	Shows fundamental defect of pernicious anemia even if patient is not in hematologic relapse
1 <sup>131</sup> -labeled rose bengal; intra-venous	Rates of removal from bloodstream and concentration in liver	An index of liver function and biliary patency

Table VII-1. Contd.		
Test	Measurement	Usefulness
1 <sup>131</sup> -labeled fat; oral	Blood, urinary, and fecal levels	Information on intestinal absorption, pancreatic function
Cr51, P32 red cell mass; [131] albumin-plasma volume, Na <sup>24</sup> , K <sup>42</sup> for Na and K spaces	All based on dilution principle; solution of isotope (or suspension of red cells tagged in vitro) given intravenously and after mixing in the body, blood concentration determined	These tests give data on red cell and plasma volumes, fluid balance and electrolyte status; may be helpful in presurgical evaluation, study of metabolic and endocrine disorders.
	Therapy	
	Patients Treated	Usefulness
Il31 as NaI; usually oral	Uncomplicated Graves' disease patients over 40; complicated Graves' disease any age	General results excellent; multiple doses needed in some cases; significant incidence of early or delayed myxedema
	Toxic nodular goiter in selected patients	Eventual control of toxicity in most cases; may require repeated doses; concern over malignancy already present at onset, plus desire for rapid result, may recommend surgery for patients who can be well prepared

$\operatorname{Treated}$	
Patients	

Thyroid carcinoma in selected patients Certain patients with angina pectoris or intractable congestive heart failure

Chronic granulocytic leukemia

P<sup>52</sup> as soluble phosphate; oral

or intravenous

Chronic lymphocytic leukemia

Polycythemia vera

Lymphosarcoma, multiple myeloma, diffuse bone metastases

Acute leukemia, Hodgkin's disease

Intracavitary in patients with effusions and ascites caused by malignant tumors

Colloidal Aul98, P32, Y90

Intravenous for granulocytic leukemia and diffuse lymphoma of liver

# Usefulness

For treatment of relatively small group of patients with nonresectable lesions showing ability to concentrate the isotope

Clinical improvement in significant percentage

Generally good results

Useful, often in conjunction with local irradiation

Good results in patients not suitable for phlebotomy alone

Occasionally some value

Rarely useful

Symptomatic value; partially supplanted by nitrogen mustards

Very limited value compared with other forms of therapy

Table VII-1. Contd.

Usefulness	Believed useful if properly used in carefully selected cases	Useful in skilled hands for properly selected patients	Highly useful and practical
Patients Treated	Direct injection into tumor in certain cases of prostate carcinoma; attempt to radiate lymphatic channels	Certain localized tumors, especially carcinoma of cervix, head, and neck	For all lesions suitable for external radiation therapy
		Co60, Cs137, radium-226, radon-222, Sr90for implantation and brachytherapy	$c_{0}60$ , $c_{s}^{137}$ for teletherapy

\*Reprinted with minor style changes with the kind permission of publishers, from G. A. Andrews, Medical students and radioisotopes, The New Physician, 9(12):25-7 (1960).

effects on patients have been tempered by the results of experience. According to Lawrence, 9 for example, cancer has not once been reported as a result of radioactive iodine therapy of thousands of hyperthyroid patients; and in 25 years, the survival rate of more than 500 polycythemia vera patients treated with P<sup>32</sup> has been about the same as that of patients with diabetes or pernicious anemia treated by standard methods. There is wide usage of radioactive iodine for toxic goiter in patients over 45, of P<sup>32</sup> for polycythemia vera, and of various isotopes for pituitary or thyroid ablations to relieve other conditions. 1

Among the radioisotopes now used in internal medicine in the largest amounts are  $I^{131}$ ,  $P^{32}$ ,  $Au^{198}$ ,  $Cr^{51}$ ,  $Fe^{59}$ , and  $Hg^{203}$ , but many others are used in smaller amounts; large quantities of  $\mathrm{Co}^{60}$  and  $\mathrm{Cs}^{137}$ are used in teletherapy units. Many radiofsotopes are being studied for potential use, and useful ones are introduced into the field as new production methods are developed and costs are lowered. Criteria for suitable isotopes include high specific activity, suitable half-life, physiological compatibility with body processes, and the ability to at least partially localize in the tissue being examined or treated. Radioiodine, for example, while not entirely thyroid-specific, is an obvious choice for thyroid conditions, as is calcium for bone. Specificity may sometimes be obtained by incorporation in compounds (e.g., radiocobalt in vitamin B12, which localizes in the liver). An isotope may be localized physically as a wire, as a colloid, or as the recently introduced ceramic beads. 10 Both the physiological and biological halflife must be considered, and selection may be made on the basis of the need for particular radiation characteristics -- from a soft beta all

the way to a hard gamma radiation. In earlier work, isotopes with a short physical half-life were found inconvenient to use, ll but they are finding application now with development of new instrumentation and sources of supply.

