



**PERSPECTIVE**  
THERAPEUTICS

**$^{203}\text{Pb}/^{212}\text{Pb}$  Image-guided alpha-particle therapy for cancer. Progress and Metrology Challenges**

February 2024

NYSE: CATX

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Forward-looking statements contained in this presentation are made as of this date, and the Company undertakes no duty to update such information whether as a result of new information, future events or otherwise, except as required under applicable law.

# Radiopharmaceuticals are a Pillar of Oncology Treatment

Unique Mechanism of Action Offers Pan-Cancer Opportunities

## Molecularly Targeted Radiation

Radioligands can precisely deliver radiation directly to cancer cells reducing off-target effects  
Proven pillar of cancer treatment

**Perspective's platform technology is optimized for greater efficacy and fewer side effects**

## Optimized Patient Selection

Molecular imaging companion diagnostics enable visualization of the therapeutic target  
Enables the selection of patients who may best respond to therapy

**Perspective's elementally matched isotopes are paired for imaging and therapy**

## Monotherapy Activity and Combination Synergies

Ability for both monotherapy and combination treatments  
Potential synergies with DNA damage response and immune checkpoint inhibitors

**Perspective's targeted alpha therapy delivers potent and immunostimulatory radiation to tumor**

## Outpatient Friendly

Modern medical isotopes enable radiopharmaceuticals to be administered outside of hospitals  
Treatments are easily-accessible globally with several hundred therapeutic locations in the U.S alone  
Perspective's short half-life isotopes simplify patient administration and waste management

## Unique Business Opportunity

Radiopharmaceutical theranostic product development is highly-specialized and technical  
Greater expertise needed than for standard medicines potentially creating higher barriers to entry  
**Perspective develops patent-protected best-in-class intellectual property**

# $\alpha$ -Particles Have Superior Tumor Killing Properties vs. $\beta$ -Particles

More Powerful Effects Than Approved  $\beta$  Therapy

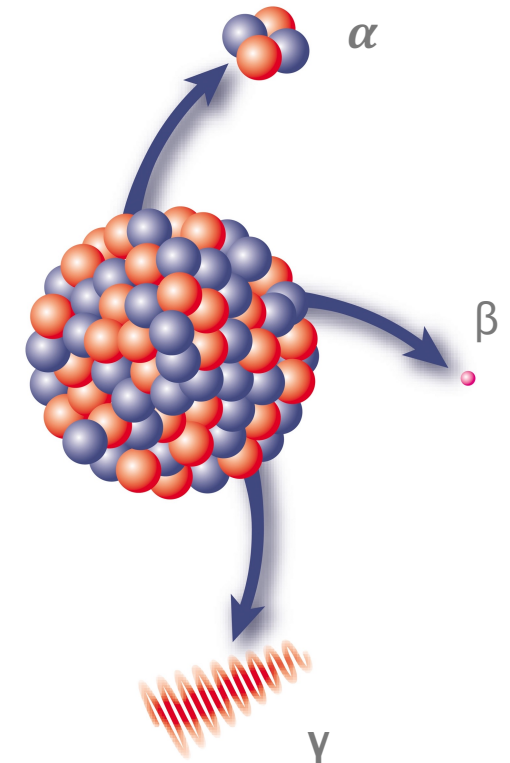
Higher atomic mass  
Lethal double-stranded DNA breaks  
DNA repair mechanisms overwhelmed

Precision Delivery Provides Targeted Cell Destruction

Deposit energy over 3-5 cell diameters vs. beta particles (up to 200 cells)

Anti-Tumor Immune Response<sup>1</sup>

Evidence for antitumor response alone or in combination with immunotherapies  
Consistent with “Abscopal effect” observed with external beam radiation therapy



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

# Management Team

Deep Experience in Radiopharmaceuticals and Oncology Drug Development



**Thijs Spoor**

Chief Executive Officer

20+ years of expertise in biotechnology companies; public and private companies; oncology and nuclear pharmacy



**Markus Puhlmann, MD MBA**

Chief Medical Officer

20+ years of oncology drug development across all phases, experience coordinating multiple regulatory filings



**Michael Schultz, PHD**

Co-Founder and Chief Science Officer

20+ years industry and research experience in radiopharmaceuticals; inventor of Perspective radiopharmaceutical products



**Jonathan Hunt**

Chief Financial Officer

20+ years of expertise in financial controls and public accounting for large and small companies across multiple industries



**Frances Johnson, MD**

Co-Founder and Chief Innovation Officer

20+ years in clinical trials execution, managing academic research programs, and start-up of CareDx, Inc.



**Amos Hedt**

Chief Business Strategy Officer

20+ years of expertise in early-stage pharmaceutical and biotech drug development; 10+ years in radiopharmaceuticals

# Platform Expansion Engine

Two Lead Programs in Clinic and Broad Proprietary Pipeline

| Program                   | Indication                        | Discovery      | Human Clinical Imaging | First in Human Therapy | Phase 1/2 | Phase 3 |
|---------------------------|-----------------------------------|----------------|------------------------|------------------------|-----------|---------|
| VMT- $\alpha$ -NET        | Neuroendocrine cancer             | [Progress bar] |                        |                        |           |         |
|                           | Pheochromocytomas, paragangliomas | [Progress bar] |                        |                        |           |         |
|                           | Small cell lung cancer            | [Progress bar] |                        |                        |           |         |
| VMT01                     | Melanoma (MC1R)                   | [Progress bar] |                        |                        |           |         |
| VMT02 (PET agent)         | Melanoma (imaging of MC1R)        | [Progress bar] |                        |                        |           |         |
| Program 3 (Novel peptide) | Multiple solid tumors             | [Progress bar] |                        |                        |           |         |
| PSV401 (Radio-hybrid)     | Prostate (PSMA imaging & therapy) | [Progress bar] |                        |                        |           |         |
| Program 5 (Novel peptide) | Prostate, Breast                  | [Progress bar] |                        |                        |           |         |
| Other Programs            | Solid and hematological tumors    | [Progress bar] |                        |                        |           |         |

