

ELECTRON LINEAR ACCELERATOR PRODUCTION OF ^{43}K

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Carrier-free radiopharmaceutical ^{43}K as KCl was produced by the $^{44}\text{Ca}(\gamma, p)^{43}\text{K}$ reaction induced by bremsstrahlung radiation from an electron linear accelerator. An irradiation of 1 hr using 5.4 kW of 45-MeV electrons passing through a thick tantalum converter produced yields in excess of 1.5 mCi/gm ^{44}CaO . The ^{43}K was separated from the target material and prepared as ^{43}KCl using ion-exchange chromatography. This radiopharmaceutical is now being prepared in millicurie quantities for myocardial imaging, electrolyte studies, and other uses for which ^{40}K and ^{42}K are unsuited.

The radionuclide ^{43}K , in the form of KCl, has been found to be an effective agent for myocardial imaging and electrolyte studies (1-4). Clinically usable quantities of this radioisotope have been produced in high-flux nuclear reactors by the $^{43}\text{Ca}(n, p)^{43}\text{K}$ reaction and in cyclotrons by the $^{40}\text{Ar}(\alpha, p)^{43}\text{K}$ reaction (1,4,5). An alternate reaction is $^{44}\text{Ca}(\gamma, p)^{43}\text{K}$ induced by bremsstrahlung radiation developed by the electron beam of a linear accelerator (6,7). In a practical application of this photonuclear reaction we have routinely produced carrier-free ^{43}K . Our method uses a moderately priced, enriched (94%) ^{44}Ca target and requires no elaborate irradiation or handling equipment to produce sufficient quantities for clinical investigations.

METHODS

A quantity of 94%-enriched ^{44}Ca was purchased as CaCO_3 and converted to the oxide to eliminate the photoproduction of ^{11}C during irradiation. For each production exposure, approximately 0.30 gm of ^{44}CaO was sealed in a quartz ampule and inserted into an air-cooled quartz target holder (Fig. 1). High-energy bremsstrahlung was produced by passing the electron beam of the AFRRRI electron

linear accelerator through a thick (0.7 radiation length), water-cooled tantalum converter. The target was placed approximately 2.5 cm from the converter exit window. An irradiation of 1 hr using 5.4 kW of 45-MeV electrons produced ^{43}K activities greater than 450 μCi in 0.3 gm of ^{44}CaO . Targets were manually transferred to a lead container after irradiation and carried to the hot lab for chemical processing.

Chemical separation of potassium from calcium was performed on a Bio-Rad AG 50W-X8 in H^+ form ion-exchange column using a procedure developed at ORNL (5), modified to incorporate a

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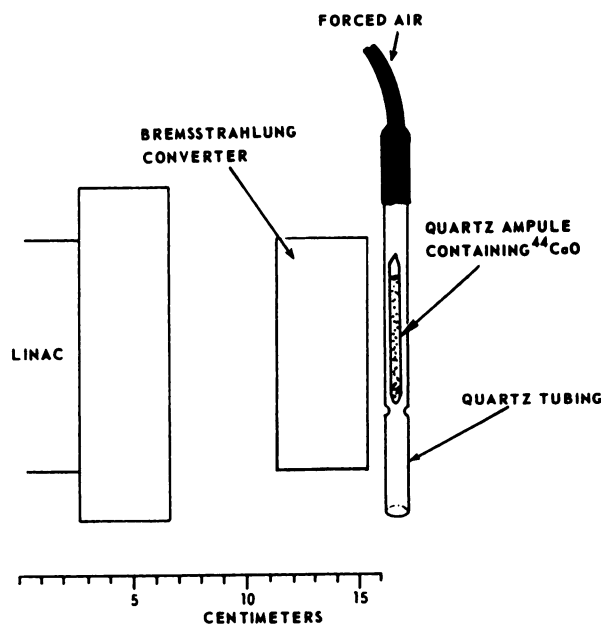


FIG. 1. Diagram of irradiation arrangement. Electron beam from Linac passes through bremsstrahlung converter, producing high-energy photons which induce photonuclear reactions in target material.

simple pneumatic pressure system to elute the ^{43}K from the column. A standard nitrogen or helium tank was regulated at about 4 psi to maintain a flow rate of 3 ml/min. The exchange column was maintained at approximately 80°C by a hot water jacket and an infrared heating lamp directed onto the column head reservoir. At this elevated temperature, the difference between the selectivity coefficients of K^+ and Ca^{2+} increases, thereby enhancing the separation of the two ions. The selectivity coefficients were unaffected by the operating pressure of the column (8). The eluted KCl was taken to dryness, dissolved in normal saline, and passed through a 0.22-micron Millipore filter. After elution from the column, the target material was then recovered by precipitation with saturated ammonium oxalate, furnace in a nitrogen atmosphere at 950° for 6 hr, and encapsulated for reactivation. Recovery of the enriched calcium was 98% or better. The recovered CaO contained minimal amounts of radionuclides derived from other stable calcium isotopes present in the target material. No special handling precautions were required.