#### Iodine

 $I^{131}$  (8.1 d. 0.33- and 0.61-Mev  $\beta$ 's, 0.36- and 0.64-Mev  $\gamma$ 's). Iodine-131 is most commonly used for diagnosis of thyroid disorders. Other uses include incorporation in human serum albumin for determination of the total circulating plasma volume, placentography, 12 estimation of mitral valve size. 13 and detection of right-to-left cardiac shunts. There has been partial success in localizing brain tumors with  $I^{131}$ -labeled compounds, and  $I^{131}$ -labeled methyl iodide has been used for detecting left-to-right cardiac shunts. 14 Rose Bengal containing  $I^{131}$  is useful for liver function tests and for blood flow tests  $I^{15}$  and has been suggested for detecting liver metastases that are not obvious from surgery. 16 Recent reports indicate the Rose Bengal-I hepatoscan to be a satisfactory technique. 17 Other 1 31-labeled compounds reported useful are Hippuran for renal tests 18 and for determining cerebral circulation time; 19 Diodrast for evaluating venous diseases of the lower extremities; 20 sulfobromophthalein for liver function tests; 21 and compounds for studying fat metabolism. 22 An I labeled compound demonstrated 200-ml pericardial effusions in a simulated system, 23 and an accurate and reproducible method for insulin assay with I has been reported. 24

In therapy, I<sup>131</sup> has been used for nontoxic<sup>25</sup> as well as toxic goiter.<sup>1</sup> It is used as a palliative in thyroid carcinoma, and an

I<sup>131</sup>-tagged fat-soluble medium, which accumulates in regional lymph nodes, is potentially useful for irradiation of neoplasms.<sup>26</sup> Thyroid ablation by I<sup>131</sup> in more than 1000 patients improved angina pectoris in 75% and congestive heart failure in 60% of those afflicted.<sup>27</sup> Fortunately, gamma irradiation of the ovaries from urinary excretion of administered I<sup>131</sup> is considered negligible.<sup>28</sup>

 $I^{125}$  (57.4 d, no  $\beta$ , 0.035-Mev  $\gamma$ ). Iodine-125, which is easily produced by neutron bombardment of Xe<sup>124</sup>, is becoming increasingly valuable in diagnosis. Irradiation of a patient by it is about half 29 that from I<sup>131</sup>, assuming that the biological half time is equal to the physical half-life. Good thyroid and liver scans have been made with it, 30 and Hippuran labeled with I<sup>125</sup> has been used for renograms. 31 As a tag for serum albumin and oleic acid and in the salt  $\operatorname{NaI}^{125}$ , it has given good results in experimental blood volume, absorption, and thyroid tests, 32, and has been suggested for indicating the extent of arteriosclerosis. 33 In a promising new technique for lipid digestion and absorption studies, a compound labeled with I and another labeled with I<sup>125</sup> are administered simultaneously.<sup>34</sup> The promising use of I<sup>125</sup> in radiography<sup>35</sup> is indicated by a photograph of the bones of a hand made with this isotope (Fig. R3, R4). Exposure to both patient and technician is low because of the ease of shielding against the 27.2 and 35.4-kev photons. Portable x-ray equipment is being developed with 1<sup>125</sup> sources. 36

 $\underline{I^{132}}$  (2.3 hr, 1.5- to 2.1-Mev  $\beta$ 's, 0.53- to 0.95-Mev  $\gamma$ 's). The use of  $\underline{I^{132}}$ , which is easily recovered from a tellurium source, would decrease patient exposure to 1-2% of that from  $\underline{I^{131}}$  and would

<sup>\*</sup>See also p..245.

allow repeated administration.<sup>37</sup> Thyroid function after  $I^{131}$  therapy may be determined by observing the absorption of  $I^{132}$ , whose higher energy gammas are easily detected in the presence of residual  $I^{131}$ .<sup>38</sup>

 $\underline{I^{123}}$  (13.3 hr, no  $\beta$ , 0.16-Mev  $\gamma$ ). With  $\underline{I^{123}}$ , exposure of a patient would be only 1-5% of that from  $\underline{I^{131}}$ . Experiments on preparation of curie amounts are projected.<sup>39</sup>

 $1^{124}$  (4 d, 1.53- and 2.13-Mev  $\beta$ 's, 0.6- and 1.72-Mev  $\gamma$ 's). Indine-124 has been suggested for greater uniformity of dose in thyroid carcinoma.