# Lead-212 ( $^{212}\text{Pb}$ ): The Optimal Therapeutic Isotope

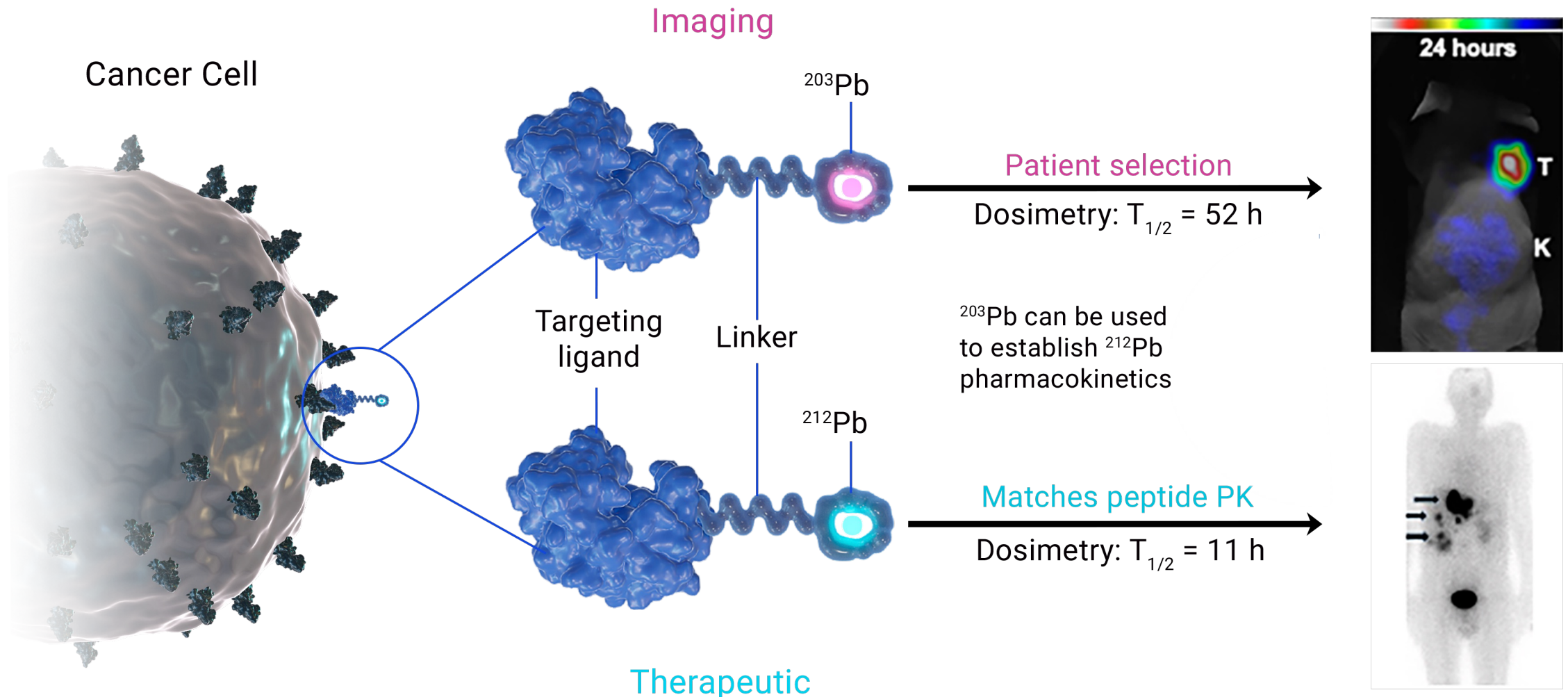
Alpha Particles Provide Numerous Benefits Over Currently Used Beta Particle Radiotherapies

- With a much higher atomic mass, **alpha ( $\alpha$ )** particles generate more energy and travel a shorter distance compared to beta ( $\beta$ ) particles, making them more cytotoxic, while reducing their off-targeting effects on healthy tissue
- Alpha radiation causes direct lethal double-stranded DNA breaks, vs indirect single-stranded breaks in beta ( $\beta$ ) radiation
- Cell death expected – NO resistance
- Greater therapeutic efficacy expected to improve outcomes with better safety

|                          | Lead ( $^{212}\text{Pb}$ ) | Iodine ( $^{131}\text{I}$ ) | Lutetium ( $^{177}\text{Lu}$ ) | Actinium ( $^{225}\text{Ac}$ ) | Implication <sup>1</sup> |
|--------------------------|----------------------------|-----------------------------|--------------------------------|--------------------------------|--------------------------|
| Emission Profile         | Alpha                      | Beta                        | Beta                           | Alpha                          | Potent                   |
| Half Life                | 0.46 days                  | 8 days                      | 6.7 days                       | 10 days                        | High dose-rate           |
| Off Target Toxicity Risk | Low                        | Very high                   | Low                            | High                           | Best                     |
| Supply                   | High                       | High                        | Low                            | Low                            | Abundant                 |
| Cost of Production       | Low                        | Low                         | High                           | High                           | High margin              |

# Pb-based Theranostics Enable Both Diagnosis and Targeted Treatment of Cancer

Identical Distribution of  $^{203}\text{Pb}$  and  $^{212}\text{Pb}$  for Imaging and Treatment, Respectively





