

# **Corporate Presentation**

May 2023

**NYSE: CATX** 



### Legal Disclaimers

Statements in this presentation about Perspective Therapeutics, Inc.'s (the "Company") future expectations, including: the functionality, capabilities, and benefits of the Company's therapies including its targeted alpha-particle radiotherapy; whether favorable results of the initial trials will continue and whether those results can be commercialized for wider use; whether it maintains its contract with the Department of Energy ("DOE") for the therapeutic isotope supply of "Th-228"; whether the Fast Track FDA approvals are maintained; expectations about the Company's addressable markets; the potential size of the commercial market for the Company's treatment programs; the Company's expectations, beliefs, intentions, and strategies regarding the future; and all other statements in this presentation, other than historical facts, are "forwardlooking statements" within the meaning of the Private Securities Litigation Reform Act of 1995 ("PSLRA"). This statement is included for the express purpose of availing the Company of the protections of the safe harbor provisions of the PSLRA. It is important to note that actual results and ultimate corporate actions could differ materially from those in such forward-looking statements based on such factors as whether additional studies are released that reinforce the results of the studies discussed in this presentation; whether the anticipated benefits of the Company's therapies including its targeted alpha-particle radiotherapy are realized; the ability to raise ongoing capital to fund added costs of research and development; the ability of the Company to manage growth and successfully integrate its businesses; whether the Company can maintain its key employees; training and use of the Company's products; market acceptance and recognition of the Company's products; the Company's ability to enforce its intellectual property rights; whether it can maintain and not default on its therapeutic isotope supply agreement with the DOE; successful completion of future research and development activities; whether we, our distributors, and our customers will successfully obtain and maintain all required regulatory approvals and licenses to market, sell, and use our products in their various forms; the procedures and regulatory requirements mandated by the FDA for additional trials, Phase I and II approvals, Fast Track approvals, and 510(k) approval and reimbursement codes; changes in applicable laws and regulations; and other risks detailed from time to time in the Company's reports filed with the SEC.

Unless required to do so by law, we undertake no obligation to publicly update or revise any forward-looking statements, whether as a result of new information, future events, or otherwise. For more information regarding risks and uncertainties that could affect the Company's results of operations or financial condition, please review the definitive Proxy Statement filed on November 7, 2022, and our Form 10-K filed on September 28, 2022, with the SEC. This presentation shall not constitute an offer to sell or the solicitation of an offer to buy any securities of the Company nor shall there be any sale of securities in any jurisdiction in which such offer, solicitation, or sale would be unlawful prior to registration or qualification under the securities laws of any such jurisdiction.





The combined focus advances precision cancer treatments using target delivery of radiation and imaging technologies; *treating cancer from the inside out* 

### Best-in class development team

- IP and innovation in drug development
- Proprietary peptide-chelator based targeting platform
- In-house <sup>212</sup>Pb isotope supply
- 2 Clinical stage programs and robust pipeline

### >\$18m in competitive NIH awards

• Science rigorously validated by independent peers

## February 6, 2023 Completed merger with Isoray Medical



 Commercially available Cs-131 for LDR brachytherapy
 Multi-indications for brachytherapy & Targeted Alpha Therapies
 Stronger balance sheet

### Management Team:

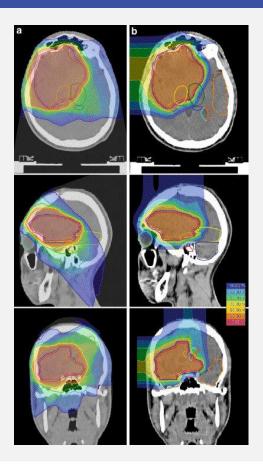
Deep Radiotherapy and Development Experience



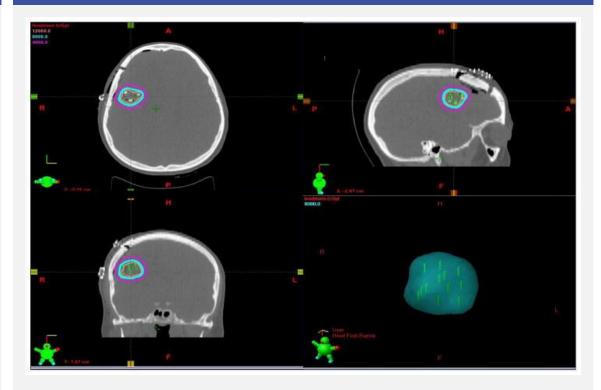


### Targeting Cancer Cells from the inside out: External Beam Radiation irradiates Healthy Tissue Along with Tumors

#### Absorbed dose 4 weeks post External Beam treatment



### Similar tumor dose post Cesium-131 implant



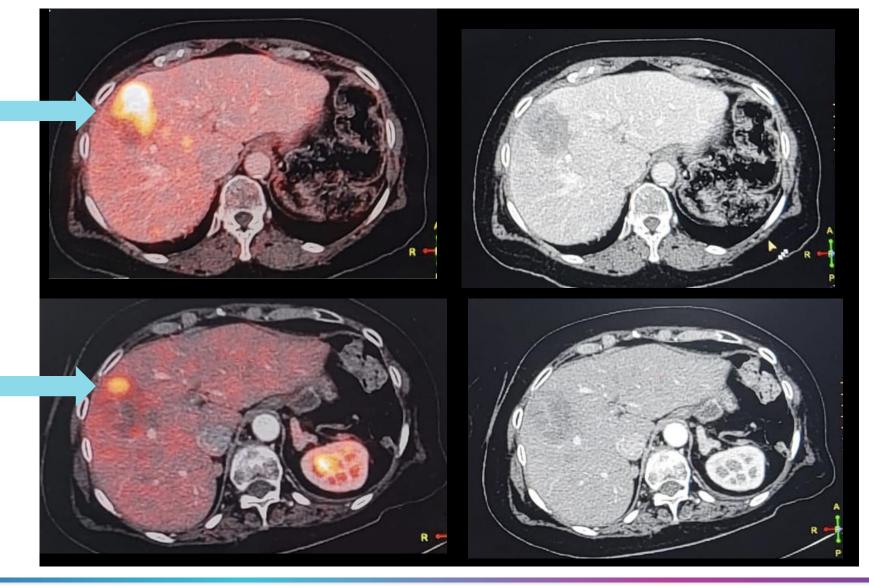


# 1<sup>st</sup> Patient with [<sup>212</sup>Pb]VMT- $\alpha$ -NET – significant tumor response after a single dose

Tumor Before Treatment

**Tumor After** 

1 Dose





Dr. Ishita B Sen Director & Head Dept. of Nuclear Med. & Molecular Imaging Fortis Memorial Research Institute, Gurgaon, India