# Phosphorus

 $P^{32}$  (14.3 d, 1.71-Mev  $\beta$ , no  $\gamma$ ). The largest use of  $P^{32}$  is in treatment of polycythemia vera and chronic leukemia. It has not been very useful in acute leukemia. Blotting paper soaked with radioactive sodium phosphate was used in early applications for skin lesions, and activated red phosphorus incorporated in polyethylene in later work. Metastatic bone cancer has been treated with  $P^{32}$  as the phosphate. 41

Colloidal chromic phosphate-P<sup>32</sup> and Au<sup>198</sup>, as well as nitrogen mustard, for control of malignant intracavity effusions are compared in a recent review<sup>42</sup> of 222 cases treated with P<sup>32</sup> and 2110 with Au<sup>198</sup>. Colloidal chromic phosphate-P<sup>32</sup> has been used for palliation of cancer, and colloidal zirconyl phosphate-P<sup>32</sup> administered intravenously to more than 250 patients with chronic leukemia and lymphomas is reported to have resulted in prolonged remissions and long treatment-free periods.<sup>43</sup>

In diagnostic applications, an in vivo test for human spermatogenesis using sodium phosphate- $P^{32}$  has been described, <sup>44</sup> and the ability of newly proliferating cells to concentrate  $P^{32}$  has been used to

differentiate benign from malignant ocular lesions. 45 Gold

Au<sup>198</sup> (2.7 d, 0.97-Mev  $\beta$ , 0.41-Mev  $\gamma$ ). Gold-198 is most commonly used as a colloid, which has 0.003- to 0.007- $\mu$ -dia. particles. When this material is administered to a patient and a scan is performed, normal liver shows an even gradation of activity, but lesions result in activity concentrations. Diagnostic uses include determination of liver blood flow, and in regions where the parasitic Taenia Echinococcus granulosus occurs it has been used for localizing hydatid liver cysts and for following up healing after drug administration or surgery.  $^{47}$ 

Colloidal gold has been used in treating both primary carcinoma and metastases. Transportation to the lymph nodes is hastened if the particles are coated with silver, but radioactive colloidal gold has been injected directly into the lymphatics. Other uses are in treating leukemia and in controlling malignancy-induced intracavity fluids.

Interstitial implants of Au<sup>198</sup> seeds have been used for hypophysectomy and for palliation of metastatic carcinoma.<sup>53</sup> The hypophysis implantation requires about 30 min. Problems in surgical and radiotherapeutic pituitary ablation are discussed in recent reviews.<sup>54</sup> Chromfum

 ${\rm Cr}^{51}$  (28 d, no  ${\rm B}$ , 0.32-MeV  ${\rm \gamma}$ ). Hexapositive  ${\rm Cr}^{51}$ , in  ${\rm Na_2Cr}^{51}{\rm O_4}$ , attaches to the red cells and may be used to determine such quantities as red cell survival time. A new instrument for accurate blood volume measurement uses  ${\rm Cr}^{51}$ -tagged red cells or  ${\rm I}^{131}$ -albumin. Such cells have been suggested for determination of cardiac output and for placental

localization (no activity is transferred to the fetal circulation), and are the basis for a simple method for quantitation of menstrual blood loss<sup>56</sup> and for spleen studies.<sup>57</sup> Tripositive chromium (as  $\text{Cr}^{51}\text{Cl}_3$ ) in solution attaches to plasma proteins and is useful for determining plasma volume. Serum albumin labeled with  $\text{Cr}^{51}$  is easily prepared and has been introduced for demonstrating gastrointestinal protein loss.<sup>58</sup>

Implants of Cr<sup>51</sup> have been recommended for cancer therapy.<sup>59</sup>
Iron