# Neuroendocrine Tumors: VMT- $\alpha$ -NET

Targeting the somatostatin receptor to treat rare neuroendocrine-type cancers

# VMT- $\alpha$ -NET Currently in Phase 1/2a Studies: Key Facts



Targeting somatostatin receptor type 2 (SSTR2) for the imaging and treatment of neuroendocrine tumors

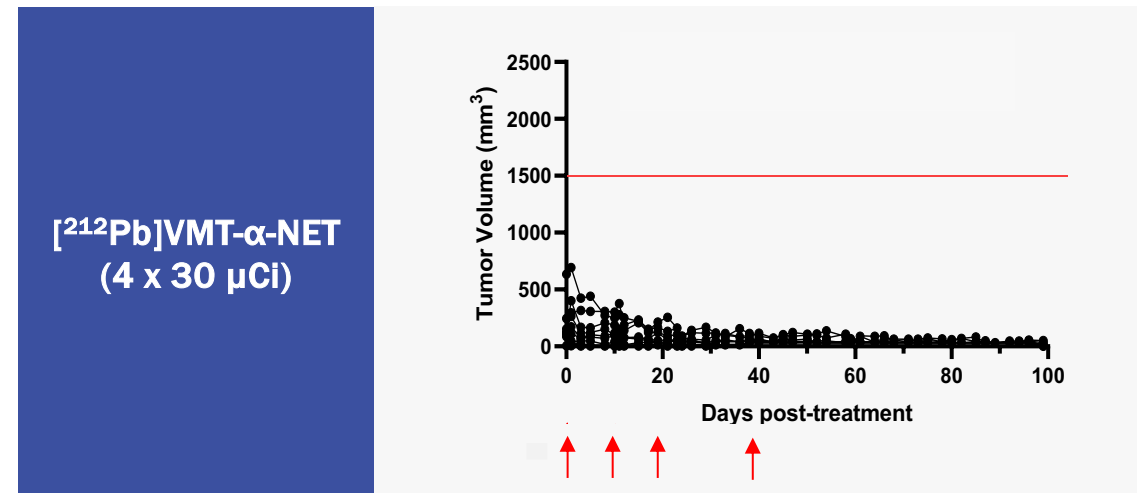
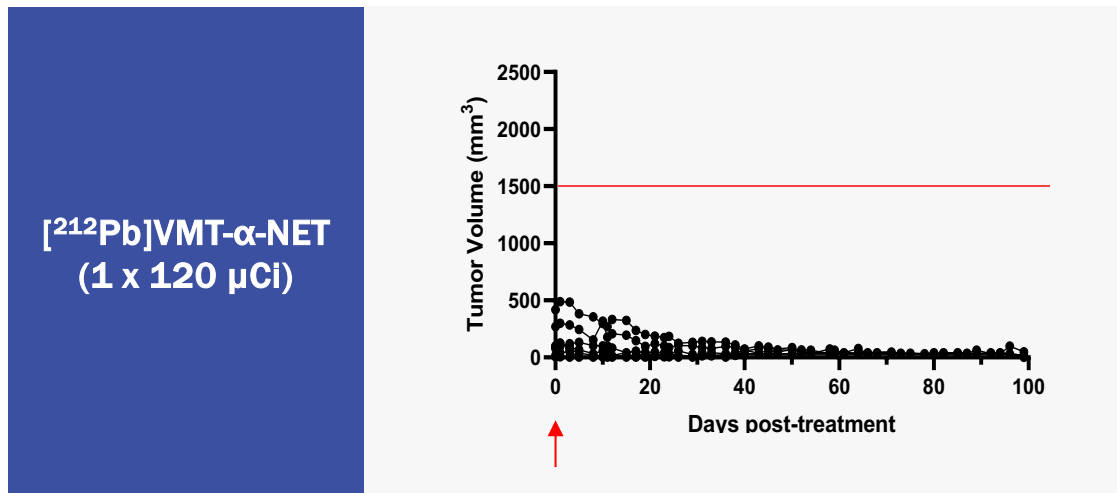
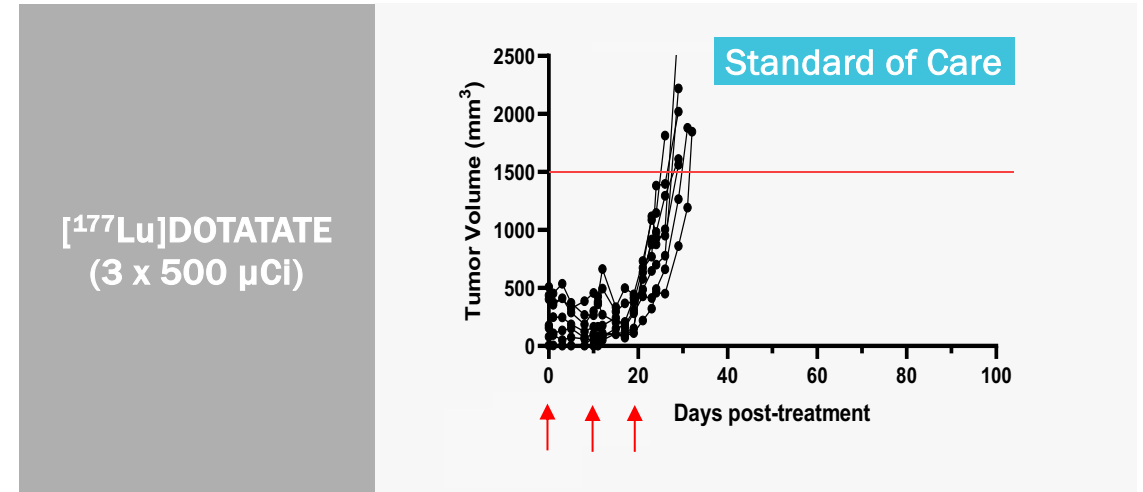
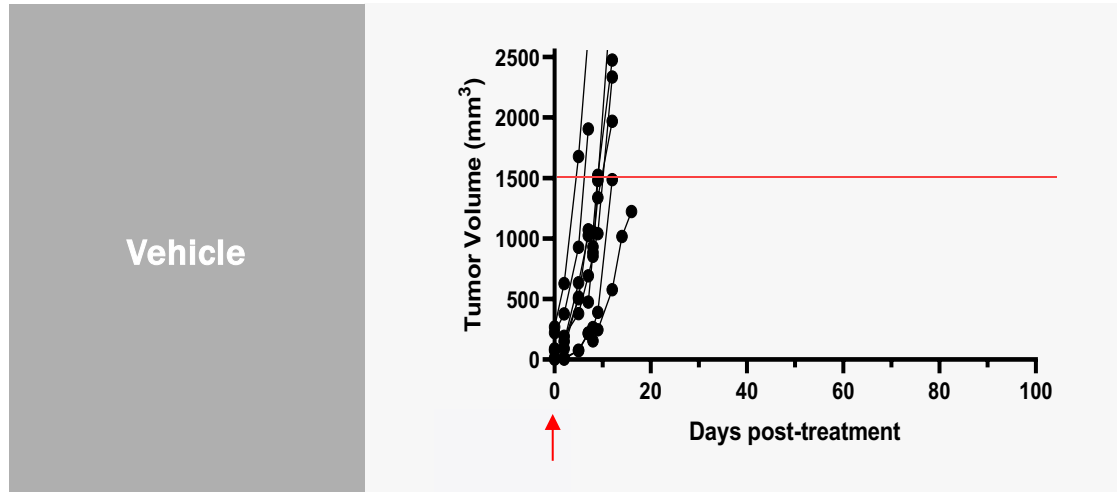
Initiated first-in-human imaging (2021) & therapy (2022) under compassionate use

