

A Simple Method for Measuring Lu-177m Contaminants in Lu-177 Based Therapeutics

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INTRODUCTION

Purpose: We present a method to measure and quantify ^{177m}Lu in ^{177}Lu -based radiopharmaceutical therapy.

Radiopharmaceutical therapy has expanded rapidly with the adoption of ^{177}Lu -based agents such as Lutathera® and Pluvicto®, ^{177}Lu has a favorable half-life (~6.64 days) and emission profile for therapy and imaging.

The production route impacts the presence of the long-lived contaminant ^{177m}Lu (~160-day half-life), which has implications for waste handling and regulatory compliance.

Production of ^{177}Lu

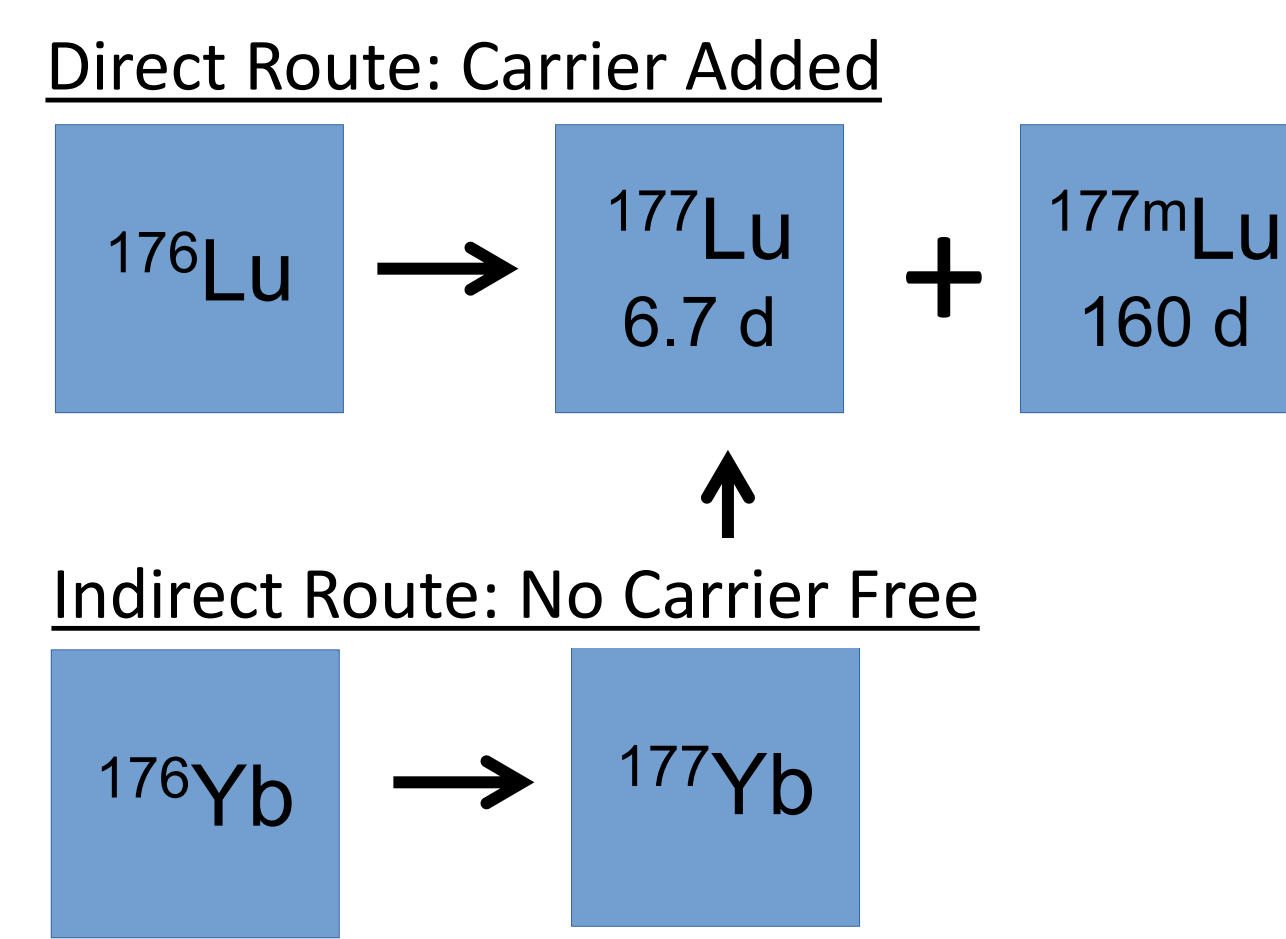


Figure 1. Direct and indirect production routes for ^{177}Lu . The direct production route includes a small cross-section for ^{177m}Lu .

