

**PSI** Center for  
Life Sciences

# Preclinical Development of $^{161}\text{Tb}$ -based Radiopharmaceuticals for Radioligand Therapy

PRISMAP Radiolanthanide Workshop

3-5 September 2024, Paul Scherrer Institute, Switzerland

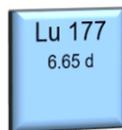
Cristina Müller, PhD

Research Group Leader at PSI, adjunct Prof. at ETH Zurich, Switzerland

# $^{177}\text{Lu}$ -based Radioligand Therapy

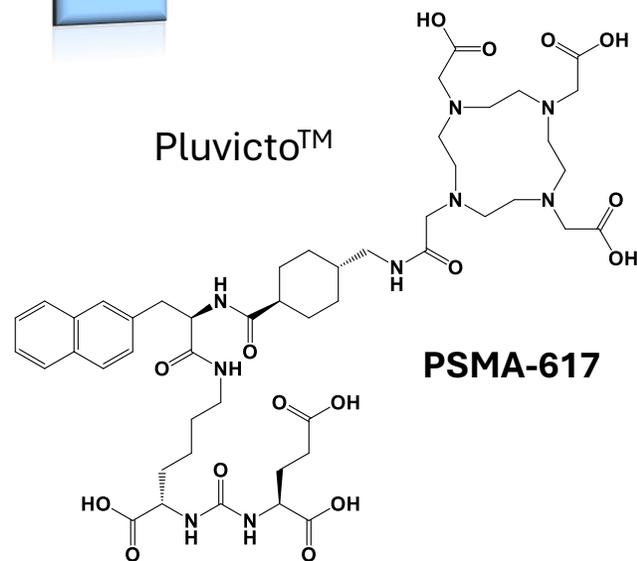


## PSMA-targeting radioligand



$\beta^-$ -Therapy

Pluvicto™



PSMA-617

FDA approved in March 2022

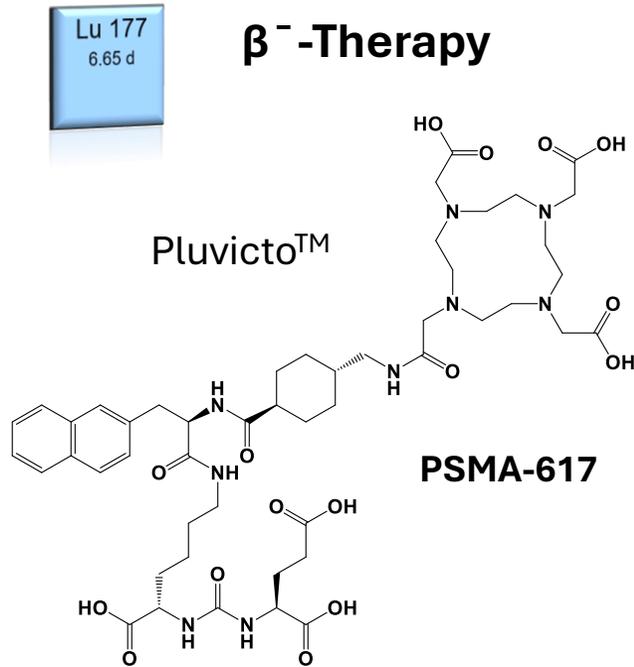
PSMA = prostate-specific membrane antigen



# <sup>177</sup>Lu-based Radioligand Therapy



## PSMA-targeting radioligand



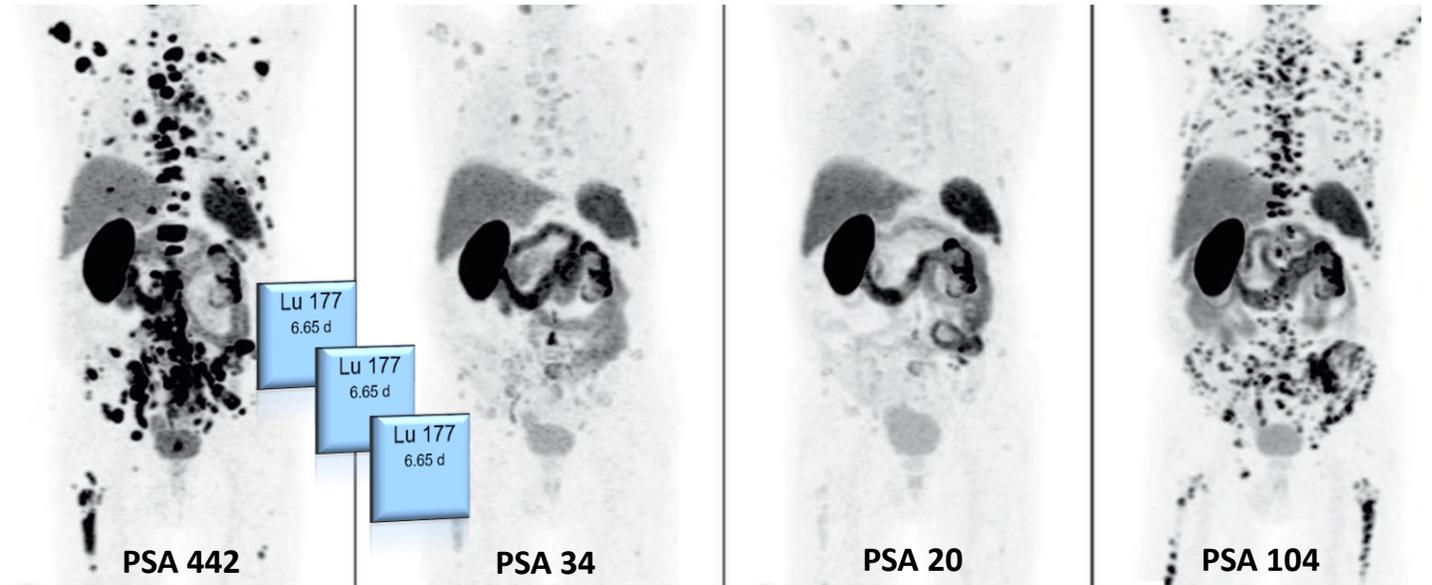
FDA approved in March 2022

Pre-therapeutic  
tumor spread

After 3 cycles  
(3 months):  
delayed progression

After 3 cycles  
(6 months):  
delayed progression

After 3 cycles  
(9 months):  
**marked progression**



Iravani et al. 2020 Prostate Cancer and Prostatic Diseases, 23:38.