Fast Track Designation for first line therapy received October 2022  
Therapeutic Trial in first line setting currently recruiting under open IND

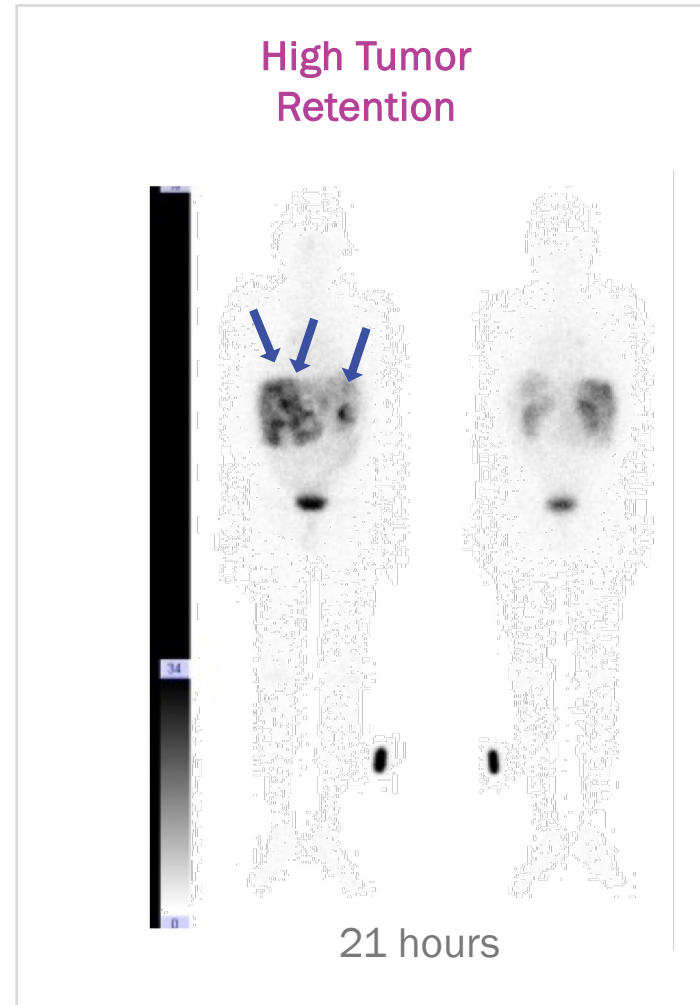
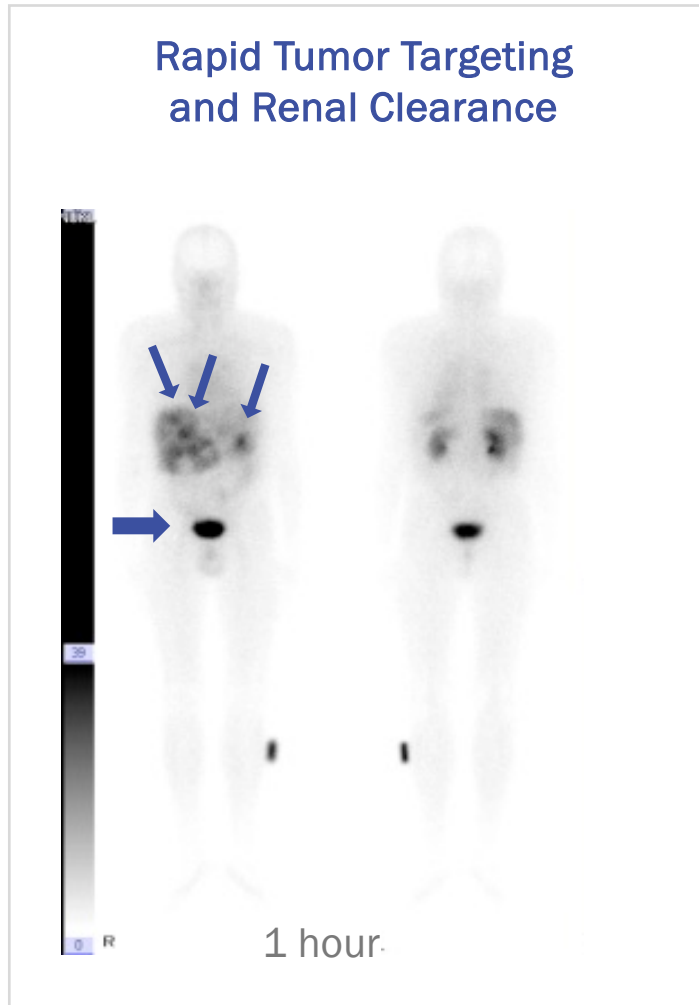
US Phase 1 prospective dosimetry study in PRRT refractory patients recruiting at the University of Iowa

