



# Fusion & AstraZeneca Strategic Collaboration

To Develop and Commercialize Next-Generation Targeted Alpha  
Therapy Radiopharmaceuticals and Combination Therapies

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Chief Scientific Officer



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**John Valliant, PhD**  
Chief Executive Officer



# Collaboration Overview and Rationale







## 1) Novel Targeted Alpha Therapies

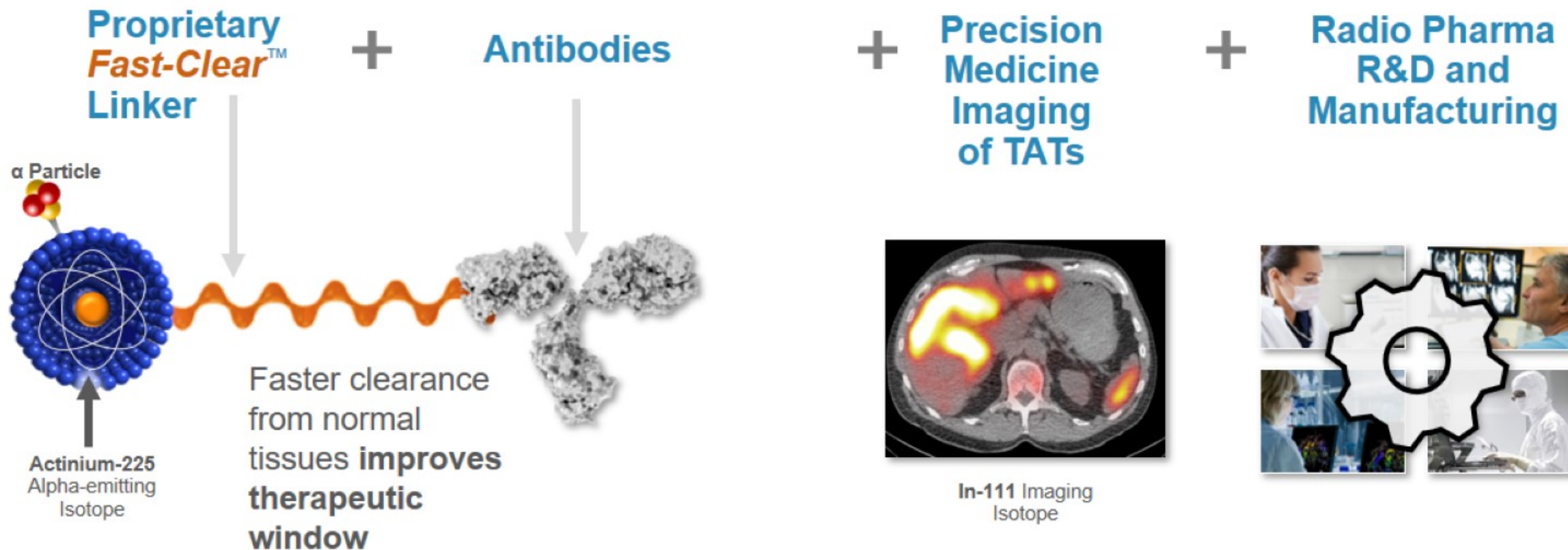
- Jointly select up to three new TATs
- Co-fund
- Co-develop
- Option to co-commercialize in U.S.

## 2) Combination Therapies with TATs

- DNA Damage Response Inhibitors (DDRIs)
- Immuno-Oncology Agents
- AZ solely funds unless Fusion opts-in



## FUSION'S TAT PLATFORM



Combining Fusion's Platform with AZ's Antibody & DDRi Portfolio





- Delivers access to breadth of AstraZeneca's world-class oncology pipeline & products, resources and expertise
- Avoids early-stage development risk
- Provides optionality around later-stage risk and expense
- Preserves significant back-end economics
- Maintains all rights to existing Fusion pipeline







**Marc Schwabish, PhD**

Senior Vice President,  
Business Development and  
US Operations

# Details of the Agreement

## Therapeutic Expansion

- Maximize utilization of our TAT platform
- Maximize potential applications of Targeted Alpha radiation Therapies
- Expand Fusion's pipeline via co-development and co-commercialization



## Business Model

- Global partnership
- Long-term agreement
- Fully-retain rights to our existing products
- Retain co-commercialization rights on new TATs
- Upfront payment
- Future development milestones
- Financial flexibility minimizes near-term risk





# 1) Overview of the Terms to Develop Novel Targeted Alpha Therapies

**Up to three new TATs will be jointly selected and then...**

- Co-developed with a 50/50 R&D cost split

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Each partner can opt-out at defined breakpoints in exchange for milestone payments to the other party and a low to high single-digit royalty on future sales

- 1st → following pre-clinical activities
- 2nd → following completion of Phase 1 study

- 
- AstraZeneca and Fusion to co-commercialize
    - Fusion has option to co-promote in the U.S.
    - AstraZeneca to commercialize in ROW
    - 50/50 profit and loss share



## 2) Agreement to Develop Novel Clinical Combination Therapies

### AstraZeneca

- Nominate for preclinical evaluation up to five targets
- Fully fund all R&D (unless Fusion opts-in to co-fund clin dev)
- AZ responsible for clinical trial operations and related financial obligations