 $\rm Fe^{59}$  (45 d, 0.26- and 0.46-Mev β, 1.1- and 1.3-Mev γ). Detailed methods for  $\rm Fe^{59}$  tests are described in a recent brochure. As the chloride, citrate, or sulfate,  $\rm Fe^{59}$  has been used for iron absorption and utilization studies. It has also been used for determining red cell survival time, but the necessity of labeling the cells in vitro makes it less convenient than  $\rm Cr^{51}$ . The advantages and disadvantages of  $\rm Fe^{59}$  and  $\rm Cr^{51}$  are discussed in a recent review.  $\rm ^{62}$ 

 ${\rm Fe}^{55}$  (2.94 y, 0.0059-Mev x ray following electron capture). Comparison of the percentage utilization of intravenously administered  ${\rm Fe}^{59}$  to that of oral  ${\rm Fe}^{55}$  has been used for iron absorption studies. Mercury

Hg  $^{203}$  (47 d, 0.21-Mev  $\beta$ , 0.28-Mev  $\gamma$ ). Besides the use of mercury isotopes in tracing kidney drugs,  $^{64}$  Hg  $^{203}$  has been used for localizing brain tumors and for determining cerebral circulation time. Neohydrin and Chlormerodrin labeled with Hg  $^{203}$  have been used for kidney examinations.  $^{66}$ 

#### Boron

B<sup>10</sup>. When the stable isotope B<sup>10</sup> is deposited in brain tumor tissue and activated by neutron bombardment, the released alpha radiation is effective for control and palliation.<sup>67</sup> By this treatment, cells of transplantable sarcomas in mice have been killed and cancer has disappeared in 2-4 weeks.<sup>68</sup> This method of treatment is not new, but basic techniques are still being investigated.<sup>69</sup>

# Fluorine

 $\frac{F^{18}}{(1.87 \text{ hr}, 0.65\text{-Mev positron emission})}$ . Fluorine-18 has been used as a replacement for Ca<sup>47</sup> and Sr<sup>85</sup> in bone scanning.<sup>70</sup> It has also found application in thyroid studies<sup>71</sup> and in locating brain tumors.<sup>72</sup>

# Selenium

Krypton

 $\rm Se^{75}$  (120 d, no  $\beta$ , 0.14- to 0.28-Mev  $\gamma$ 's). The high specificity of  $\rm Se^{75}$ -selenomethionine for the pancreas suggests that this compound might be useful in early detection of pancreas carcinoma. 73

Kr<sup>85</sup> (10.6 y, 0.67-Mev β). Because of its low solubility, injected or inhaled Kr<sup>85</sup> is useful in determining cerebral and coronary blood flow and cardiac and pulmonary shunts.<sup>74</sup> For example, if a right-to-left cardiac shunt is present, activity in the arterial system rises rapidly when Kr<sup>85</sup> is given intravenously.<sup>75</sup> If a left-to-right shunt is present, the Kr<sup>85</sup> concentration in the pulmonary artery reaches a value 20 times that in a systemic artery following inhalation of this gas.<sup>76</sup> A simple technique for studying pulmonary function with Kr<sup>85</sup> has been devised,<sup>77</sup> and a new technique for rapid and accurate

measurements of serial heart blood flow was reported recently. 78
Strontium and Yttrium

 $Sr^{90}$  (28 y, 0.54-Mev  $\beta$ , 2.27-Mev  $\beta$  from  $Y^{90}$  daughter, no  $\gamma$ ). In an external source containing  $Sr^{90}$  designed for use in ophthalmology, the  $Sr^{90}$  beta particles are filtered out so that only the  $Y^{90}$  betas actually reach the lesion. The lens is not irradiated.

"Radiation curettage," destruction of the uterine mucosa by radiation from a  $\rm Sr^{90}$  source placed in the uterine cavity, has been recommended for control of endometrial bleeding when no malignancy is indicated. <sup>79</sup>

 $\mathrm{Sr}^{85}$  (65 d, no  $\beta$ , 0.5-Mev  $\gamma$ ). Strontium-85 has been used for localizing bone tumors <sup>80</sup> and for studying bone metabolism. Because of its shorter half-life, irradiation of the body is less than with  $\mathrm{Sr}^{90}$ . Its gamma radiation can be detected externally.

