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Name/Org: Kevin J. Hart Date: November 1, 2019
Guidance (if applicable): DOE Isotope Program PM Office





Agenda

11:00 – 11:10 AM **Roy Copping**, Oak Ridge National Laboratory (Moderator) Introduction – "Oak Ridge Ra-224/Pb-212 generator production"

11:10 – 11:20 AM **Mengshi Li**, Viewpoint / University of Iowa "Preclinical evaluation of synergistic anti-tumor effect from combination of ²¹²Pb alpha-radiation and immune checkpoint inhibitors"

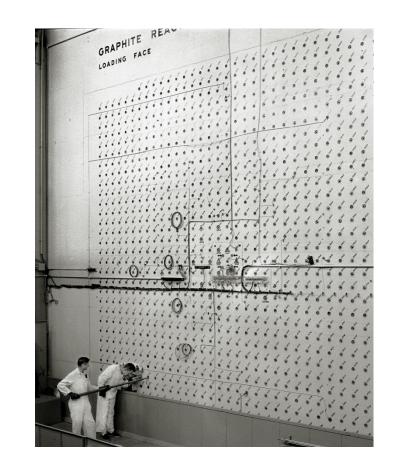
11:20 – 11:30 AM **Sangeeta Ray**, Johns Hopkins University "Preclinical evaluation of 212Pb-based radiopharmaceutical therapy of prostate cancer"

11:30 – 11:40 AM **Ebrahim Delpassand**, Chairman & CEO, RadioMedix Inc. "²¹²Pb-AlphaMedixTM Targeted Alpha Therapy (TAT): A potential breakthrough in treatment of metastatic SSTR expressing NET"

11:40 – 11:50 AM **Matt O'Hara**, Pacific Northwest National Laboratory "Recent activities in ²¹²Pb generator development at PNNL"

11:50 – 12:30 PM Q&A Segment

Isotope production, enrichment and distribution began at Oak Ridge in 1946





ORNL has a Rich History in Medical Radioisotopes

1946



1st 14C shipment to Barnard Free Skin and Cancer Hospital, St. Louis.

1946-1963: 1000's of shipments of up to 60 different radioisotopes 1947-2009



Large-scale mouse genetics Disease Treatment project to study the effects of radiation on mammals

Radiation Protection and Effects

- Broad biological research program

1993



Cancer and Heart

²²⁵Ac /²¹³Bi generator

1997



²²⁵Ac & production 1998-2011



Office of Biological and Environmental Research: Low Dose Radiation Research

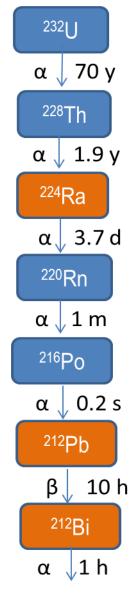
2017



²²⁷Ac/²²³Ra treatment for prostate cancer patients



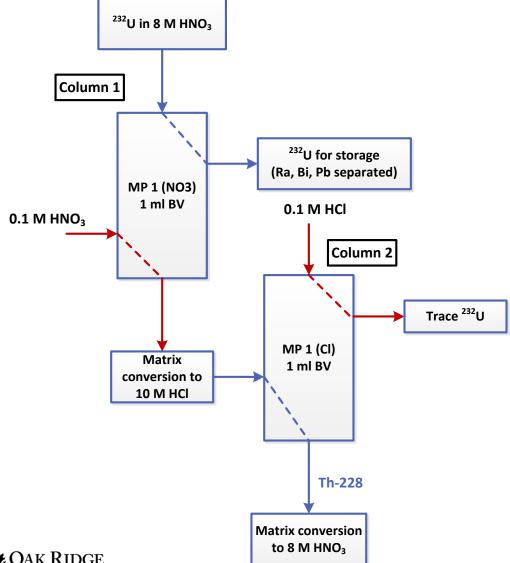
²²⁴Ra/²¹²Pb Generator production



- Production established in 2014
- ²²⁴Ra (3.6 d) separated from parent ²²⁸Th and loaded on cation resin
- "Generator" (<16 mCi) can be milked periodically for $^{212}{\rm Pb}$ (t $_{1/2}$ = 10 hrs) and $^{212}{\rm Bi}(1~{\rm hr})$ and used in TAT applications
- Shipments every three weeks
- ²²⁸Th cow periodically supplemented from ²³²U parent