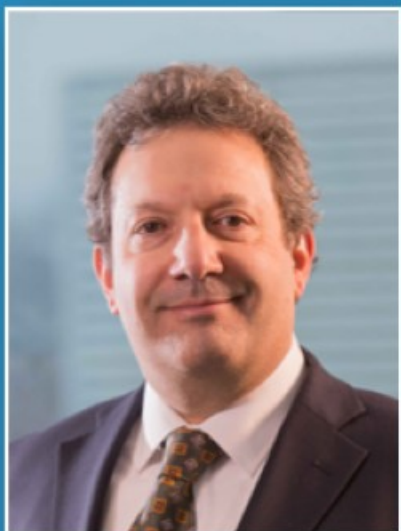
### Fusion

- Responsible for TAT drug supply
- Can opt-in to co-fund post-P1 development (50/50)
- If Fusion does not opt in:
  - Fusion obligated to pay back share of R&D expense plus risk premium

- Both companies retain full rights to their respective compounds
- Equal representation on Joint Steering and other committees







**Eric Burak, PhD**

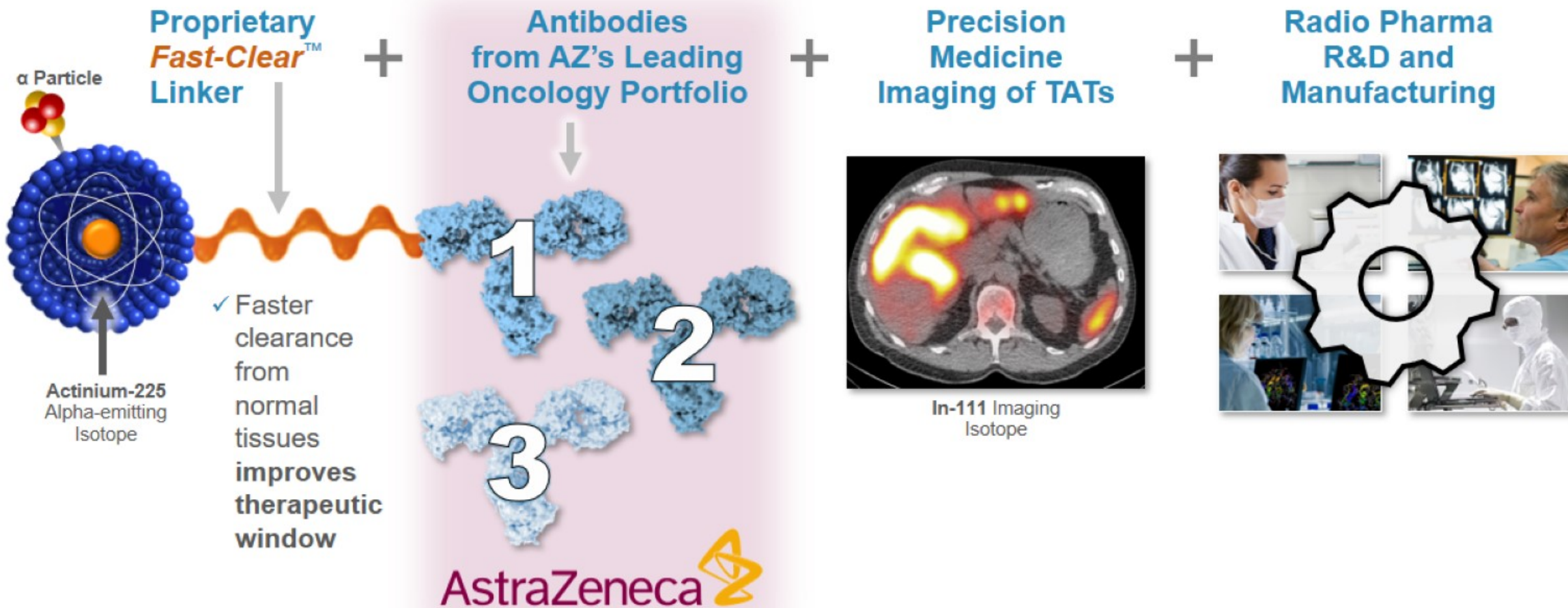
Chief Scientific Officer



# Scientific Rationale

# 1) Using Antibodies in AstraZeneca's Leading Oncology Portfolio To Discover, Develop and Commercialize Up to Three Novel TATs