# VMT- $\alpha$ -NET Shows Significant Improvement vs Standard of Care in Preclinical Models

Superior Efficacy with Single Dose or Multiple Administrations



# $^{203}\text{Pb}$ SPECT Imaging Reveals Favorable VMT- $\alpha$ -NET Properties<sup>1</sup>

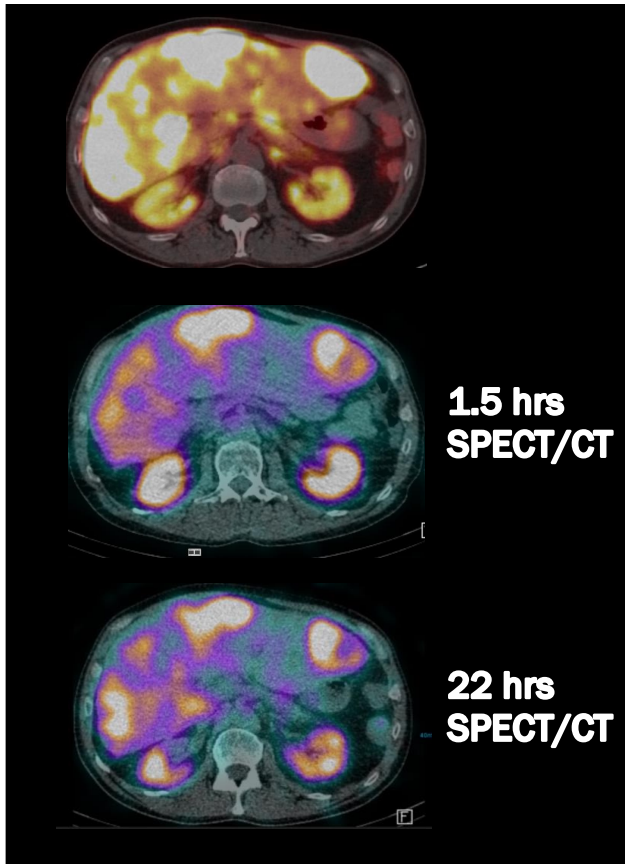


- Tumors visible within 1 hour indicates rapid binding to SSTR2 target
- High intensity above background implies excellent therapeutic window
- Unbound drug in bladder within 1 hour for excretion
- Low renal retention due to neutral charge on proprietary Pb-specific chelator

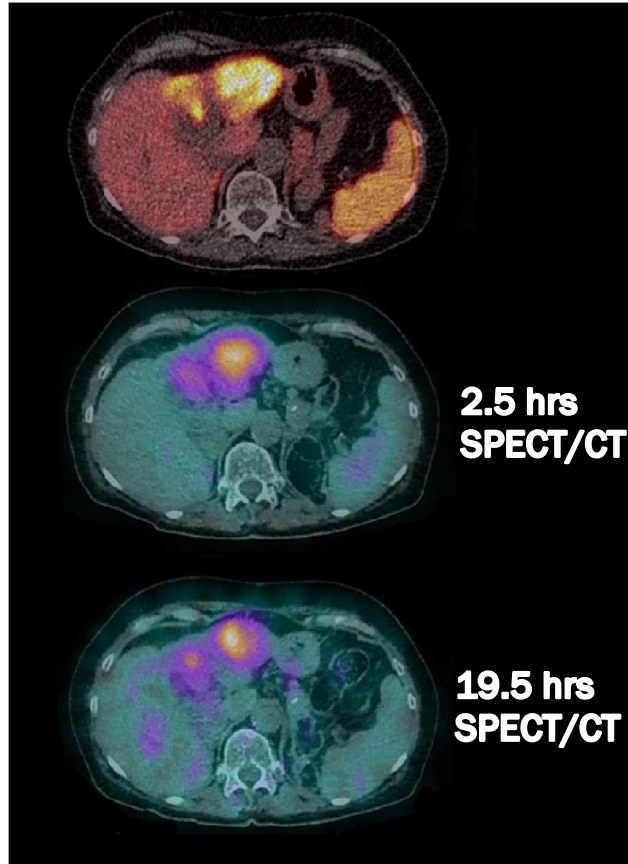
# $^{212}\text{Pb}$ SPECT/CT Imaging Confirms VMT- $\alpha$ -NET Tumor Uptake