 $\frac{\text{Sr}^{87\text{m}}}{\text{(2.8 hr, 0.39-Mev }\gamma)}$ . This isotope, which can be obtained quickly, as needed, from 80-hr  $\chi^{87}$ , has been mentioned  $^{81}$  for possible use in obtaining information available in a short time; for example, circulation time, cardiac output, and various clearances, and in radiography.

 $\underline{Y}^{90}$  (64.5 hr, 2.27-Mev  $\beta$ ). Injected ceramic  $\underline{T}^{90}$  particles of  $\sim 60$   $\mu$  dia., which are trapped in capillaries of the lung, liver, and other organs, have been used for cancer therapy. Thirty-five patients with Parkinson's disease were improved by brain implants of ceramic  $\underline{T}^{90}$  beads. Carrier-free  $\underline{T}^{90}$  may be eluted with citrate from a cation exchange resin preparation of  $\underline{Sr}^{90}$ . It may be infiltrated as a solution, implanted in a plastic filament, or applied in a bag or absorbed in

blotting paper.85

Because it is a pure beta emitter, colloidal Y<sup>90</sup> results in less exposure of hospital personnel than Au<sup>198</sup>, and it has been used for control of malignancy-induced effusions.<sup>86</sup> Complete pituitary destruction has been reported as a result of packing the fossa, following surgery, with powdered Y<sup>90</sup> dispersed in wax.<sup>87</sup> For superficial radiation therapy, Y<sup>90</sup> has been incorporated in small plates, which can be selected to cover an irregular area.<sup>88</sup> Transperineal injection of Y<sup>90</sup> has been used for prostatic cancer.<sup>89</sup> Intravenous Y<sup>90</sup> gave good results in eight polycythemia vera patients.<sup>90</sup>

# Calcium

 $\underline{\text{Ca}^{45}}$  (164 d, 0.25-Mev  $\beta$ , no  $\gamma$ ). Calcium-45 has been used for studying calcium metabolism, Paget's disease of bone, and hypoparathyroidism. 91

 $\text{Ca}^{47}$  (4.7 d, 0.66-and 1.94-Mev  $\beta$ 's, 0.48- to 1.3-Mev  $\gamma$ 's; accompanied by 3.5 d Sc  $^{47}$  daughter with 0.46- and 0.62-Mev  $\beta$ 's and 0.175- Mev  $\gamma$ ). Possible medical uses of the now-available  $\text{Ca}^{47}$ , which is of interest because of its high gamma energy, are being extensively investigated.  $^{92}$ 

# CobaIt

 $^{60}$  (5.3 y, 0.31-Mev  $\beta$ , 1.17- and 1.33-Mev  $\gamma$ ). By far the largest use of  $^{60}$  is in teletherapy units,  $^{1b}$ ,  $^{6}$ ,  $^{93}$  descriptions of which may be obtained from various equipment manufacturers (see also book review section of this issue). Such units were widely distributed by 1959 (Table VI-2).  $^{94}$  In 1962 they were reported to number 1400, of which 550 are in the U.S.A. and Latin America.  $^{95}$  They are reported

Footnotes to Table VI-2.

a Reprinted, with minor style changes, from reference 94.

bThese figures represent the number of licenses issued as of August 1, 1959; they do not include such teletherapy installations as those at Oak Ridge Institute of Nuclear Studies and Argonne Cancer Research Hospital, for which no license was required.

<sup>C</sup>Estimated, mainly low curie units.

d\_Including only countries for which information is available.

Table VI-2. Radioisotope Teletherapy Source Distribution

T.o.o.		Unit		\$C [		Unit	
	$_{ m Ir}^{ m 192}$	$c_s^{137}$	09 <sup>ဝ၁</sup>	הסכם כדי סוו	$^{ m 192}$	$c_{\rm s}^{137}$	09 ၀၁
Africa Algeria Union of South Africa	1 1	1 1	ਜਿਜ	Tebanon Philippines Thailand	1 1 1	1 1 1	<b>н</b> #н
America Argentina Brazil Canada	1 1 1	1 1 0	~≈4	Europe Austria Belgium Denmark	1 1 1	1 1 1	0 0 0
Chile Cuba	1 1 ·	·1 1 ·	a a .	ļ	1 1	ı <del>d</del>	40,
El Salvador Mexico Peru	1 1 1	1 1 1	12 1	Germany (Fed. Republic) Greece Hungary		1 1 1	4 ろエ
Puerto Rico United States Umbanav	1 1 1	1 <sup>9</sup> 1	1 01 44 0	Italy Monaco	1)1	, , ,	19, 14
Venezuela Asia		t	110.	Norway Portugal Spain		H 1 1	+ 1 4 7V E
burma Ceylon China (Taiwan) Hong Kong	1 1 1 1	1 1 1 1	<b>ਜਜਜ</b> ਜ	Sweden Switzerland United Kingdom Yugoslavia	וולו	ιιωι	てどれり
India Indonesia Iran Israel Japan	1 1 1 1 1	1 1 1 1 1	500° 500°	Oceania Australia New Zealand Total <sup>d</sup>	।। न	18	4 5 689