## FUSION'S TAT PLATFORM

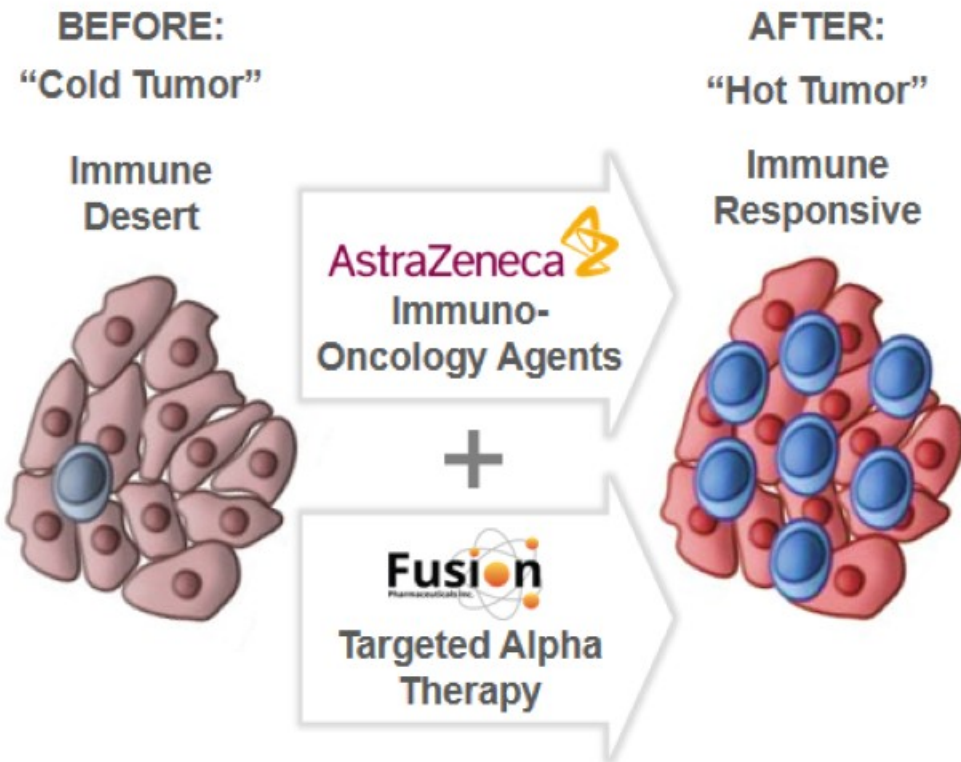




## 2) Evaluate Up to Five Combination Therapy Programs Using TATs with AZ's Portfolio of Immuno-oncology Agents & DDRis

### Immuno-Oncology Combo Therapy

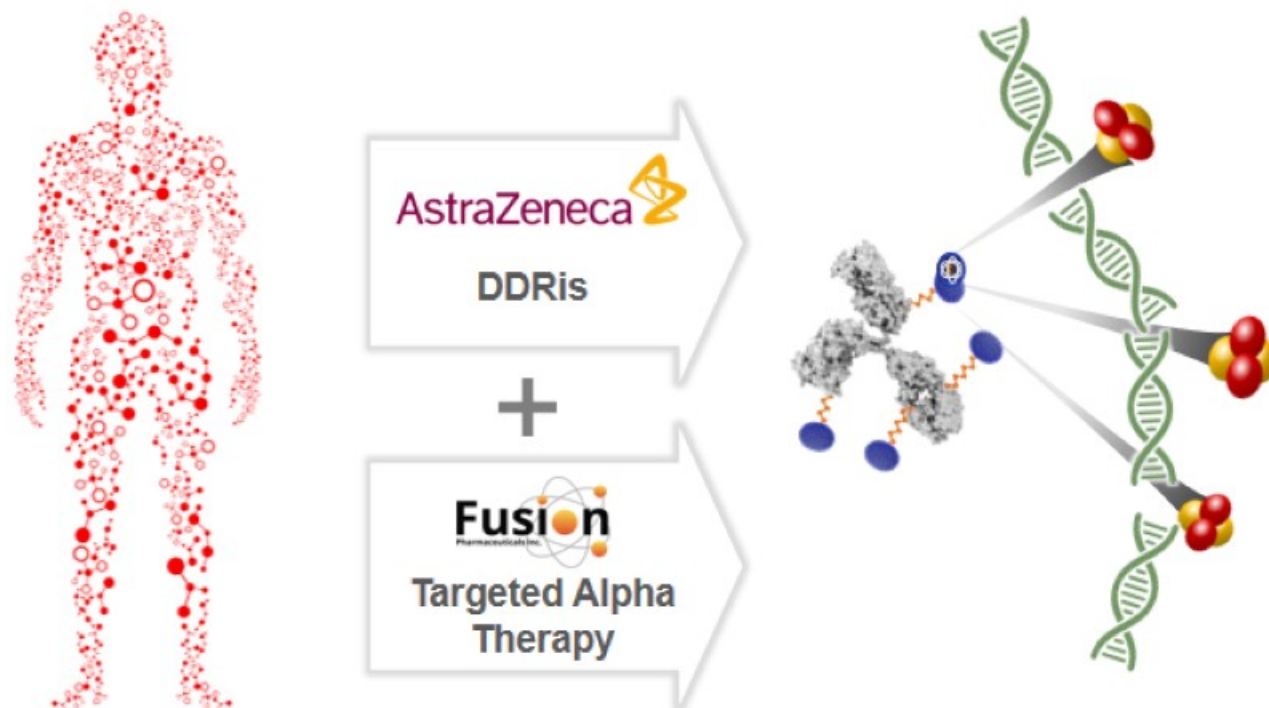
Turning I/O "resistant" tumors into I/O "sensitive"