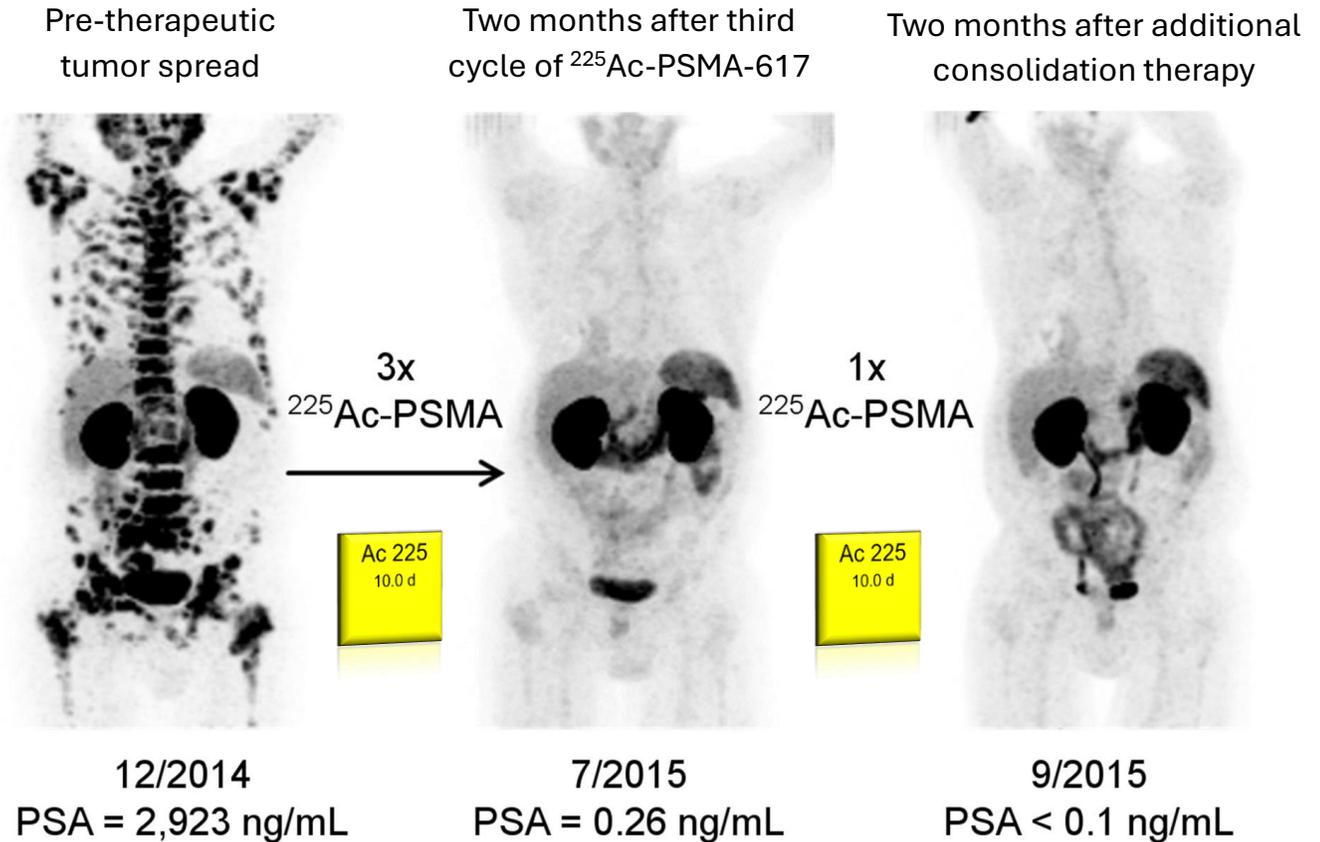
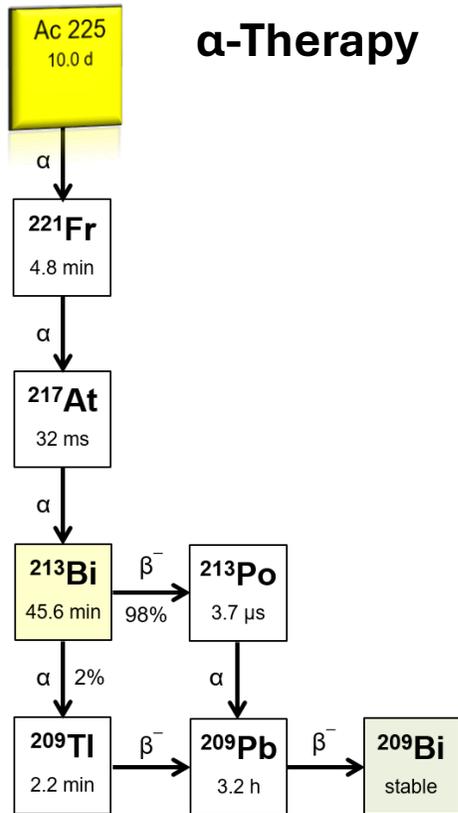


*<sup>177</sup>Lu is not sufficiently effective to eliminate single cancer cells and cancer cell clusters.*

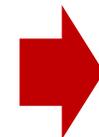


# $^{225}\text{Ac}$ -based Radioligand Therapy

## $^{225}\text{Ac}$ decay chain



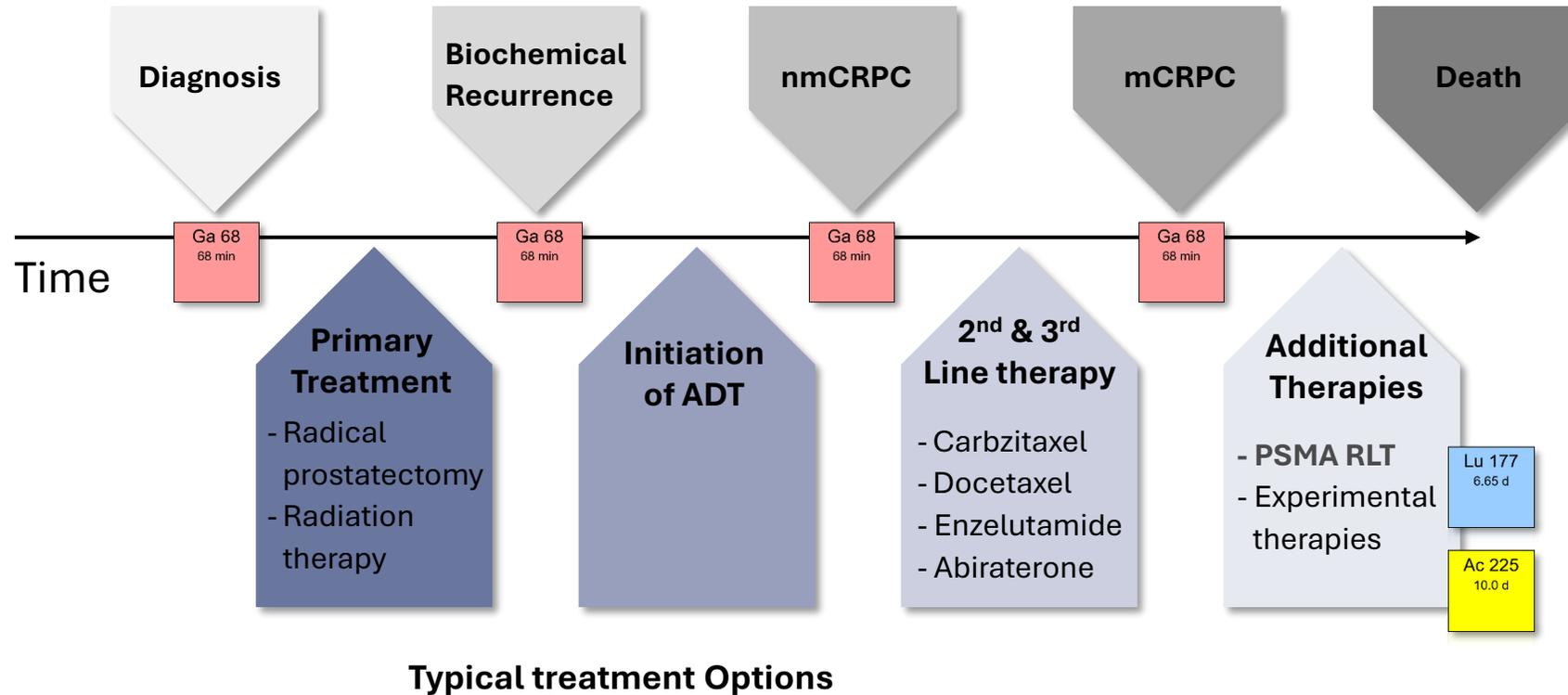
Kratochwil et al. 2016, J Nucl Med Mol Imaging 57:1941.



