

RADIOPHARM VENTURES INSTITUTIONAL & VENTURE CAPITAL PRESENTATION

Sydney, Australia – 22 April 2024 – Radiopharm Theranostics (ASX:RAD, “Radiopharm” or the “Company”), a developer of a world-class platform of radiopharmaceutical products for both diagnostic and therapeutic uses, is pleased to announce a presentation on the progress of Radiopharm Ventures, the joint venture between RAD and MD Anderson, to multiple US biotech specialised funds and venture capital firms.

The presentation was given at the ‘Oncology/Cell Tx Innovation: The Texas Trifecta’ event, featuring MD Anderson Cancer Center (MDA), Baylor College of Medicine (BCM) & Rice University (RU) and organized by Truist Securities Life Sciences.

RAD CEO & Managing Director Riccardo Canevari and Professor David Piwnica-Worms presented on the lead agent for Radiopharm Ventures, RV 01, a B7H3 targeted radiopharmaceutical therapy designed with strong affinity for the 4IG isomer of B7H3 that is highly expressed in the tumor and not in the healthy tissues. The presentation also covered detail on the new surfaceomes platform.

“It was an excellent opportunity for RAD and Radiopharm Ventures to be invited to present at this event, and our presentation generated significant interest from the attendees,” said Mr Canevari.

The presentation slides used at the event are attached below.

About Radiopharm Theranostics

Radiopharm Theranostics is a clinical stage company developing a world-class platform of innovative radiopharmaceutical products for diagnostic and therapeutic applications in areas of high unmet medical need. Radiopharm has been listed on the ASX (RAD) since November 2021. The company has a pipeline of distinct and highly differentiated platform technologies spanning peptides, small molecules and monoclonal antibodies for use in cancer, in pre-clinical and clinical stages of development from some of the world’s leading universities and institutes. The pipeline has been built based on the potential to be first-to-market or best-in-class. The clinical program includes one Phase II and two Phase I trials in a variety of solid tumour cancers including lung, pancreas and brain. Learn more at Radiopharmtheranostics.com.

Authorized on behalf of the Radiopharm Theranostics Board of Directors by Executive Chairman Paul Hopper.

For more information:

Riccardo Canevari
CEO & Managing Director
P: +1 862 309 0293
E: rc@radiopharmtheranostics.com

**ASX ANNOUNCEMENT
22 APRIL 2024**

Matt Wright
NWR Communications
P: +61 451 896 420
E: matt@nwrcommunications.com.au

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Any opinions expressed reflect the Company’s position at the date of this presentation and are subject to change.



RADIOPHARM
VENTURES

A Joint Venture between

THE UNIVERSITY OF TEXAS
MDAnderson
~~Cancer~~ Center
Making Cancer History®

 **RAD**
RADIOPHARM THERANOSTICS

Company Presentation – *April 2024*

COMPANY VISION & STRATEGY

B7H3 FRANCHISE & NOVEL TARGETS FOR RADIOPHARMACEUTICAL THERAPIES



Private Company created in Q4 2022 between MD Anderson Cancer Center (MDACC) and RAD

- Four technologies licensed from MDACC
- R&D costs currently covered by Radiopharm Theranostics (RAD: ASX)



Intellectual Property

- B7H3 targeting mAb with patent protection 2044+ (Global rights, exclusivity, any therapeutic Isotope)
- Three additional technologies not yet disclosed, with potential indications in multiple solid tumors, targeting novel pathways



Deep Expertise in Radiopharmaceuticals

- Executive team includes top MD Anderson scientists and Radiopharm Theranostics Leadership Team
- Radiopharm Ventures has access to MD Anderson preclinical labs (in vitro and animal models), nuclear medicine facilities (radiolabeling) & clinical development experts (Phase I-II trial and protocol design)

LEADERSHIP TEAM and JV SCIENTIFIC COMMITTEE



**Prof. David
Piwnica-Worms**

- Inventor B7H3 targeting molecule
- Department Chair, Department of Cancer Systems Imaging, Division of Diagnostic Imaging, MDACC
- Professor, Department of Cancer Systems Imaging, Division of Diagnostic Imaging, MDACC
- Executive Director, Quantitative Imaging Analysis Core (QIAC), Division of Diagnostic Imaging, MDACC
- Gerald Dewey Dodd, Jr., Endowed Distinguished Chair in Diagnostic Imaging, Division of Diagnostic Imaging, MDACC



Prof. Sam Hanash

- Inventor three technologies (undisclosed)
- Director, Department of Red and Charline McCombs Institute for the Early Detection and Treatment of Cancer, MDACC
- Co-Director, Department of Center for Global Cancer Early Detection, MDACC
- Distinguished Chair, Department of Evelyn & Sol Rubenstein Distinguished Chair for Cancer Prevention, MDACC
- Professor, Department of Clinical Cancer Prevention - Research, Division of OVP, Cancer Prevention and Population Sciences, MDACC



Riccardo Canevari

- Radiopharm Theranostics CEO since 2021
- Previously, Chief Commercial Officer of Novartis Company Advanced Accelerator Applications S.A.
- Lead for Lutathera in-market growth strategy and Pluvicto launch strategy
- Senior Vice President & Global Head, Breast Cancer Franchise, for Novartis Oncology
- Lead for CDK4/6 (Kisqali) global launch in HR+ Breast Cancer
- Lead for PI3K inhibitor (Piqray) global launch in HR+ Breast Cancer

COMPANY PIPELINE

CODE	TARGET & MOLECULE	INDICATION	Dx/Tx	ISOTOPE	NOTES
RV 01	B7H3 mAb	Prostate, Lung, Hepatocellular Carcinoma, Pancreatic, Colorectal, Head & Neck, Breast	Therapy	Lu177 (β)	<p>CMC GMP production ongoing Preclinical studies completed</p> <p>Next steps: complete CMC, and conduct IND enabling GLP Tox and Biodistribution studies</p> <p>Entering Phase 1 in the first half of 2025</p>
RV02	Targets UNDISCLOSED				
RV03	BINDING MOIETY: Nanobody platform	Multiple solid tumors	Therapy	Lu177(β)	Candidate selection ongoing
RV04				Tb161 (β+Au) Ac225 (α)	