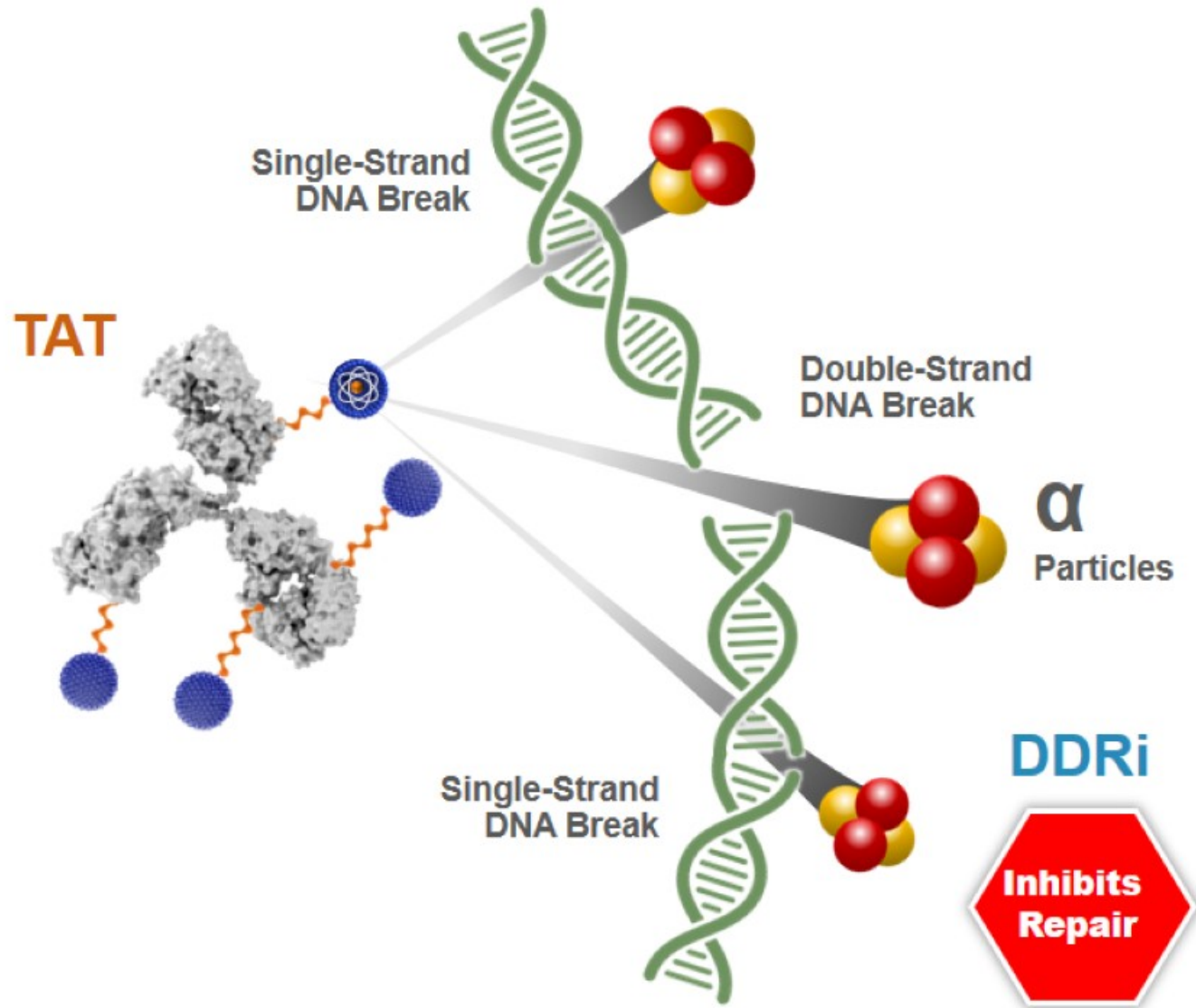
Enhancing antigen presentation and stimulating T-cell recruitment

### DNA Damage Response Inhibitors

TAT = DNA Damage : DDRis