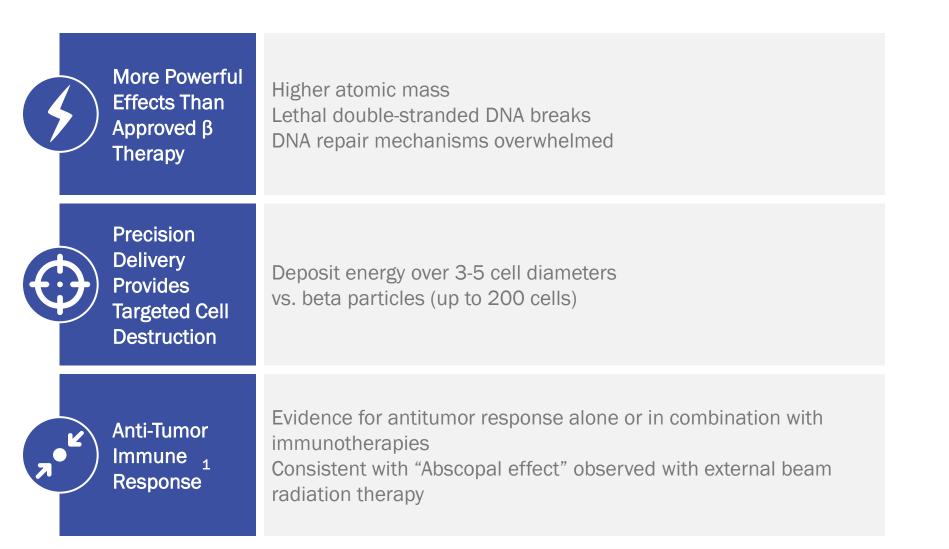


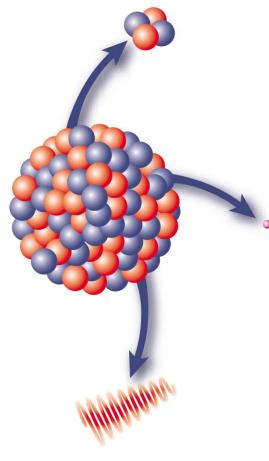
## Radiopharmaceuticals Are a New Pillar of Oncology Treatment with Pan-Cancer Opportunities

Molecularly targeted radiation	Radiotherapy is a proven pillar of cancer treatment but lacks precision. By precisely delivering radiation directly to cancer cells with radioligands, the power of radiotherapy can be realized while reducing the off-target effects.
Optimized patient selection	Molecular imaging companion diagnostics enable visualization of the therapeutic target to optimize the selection of patients who may best respond to therapy.
Monotherapy activity and combination synergies	Many opportunities exist to develop radiopharmaceuticals as monotherapies and additional options exist to extend the therapeutic uses by combining with other precision treatments, such as DNA damage response and immune checkpoint inhibitors.
Outpatient friendly	Modern medical isotopes enable radiopharmaceuticals to be administered outside of hospitals, making them easily accessible globally - over 1,400 outpatient locations in the U.S alone.
Limited competitors + deep moat preventing new entrants = unique business opportunity	Radiopharmaceutical theranostic product development is more highly specialized and technical than for standard medicines, requiring deep knowledge and expertise across multiple disciplines.