Supply Chain Enhancement



RV01: Lead Candidate

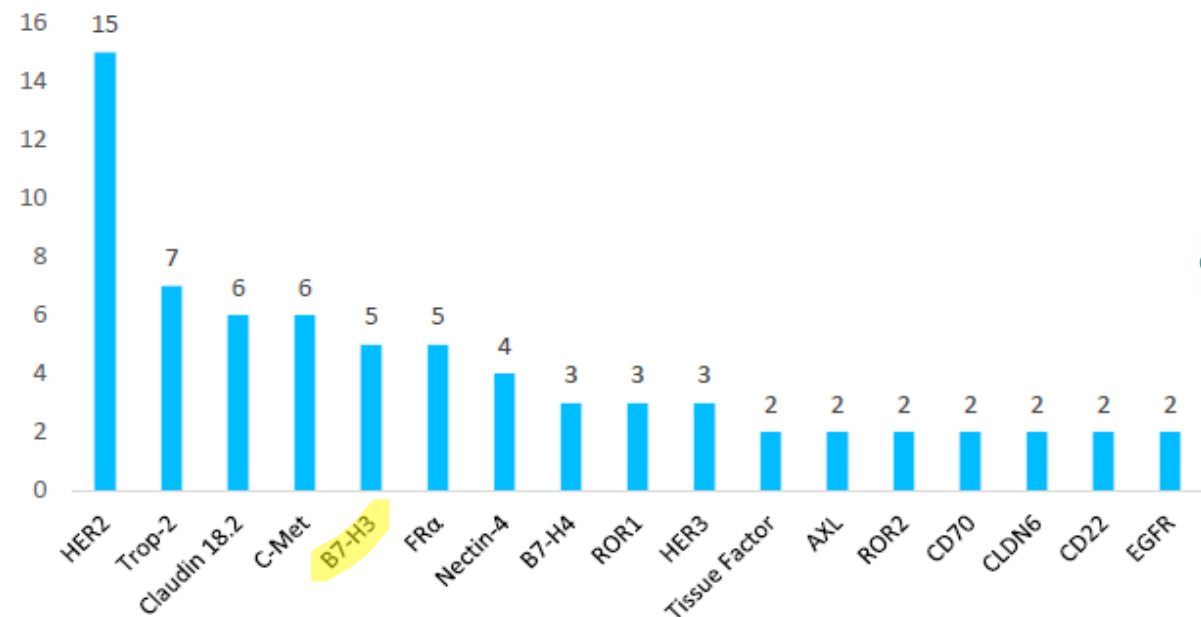
**B7H3-Targeting Monoclonal Antibody (mAb) for Beta-Radioligand
Therapy**



FIVE B7H3 ANTIBODY DRUG CONJUGATES IN DEVELOPMENT

PHASE 2 READOUT WITH ADC IN PROSTATE CANCER EXPECTED SOON

Clinical Stage Targets in Development (At least 2 or more)



Source: BioMed Tracker, Leerink Partners Research

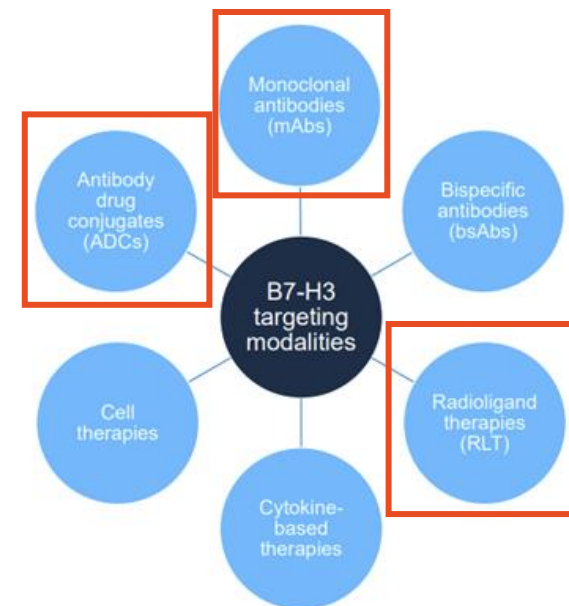


Ph2 TAMARAK B7H3 study in mCRPC*
Interim efficacy data May 31st, 2024



Ph3 Ideate-2 B7H3 study in SCLC**
Initiation in 2024

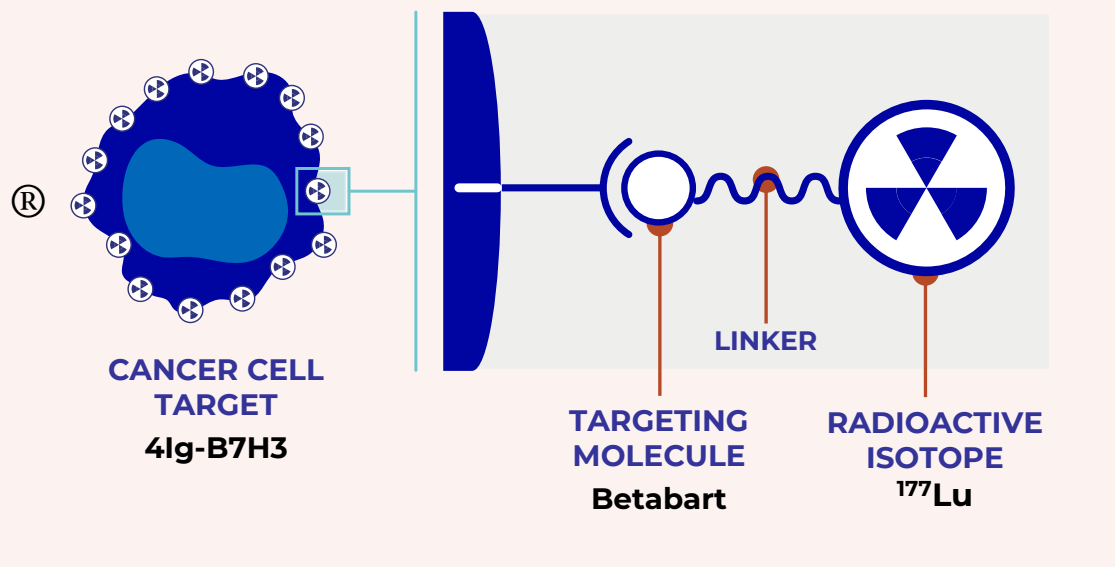
DIFFERENT MODALITIES TO TARGET B7H3



*mCRPC: Metastatic Castration Resistant Prostate Cancer. NCT05551117

**SCLC: Small Cell Lung Cancer. NCT06203210

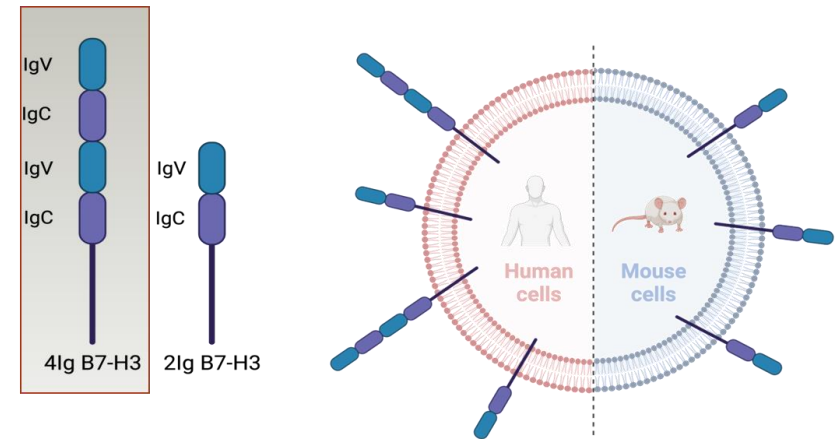
FIRST AND ONLY SELECTIVE B7H3 RADIOPHARMACEUTICAL IN DEVELOPMENT



BETABART[®]
Isoform-Selective Targeting of 4lg-B7-H3
for PET Imaging and Beta-Radioligand Therapy

THERAPY


- Multi-indication potential in B7H3+ solid tumors (Prostate, Pancreatic, Hepatocellular Carcinoma, Colorectal, Breast, H&N, Lung, Ovarian, ...)
- Phase I planned in early 2025



- A soluble 2lg-B7H3 isoform circulating in the blood is a potential pseudo-target decoy (sink) not widely appreciated as a confounding factor in therapy.