See following page for footnotes.

to be in use in Communist China, and have been referred to as the most important product of Atomic Energy of Canada Ltd.  $^{97}$  A nonroutine use of  $\mathrm{Co}^{60}$  teletherapy is a possible new treatment for leukemia in which blood, circulated outside the body through a tube, is exposed to  $\mathrm{Co}^{60}$  radiation.  $^{98}$ 

Cobalt-60 needles have been implanted in tumors, and seeds enclosed in nylon tubing have been used in interstitial therapy and in wounds after surgery for a malignant condition. Spheres of Co<sup>60</sup>, gold-plated to prevent dusting, may be placed in hollow organs such as the uterus or bladder. Cobalt-60 may be incorporated in plastic and molded to fit irregular areas such as the oral cavity.

Vitamin  $B_{12}$  labeled with  ${\rm Co}^{60}$  has been used in diagnosing pernicious anemia  $^{102}$  and localizing liver tumors.  $^{103}$ 

Other Cobalt Isotopes.  $\text{Co}^{56}$  (77.3 d, 0.44- and 1.46-Mev  $\beta$ , 0.85-1.75-Mev  $\gamma$ );  $\text{Co}^{57}$  (270 d, no  $\beta$ , 0.014- and 0.12-Mev  $\gamma$ );  $\text{Co}^{58}$  (71 d, 0.48-Mev  $\beta$ , 0.81-Mev  $\gamma$ ). Various other cobalt isotopes have been used to label vitamin  $\text{B}_{12}$ . Cobalt-57 is reported to give the lowest irradiation of a patient. Labeled vitamin  $\text{B}_{12}$  ( $\text{Co}^{57}$ ) has been shown to concentrate in the parathyroid in dogs of and is therefore potentially useful in parathyroid conditions.

# Cesium

 ${\rm Cs}^{137}$  (30 y, 0.51- and 1.18-Mev  $\beta$ , 0.662-Mev  $\gamma$  from daughter  ${\rm Ba}^{137}$ ). This isotope, which is recovered as a by-product from the processing of nuclear reactor fuel, is a relatively new isotope, but many  ${\rm Cs}^{137}$  teletherapy units are in use (Table VII-2). The long half-life and low cost of the activity are the principal advantages over  ${\rm Co}^{60}$ . Typical

units have been described elsewhere. lc,93a,106 (See also book review section of this issue.)

# Iridium

 ${\rm Ir}^{192}$  (74 d, 0.67-Mev  $\beta$ , 0.30- to 0.47-Mev  $\gamma$ 's). Radioiridium teletherapy is an old idea. <sup>107</sup> A new 100-curie  ${\rm Ir}^{192}$  irradiation unit weighs only 38 lb; a 10-curie source weighing 28 lb is also available. <sup>108</sup> A unit with interchangeable ends of different sizes and shapes, for interorificial therapy, has been described recently. <sup>109</sup>

The half-life of Ir<sup>192</sup> is about 5% that of Co<sup>60</sup>, and its less penetrating gamma rays allow greater local absorption and necessitate less protection for hospital personnel. Internally, it has been used as seeds embedded in nylon ribbon or as needles. Experimentally, an Ir<sup>192</sup>-labeled resin suspension has been used with external counting for determining gastric emptying rates.

# Arsenic and Copper

As  $^{74}$  and Cu  $^{64}$  (17.4 d and 12.8 hr, respectively; positron emitters). These isotopes have been used for localizing brain tumors. 112

 $_{\rm As}^{76}$  (26.5 hr) and  $_{\rm As}^{72}$  (26 hr). Arsenic-76 has been used for brain tumor localization, as has  $_{\rm As}^{72}$ , which yields nearly three times as many positrons per average disintegration  $_{\rm As}^{113}$  as  $_{\rm As}^{74}$ .