### *α*-Particles Have Superior Tumor Killing Properties vs. β-Particles



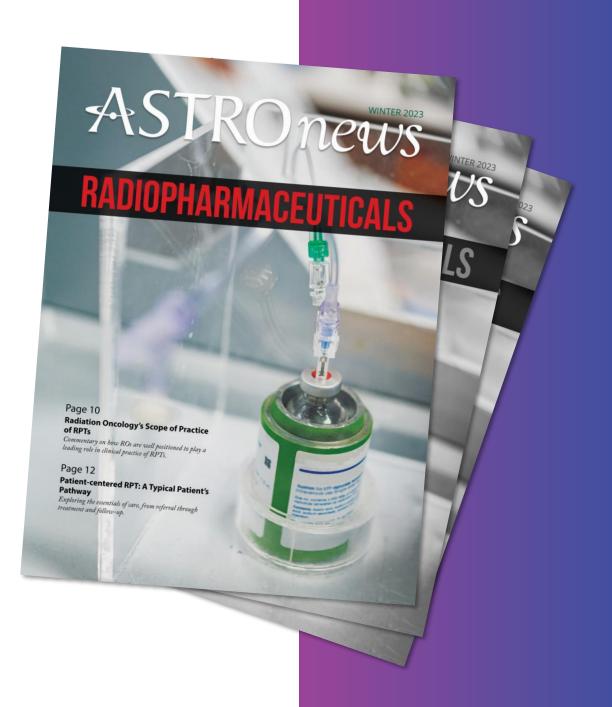


 $\alpha$ -particles are >7,000-fold greater in atomic mass



# Targeted α-Particle Radiotherapy

A New Class of Oncology Therapeutics



# Lead-212 (<sup>212</sup>Pb): The Optimal Therapeutic Isotope

Superior\* Therapeutic Energy Expected to Improve Outcome with Better Safety

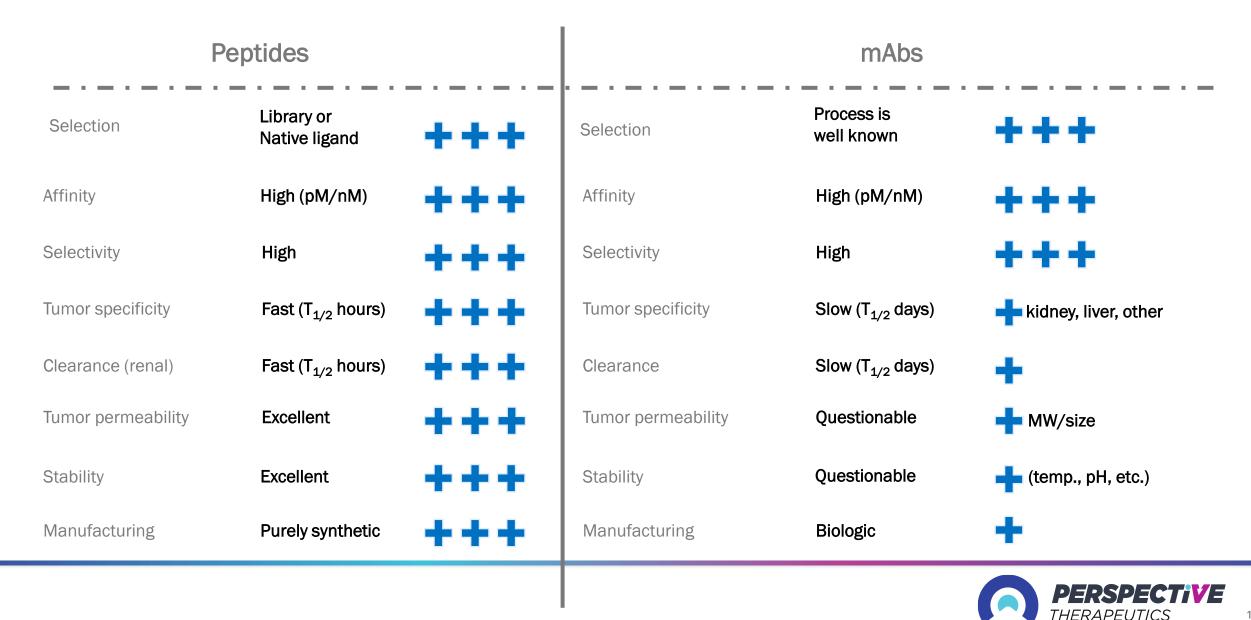
	lodine ( <sup>131</sup> l)	Lutetium ( <sup>177</sup> Lu)	Actinium ( <sup>225</sup> Ac)	Lead ( <sup>212</sup> Pb)	Implication*
Emission Profile	Beta	Beta	Alpha	Alpha	Potent
Half Life	8 days	6.7 days	10 days	0.46 days	Rapid Clearance
Off Target Toxicity Risk	Very high	Low	High	Low	Best
Supply	High	Low	Low	High	Abundant
Cost of Production	Low	High	High	Low	High margin



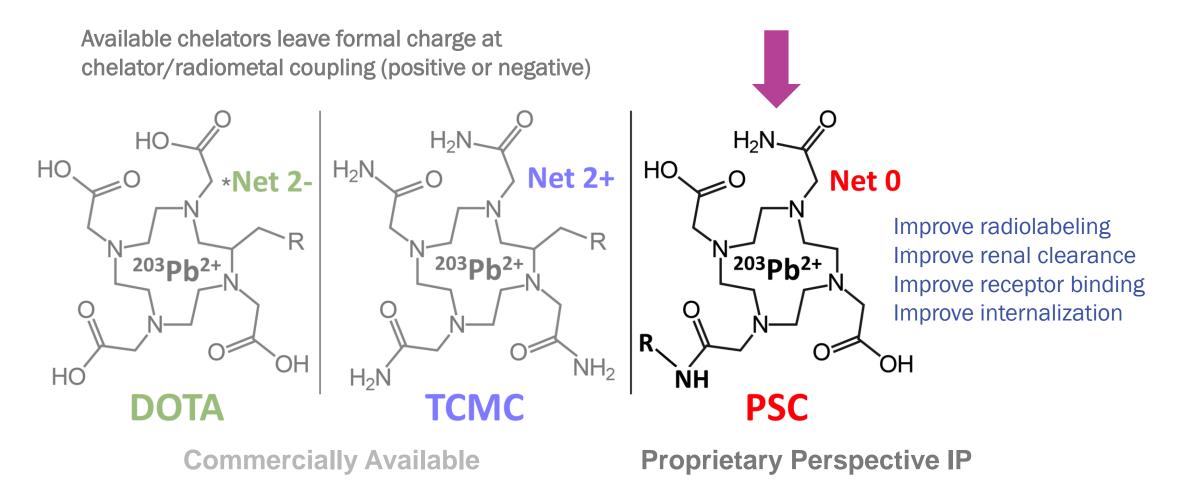


\*Source: Management estimates and assumptions based on current literature and known physical constants

### Peptides emerging as ideal ligands for radiopharmaceutical therapy



### Finding an optimized <sup>212/203</sup>Pb Chelator





### Lead (Pb): The Ideal Theranostic Isotope

#### Ideal Theranostic

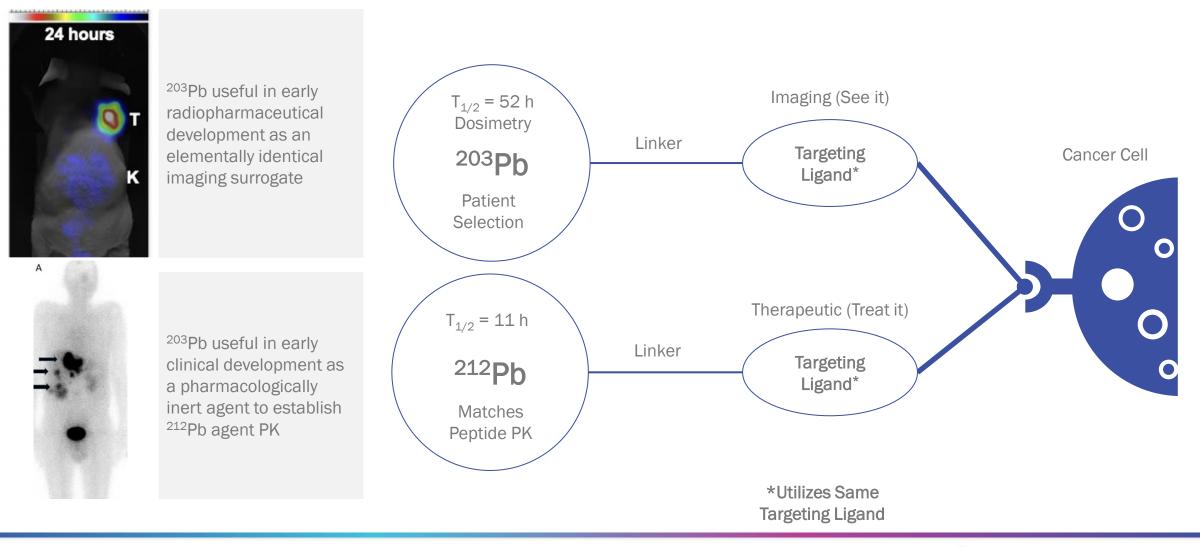
- Ideal agreement between imaging and therapeutic compounds
- Readily available isotope
- Ideal chelator
- Rapid clearance from blood
- High tumor retention @24 hours
- Short t-<sup>1</sup>/<sub>2</sub> gives rapid effect while minimizing environmental impact
- No unsafe daughter isotopes