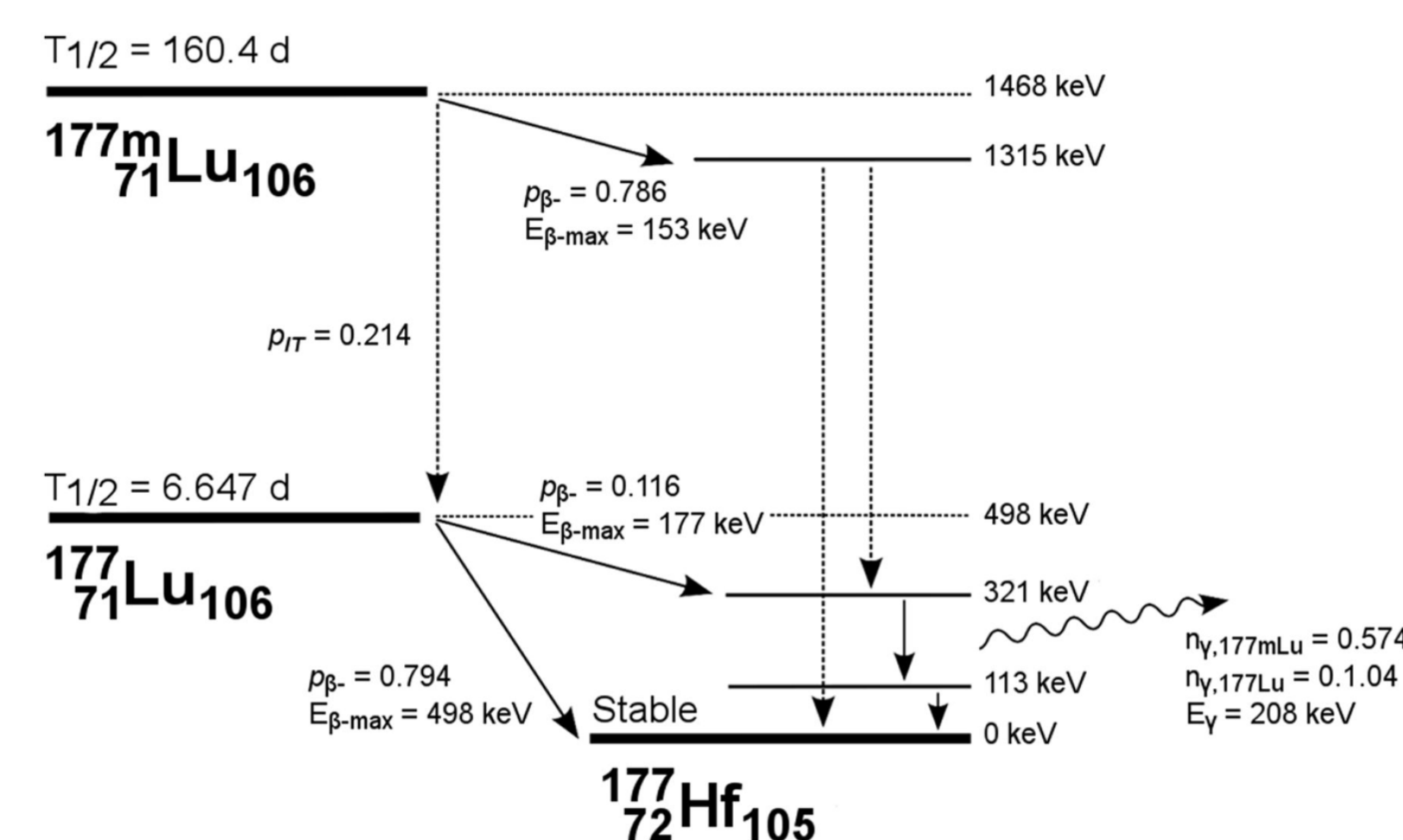


Figure 2. Decay scheme for $^{177m}\text{Lu} \rightarrow ^{177}\text{Lu} \rightarrow ^{177}\text{Hf}$. Adapted from ref. 3

METHODS AND MATERIALS

- 24 samples were collected from unused patient vials (14 Lutathera®, 10 Pluvicto®)
- 1 mL aliquots were analyzed on the following detectors:
 - HPGe detector
 - Nal well detector
 - Liquid scintillation counter (LSC)
- All detector systems were calibrated using traceable standards
- Spectra were collected on each detector system and integrated over the ^{177m}Lu photopeaks of 108 keV, 113 keV, and 120 keV