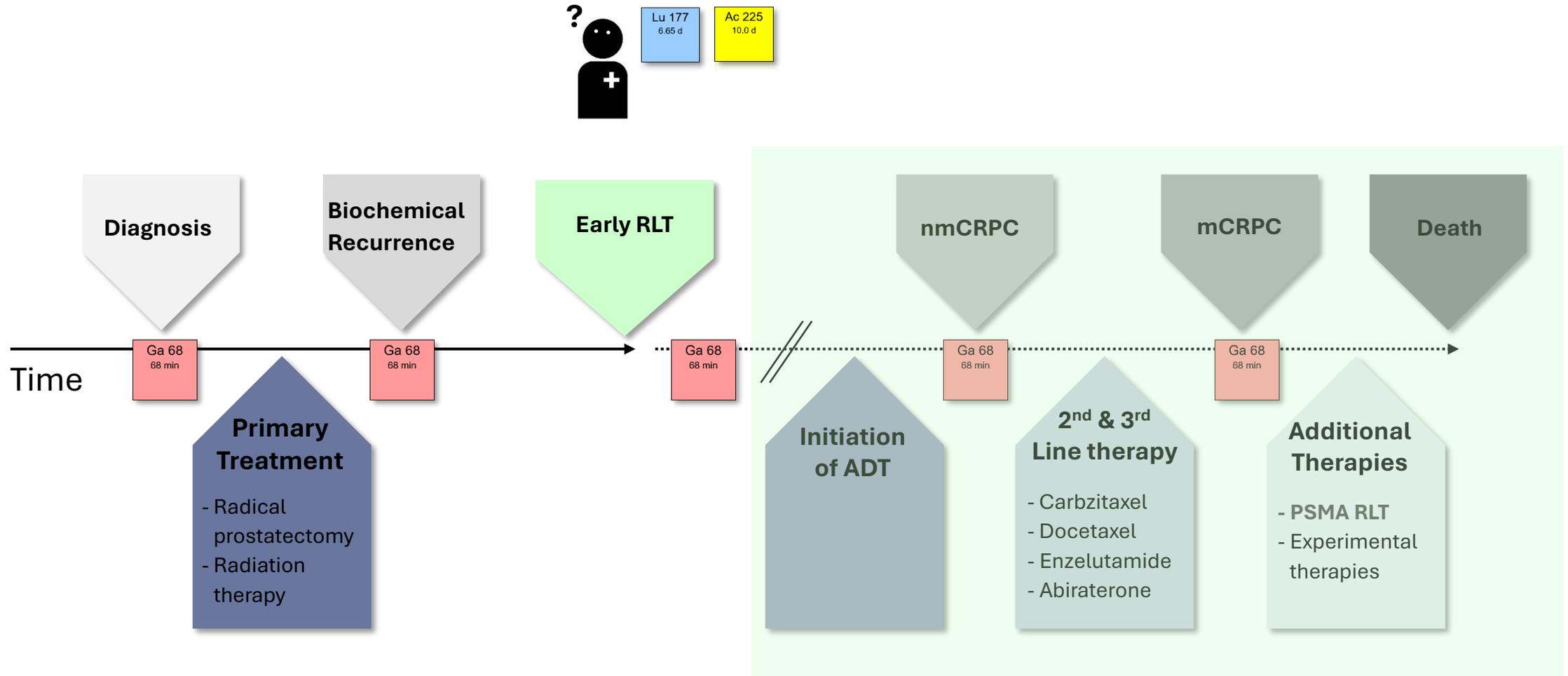
$^{225}\text{Ac}$  is effective to eliminate micrometastases but may cause severe side effects.

# Current Application of PSMA Radioligands

## Key clinical events during the course of prostate cancer



# Application of RLT at an Earlier Disease Stage



# Comparison of $^{161}\text{Tb}$ and $^{177}\text{Lu}$

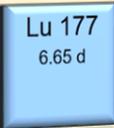
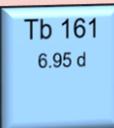
## Decay characteristics

Nuclide	$T_{1/2}$	$\beta^-$ - energy (mean)	$\gamma$ radiation; energy (%)	Conversion & Auger electrons
	6.65 days	134 keV	54 keV (4%) 113 keV (6%) 208 keV(10%)	No
	6.95 days	154 keV	45 keV (18%) 49 keV (17%) 75 keV (10%)	<b>Yes!</b>

\*Auger electrons: energy: 20 eV-1 keV; tissue range: 2-500 nm; LET: 4-26 keV/ $\mu\text{m}$

# Treatment of Macro- and Micrometastases

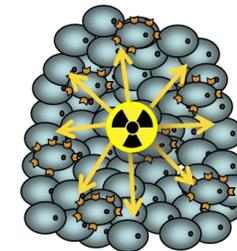
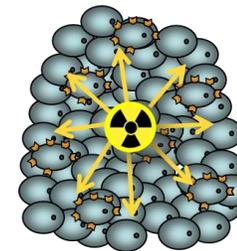
## Decay characteristics

Nuclide	$T_{1/2}$	$\beta^-$ - energy (mean)	$\gamma$ radiation; energy (%)	Conversion & Auger electrons
 Lu 177 6.65 d	6.65 days	134 keV	54 keV (4%) 113 keV (6%) 208 keV(10%)	No
 Tb 161 6.95 d	6.95 days	154 keV	45 keV (18%) 49 keV (17%) 75 keV (10%)	<b>Yes!</b>