### <sup>212</sup>Pb & Viewpoint Chelator

- <sup>203</sup>Pb and <sup>212</sup>Pb matched pair
- Generator produced
- PSC carries 0 net charge
- Peptide targeting
- High binding
- Low hospital and patient impact for radiation safety
- Decays to cold Pb



### *Theranostics: Understand Which Patients Will Respond to Therapy*





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## Discovery Platform Gives Broad Proprietary Pipeline

Program	Indication	Discovery	Lead Optimization	Development Candidate	Clinical Imaging, IND enabling	Clinical Trials (Phase 1-3)
	Neuroendocrine					
VMT-α-NET	Pheochromocytomas, paragangliomas					
	Small cell lung cancer					
VMT-01	Melanoma (MC1R)					
VMT-02 (PET agent)	Melanoma					
Program A (Novel peptide)	Multiple solid tumors					
Program B (Novel peptide)	Prostate					
Program C (Novel small molecule)	Prostate					
Program D (Novel peptide)	Breast cancer					
Program E (Novel Small Molecule)	Melanoma					
Other programs (Peptides)	Multiple Solid Tumors					



# Neuroendocrine Tumors: VMT-alpha-NET

Using the somatostatin receptor to target rare and neuroendocrinetype cancers

### *VMT-α-NET*

### **Key Facts**



Targeting somatostatin receptor type 2 (SSTR2)

Currently in Phase 1/2a studies for the imaging and treatment of neuroendocrine tumors



Initiated first-in-human imaging (2021) & therapy (2022) under compassionate use



US Phase I imaging study enrolling at the University of Iowa. Therapeutic IND open.



Received Fast Track Designation October 2022 Opportunity for Orphan Drug and Rare Pediatric Disease Designations



### Gastroenteropancreatic Neuroendocrine Tumors (GEP-NETs)

Malignancies of neuroendocrine cells, which are specialized cells that secrete hormones and other bioactive substances which are found throughout the body Often grows in the pancreas or other areas of the gut, • such as the stomach, small intestine, rectum, colon or appendix Significant unmet need with a \$5 billion market opportunity<sup>1</sup>

~12K new diagnoses annually<sup>2</sup>

~175,000 people are living with this diagnosis<sup>2</sup>

Treatment depends on the type of tumor. Some approaches may include surgery, radiation, and chemotherapy



## Preclinical Imaging Demonstrates Superiority of Perspective's Platform Technology

#### Key Takeaways



Tumor model expressing somatostatin receptor 2 (SSTR2)



8-fold improved tumor uptake with decreased kidney retention

#### CT SPECT Scan – 24 Hours After Injection





Colored area is retention of drug in tumor (T) and kidney (K)

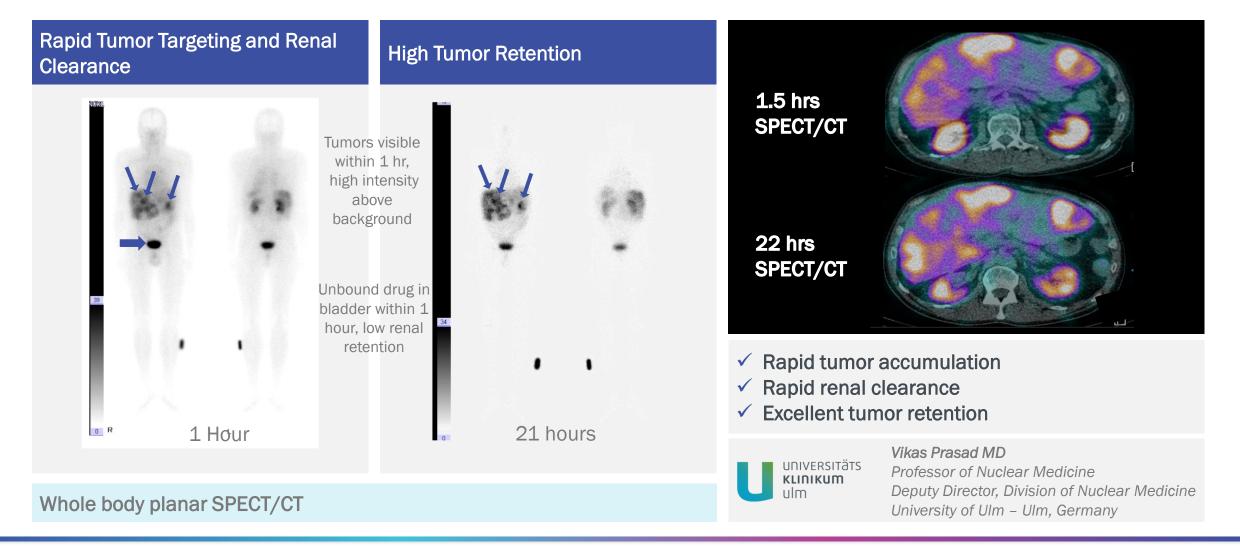


### <sup>203</sup>Pb-VMT-α-NET

Warm hues indicate greater retention of drug in tumor (T) Faint retention in kidney (K)



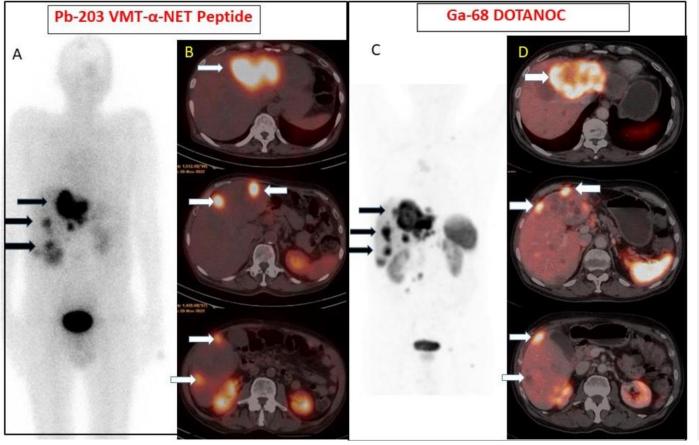
## *Imaging Reveals Favorable VMT-α-NET Properties*