CLINICAL DEVELOPMENT & REGULATORY STRATEGY

- Preclinical models validated selective targeting of tumor cells
- Significant tumor reduction and increased survival in the animal model
- GMP CMC production ongoing. GMP batch in Q3 2024
- GLP TOX and Biodistribution studies in Q3 –Q4 2024
- Phase I trial in the first half of 2025

Preclinical	PHASE I
	Basket trial in multiple indications
CMC GMP, GLP Tox, BioD	25 pts
completed by end 2024	Opening in first half 2025

Targeting B7H3 in Advanced Prostate Cancer may Optimize Clinical Outcomes, Reduce Off-Tissue Toxicities & Improve Quality of Life

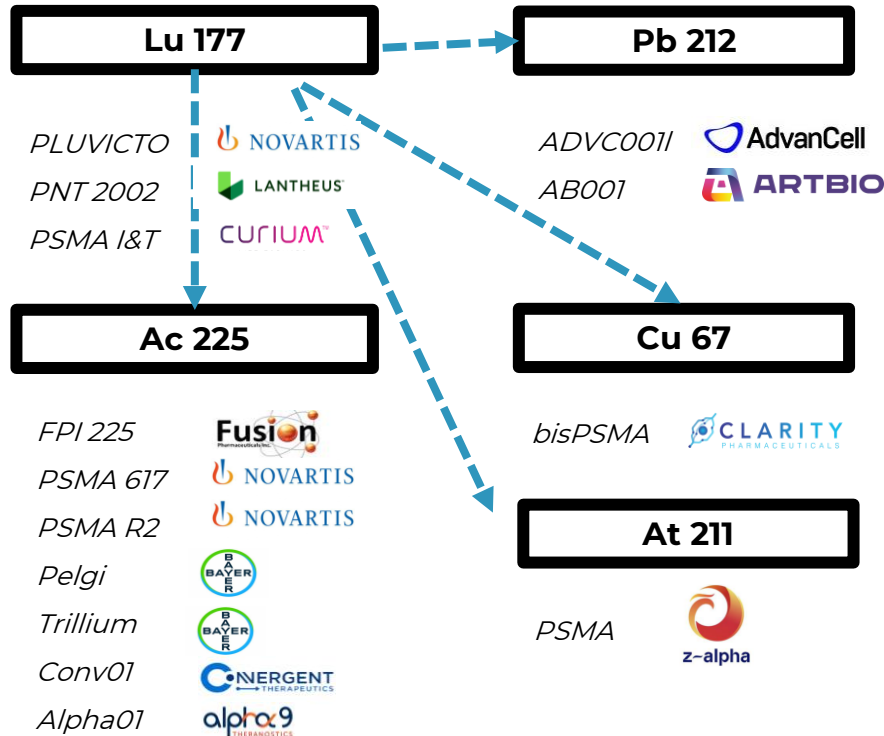
Current SoC Treatments: ARIs, PARPs, PSMA RLT*

NEW Mechanisms of Action

NOVEL ARIs & PARPs

- Abiraterone
- Apalutamide
- Darolutamide
- Enzalutamide
- Olaparib
- Rucaparib

PSMA RLT



B7H3

MACROGENICS
Most advanced B7H3 ADC in development

RADIOPHARM VENTURES
First and only selective B7H3 radiopharmaceutical in development

CD46 ADC mAb (FORTIS THERAPEUTICS)

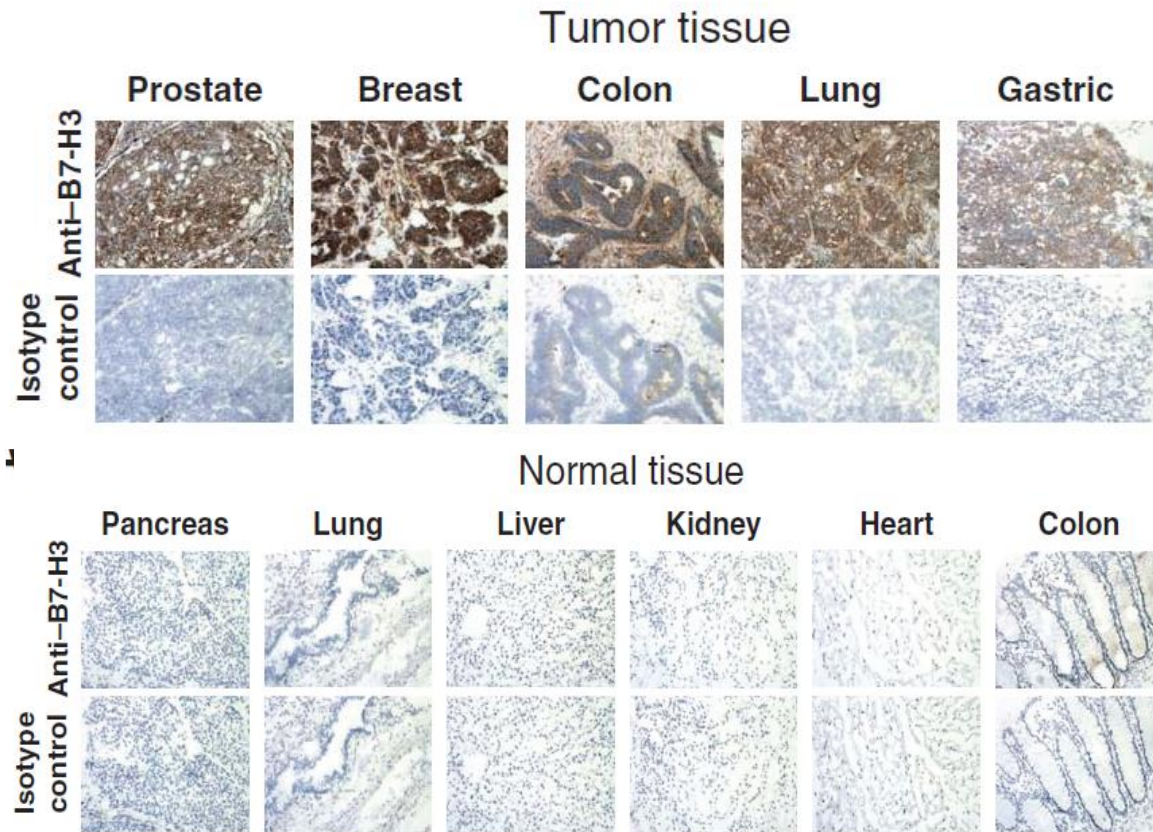
STEAP1 T-cell engager (AMGEN)

*SoC: Standard of Care
ARIs: Androgen Receptor inhibitors
PARPs: Poly (ADP-ribose) Polymerase
RLT: Radioligand Therapies

PRECLINICAL DATA DEEP DIVE

*Isoform-Selective Targeting of 4Ig-B7H3
for Imaging and Beta-Radioligand Therapy*