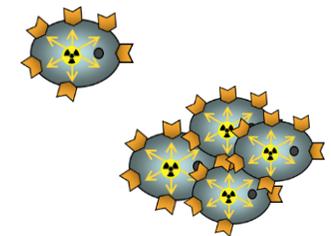
## Hypothesis

Macrometastases

Single Cancer Cells & Cancer Cell Clusters

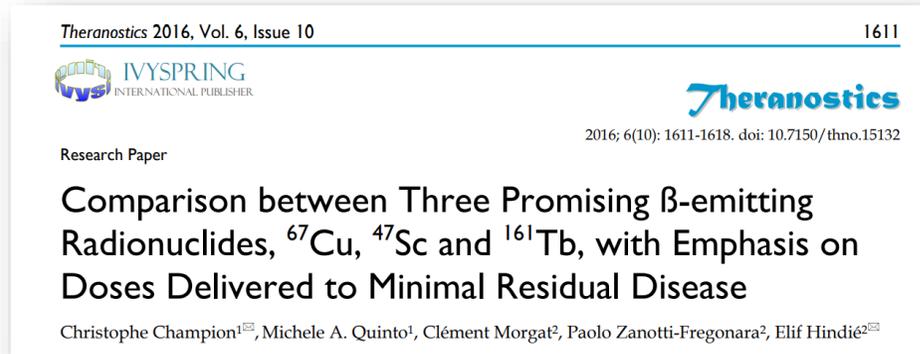


+



\*Auger electrons: energy: 20 eV-1 keV; tissue range: 2-500 nm; LET: 4-26 keV/ $\mu$ m

## Dose calculations



### Theoretical dose calculations:

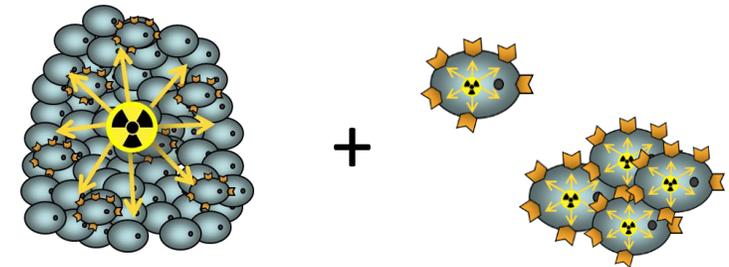
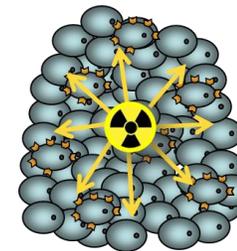
The absorbed dose in single cancer cells or cell monolayers is 3-4-fold increased when using  $^{161}\text{Tb}$  as compared to  $^{177}\text{Lu}$ .

Champion et al. 2016, *Theranostics*

## Hypothesis

Macrometastases

Single Cancer Cells & Cancer Cell Clusters



# More than a Decade of Preclinical Work with $^{161}\text{Tb}$



Zentralklinik Bad Berka

Baum et al.  
 $^{161}\text{Tb}$ -DOTATOC:  
First-in-human  
**JNM 2021**

Tschan et al.  
 $^{161}\text{Tb}$ -SibuDAB &  
 $^{161}\text{Tb}$ -PSMA-I&T:  
Therapy  
**JNM 2023**

Universitätsspital  
Basel

Fricke et al.  
 $^{161}\text{Tb}$ -DOTA-LM3  
Clinical Image  
**EJNMMI 2024**

Müller et al.  
 $^{149}\text{Tb}$ -,  $^{152}\text{Tb}$ -,  
 $^{155}\text{Tb}$ -,  $^{161}\text{Tb}$ -  
folate  
**JNM 2012**

Müller et al.  
 $^{161}\text{Tb}$ -folate &  
 $^{177}\text{Lu}$ -folate:  
Therapy  
**EJNMMI 2014**

Müller et al.  
 $^{161}\text{Tb}$ -PSMA-617:  
Therapy  
**EJNMMI 2019**

Borgna et al.  
 $^{161}\text{Tb}$ -DOTA-LM3 &  
 $^{161}\text{Tb}$ -DOTATOC:  
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**EJNMMI 2022**

Busslinger et al.  
 $^{161}\text{Tb}$ -DOTA-LM3 &  
 $^{161}\text{Tb}$ -DOTATATE  
Tolerability  
**EJNMMI 2024**



EANM'18  
Düsseldorf



2011    2012    2014    2016    2018    2020    2022    2023    2024

Tb 161  
6.95 d

The articles mentioned above are a selection; additional articles about production and preclinical research have been published.

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EANM'18  
Düsseldorf



2011 2012 2014 2016 2018 2020 2022 2023 2024

Tb 161  
6.95 d

The articles mentioned above are a selection; additional articles about production and preclinical research have been published.

# Preclinical Work with $^{161}\text{Tb}$ : Somatostatin Analogues



  
Zentralklinik Bad Berka

 Universitätsspital  
Basel

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EANM'18  
Düsseldorf



2011 2012

2014

2016

2018

2020

2022

2023

2024

**Somatostatin Analogues**

# Application of $^{161}\text{Tb}$ with Somatostatin Analogues



Application of  $^{161}\text{Tb}$  in combination  
with somatostatin analogues?





Application of  $^{161}\text{Tb}$  in combination with somatostatin analogues?



**NETRF Petersen Award 2018** to investigate the utility of  $^{161}\text{Tb}$  in combination with DOTATOC



**R. Schibli**  
R.P. Baum  
N.P. van der Meulen  
C. Müller



**Goal:** Further development of  $^{161}\text{Tb}$  and translation of  $^{161}\text{Tb}$ -DOTATOC to a first-in-human application.

# First Patient Image of a $^{161}\text{Tb}$ -based Radiopharmaceutical



## “First-in-human” application

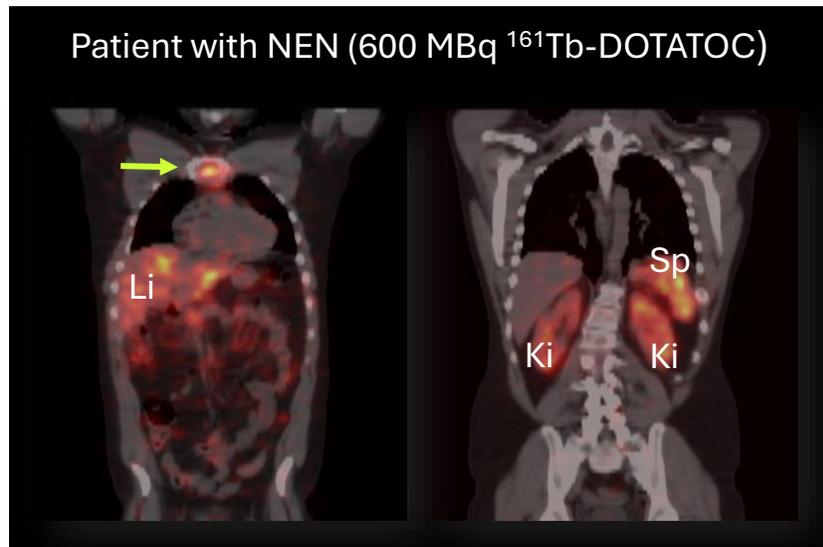


Zentralklinik Bad Berka, Germany

R. Baum



P. Bernhardt



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# Application of $^{161}\text{Tb}$ with Somatostatin Receptor Antagonists?