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## *Imaging VMT-α-NET Clearly Identifies Lesions*



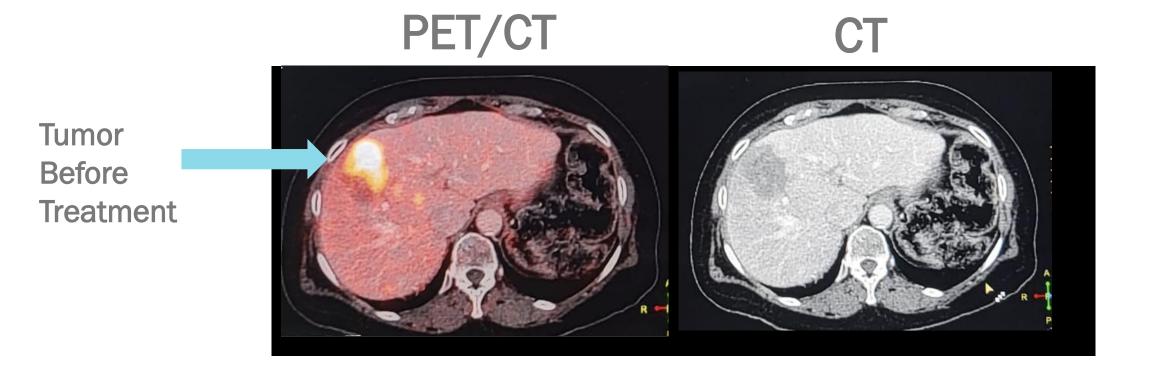
 Whole body Planar (A) and transverse SPECT CT (B) images acquired at 2 hours post injection of <sup>203</sup>Pb-VMT- α-NET showed excellent uptake of Pb-203 VMT-α-NET peptide in the liver metastatic sites in a patient of metastatic neuroendocrine tumor

 A comparison with <sup>68</sup>Ga DOTANOC PET/CT scan demonstrated similar distribution of tracer in the metastatic liver lesions-Maximum intensity projection (C) and Fused transverse PET/CT slices (D).

Clear uptake of tracer at sites with hepatic metastases



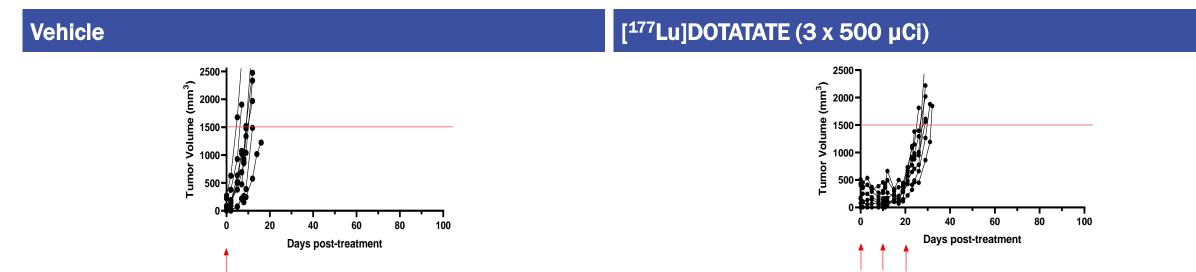
1<sup>st</sup> Patient with [<sup>212</sup>Pb]VMT- $\alpha$ -NET – NET patient, liver metastasis.





### **Preclinical Data**

**Drug Administered** 



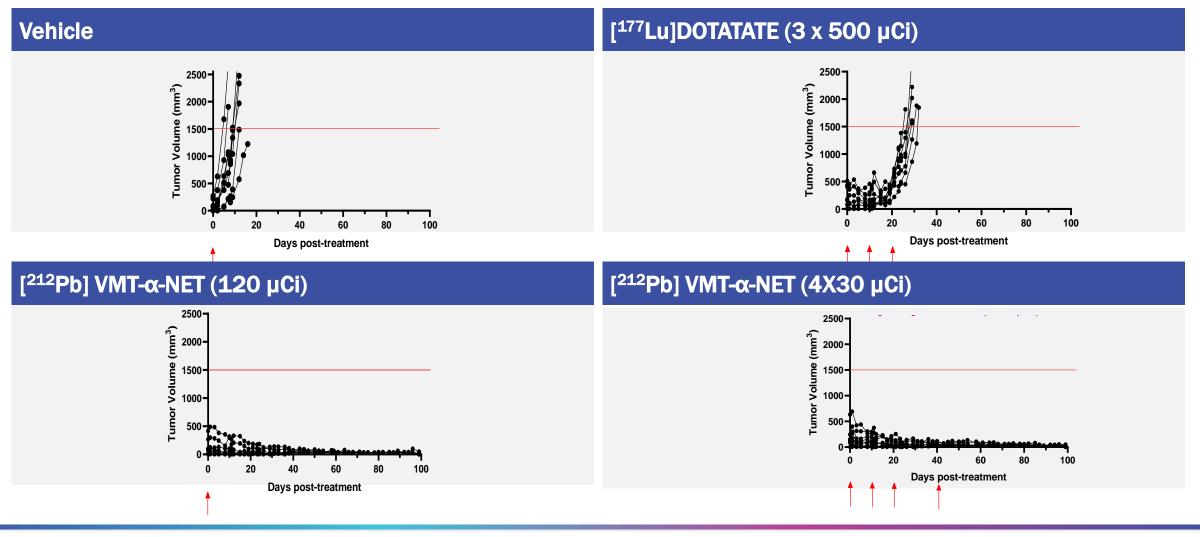
- Animal study in mice showing tumor growth curves of implanted neuroendocrine tumors
- Mice were treated with placebo vs. an FDA approved beta-emitter
- Targeted radiation shows improvements over placebo