B7H3 PATHWAY HIGHLY ATTRACTIVE PAN-TUMOR TARGET



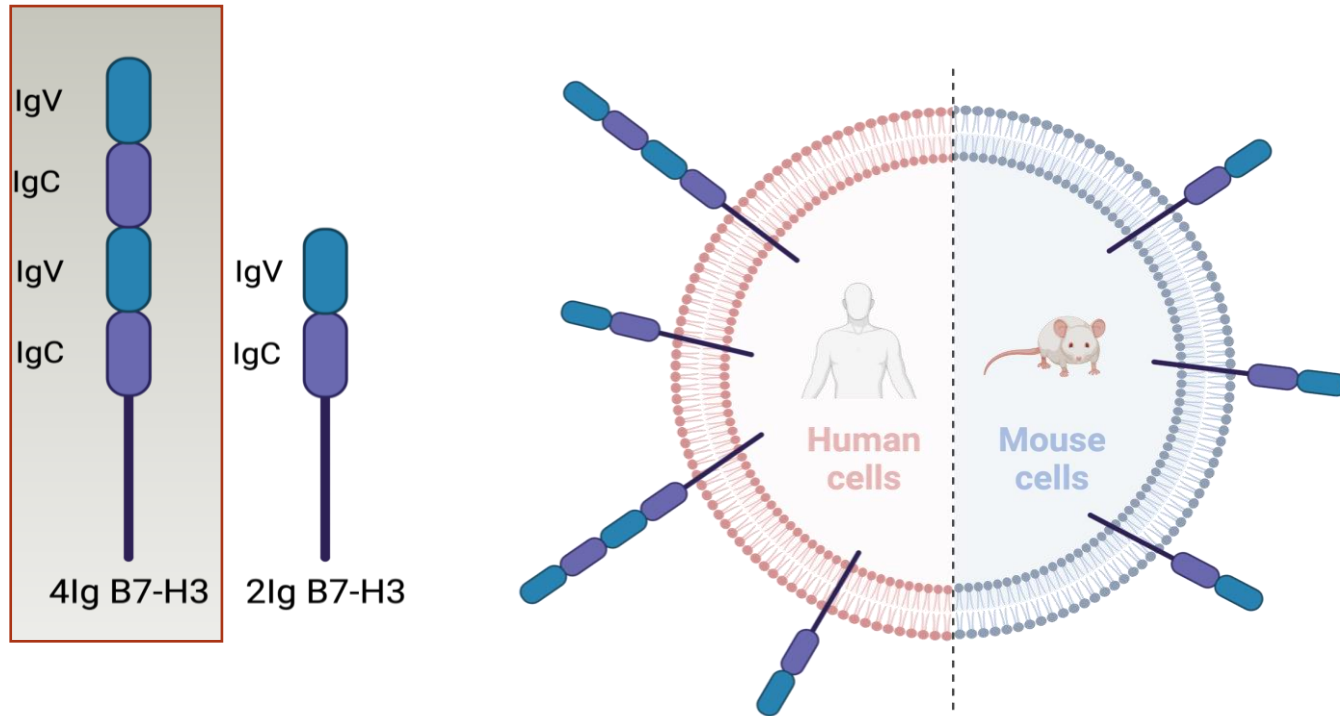
Loo, D. et. al. *Clinical Cancer Research* (2012).

High B7-H3 Expression Levels in Solid Tumors

Potential Indications	B7-H3 Positive*		2+ or Above	
Head and Neck Cancer	19/19	100%	19/19	100%
Kidney Cancer	77/78	99%	75/78	96%
Glioblastoma	65/66	98%	63/66	95%
Thyroid Cancer	34/35	97%	33/35	94%
Mesothelioma	41/44	93%	39/44	89%
Melanoma	132/146	90%	94/146	64%
Prostate Cancer	88/99	89%	51/99	52%
Pancreas Cancer	69/78	88%	45/78	58%
Bladder Cancer	134/156	86%	123/156	79%
Lung Cancer	324/379	85%	300/379	79%
Breast Cancer	189/249	76%	156/249	63%
Ovarian Cancer	59/79	75%	36/79	46%

*B7-H3 positivity reflects any grade staining (1-3+) via FFPE tumor microarray (cytoplasmic, membrane, and vasculature staining); B7-H3 is expressed on tumor as well as tumor vasculature.

B7H3 IS EXPRESSED AS MULTIPLE ISOFORMS

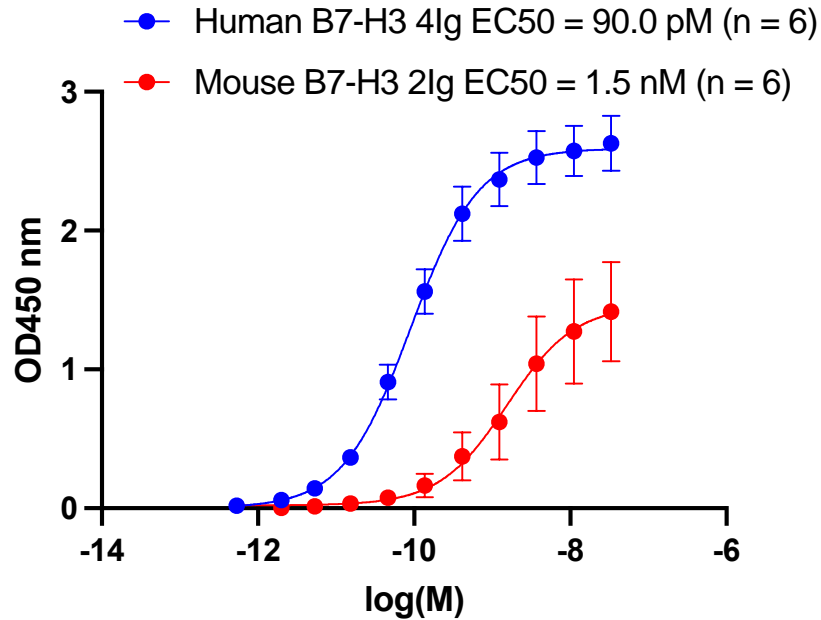


- The 4Ig-B7H3 isoform is the dominant isoform in human cancers.
- A soluble 2Ig-B7H3 isoform circulating in the blood is a potential pseudo-target decoy (sink) not widely appreciated as a confounding factor in therapy.

GOAL: DEVELOP A HIGH-AFFINITY 4IG-ISOFORM-SPECIFIC ANTI-B7H3 TUMOR-TARGETING ANTIBODY

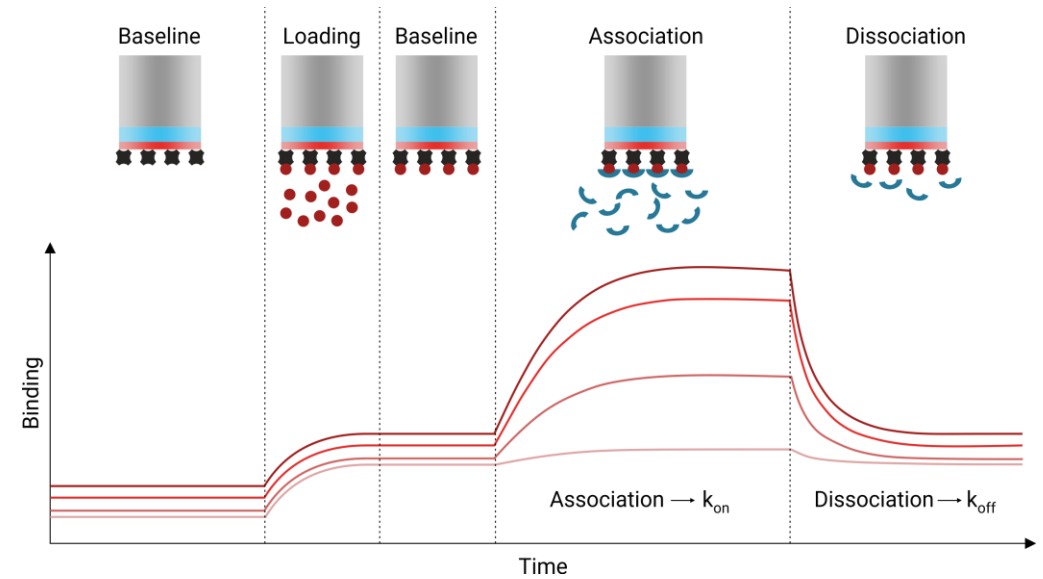
MILL33B, THE MURINE PREDECESSOR MAB CONTAINING THE CDR*, AND OUR HUMANIZED MAB (BETABART®) DEMONSTRATE HIGH AFFINITY & SELECTIVITY FOR HUMAN 4IG-B7H3

