

UNITED STATES ATOMIC ENERGY COMMISSION

WASHINGTON, D.C. 20545

JUN 1 2 1974



Honorable Gaylord Nelson Subcommittee on Monopoly Select Committee on Small Business United States Senate

Dear Senator Nelson:

This is in response to your letter of May 8, 1974, requesting information on the Atomic Energy Commission's contribution in the development of drugs during the last 15 years.

One of the principal charges to the Commission is the development of new radiopharmaceuticals for the diagnosis and treatment of disease. Ever since its inception the AEC has played a prominent role in supporting research and development, including in many cases pre-clinical and clinical testing of radiopharmaceuticals. A new medical discipline, nuclear medicine, has come into being largely as a result of advances of AEC-supported research.

Radioisotopes have been made available by AEC laboratories for use in medical research since August 1946 from AEC-operated specialized research facilities. In addition, the Commission has supported research through contracts with medical schools, universities, hospitals and other research organizations.

Almost all radiopharmaceuticals used in various diagnostic and therapeutic applications in nuclear medicine were developed in AEC-supported laboratories or by AEC-supported scientists. As an example, it is reported that out of some 30 different radioisotopes currently used, technetium-99m is administered better than 50 percent of the time. At present, technetium-99m is available in a variety of chemical forms from most radiopharmaceutical manufacturers.

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A measure of the impact of nuclear medicine research is reflected in the growth of diagnostic nuclear medicine. Stanford Research Institute reported in 1959 that 400,000 doses of radioisotopes were administered to humans. In 1966 this number had grown to approximately 1,743,000 administrations. For 1974 the Society of Nuclear Medicine estimates that 10,000,000 administrations will be made.

The enclosed list of radiopharmaceuticals currently in routine use provides an indication of the broad spectrum of medical uses.

With regard to the total support of this program, it is necessary to point out that nuclear medicine research or research on the development of individual radiopharmaceuticals is not identified as a line item in the budget. Instead, the research and development of radiopharmaceuticals is part of a larger program which includes all beneficial uses of radiation and radioisotopes. In the past 15 years approximately \$30,000,000 has been spent in support of radiopharmaceutical research. In 1959 about \$1,500,000 was spent with increases over the years to the current level of approximately \$3,000,000.

In addition to radiopharmaceutical research, AEC has supported drug research initiated in the mid-1960's by Dr. George C. Cotzias and his associates at the Brookhaven National Laboratory which has led to the use of the drug L-dopa (L-3, 4-dihydroxyphenylalanine) in the treatment of Parkinson's disease. When administered daily in large oral doses, the drug induces substantive symptomatic improvement in most Parkinsonian patients and dramatic results in some. L-Dopa has been found to be well tolerated and can generally be administered for protracted periods of time. Though not severe in most cases, undesirable side effects have been encountered.

On the basis of clinical studies involving hundreds of subjects and clinical experience with many more, as reported by a number of medical authorities, it appears that L-dopa gives beneficial results in about two out of every three Parkinsonian patients—the remainder being either unresponsive to the drug or unable to tolerate it.

L-Dopa therapy is now recognized as the best approach to the management of Parkinson's disease. However, Dr. Cotzias' main contribution has been that of a better understanding of neurologic disorders and a sounder basis for their control via drug therapy.

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By now, more than 6,000 victims of Parkinson's disease have benefited from L-dopa therapy. Since there are in this country an estimated 500,000 - 1,500,000 persons afflicted with this disease, many remain to be treated. Now that L-dopa has become readily available and has been approved by the Food and Drug Administration for general medical use, the full benefit of the drug should soon be realized in this country. Many victims of manganism, in Chile and elsewhere, will likewise be helped by L-dopa. Total AEC support to date of research directly related to the development of L-dopa is \$4,700,000.

I trust that the information provided has adequately answered your questions. If I may be of further assistance, please contact me.

Sincerely,

Original signed by Dixy Lee Ray **Chairman**

Enclosure: Radiopharmaceutical List

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Form AEC-318 (Rev. 9-53) AECM 0240