### **Preclinical Data**

VMT- $\alpha$ -NET shows improvements with single or multiple doses

**Drug Administered** 

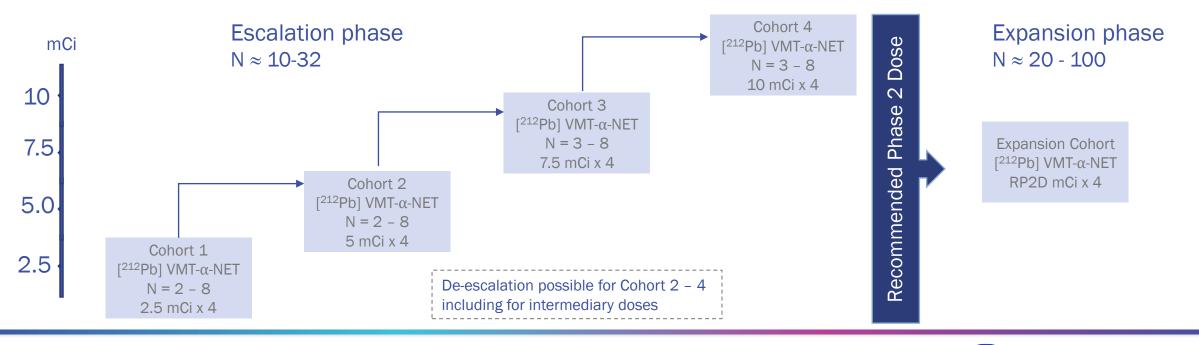




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# *Phase I/IIa* [<sup>212</sup>*Pb*]*VMT-α-NET mTPI* [*NCT05636618*]

Primary Objective	To determine the MTD/MFD of [ <sup>212</sup> Pb]VMT- $\alpha$ -NET (RP2D)	Imaging	FDA approved SSTR2 PET/CT
Population	<ul> <li>Unresectable or metastatic SSTR2-positive neuroendocrine tumors</li> <li>PRRT naïve</li> </ul>	Therapeutic Dose	2.5–10 mCi dose escalation with fixed dosing every 8 weeks for up to 4 cycles
Escalation n≈10-32 Expansion n≈20 - 100		Estimated Time to Primary Completion	~18 months
Design Methodology	Bayesian mTPI2 based on iterative toxicity probability monitoring	Dosimetry	To be assessed during screening for cohorts 1 & 2 using 5-7 mCi therapeutic surrogate[ <sup>203</sup> Pb]VMT- $\alpha$ -NET





### *Clinical Development Plan Data expected 2023*

### VMT- $\alpha$ -NET All neuroendocrine tumor types

Compassionate Image-Guided Therapy Phase I/IIa Dose Escalation Study **Provisional Results Expected in 2023** Expected to Commence Enrollment Q2 2023 **Clinical Sites:** Multiple global sites Clinical Site: Multiple in US; Mayo, UI and UW + **Subjects:** Up to 7 with refractory NETs Subjects: ~30; refractory ALL NETs, radiotherapy naive **Phase:** 0/1 First-in-human, compassionate use **Phase:** Phase I/IIa; safety, mTPI-2\* dose-ranging **Endpoints:** Safety and tumor targeting by imaging **Endpoints:** Safety and tumor targeting by imaging **Secondary:** Efficacy signal by serial imaging **Secondary:** Efficacy signal by serial imaging **Status:** 3 subjects dosed with  $[^{203/212}Pb]VMT- \alpha$ -NET **Status:** IND given "safe to proceed" from FDA



# Melanoma Program: VMT01/02

Using the melanocortin receptor MC1R to target melanoma for imaging and therapy

## *VMT01/02*

### **Key Facts**



Targeting melanocortin 1 receptor (MC1R)

Currently in Phase 1/2a studies for the imaging and treatment of metastatic melanoma



Preclinical combination data (published) resulted in \$2m NIH SBIR Grant



Provisional results from completed Phase 1 imaging study expected H1 2023 Studies being conducted at the Mayo Clinic Rochester

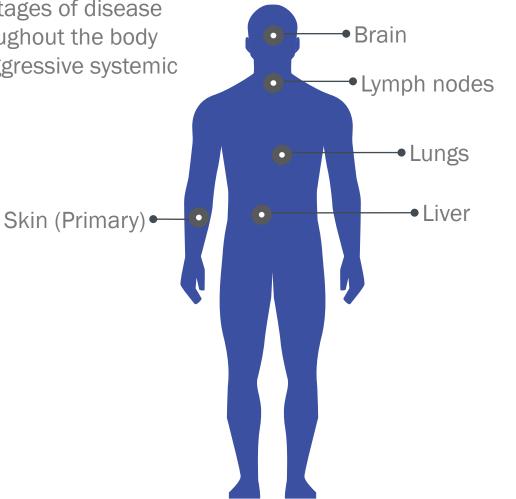


Pending Orphan Drug Designation Application



### Metastatic Melanoma

Advanced stages of disease occurs throughout the body requiring aggressive systemic treatment



Significant unmet need with an \$8 billion market opportunity<sup>1</sup>

~100K new diagnoses of melanoma annually<sup>2</sup>

~6,850 deaths annually from metastatic melanoma<sup>2</sup>

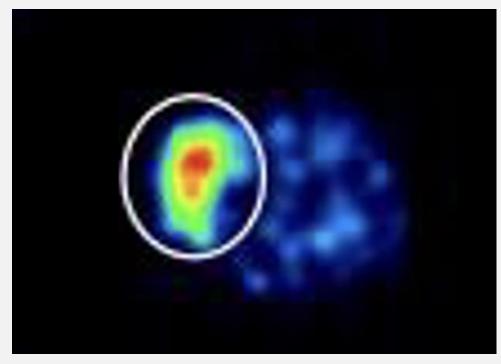
In most cases, can't be cured but treatment can support a longer life