MIL33B ELISA



**MILL33B Affinity Ratio
4Ig to 2Ig : ~17**

BIO-LAYER INTERFEROMETRY ANALYSIS

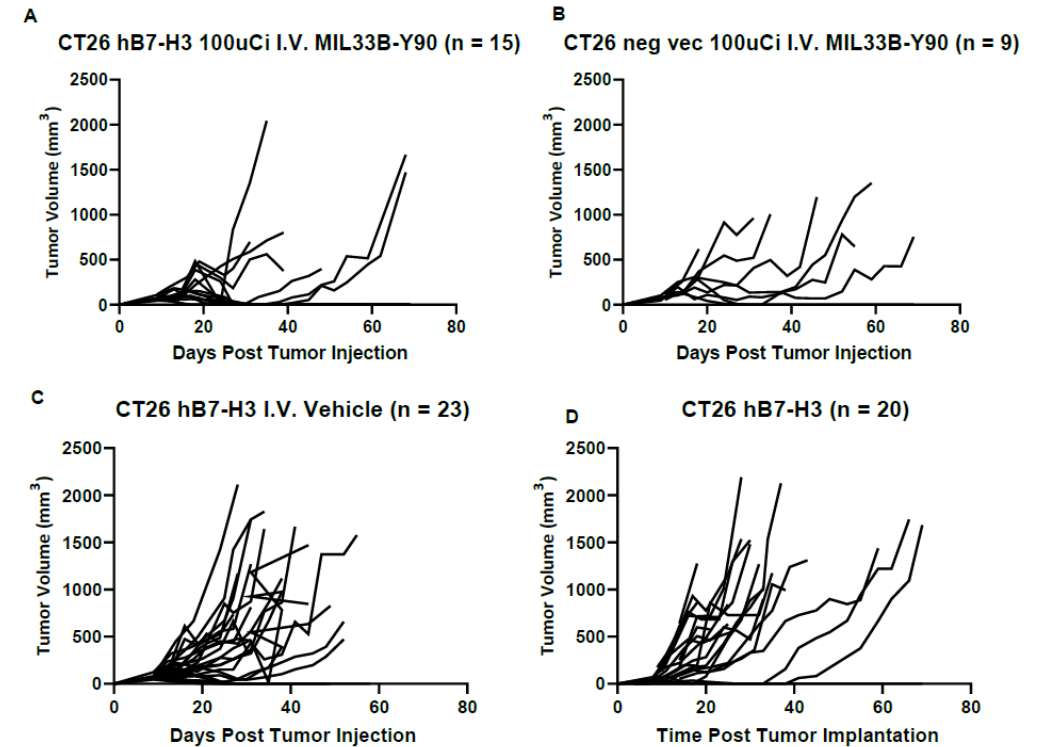
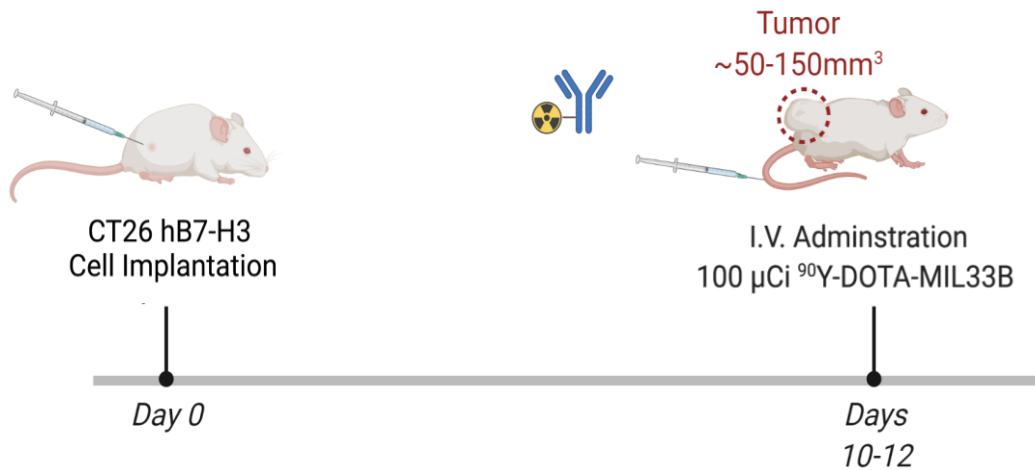