## “First-in-human” application



Zentralklinik Bad Berka, Germany

R. Baum



P. Bernhardt

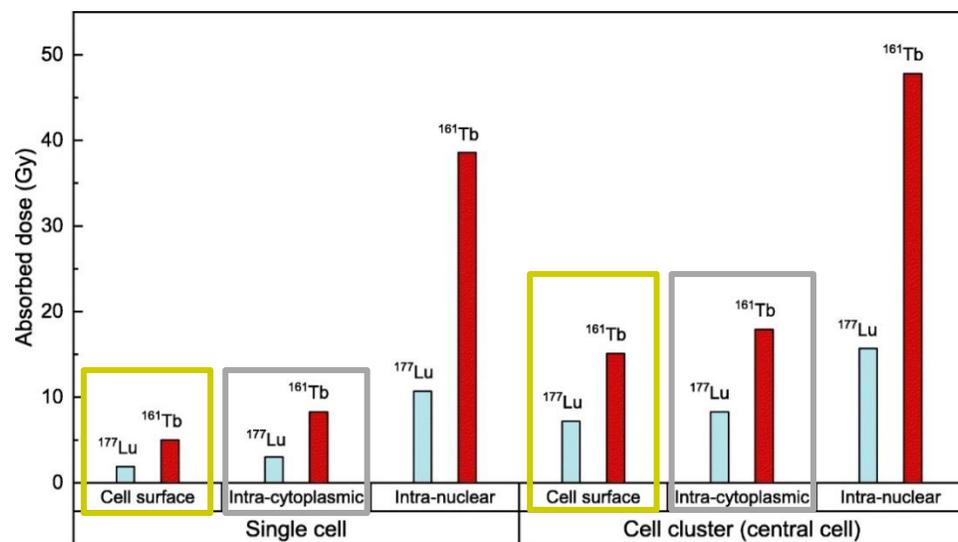


Application of  $^{161}\text{Tb}$  in combination with SST receptor **antagonists**?



# $^{161}\text{Tb}$ Outperforms $^{177}\text{Lu}$ also when Localized at the Membrane

## “Dose calculations”



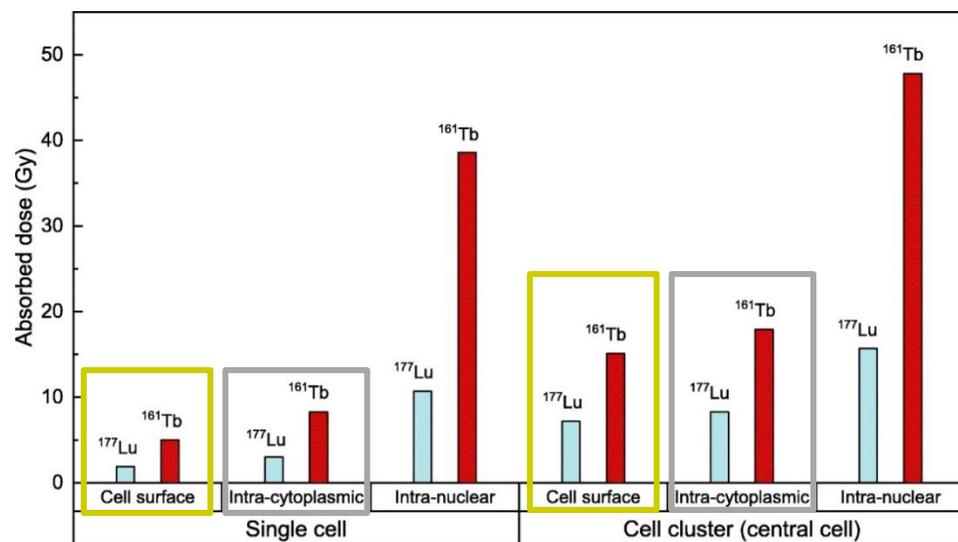
*$^{161}\text{Tb}$  should be a better candidate than  $^{177}\text{Lu}$  for irradiating single tumor cells and micrometastases, **regardless of the radionuclide distribution.***



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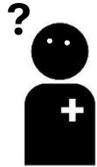


*$^{161}\text{Tb}$  should be a better candidate than  $^{177}\text{Lu}$  for irradiating single tumor cells and micrometastases, **regardless of the radionuclide distribution.***

## Enhancement factor (single cell)

	Cell surface	Intra-cytoplasmic	Whole cell	Intra-nuclear
$^{177}\text{Lu}$	1.9	3.0	5.8	10.7
$^{161}\text{Tb}$	5.0	8.3	19.5	38.6
$^{161}\text{Tb}/^{177}\text{Lu}$	2.6	2.8	3.4	3.6

# The Use of SSTR Antagonists with $^{161}\text{Tb}$ makes Sense



Application of  $^{161}\text{Tb}$  in combination with SST receptor **antagonists**?



The comparison of SST receptor agonists (internalizing peptide) and SST receptor antagonists (non-internalizing peptide) with terbium-161 makes sense.

## Enhancement factor (single cell)

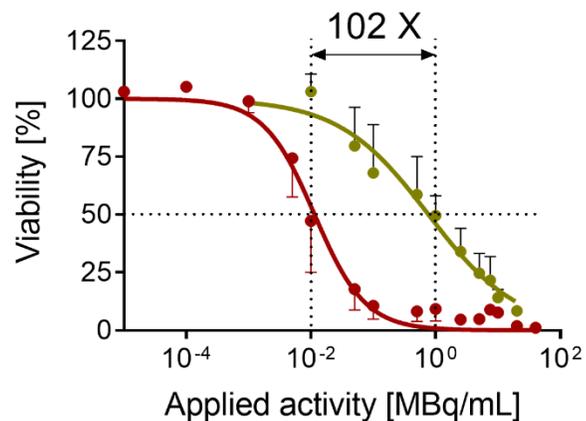
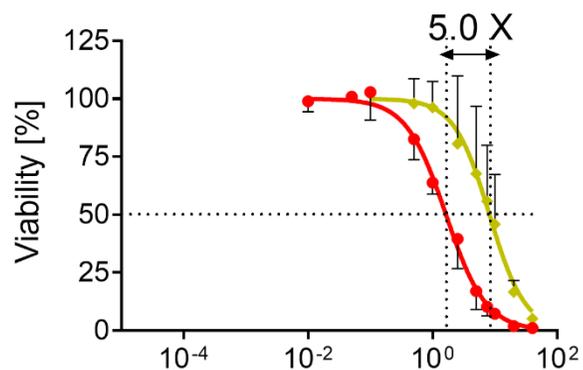
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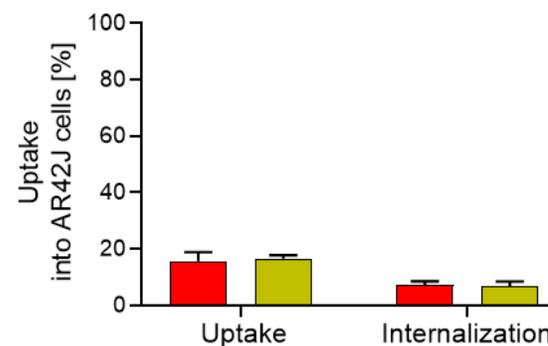