METHODS AND MATERIALS

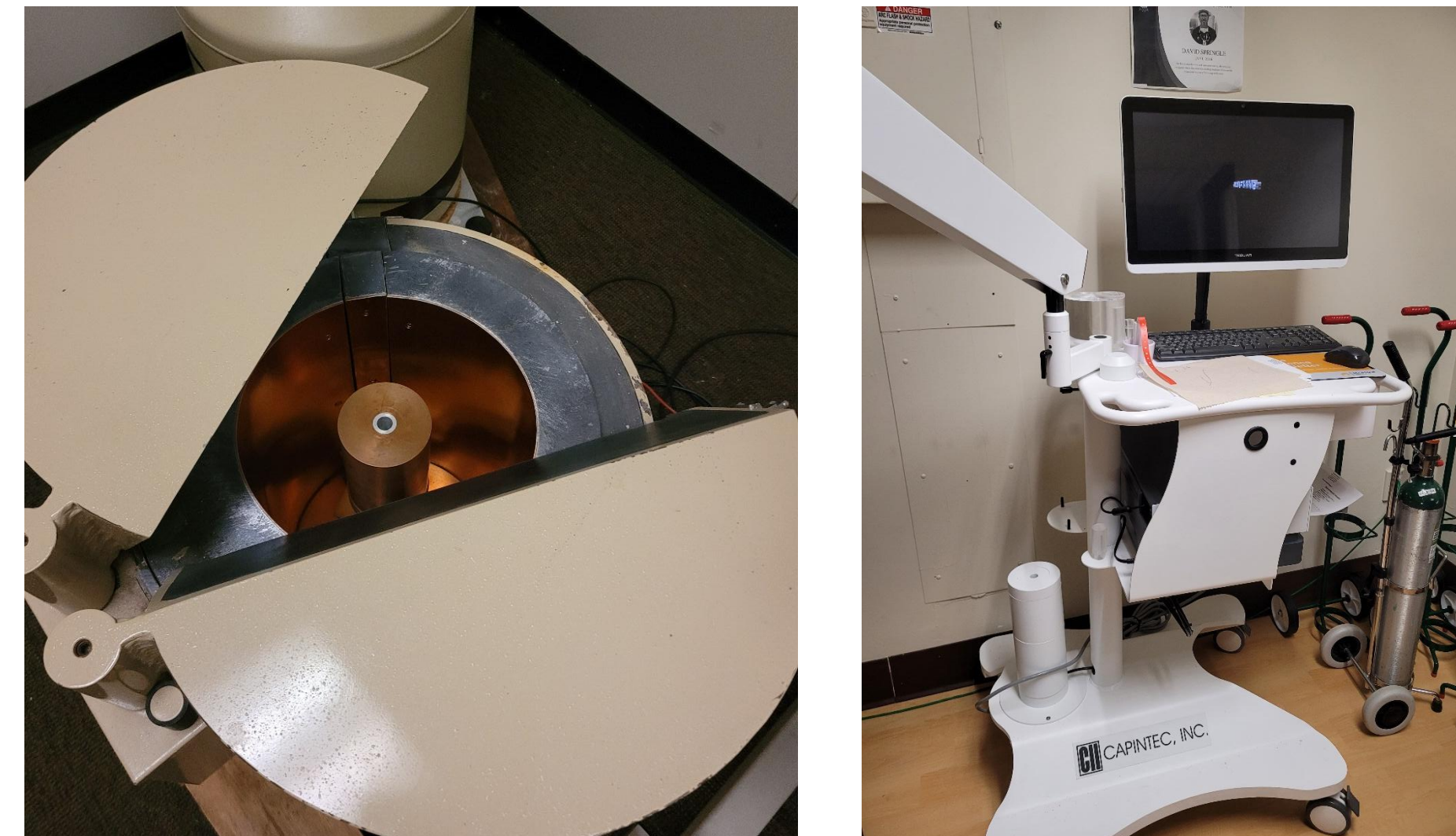


Figure 3. (left) HPGe well in shielding enclosure at the University of South Carolina and (right) Nal well in the nuclear medicine department at the author's institution.

- Individual peaks of 108 keV, 113 keV, and 120 keV could not be resolved on the Nal well and these peaks were combined for all measurements