Diagnostic and Therapeutic Show Same Uptake and Retention Characteristics

$^{203}\text{Pb}$  SPECT/CT Imaging<sup>1</sup>  
Pt#001



$^{212}\text{Pb}$  SPECT/CT Imaging<sup>2</sup>  
Pt#009

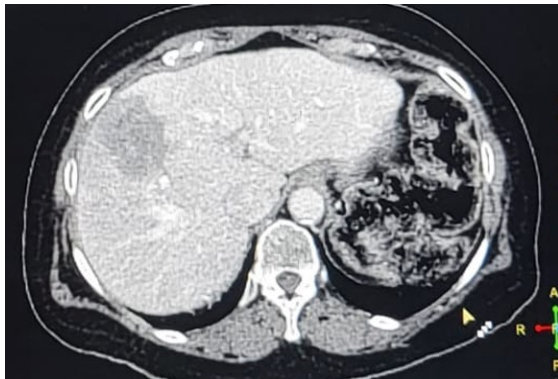
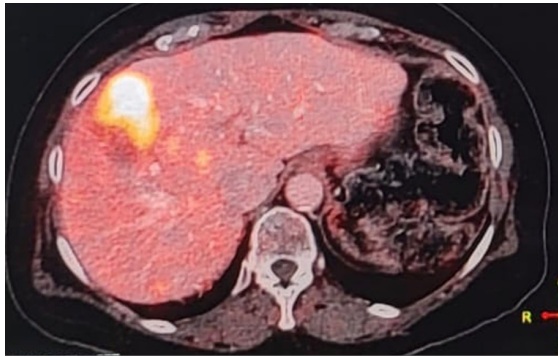


- Both  $^{203}\text{Pb}$  and  $^{212}\text{Pb}$  can be imaged directly using SPECT
- SPECT/CT shows very rapid tumor uptake and retention of [ $^{212}\text{Pb}$ ]VMT- $\alpha$ -NET
- After 24 hours more than 80% of alpha particles will be generated
- This high alpha dose rate is ideally matched to the biological clearance of the VMT- $\alpha$ -NET peptide

# Almost Complete Response After 3 Doses of [<sup>212</sup>Pb]VMT-α-NET

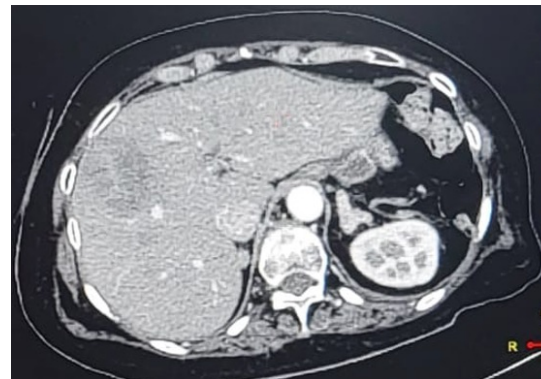
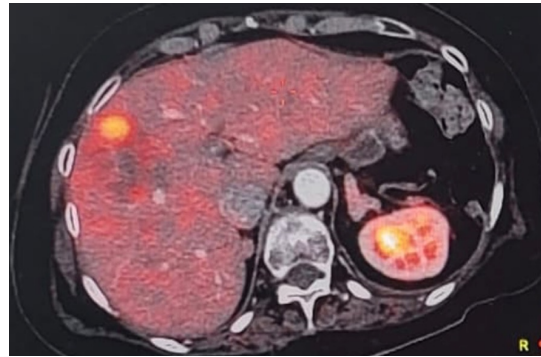
Metastatic NET Pancreas with Adrenal Crisis – PET/CT

Tumor Before Treatment



(S.ACTH)<sup>1</sup>– 790 pg/ml

Tumor After 1 Dose



Tumor After 3 Doses



S.ACTH – 96 pg/ml



# $^{212}\text{Pb}$ is Plentiful, Storable, Scalable & Suitable for Distributed Logistics

The supply chain is lower-risk and more robust than other therapeutic isotopes

## Isotope Source



Naturally occurring in mining waste  
Also produced in industrial nuclear processes  
Can be made on demand if needed

All other therapeutic isotopes require capital-intensive infrastructure manufacturing processes (irradiation)

## Isotope Purification



Parent isotope Thorium-228 can be stored (2 yr half-life)  
 $^{212}\text{Pb}$  purified from  $^{228}\text{Th}$  or  $^{224}\text{Ra}$  source in simple separation step

VMT- $\alpha$ -GEN enables shipping of isotope and purification of  $^{212}\text{Pb}$  in one package