Prevent DNA Damage Repair



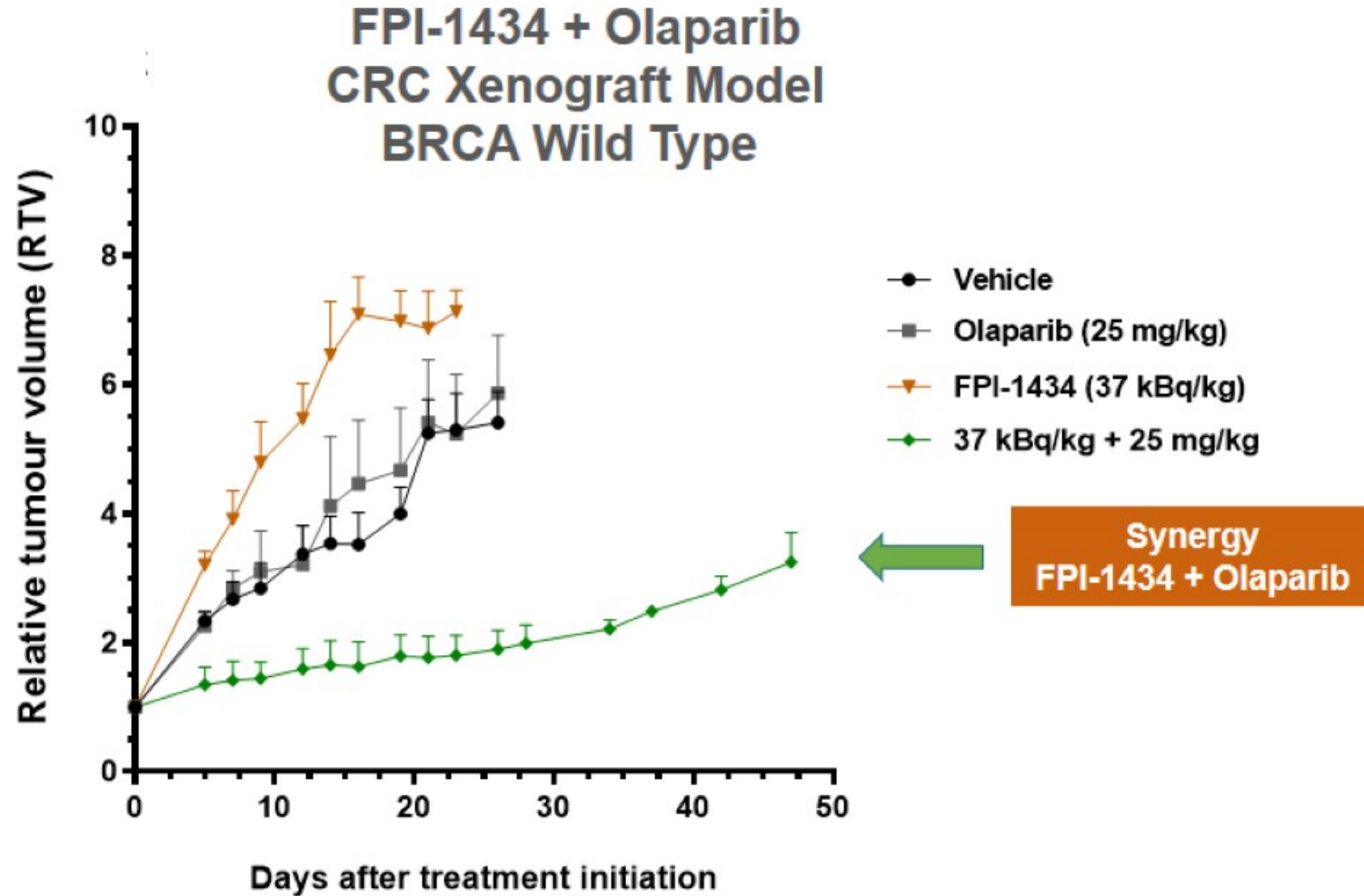
Potential Future Market Expansion, and Earlier Stage Treatment with Lower Required Doses