# In Vitro Cell Uptake Study: Agonist vs. Antagonist

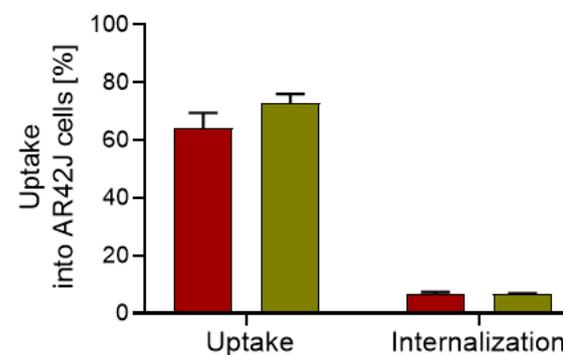
## Tumor cell viability (MTT)



## Cell uptake & internalization



161Tb-DOTATOC  
177Lu-DOTATOC

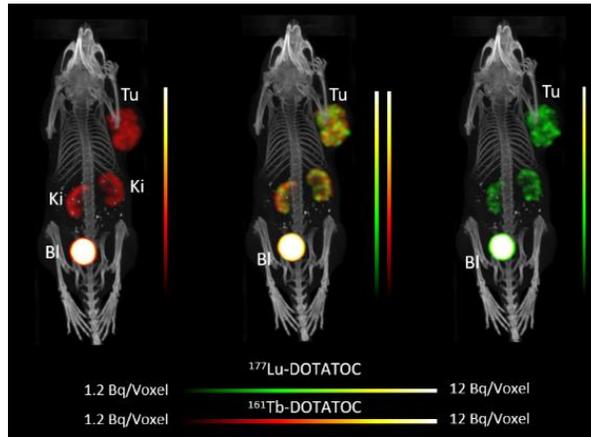


161Tb-DOTA-LM3  
177Lu-DOTA-LM3

# Equal Tissue Distribution of $^{161}\text{Tb}$ - & $^{177}\text{Lu}$ -based Radiopeptides

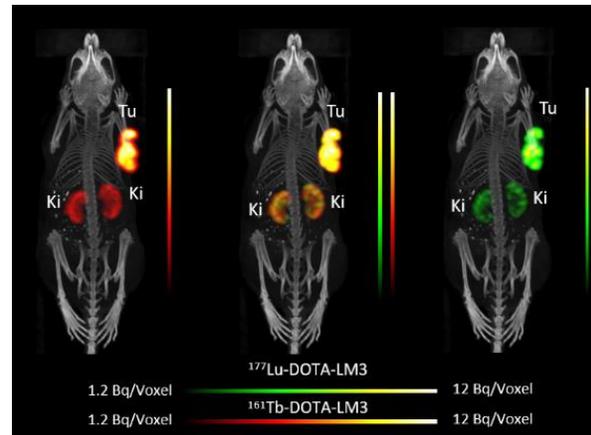


## Dual-isotope SPECT imaging



### DOTATOC

15 MBq  $^{161}\text{Tb}$   
& 15 MBq  $^{177}\text{Lu}$ ;  
1 nmol/mouse

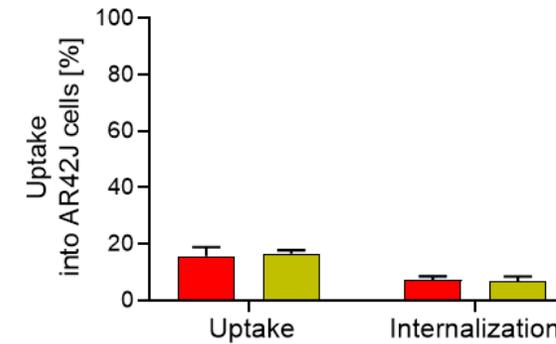


### DOTA-LM3

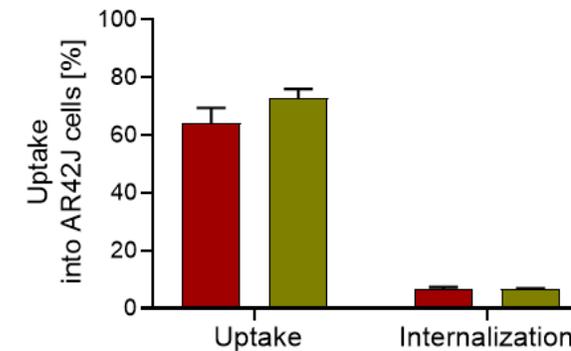
15 MBq  $^{161}\text{Tb}$   
& 15 MBq  $^{177}\text{Lu}$ ;  
1 nmol/mouse

AR42J tumor  
bearing mice

## Cell uptake & internalization



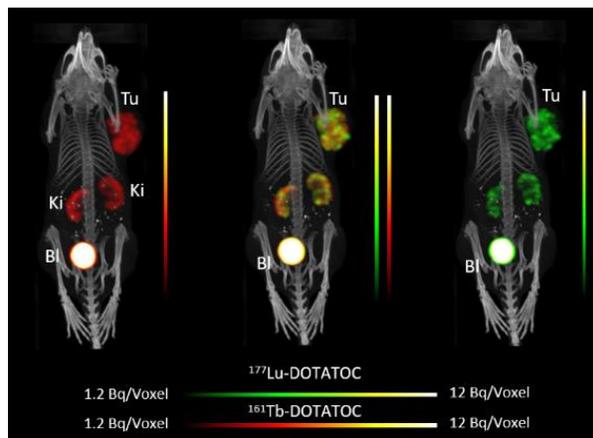
■  $^{161}\text{Tb}$ -DOTATOC  
■  $^{177}\text{Lu}$ -DOTATOC



■  $^{161}\text{Tb}$ -DOTA-LM3  
■  $^{177}\text{Lu}$ -DOTA-LM3

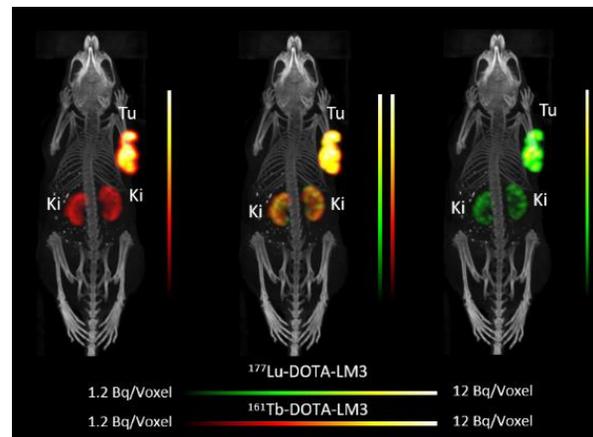
# Preclinical Therapy – $^{161}\text{Tb}$ vs. $^{177}\text{Lu}$ & Agonist vs Antagonist