**BETABART® Affinity Ratio
4Ig to 2Ig : >300**

*CDR: Complementarity-Determining Region
<https://2bind.com/bli/>

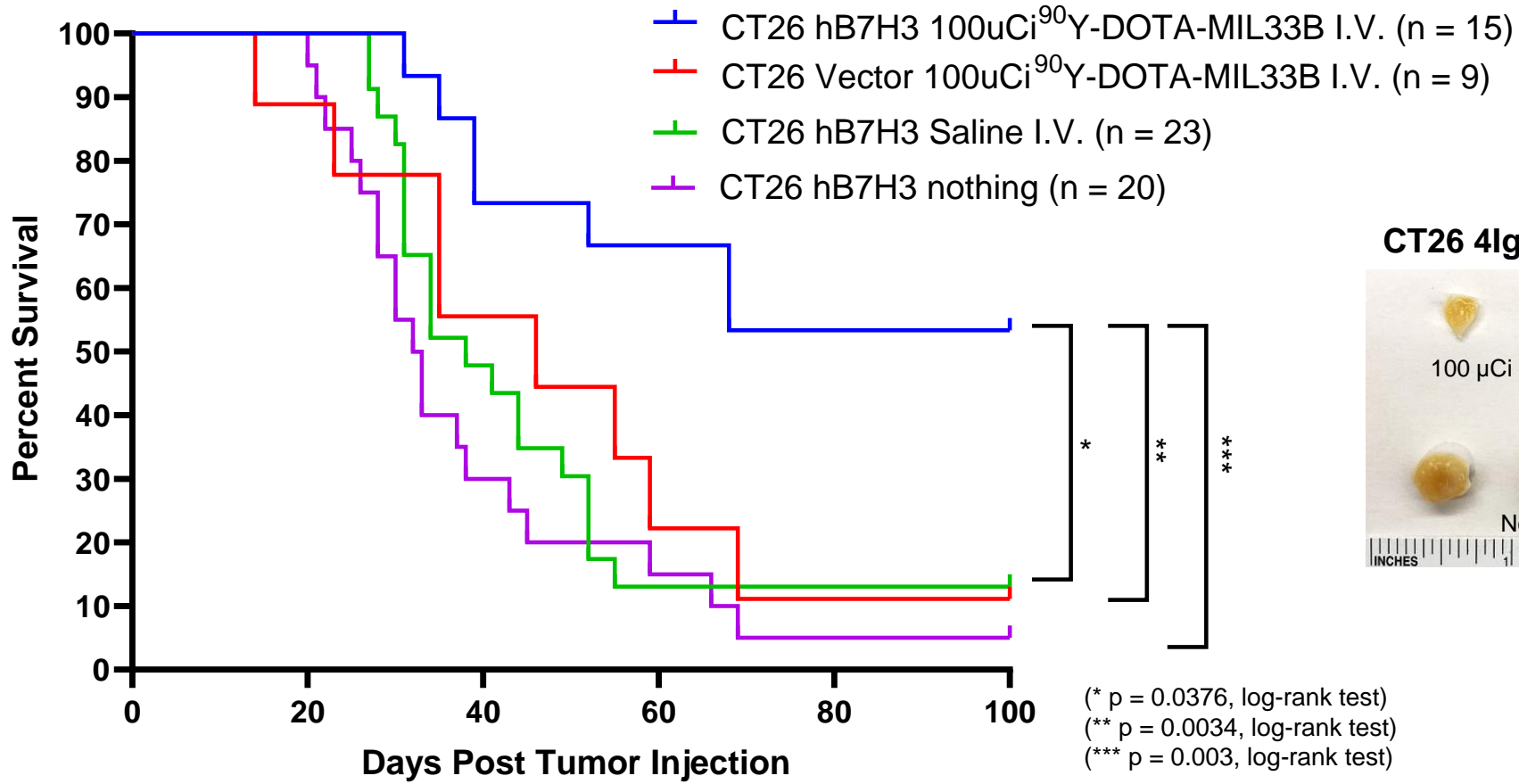
THERAPEUTIC EFFICACY: TUMOR REDUCTION

I.V. TREATMENT OF ESTABLISHED IR-RESISTANT CT26 TUMORS WITH I.V. ^{90}Y -DOTA-MIL33B



Supplemental Figure 8: Caliper measurements of mice treated with ^{90}Y -DOTA-MIL33B. 15 mice harboring CT26 (h)4Ig-B7-H3 tumors received 100 µCi ^{90}Y -DOTA-MIL33B i.v. leading to partial regression of 3 tumors and complete regression of 8 tumors (**A**). 9 mice harboring CT26 neg vec tumors received 100 µCi ^{90}Y -DOTA-MIL33B i.v. leading to partial regression of 1 tumor and complete regression of one tumor (**B**). We also observed regression of 3 out of 23 CT26 4Ig-B7-H3 tumors that received i.v. saline (**C**) and spontaneous regression of 1 out of 20 untreated CT26 4Ig-B7-H3 tumors (**D**).

THERAPEUTIC EFFICACY: 56% SURVIVAL WITH SINGLE INJECTION

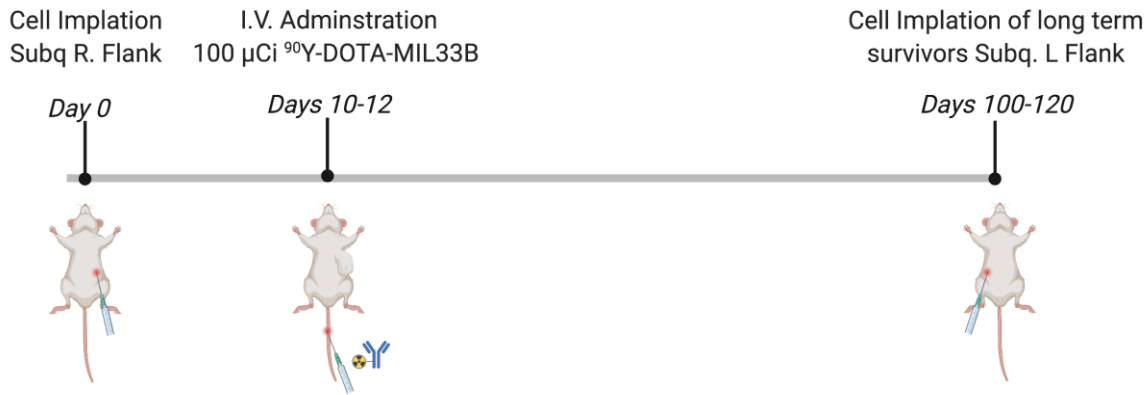


CT26 4Ig-B7H3 Tumors

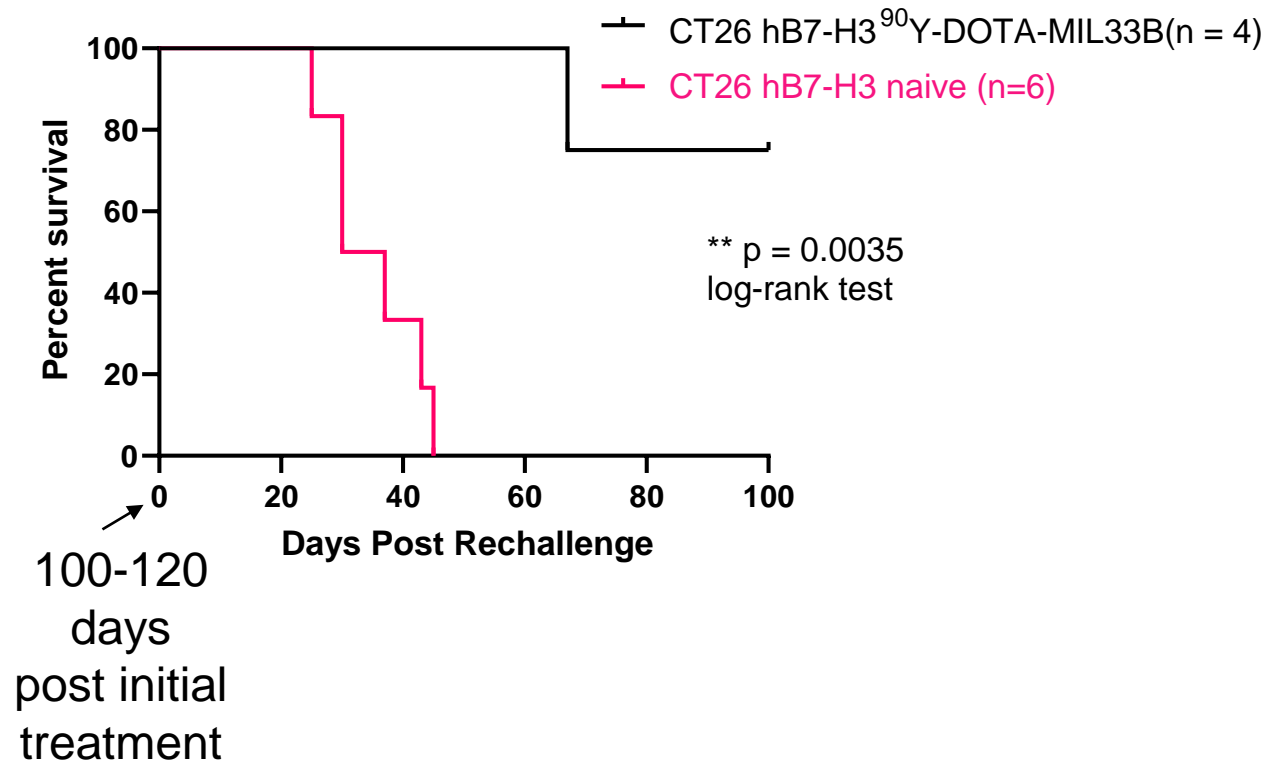


BETABART PREDECESSOR INDUCES IMMUNOLOGICAL MEMORY (TUMOR REJECTION FROM RADIOTHERAPY PRE-TREATMENT)