- TATs + DDR inhibitors are a logical combination:
  - TAT creates tumor DNA damage
  - DDRi prevents DNA repair
- Potential for increases in response rates and broader therapeutic utility



# Scientific Rationale: Combination of Sub-Therapeutic Single Agent Doses Resulted in Enhanced Anti-Tumor Activity



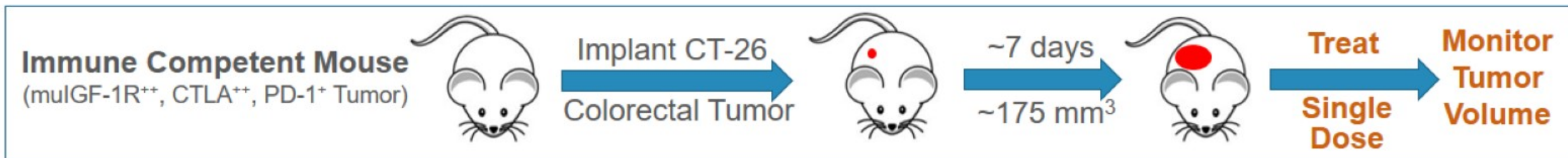
## FPI-1434 + Olaparib Combination

- Shows synergistic effects when compared to low-dose single agents alone

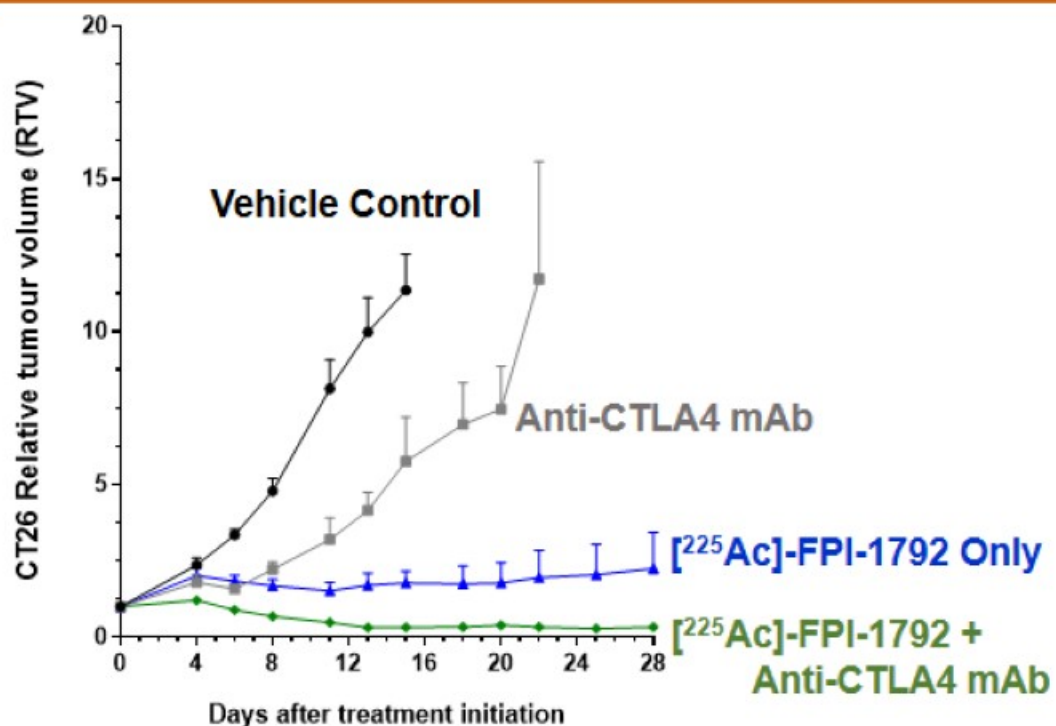
## Synergistic Efficacy

- Has been observed in multiple tumor xenograft models

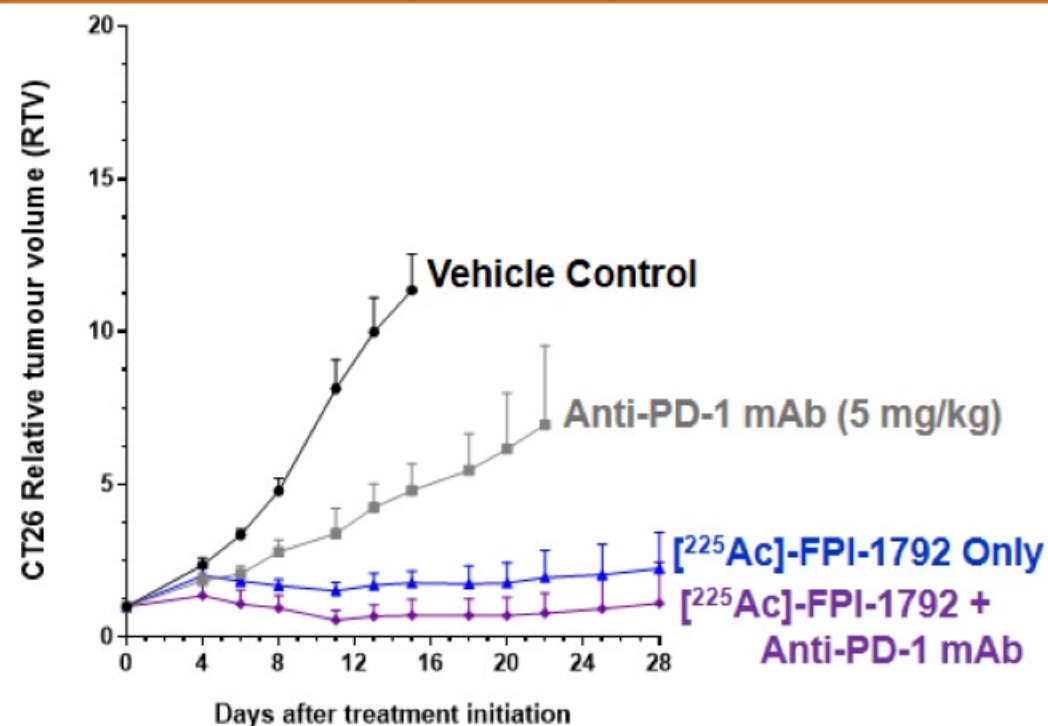
# Scientific Rationale: [<sup>225</sup>Ac] TAT + Checkpoint Inhibitors Demonstrate Enhanced Efficacy in a Mouse Syngeneic Tumor Model



## FPI-1792\* with anti-CTLA-4



## FPI-1792\* with anti-PD-1



\*Murine version of the IGF-1R antibody connected to <sup>225</sup>Ac through Fusion's Fast-Clear linker



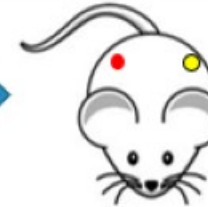
# Reduced Tumor Growth Observed in All Dose Groups Following Re-Challenge in Contralateral Flank

**Immune Competent Mouse**  
CT-26 Colorectal Tumor  
(mulGF-1R<sup>++</sup>, CTLA<sup>++</sup>, PD-1<sup>+</sup> Tumor)



Treat  
(per previous slide)