DISCUSSION

No direct cross-section data appear to be available for the reaction $^{44}\text{Ca}(\gamma, p)^{43}\text{K}$. Brix, et al (6) have reported a value for the integrated cross section of 120 MeV-mb over the energy interval 12.2–31 MeV. Wu, et al (9) measured the differential cross section at 90° for the reaction $^{40}\text{Ca}(\gamma, p)^{39}\text{K}$ finding a maximum at approximately 20 MeV and a half-width of about 4 MeV. We assumed these parameters to describe the giant resonance peak for the $^{44}\text{Ca}(\gamma, p)^{43}\text{K}$ reaction. The bremsstrahlung converter thickness was optimized for the production of photons of energy 15–25 MeV by a computer simulation technique, making use of bremsstrahlung formulas suggested by Hansen and Fultz (10). A new tungsten converter of 1.3 radiation length thickness is being constructed. Using 45-MeV electrons, it is expected to increase the radionuclide yields by more than 40%.

Irradiation of ^{44}Ca with photons of energy greater than 20 MeV produces ^{42}K by the competing γ, np reaction. The gamma and beta energies from decay of ^{42}K are higher than those from ^{43}K , resulting in a higher body dose and hindering conventional gamma scanning techniques. It is thus desirable to reduce the level of ^{42}K in most preparations of ^{43}K . The $^{44}\text{Ca}(\gamma, np)^{42}\text{K}$ reaction threshold has been computed to be higher than that for the $^{44}\text{Ca}(\gamma, p)^{43}\text{K}$ reaction (11). It is therefore possible to reduce the ratio of ^{42}K to ^{43}K activities by reducing the energy of the electron beam although the reduction also

TABLE 1. YIELDS OF ^{43}K AND ^{42}K FOR SEVERAL ELECTRON BEAM ENERGIES

Beam energy (MeV)	Yield ($\mu\text{Ci}/\text{gm}^{44}\text{CaO-hr}$)		
	^{43}K	^{42}K	Ratio $^{42}\text{K}/^{43}\text{K}$
30.0	370	23	0.062
35.7	770	92	0.12
40.4	1100	230	0.21
44.5	1500	390	0.26

reduces the yield of ^{43}K as shown in Table 1. Myocardial scans on beagles were performed at the AFRRI using 100–150 μCi per visualization. For such use, 0.30 gm of ^{44}CaO exposed to 35.7-MeV bremsstrahlung for 1 hr produced ample ^{43}K with only 12% ^{42}K contamination. Much larger amounts of ^{43}K may be produced by using larger amounts of ^{44}CaO target material although targets greater than 1 gm would require a longer ion-exchange column for separation.

REFERENCES

1. HURLEY PJ, COOPER M, REBA RC, et al: ^{43}KCl : a new radiopharmaceutical for imaging the heart. *J Nucl Med* 12: 516–519, 1971
2. ZOLLINGER RM, VAN DEWATER JM, MALETSKOS CJ, et al: Exchangeable potassium in man using a new radioisotope ^{43}K . *Surg Forum* 21: 213–215, 1970
3. JOHNSON JE, HARTSUCK JM, ZOLLINGER RM, et al: Radiopotassium equilibrium in total body potassium: studies using ^{43}K and ^{42}K . *Metabolism* 18, 663–668, 1969
4. SKRABAL F, GLASS HI, CLARK JC, et al: A simplified method for simultaneous electrolyte studies in man utilizing potassium-43. *Int J Appl Radiat Isot* 20, 677–681, 1969
5. POGGENBURG JK: Neutron products at ORNL. In *Radioisotope Production Development Meeting*, CONF-700646, Springfield, Va, National Technical Information Service, 1970, pp 19–22
6. BRIX P, HEGEL U, LINDENBERGER KH, et al: Relativmessung integrierter Wirkungsquerschnitte für den Kernphotoeffekt: Die Reaktion $\text{Ca}^{44}(\gamma, p)\text{K}^{43}$. *Z Phys* 150: 461–473, 1958
7. OKA Y, KATO T, NOMURA K, et al: A study on the yield of (γ, p) reactions with 20 MeV bremsstrahlung. *Bull Chem Soc Jap* 41: 380–384, 1968
8. HELFFERICH FG: *Ion Exchange*. New York, McGraw-Hill, 1962, pp 166–168
9. WU CP, BAGLIN JEE, FIRK FWK: Correlations in the energies and widths of structure observed in the reactions $^{40}\text{Ca}(\gamma, n_0)^{39}\text{Ca}$ and $^{40}\text{Ca}(\gamma, p_0)^{39}\text{K}$. *Phys Lett* 29B: 359–361, 1969
10. HANSEN NE, FULTZ SC: *Cross Sections and Spectra for Negative Electron Bremsstrahlung*. University of California, Lawrence Radiation Laboratory Report UCRL-6099, 1960
11. HOWERTON RJ, BRAFF D, CAHILL WJ, et al: *Thresholds for Gamma-Induced Reactions*. University of California, Lawrence Radiation Laboratory Report UCRL-14006, 1964