LONG-TERM SURVIVORS OF 90Y-DOTA-MIL33B TREATMENT RECHALLENGED WITH CT26 TUMOR CELLS

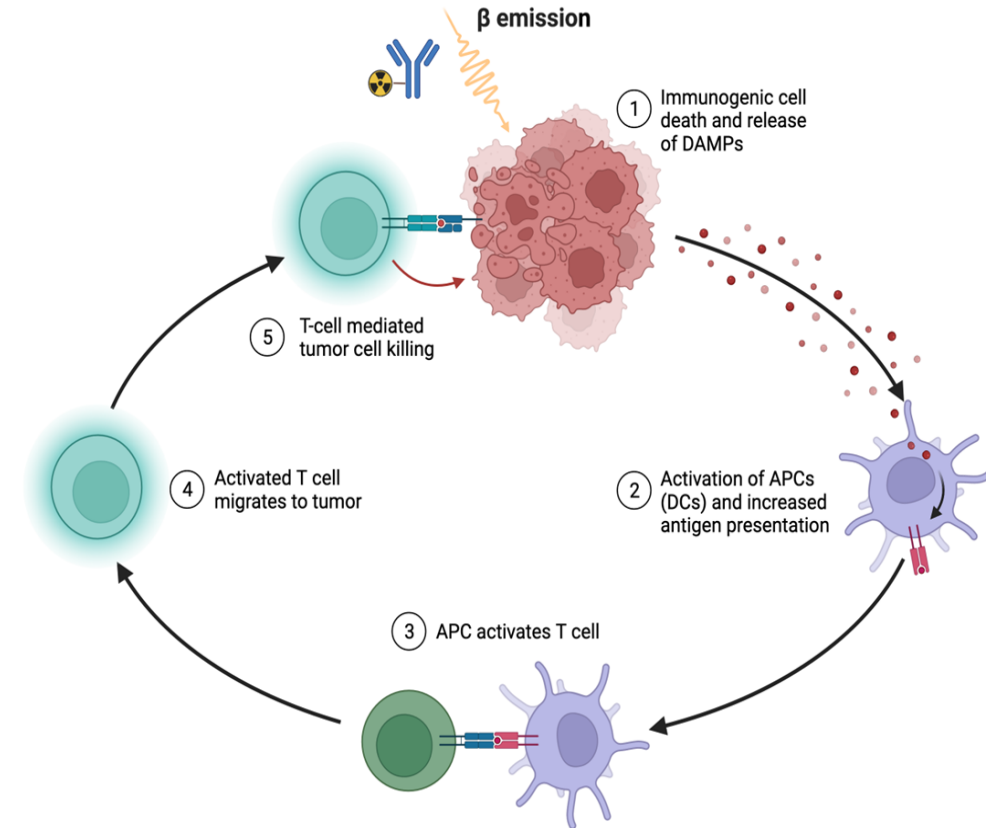


RECHALLENGE SURVIVAL CURVES: PRE-TREATED SURVIVORS VERSUS TREATMENT-NAIVE MICE



CONCLUSIONS: FIRST AND ONLY SELECTIVE B7H3 RADIOPHARMACEUTICAL THERAPY IN DEVELOPMENT

- High affinity (72 pM) antibody specific for 4Ig-B7H3.
- Treatment with a single I.V. dose leads to complete regression of established solid tumors and long-term survival of 56% in our preclinical model
- Initial promising evidence of immune system stimulation and ability to confer immune memory.
- Pre-IND studies to be completed in 2H 2024
- Phase I therapeutic trial to be started in 1H 2025



PIPELINE FOLLOW-ON ASSETS

Surfaceomes (or Cell surface) Platform

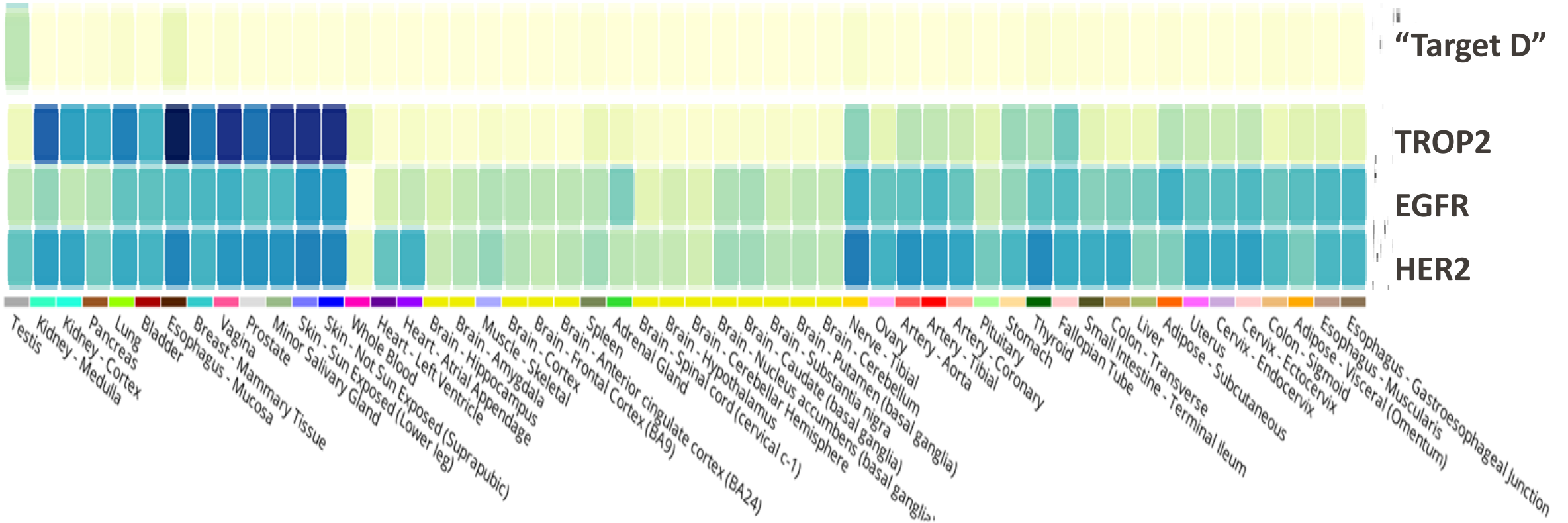
THREE NOVEL “SURFACEOMES” TARGETS IDENTIFIED BY IN-DEPTH PROFILING OF TUMOR CELLS

HIGHLY RELEVANT FOR RADIOPHARMACEUTICAL THERAPIES

- Cell surface expression - critical for radiopharmaceutical effective mechanism of actions
- Target characteristics:
 - a. Testis antigens
 - b. Expressed almost exclusively in the cancer cells;
 - c. Minimal / no expression in healthy tissues
 - d. Not currently in development with any modality (targeted therapy, ADC, RLT)