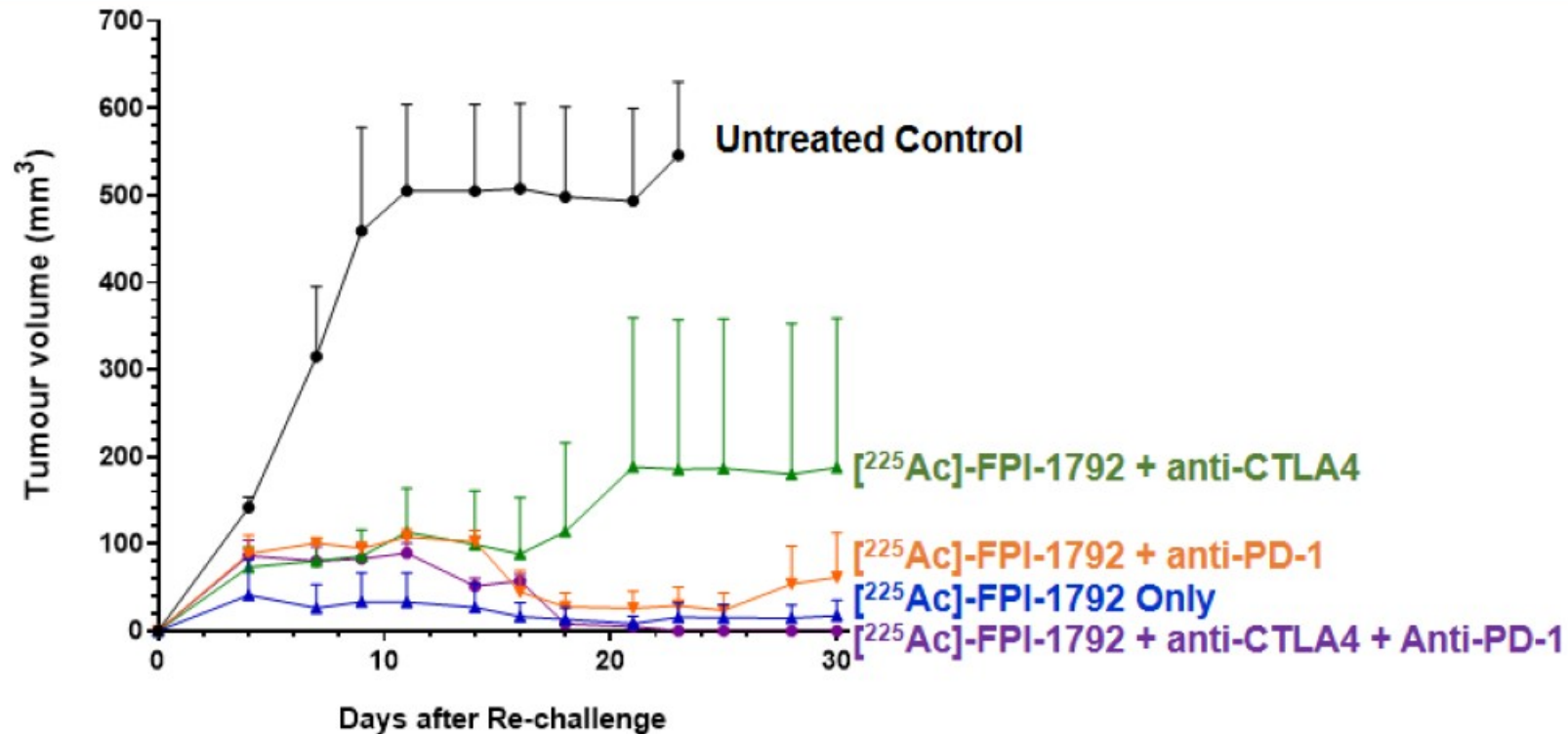
**Re-Challenge**  
**CT-26**



No Treatment

Monitor  
Tumor  
Volume

**Vaccine Study: 13 of 15 Animals Showed No Growth of a Secondary Tumor**



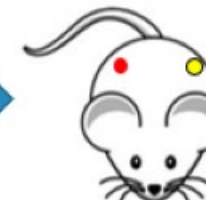
# Infiltration of Antigen-Specific CD8+ T Cells Is Responsible for “Vaccine Effect”

**Immune Competent Mouse**  
CT-26 Colorectal Tumor  
(mulGF-1R<sup>++</sup>, CTLA<sup>++</sup>, PD-1<sup>+</sup> Tumor)



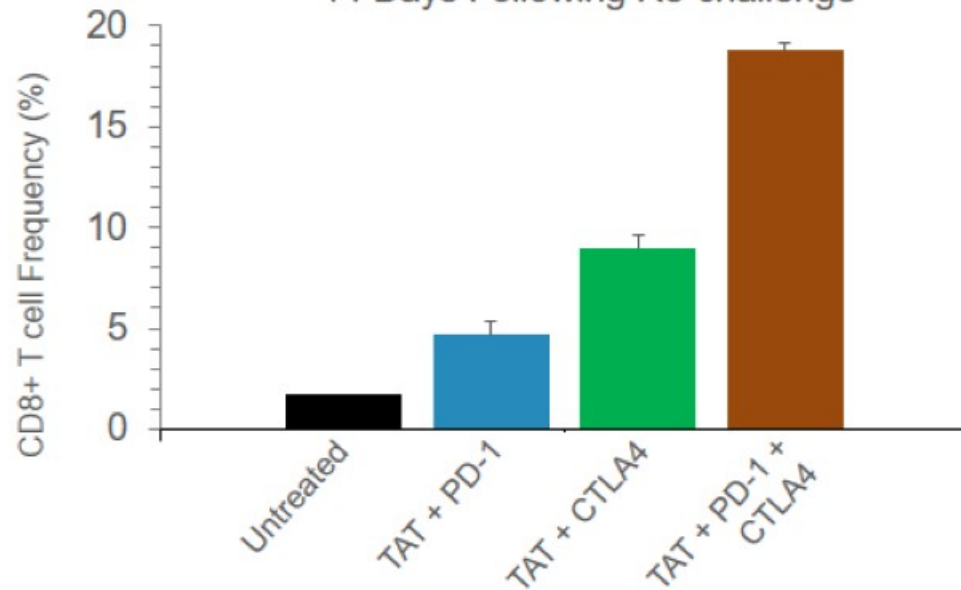
**Treat**  
(per previous slide)