## Dual-isotope SPECT imaging



### DOTATOC

15 MBq  $^{161}\text{Tb}$   
& 15 MBq  $^{177}\text{Lu}$ ;  
1 nmol/mouse



### DOTA-LM3

15 MBq  $^{161}\text{Tb}$   
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1 nmol/mouse

AR42J tumor  
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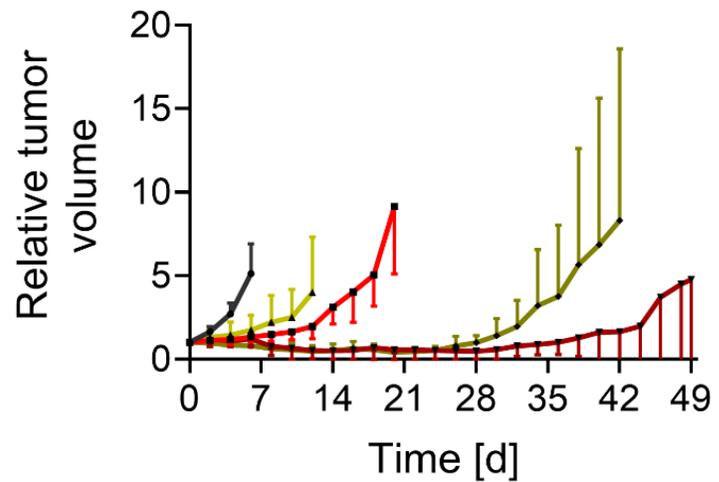


How does  $^{161}\text{Tb}$  perform compared to  $^{177}\text{Lu}$  for SST receptor targeted therapy?



Preclinical therapy study in AR42J-tumor-bearing mice injected with 2 x 10 MBq of the respective SST analogue.

## Tumor growth curves



- Control (untreated)
- <sup>161</sup>Tb-DOTATOC (2 x 10 MBq)
- <sup>161</sup>Tb-DOTA-LM3 (2 x 10 MBq)
- <sup>177</sup>Lu-DOTATOC (2 x 10 MBq)
- <sup>177</sup>Lu-DOTA-LM3 (2 x 10 MBq)



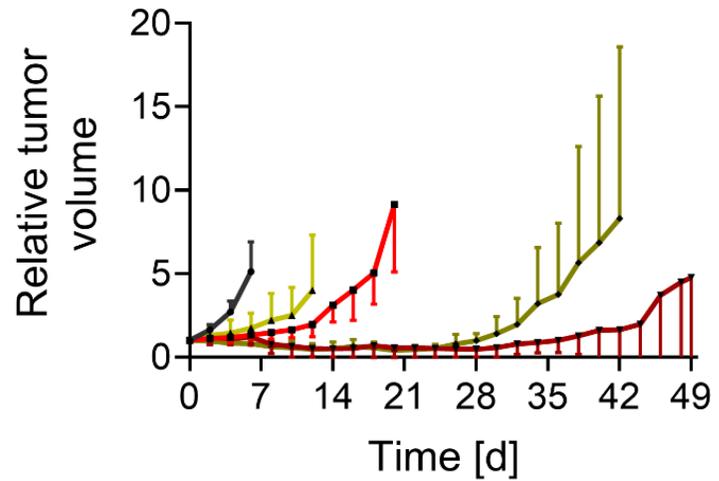
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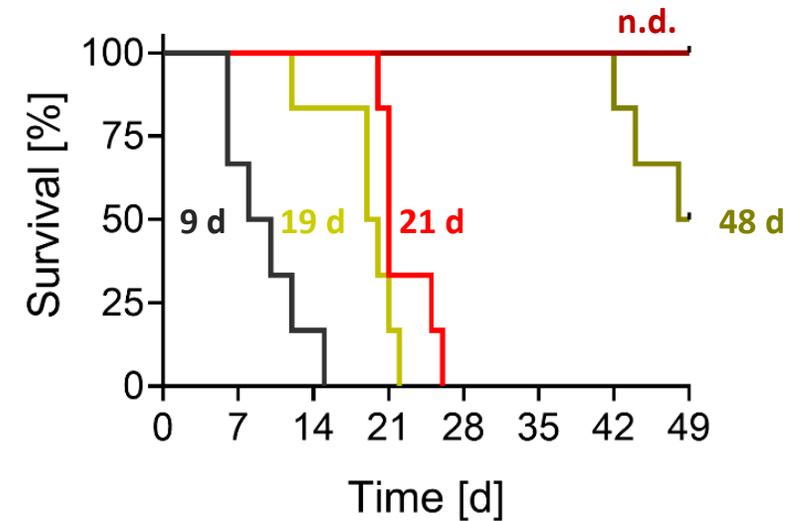
# Preclinical Therapy – Survival Curves

## Tumor growth curves



- Control (untreated)
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- <sup>177</sup>Lu-DOTA-LM3 (2 x 10 MBq)

## Survival curves

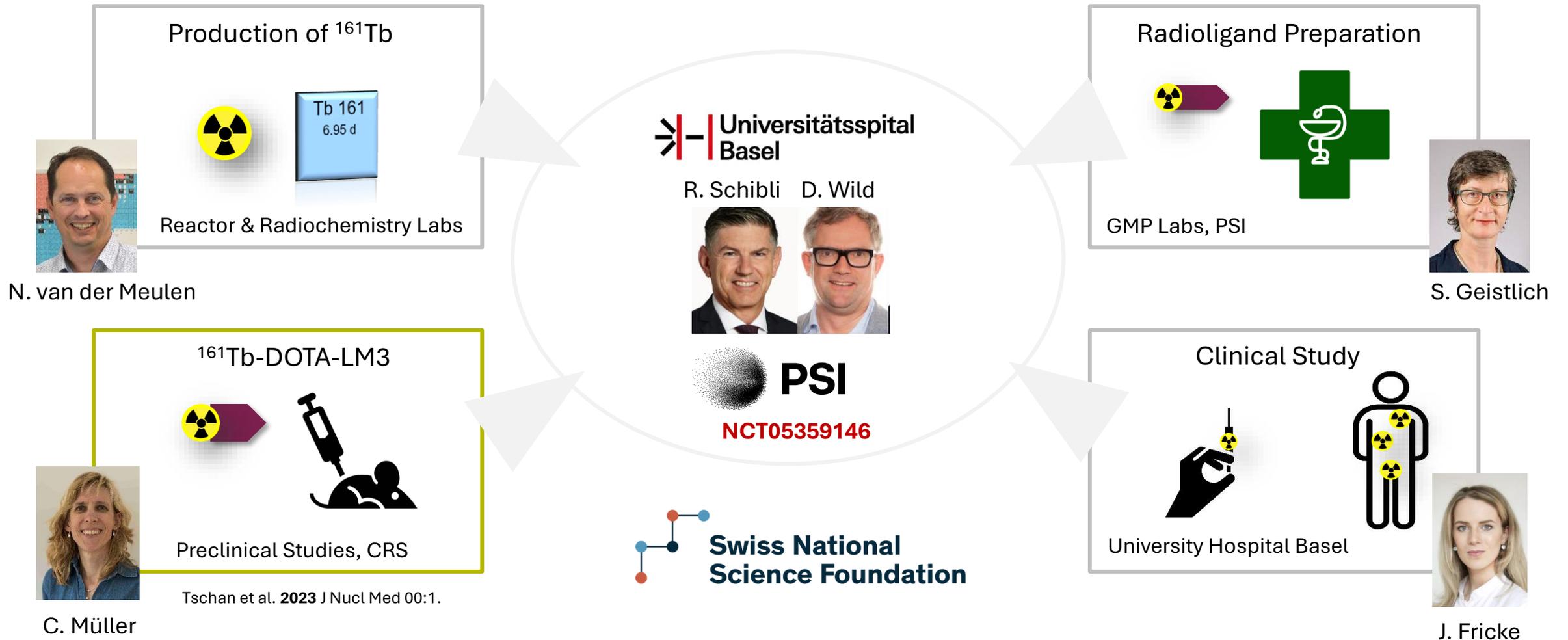


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# «BETA PLUS» Clinical Translation of $^{161}\text{Tb}$ -DOTA-LM3