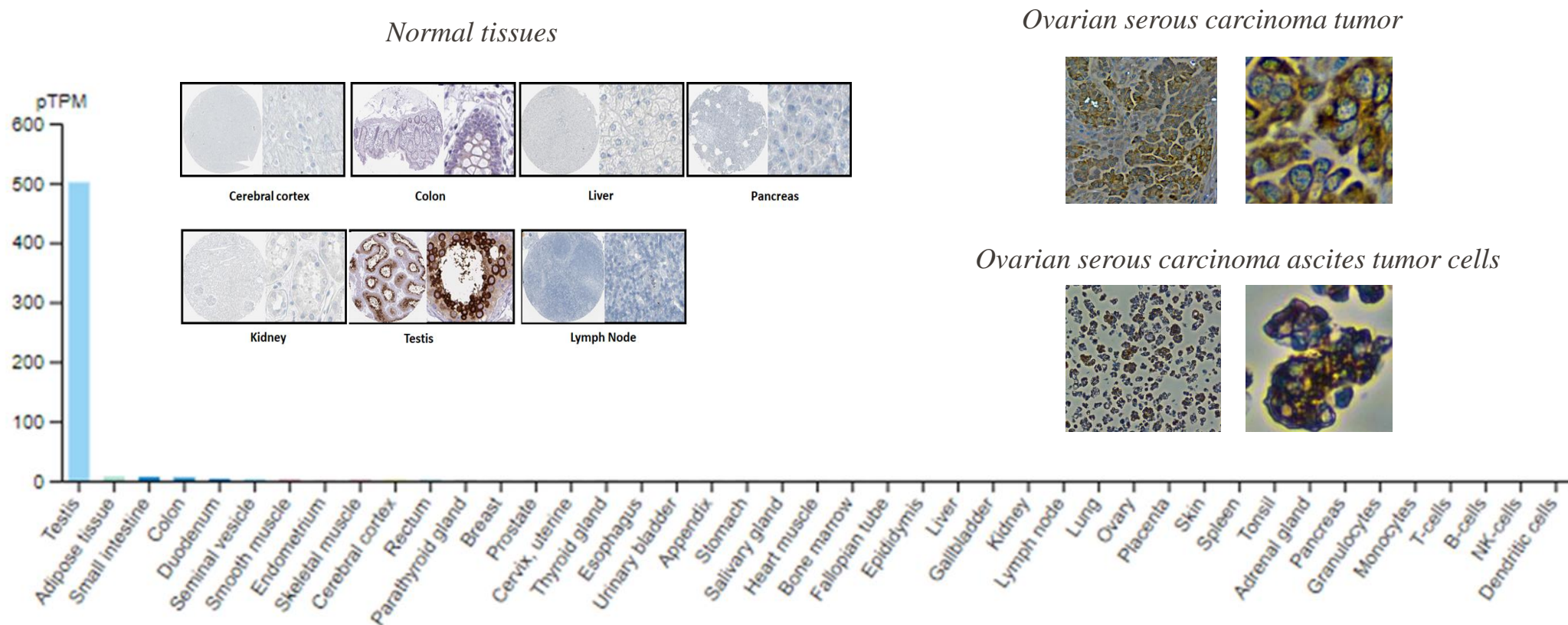
DIFFERENTIATION VS CURRENTLY KNOWN TARGETS

CONSIDERABLY MORE RESTRICTED GENE EXPRESSION IN NORMAL TISSUES

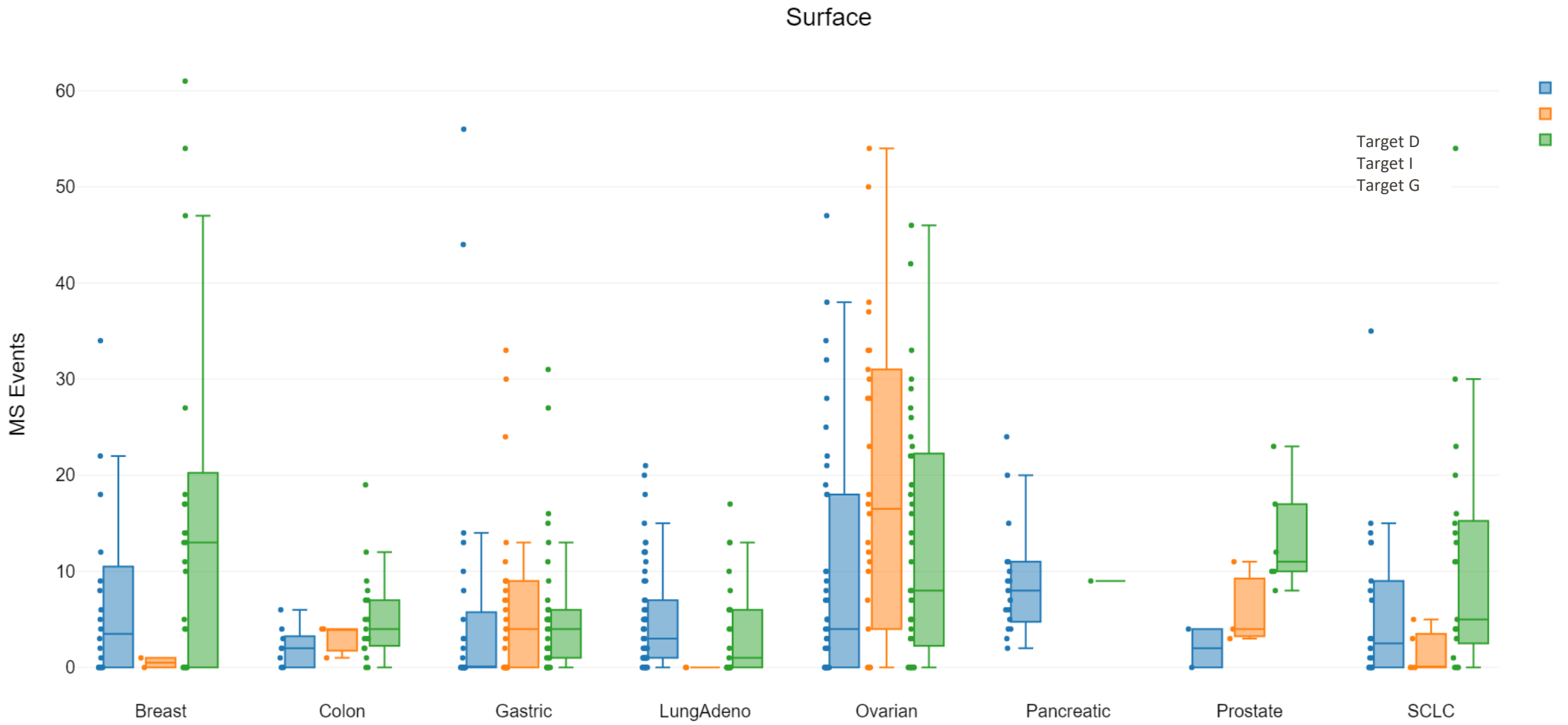


EXAMPLE “TARGET D” IN PRECLINICAL DEVELOPMENT

- Description: Testis antigen
- Expression in normal tissues: largely limited to testis
- Cancer types with surface expression : Highly expressed in TNBC, ovarian, SCLC, gastric



MULTI-INDICATION POTENTIAL FOR EACH SELECTED TARGET





RADIOPHARM
VENTURES

A Joint Venture between

CONTACT:

RICCARDO CANEVARI

rc@radiopharmtheranostics.com

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