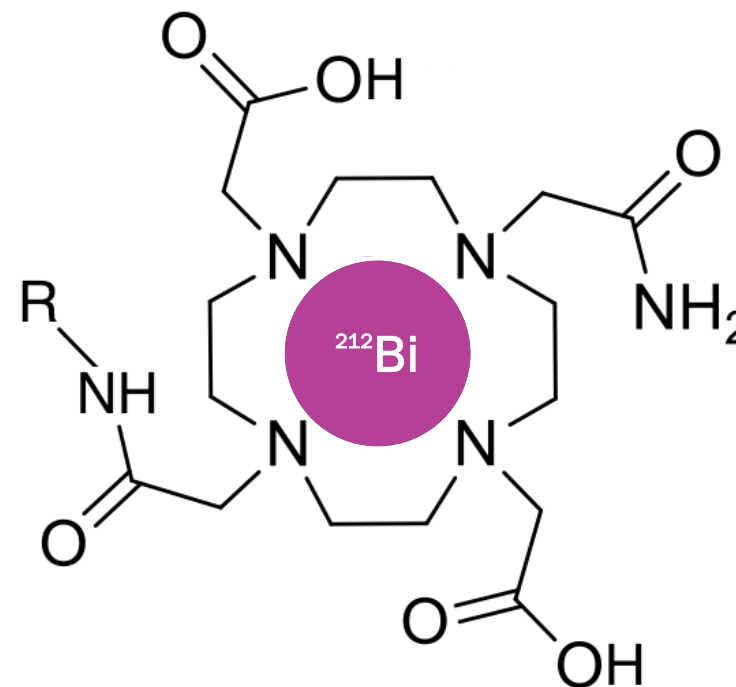
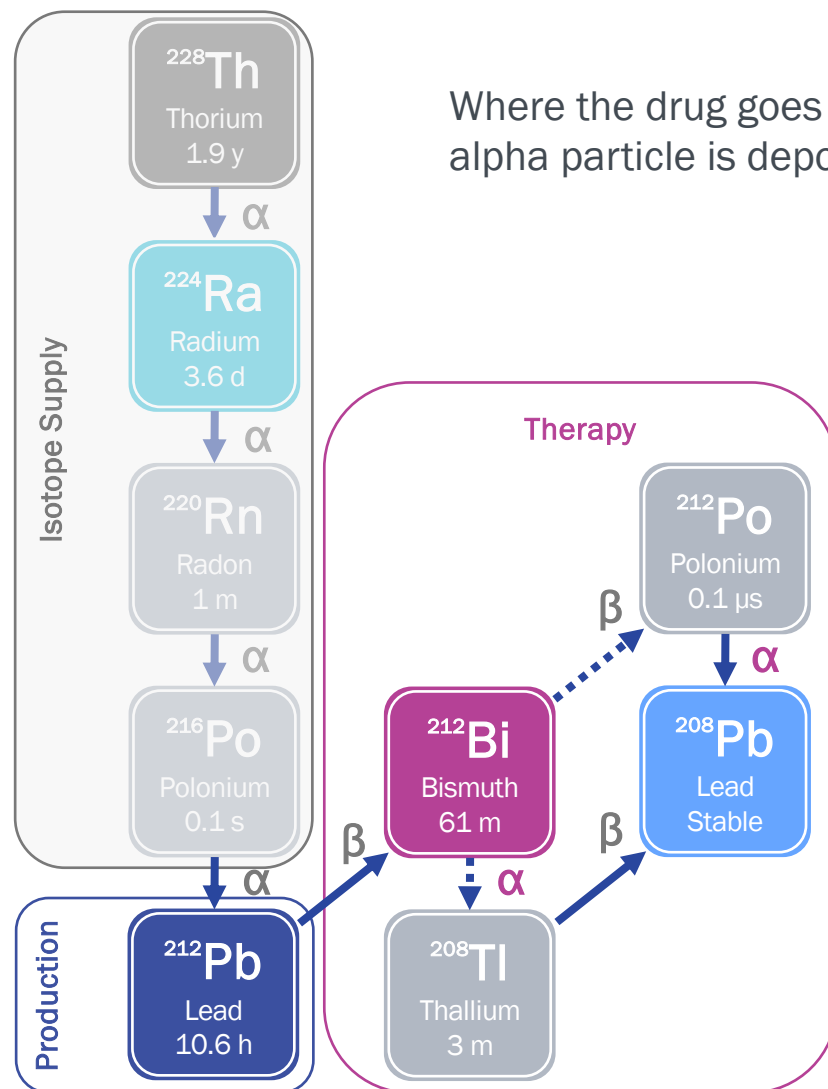
## Product Manufacturing



VMT- $\alpha$ -GEN  $^{212}\text{Pb}$  generator technology scales for commercial production  
Extremely pure isotope allows straight forward manufacturing process

10.5 hr half life of  $^{212}\text{Pb}$  allows for robust regional distribution of finished radiopharmaceuticals