**Re-Challenge**  
CT-26

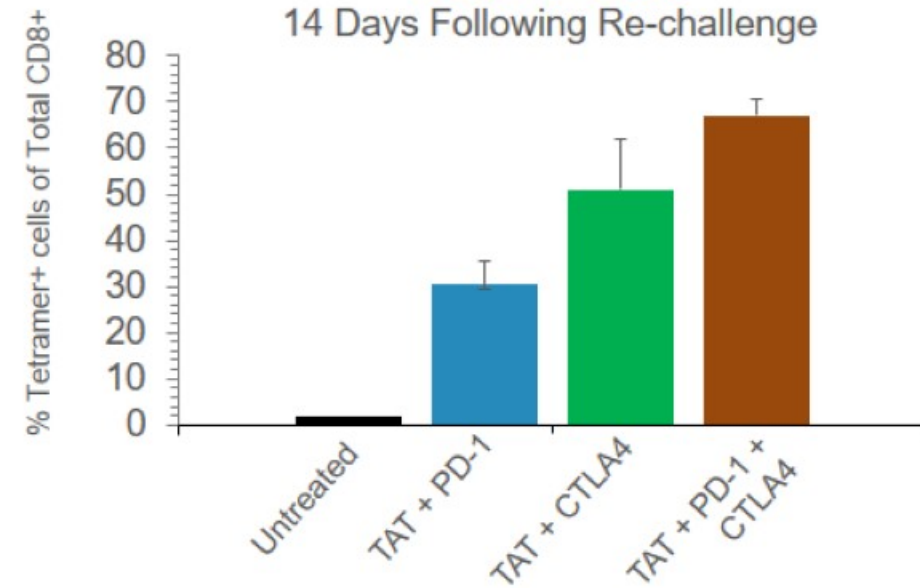


**No Treatment**  
**Tumor/  
Spleen  
Assessment**

**Tumor CD8+ T cell Recruitment**  
14 Days Following Re-challenge



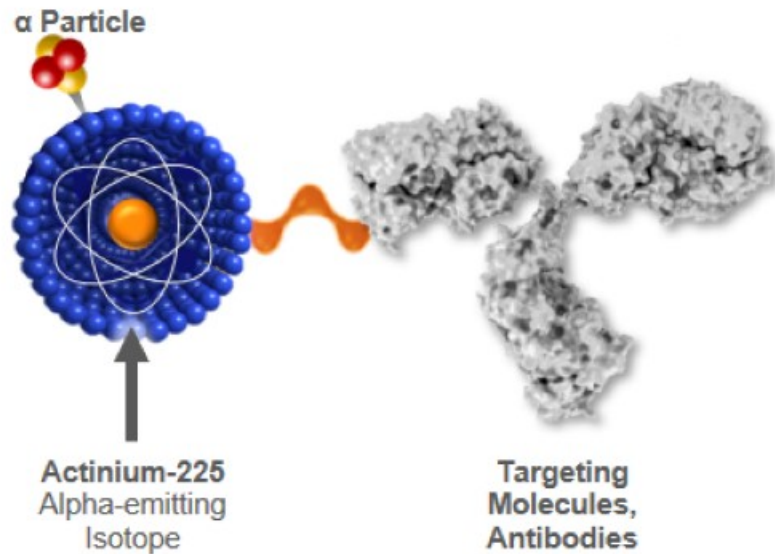
**Detection of Antigen-specific AH1 CD8+ T cells**  
14 Days Following Re-challenge



**Obtained Biochemical Evidence of T cell Memory**

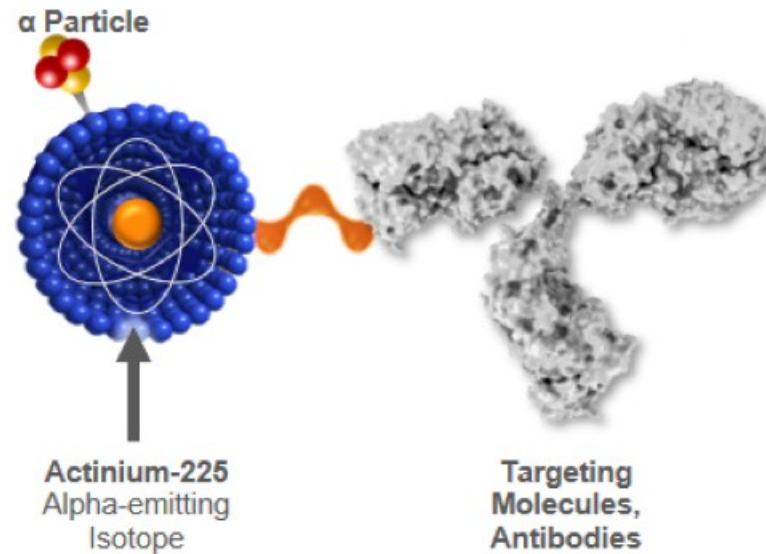


## 1) Novel Targeted Alpha Therapies



New precision medicines

## 2) Combination Therapies with TATs



- Immuno-Oncology Agents

...OR...

- DNA Damage Response Inhibitors (DDRIs)

Broader utility for DDRIs and TATs, reduced doses of TATs



**John Crowley**  
Chief Financial Officer

# Financials



# Agreement Drives Long-Term Value Creation



	Combinations (up to 5)	Novel TATs (up to 3)
Milestones (\$5M upfront)	Up to \$40M to Fusion	-
Development Expense	100% AZ (unless Fusion opts-in)	50/50
Profit Share	Profit from Respective Compounds (minus payback and risk premium if not opted-in)	50/50

**Opt Out for  
Each Novel  
TAT**

Up to  
\$145M in  
milestones  
from developer

-

Low-to-high  
single-digit  
royalty to non-  
developer

\*as of 6/30/20

**\$319M\*** for Operations into 2024

**Capacity to Support Multiple Programs**

**Scalable Platform**



- Delivers access to breadth of AstraZeneca's world-class oncology pipeline & products, resources and expertise
- Avoids early-stage development risk
- Provides optionality around later-stage risk and expense
- Preserves significant back-end economics
- Maintains all rights to existing Fusion pipeline









**Thank You**

**Fusion and AstraZeneca to Develop and Commercialize Next-Generation Targeted Alpha Therapy Radiopharmaceuticals and Combination Therapies**

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