Approved Pharmaceuticals

Isotope	Chemical Form	Use		
Chromium-51	Chromate	Spleen scans, placenta locali- zation, red blood cell labeling and survival studies.		
Chromium-51	Labeled human serum albumin	Gastrointestinal protein loss studies, placenta localization.		
Chromium-51	Labeled red blood cells	Placenta localization		
Cobalt-58 or Cobalt-60	Labeled cyanocobalamin	Intestinal absorption studies		
Fluorine-18	Fluoride	Bone imaging		
Gold-198	Colloidal	Liver scans, intracavitary treat- ment of pleural effusions and/or ascites, interstitial treatment of cancer.		
Iodine-131	Iodide	Diagnosis of thyroid functions, thyroid scans, treatment of hyper- thyroidism and/or cardiac dysfunc- tion, treatment of thyroid carcin- oma.		
Iodine-131	Iodinated human serum albumin	Blood volume determinations, cisternography, brain tumor localization, cardiac scans for determination of pericardial effusions.		
Iodine-131	Rose bengal	Liver function studies, liver scans.		
Iodine-131	Idopyracet, sodium iodo- hippurate, sodium diatrizoate, diatri- zoate methyglucamine, sodium diprotrizoate, sodium acetrizoate, or sodium iothalamate	Liver scans, kidney function studies and kidney scans.		
Iodine-131	Labeled fats and/or fatty acids	Fat absorption studies		
Iodine-131	Iodipamide	Cardiac scans for determination of pericardial effusions		
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Isotope	Chemical Form	Use		
Iodine-131	Macroaggregated iodinated human serum albumin	Lung scans		
Iodine-131	Colloidal microaggregated human serum albumin	Liver scans		
Iodine-125	Iodide	Diagnosis of thyroid function		
Iodine-125	Iodinated human serum albumin	Blood volume determinations		
Iodine-125	Rose bengal	Liver function studies		
Iodine-125	Iodopyracet, sodium iodo- hippurate, sodium diatri- zoate, diatrizoate methy- lglucamine, sodium diprotrizoate, sodium acetrizoate, or sodium iothalamate	Kidney function studies		
Iodine-125	Labeled fats and/or fatty acids	Fat absorption studies		
Iron-59	Chloride, citrate and/or sulfate	Iron turnover studies		
Krypton-85	Gas	Diagnosis of cardiac abnor- malities		
Mercury-197	Chlomerodrin	Kidney scans, brain scans		
Mercury-203	Chlomerodrin	Kidney scans, brain scans		
Phosphorus-32	Soluble phosphate	Treatment of polycythemia vera, treatment of leukemia and bone metastasis		
Phosphorus-32	Colloidal chronic phosphate	<pre>Intracavity treatment of pleural effusions and/or ascites, interstitial treatment of cancer</pre>		
Potassium-42	Chloride	Potassium space studies		
Selenium-75	Labeled methionine	Pancreas scans		
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Isotope	Chemical Form	Use		
Strontium-85	Nitrate or chloride	Bone scans on patients with diagnosed cancer		
Technetium-99m	Aggregated albumin (Human)	Lung imaging		
Technetium-99m	DTPA (iron-ascorbate)	Kidney imaging		
Technetium-99m	DTPA (tin)	Brain imaging		
Technetium-99m	Human serum albumin microspheres	Lung imaging		
Technetium-99m	Pertechnetate	Brain scans, thyroid scans, placenta localization, blood pool scans, salivary gland scans		
Technetium-99m	Sodium etridonate	Bone imaging		
Technetium-99m	Stannous polyphosphate	Bone imaging		
Technetium-99m	Sulfur colloid	Liver and spleen imaging		
Xenon-133	Gas	Diagnosis of cardiac abnor- malities, cerebral blood-flow studies, pulmonary function studies, muscle blood-flow studies		

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TO (Name and unit) John Erlewine	INITIALS	REMARKS		Record Copy	
George Gardes John Griffin	DATE	Attached is a response to inquiry on the			
Roger Mattson		study of cancer incidence among former			
TO (Name and unit) Alan Rothenberg	INITIALS	REMARKS Hanford employees by Dr. Samual Milham of			
Mike Hawkins L. Manning Muntzi	DATE 18	the Washington State Health Services			
James Liverman		Division. We have no press inquiries at			
TO (Name and unit) Robert O'Neill	INITIALS	this time but a	as I mention	ed in my note to	
Paul Bender	DATE	the Commissioners yesterday, Dr. Milham			
		apparently has	made some d	istribution among	
John A. Harris	REMARKS	medical author	ities in Was	hington.	
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We have been informed of Dr. Milham's paper. While we appreciate his efforts to assess possible health risks for workers at Hanford, his conclusion—that there have been an excessive number of cancer deaths among Hanford workers—is contrary to a wealth of information which has been collected and analyzed over many years. Reports on this work are available for anyone to examine.

Apparently unknown to Dr. Milham when he did his work, AEC has, for about 10 years, funded a carefully structured study of health effects among nuclear workers, principally at Hanford, Washington, and Oak Ridge, Tennessee, by Dr. Thomas Mancuso of the University of Pittsburgh. Findings thus far include 32,000 Hanford employees; their records are compared to some 20,000 siblings of the same sex who did not work at Hanford.

The Hanford information indicates that (1) there is no apparent significant difference in radiation-related health effects between Hanford employees and their siblings; and (2) based on scrutiny of some 4,000 death certificates, there is no evidence of cancer, or other deaths attributable to ionizing radiation, occurring more often among Hanford workers than among their non-Hanford brothers and/or sisters.

We understand that a copy of Dr. Milham's paper has been sent to Dr. Mancuso for review.



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Assistant to the Chairman

REPLY TO SENATOR NELSON'S LETTER CONCERNING THE COMMISSION'S CONTRIBUTION TO THE DEVELOPMENT OF NEW DRUGS

The reply states the Commission's role in research and development of radiopharmaceuticals and L-dopa, the significance of these drugs for improved health care to the general public, and estimates of funding for this research.

James L. Liverman, Director hivision of Biomedical and Environmental Research

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