## [<sup>203</sup>Pb]VMT-01 Gives Clear Diagnostic Images in Melanoma

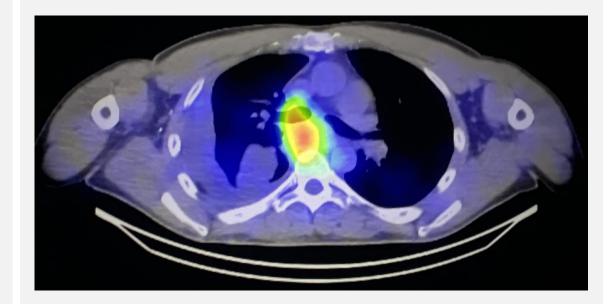
#### Mouse Melanoma

High intensity in implanted tumor above background



#### Human Melanoma

High intensity in esophageal tumor above background



#### Preclinical microSPECT

Whole body planar SPECT/CT



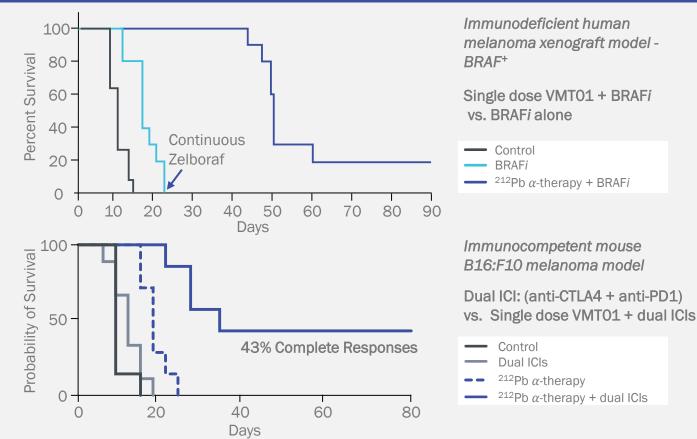
### Demonstrated Complete Responses in Multiple Animal Tumor Models

## VMT01/02

#### Key Takeaways

- High response rates in multiple tested models
- 43% complete and durable response if combined with immunotherapy in a highly resistant model
- Combination with immune checkpoint inhibitors induced synergistic antitumor effect

Li et al., Mol. Pharm., Sep 3;16(9), 2019 Li et al., Cancers, Jul 22;13(15), 2021 Single dose of VMT01 significantly arrested melanoma tumor growth and extended survival





# *Phase I/IIa* [<sup>212</sup>*Pb*]*VMT01-T101 mTPI* [*NCT05655312*]

Primary Objective	To determine the MTD/MFD of [ <sup>212</sup> Pb]VMT01-T101 (RP2D)	Imaging	[ <sup>203</sup> Pb]VMT01 SPEC/CT or [ <sup>68</sup> Ga]VMT02 PET/CT 3 – 15 mCi dose escalation with fixed dosing every 8 weeks for up to 3 cycles ~18 months		
Population		Therapeutic Dose			
Screen 120 subjects Enroll 52 subjects	Unresectable or metastatic MC1R-positive Melanoma	Estimated Time to Primary Completion			
Design Methodology         Bayesian mTPI2 based on iterative toxicity probability monitoring		Dosimetry	To be assessed using 15 - 25 mCi therapeutic surrogate [ <sup>203</sup> Pb]VMT01		
N ≈ 32 10- 5- 3 Cohort 1	Cohort 2 Cohort 1 $[^{212}Pb] VMT01-T101$ N = 2 - 8 Cohort 1 $[^{212}Pb] VMT01-T101$ N = 2 - 8 De-escalation possible		Expansion phase N ≈ 20 Expansion Cohort [ <sup>212</sup> Pb] VMT01-T101 RP2D mCi x 3		



## *Clinical Development Plan Data expected in 2023*

# VMT01 Metastatic Melanoma

Completed Phase 1 Imaging Study	Phase I/IIa Dose Escalation Study		
Provisional Results Expected in 2023	Expected to Commence Enrollment Q2 2023		
Clinical Site: Mayo Clinic (Rochester, MN)	Clinical Site: 5 US sites +/- expansion to 15		
Subjects: 7; Stage III/IV unresectable melanoma	Subjects: ~30; Stage III/IV progressive melanoma		
Phase: Phase 1 First-in-Human, cross-over design	Phase: Phase I/IIa; safety, mTPI-2* dose-ranging		
Endpoints: Safety and biodistribution	Endpoints: Safety and tumor targeting by imaging		
Secondary: Molecular target validation, image quality	Secondary: Efficacy signal by RECIST 1.1		
Status: Closed for enrollment, IND open	Status: IND safe to proceed		



# Supporting Program: VMT-α-GEN

Controlling the Therapeutic Isotope Supply

## <sup>212</sup>Pb Supply via Reusable Desktop Isotope Generator

### VMT-α-GEN

- "Limitless" feedstock from nuclear and mining waste material
- Long-term supply contract secured (US DOE)
- On demand daily doses
  - Auto-regenerates overnight
  - $\simeq 1$  week shelf life

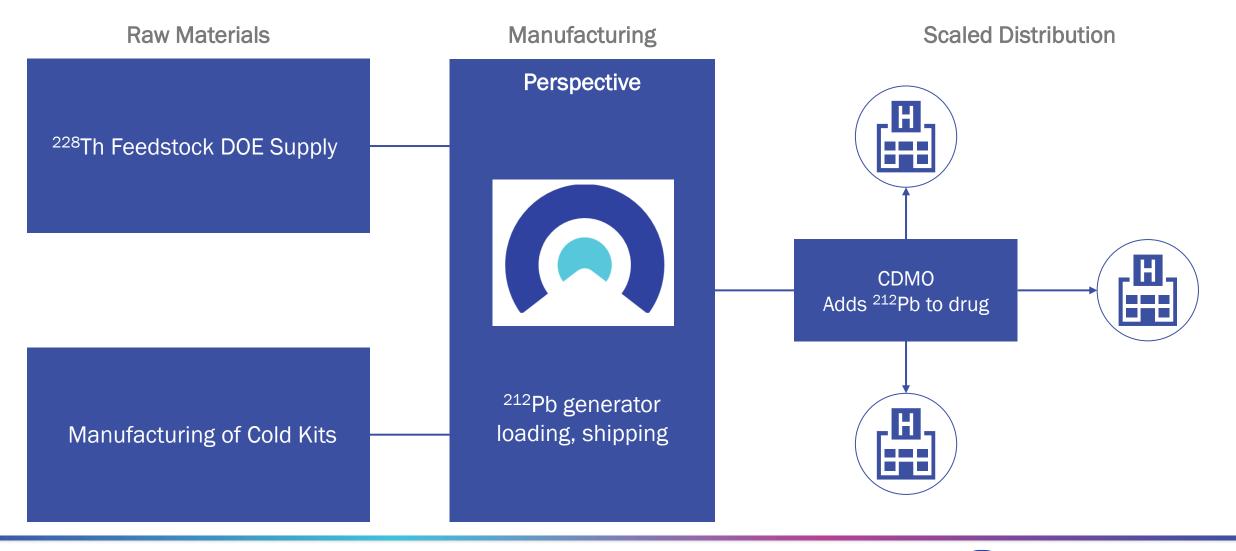


### Small, Elegant <sup>212</sup>Pb Isotope Generator

- Integrated lead shielded containment
- Simple inlet and outlet ports
- Radioactive feedstock for nearly 300 generators fits in a small vial