Chemistry

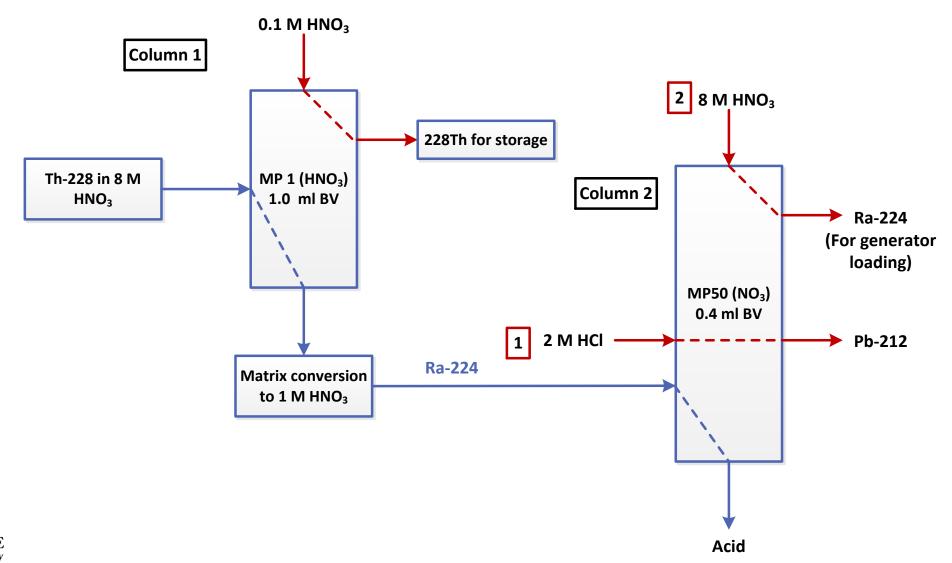
• ²²⁸Th parent recovered from ²³²U annually





Chemistry

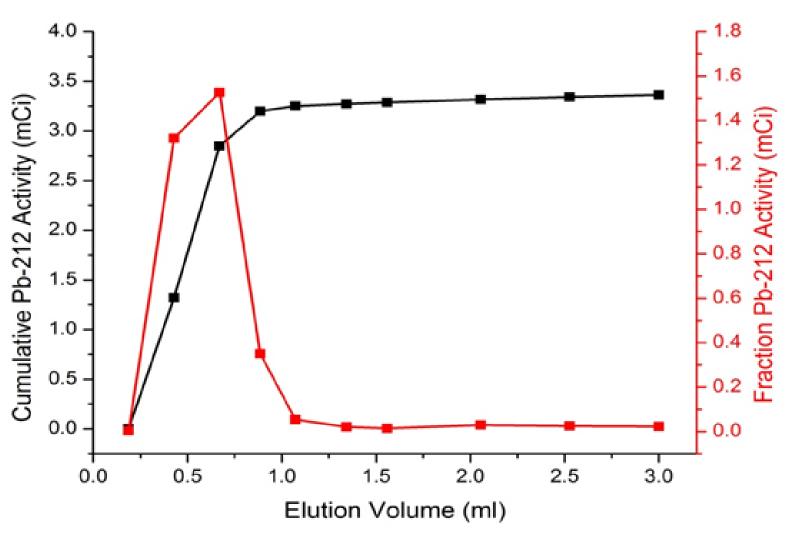
• ²²⁸Th separated and recycled during ²²⁴Ra separation for generator loading



²¹²Pb Elution Profile

212Pb can be eluted from the generator in 2 M HCl





²²⁴Ra^{/212}Pb Generator







²²⁴Ra^{/212}Pb Generator Future Developments

- ²²⁸Th available as a bi-product of Ac-227 production
- Expanded production envisaged in FY20/21
- Glovebox shield under construction to minimize worker dose and increase loading activity
- Can work with customer on schedule/activities