#### Gallium

Gallium-72 has been used to localize malignant bone tumors by external measurements, but the short half-life results in a concentration of this chemically toxic material which may perhaps be higher than is warranted by the radiation delivered.  $^{114}$ 

 $\underline{\text{Ga}}^{68}$  (68 min, 1.9-Mev  $\beta$ , 1.1-Mev  $\gamma$ ). Gallium-68 has been suggested for such applications as brain tumor localization by annihiscopy. A source from which  $\underline{\text{Ga}}^{68}$  can be easily and rapidly obtained has been designed. 115

#### Palladium

 $Pd^{109}$  (13.5 hr, 1.03-Mev  $\beta$ ) and  $Pd^{103}$  (17 d). A mixture of these two isotopes has been used for interstitial irradiation. 116

# Tantalum

Ta<sup>182</sup> (115 d, 0.36- to 0.51-Mev  $\beta$ 's, 1.12- and 1.22 Mev  $\gamma$ 's). Tantalum-182 wires have been used as tissue implants in control of malignant conditions. 117

# Ruthenium-Rhodium

Intragastric beta irradiation with Ru<sup>106</sup>-Rh<sup>106</sup> has been tried. Lutetium

Lu<sup>177</sup> (175 d; also 6.8 d, 0.50-Mev  $\beta$ , 0.18- and 0.38-Mev  $\gamma$ ). Lutetium-177 has been suggested for infiltration to lymph nodes involved in malignant disease. 119

# Sulfur

 $s^{35}$  (87 d, 0.167-Mev  $\beta$ , no  $\gamma$ ). Sulfur-35 has been used for determining extracellular water and as a therapeutic agent in chondrosarcoma and mycosis fungoides. Labeling of Congo red dye with  $s^{35}$  decreases the amount of dye needed for diagnosis of amyloidosis, eliminating the side effects that occur in some patients. Methionine labeled with  $s^{35}$  has been suggested for use in detecting pancreas tumors.  $s^{122}$ 

 $Na^{22}$  (2.6 y, 0.54-Mev  $\beta$ , 1.3-Mev  $\gamma$ ) and  $Na^{24}$  (15 hr, 1.4-Mev  $\beta$ ,

Sodium

2.7- and 1.4-Mev  $\gamma$ 's). These isotopes have been used for measuring extracellular water, sodium space, and exchangeable sodium; for determining when skin grafts are complete; and for cancer therapy. 1,123 Because of its shorter half-life, Na<sup>24</sup> has advantages for some uses. Potassium

 $K^{42}$  (12.4 hr, 3.6-Mev β, 1.5-Mev γ) and  $K^{43}$  (22 hr, 0.3- and 0.27-Mev β's, 0.37- to 0.62-Mev γ's). These isotopes have been used to localize brain and breast tumors, measure exchangeable potassium, and study potassium uptake by red blood cells. 1,124 Less shielding is re-

#### Rubidium

 ${
m Rb}^{86}$  (19 d, 1.78- and 0.72-Mev  ${
m B's}$ , 1.1-Mev  ${
m \gamma}$ ). Since rubidium acts similarly to potassium and has a more convenient half-life than  ${
m K}^{42}$ , it has been suggested for use in determining exchangeable potassium. 124c However, it must be remembered that  ${
m Rb}^{86}$  is not a potassium isotope. Myocardial scanning tests have been made with  ${
m Rb}^{86}$  in dogs. 125 Bromine and Chlorine

 $Br^{82}$  (35.7 hr, 0.44-Mev  $\beta$ , 0.55- to 0.78-Mev  $\gamma$ 's). Bromine-82 ratios in blood and spinal fluid have been used to differentiate between tuberculous and nontuberculous meningitis. For determining

Ag<sup>111</sup> was one of the early radioactive colloids used. <sup>128</sup> Silver-111, injected into the bloodstream as a protein-silver complex, is removed less rapidly by a cirrhotic liver than by a normal one. <sup>129</sup> Zinc

 ${\rm Zn}^{63}$  (38 min, 1.38- to 2.34-Mev  $\beta$ 's, 0.67- and 0.96-Mev  $\gamma$ 's). Zinc-63 was used in early work for control of malignant effusions. Its use in studying zinc metabolism was reported recently. 130

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VIII. SAFETY