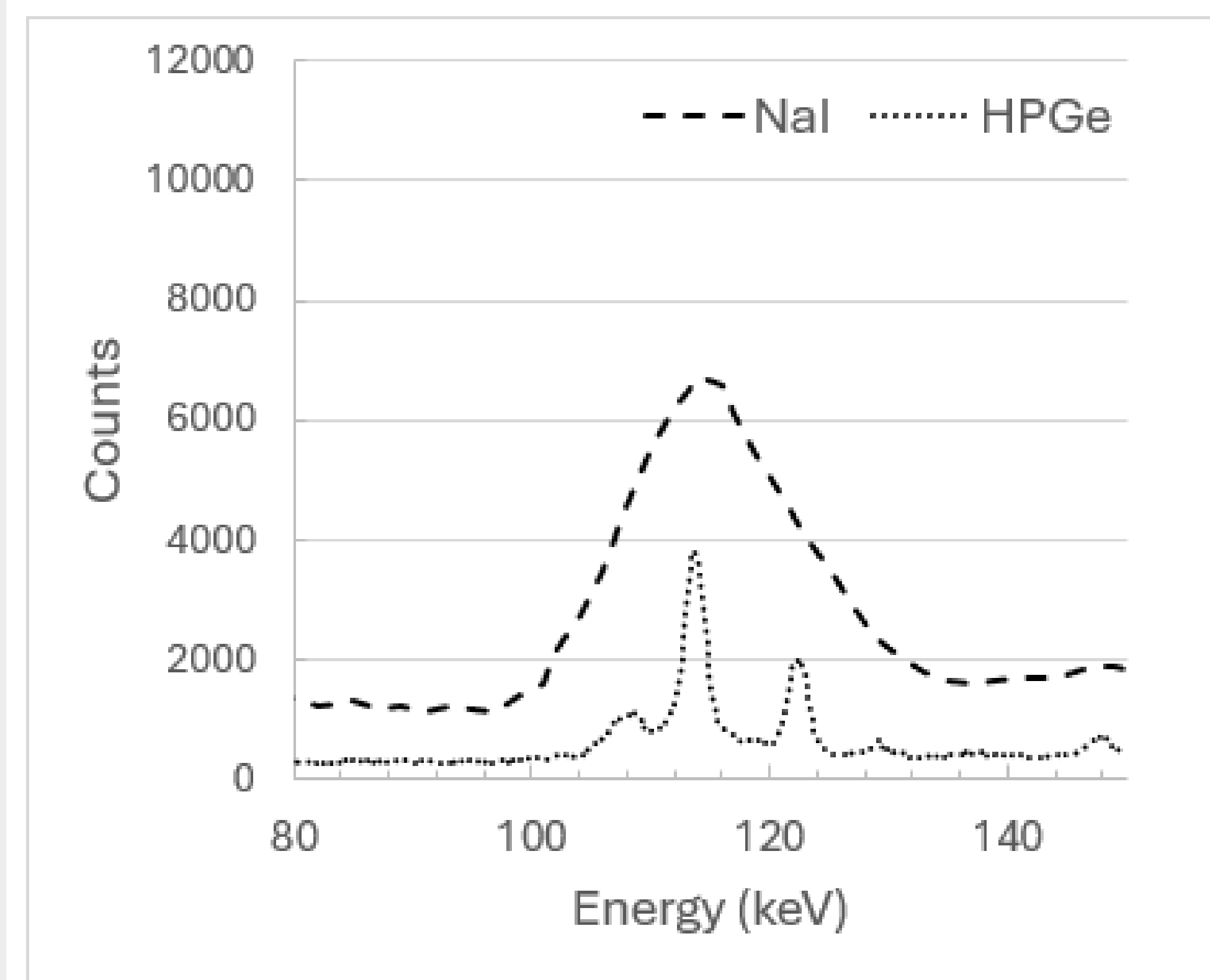


Figure 3. Comparison of HPGe and Nal spectra for the 108 keV, 113 keV, and 120 keV photopeaks collected on a Lutathera® sample.

- The minimum detectable activity (MDA) was calculated from the background under each photopeak.
- Measurements are reported as % ratio of $^{177m}\text{Lu} / ^{177}\text{Lu}$

RESULTS

- Results are shown in table 1.
- There was no statistical difference between $^{177m}\text{Lu} / ^{177}\text{Lu}$ ratios for the HPGe and Nal well detectors
- The LSC overestimated the $^{177m}\text{Lu} / ^{177}\text{Lu}$ ratio due to contributions from multiple beta decay transitions in the sequence $^{177m}\text{Lu} \rightarrow ^{177}\text{Lu} \rightarrow ^{177}\text{Hf}$ that could not be resolved.
- The $^{177m}\text{Lu} / ^{177}\text{Lu}$ ratio was above the MDA estimate for Lutathera® confirming the presence of ^{177m}Lu
- The $^{177m}\text{Lu} / ^{177}\text{Lu}$ ratio for Pluvicto® did not exceed the MDA estimate suggesting the absence of ^{177m}Lu

Table 1. Percent ratio of $^{177m}\text{Lu} / ^{177}\text{Lu}$ measured from the HPGe, Nal well and LSC detector systems.

	Lutathera (n=14)		
	HPGe	Nal	LSC