## Controlling the Therapeutic Isotope Supply





# **Corporate Overview**

## Intellectual Property

### Fully Licensed University IP

3 provisional patents

- Composition of Matter and Use VMT- $\alpha\text{-NET},$  Chelator
  - USA, EU, Australia
- 3 issued patents
- Composition of matter and use on melanoma targeting peptides
  - USA
- Composition of Matter and Use VMT01
  - USA, EU, Australia



Potential for Orphan Drug Designation



### Potential for U.S. FDA Priority Review Voucher

 VMT-α-NET is a candidate for pediatric neuroblastoma indication



### **Clinical Trial Principal Investigators**

Renowned Experts in Radiotherapy Development



Geoffrey B. Johnson MD, PhD Chair, Division of Nuclear Medicine

Chair, PET/MR R&D Associate Professor Departments of Radiology and Immunology Mayo Clinic – Rochester, MN



Yusuf Menda MD Professor of Radiology

# IOWA

Chair, Division of Nuclear Medicine Project Leader Neuroendocrine Tumor SPORE University of Iowa – Iowa City, IA



Vikas Prasad MDWashington University in St. LouisProfessor of Radiology

MAYO CLINIC

Associate Professor Radiology, Division of Nuclear Medicine

Washington University in St Louis – St Louis, MO



## *Clinical Trial Principal Investigators: Renowned Experts in Radiotherapy Development*



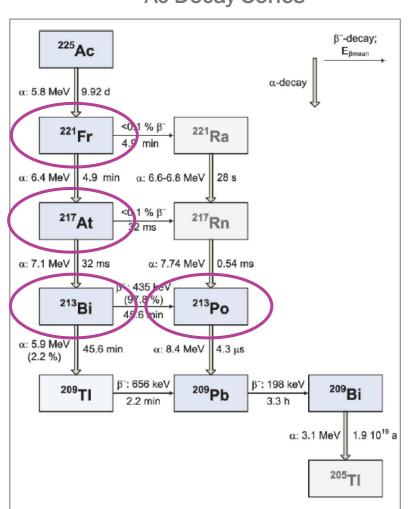




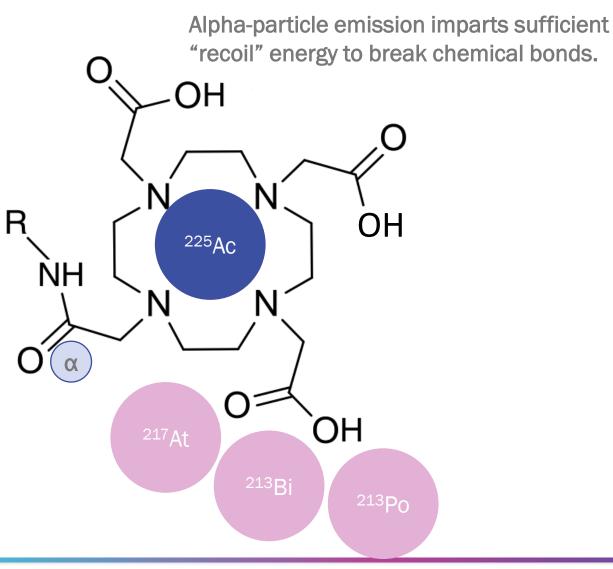
# Appendix



### *Isotope: Decay chain – Biological implications*



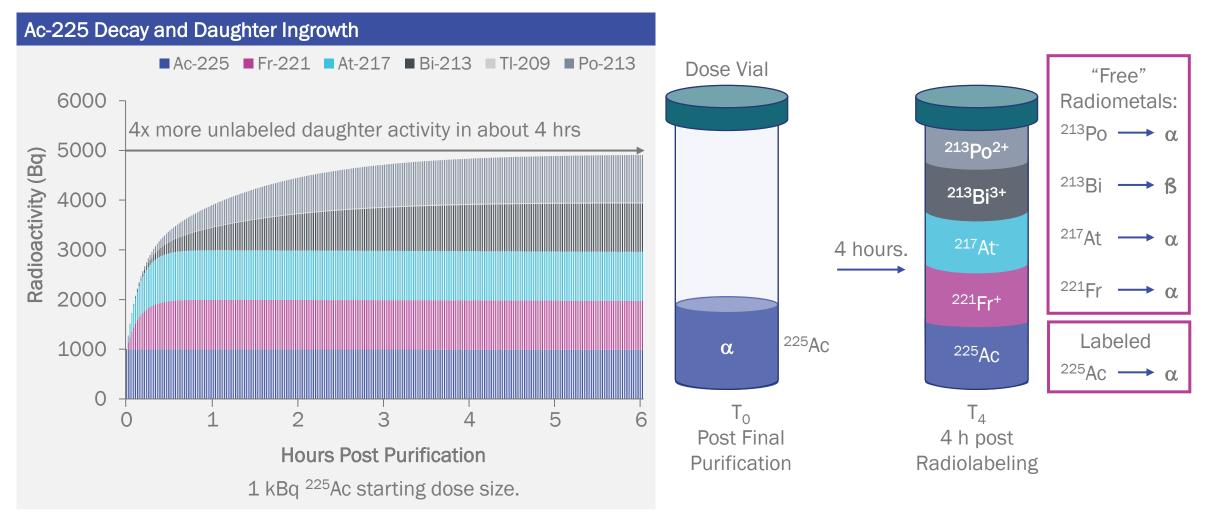
<sup>225</sup>Ac Decay Series





### *Isotope: Decay chain – Product implications*

Post final radiolabeling and purification, alpha and beta emitting daughters of <sup>225</sup>Ac build up fast.





### Isotope: Decay chain and chelator selection

<sup>212</sup>Pb Decay Series