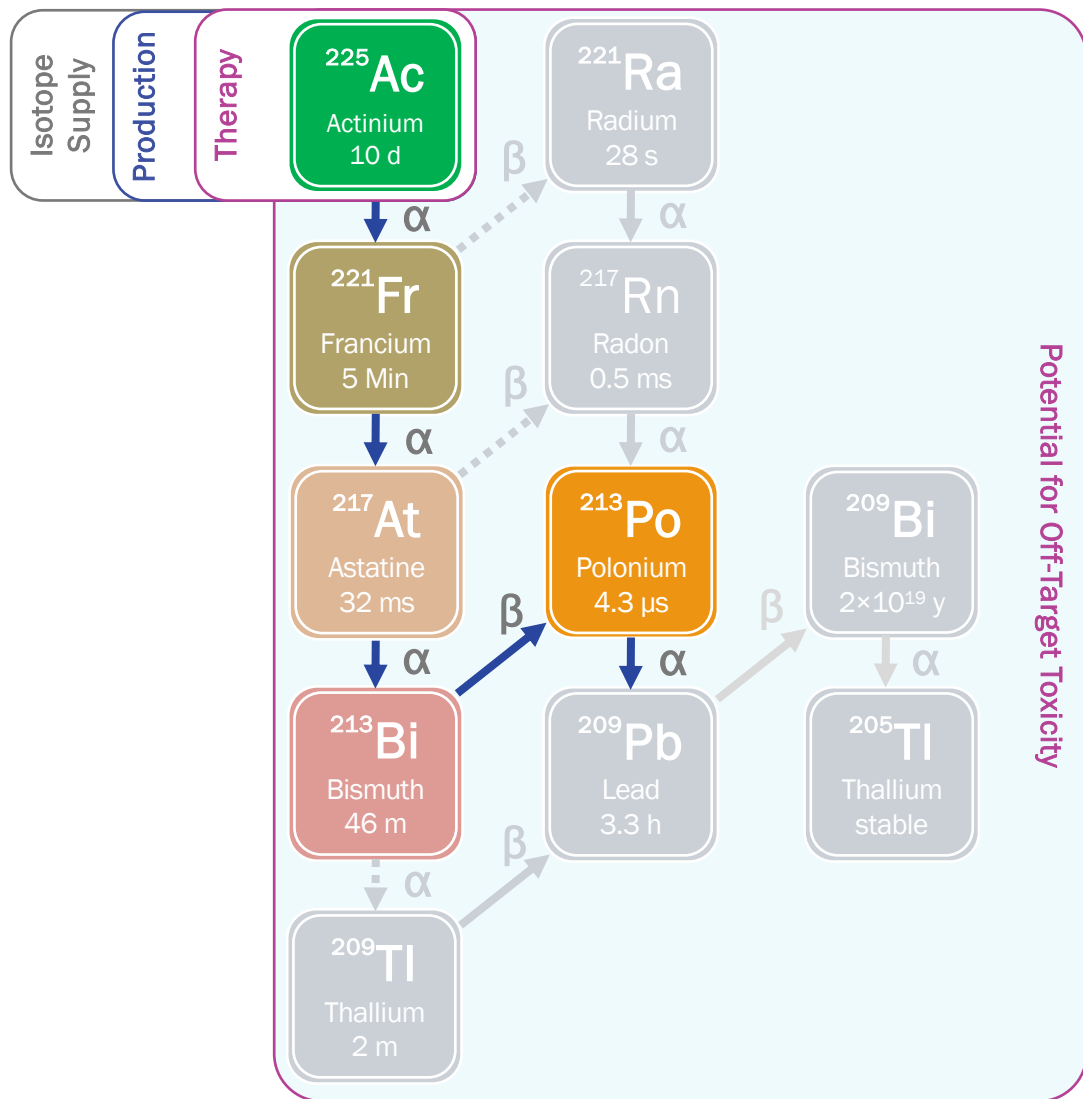
# $^{212}\text{Pb}$ Isotope Decay Chain and Importance of the Pb-Specific Chelator



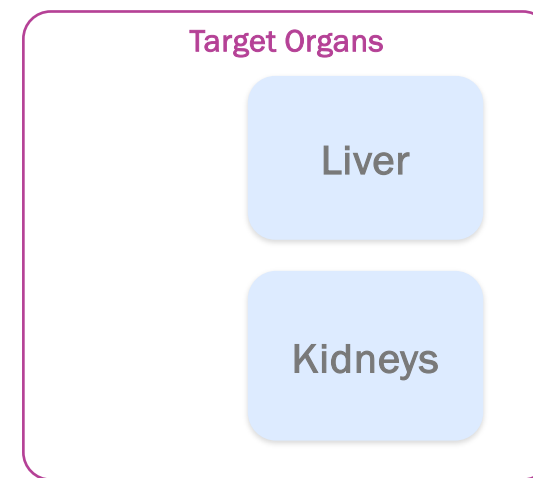
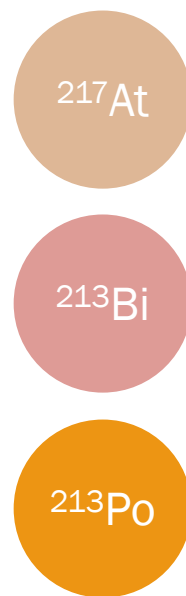
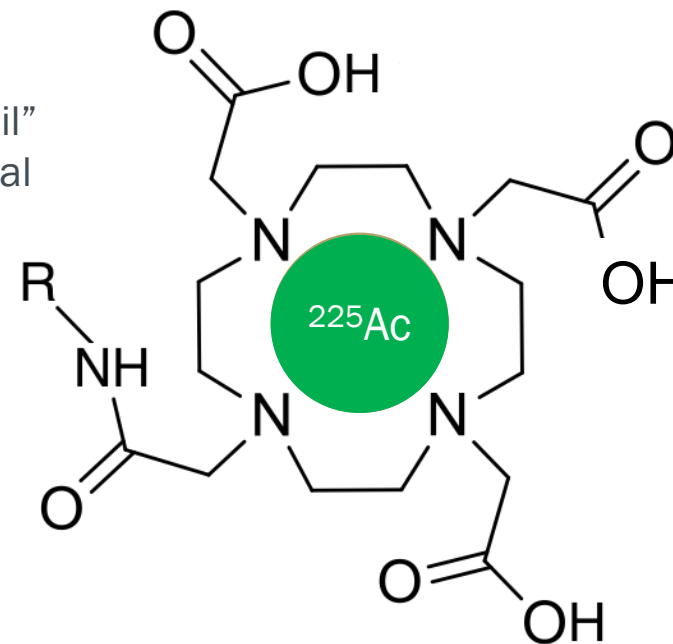
- Perspective's proprietary chelator retains 98% of  $^{212}\text{Bi}$  after transition in drug formulation
- Generic chelators leak the  $^{212}\text{Bi}$  alpha-emitting daughter up to 36%<sup>1</sup>



# $^{225}\text{Ac}$ Isotope Decay Chain and Potential for Off-Target Toxicity



Alpha-particle emission imparts sufficient “recoil” energy to break chemical bonds



# Summary

Thank you!

- **Cancer targeted alpha-particle radionuclide therapy for cancer emerging as a potent approach to cancer treatment**
- **$^{212}\text{Pb}$  has ideal properties for cancer therapy**
- **Imaging of  $^{203}\text{Pb}$  and  $^{212}\text{Pb}$  are powerful tools for radiopharmaceutical development and patient care**
- **Decay series of alpha-particle emitters has presented a measurement challenge to the metrology community**