Mean (MDA)	5.86×10^{-3} (1.56×10^{-4})	6.22×10^{-3} (1.19×10^{-4})	8.48×10^{-3} (7.30×10^{-10})
SD	1.73×10^{-3}	1.32×10^{-3}	1.96×10^{-3}
	Pluvicto (n=10)		
	HPGe	Nal	LSC
Mean (MDA)	4.97×10^{-7} (4.43×10^{-6})	6.94×10^{-7} (1.09×10^{-5})	2.45×10^{-5} (7.87×10^{-9})
SD	8.53×10^{-7}	1.12×10^{-6}	1.65×10^{-5}

MDA = Minimum Detectable Activity

DISCUSSION

- HPGe and Nal well detectors provide sufficient sensitivity to measure and quantify trace amounts of ^{177m}Lu
- LSC systems can detect ^{177m}Lu activity but lack specificity for quantification
- All Lutathera® samples in this study had measurable amounts of ^{177m}Lu . These amounts were consistent with other literature reports
- None of the Pluvicto® samples had ^{177m}Lu that exceeded the MDA
- The presence of ^{177m}Lu likely reflects the production method used to source ^{177}Lu .

CONCLUSIONS

- A practical hospital-based methods for detecting ^{177m}Lu was demonstrated
- Some, but not all, ^{177}Lu -based therapeutics contain measurable amounts of ^{177m}Lu
- HPGe and Nal spectroscopy are preferred for detection and quantification
- LSC is not suitable for quantification due to beta decay interference.

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