Combined Beta- Plus Auger Electron Therapy Using a Novel Somatostatin Receptor Subtype 2 Antagonist  
Labelled with Terbium-161 ( $^{161}\text{Tb}$ -DOTA-LM3)



# Tolerability Study in Mice (without Tumors)



## Study design

Immunocompetent, female mice (FVB)

■ Control (untreated)

■  $^{161}\text{Tb}$ -DOTA-LM3 (20 MBq) ■  $^{161}\text{Tb}$ -DOTATATE (20 MBq)

Peptide mass: 0.2 nmol peptide per mouse

- Blood was taken from the sublingual vein to determine blood cell counts and prepare blood smears at:  
Day 10, Day 28 and Day 56 after application
- At Day 56, full necropsy was performed:
  - Mass of liver, kidneys, spleen and brain and respective ratios
  - blood plasma parameters (BUN, ALP, TBIL, ALB)



# Tolerability Study: Blood Cell Counts

## Study design

Immunocompetent, female mice (FVB)

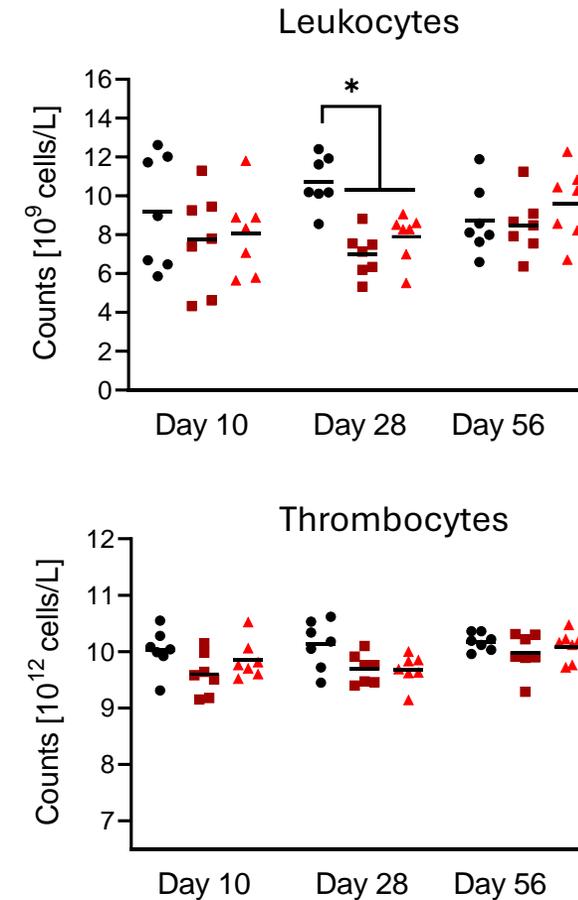
■ Control (untreated)

■  $^{161}\text{Tb}$ -DOTA-LM3 (20 MBq) ■  $^{161}\text{Tb}$ -DOTATATE (20 MBq)

Peptide mass: 0.2 nmol peptide per mouse

- Blood was taken from the sublingual vein to determine blood cell counts and prepare blood smears at: Day 10, Day 28 and Day 56 after application
- At Day 56, full necropsy was performed:
  - Mass of liver, kidneys, spleen and brain and respective ratios
  - blood plasma parameters (BUN, ALP, TBIL, ALB)

## Blood cell counts

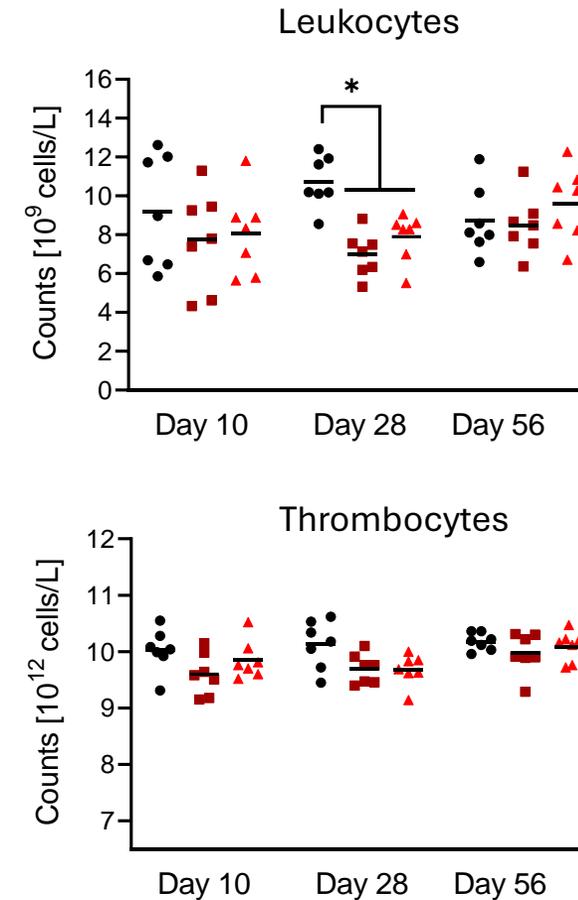


# Conclusions of the Tolerability Study

## Conclusion:

- At **20 MBq** radiopeptide per mouse, no changes were determined as compared to control mice.
- At **100 MBq** radiopeptide per mouse, the blood cell counts dropped, but mostly recovered over time.
- The drop was more pronounced for the **antagonist than for the agonist** irrespective of the radionuclide.
- $^{161}\text{Tb}$ -labeled radiopeptides seemed to be well tolerated.**
- Based on this study, it can be assumed that  **$^{161}\text{Tb}$ -DOTA-LM3 can be used at equal activities as  $^{177}\text{Lu}$ -DOTA-LM3** (which is lower than for the agonists)

## Blood cell counts



# Preclinical Work with $^{161}\text{Tb}$ : PSMA Ligands



Zentralklinik Bad Berka

Universitätsspital Basel

Baum et al.  
 $^{161}\text{Tb}$ -DOTATOC:  
First-in-human  
**JNM 2021**

Tschan et al.  
 $^{161}\text{Tb}$ -SibuDAB &  
 $^{161}\text{Tb}$ -PSMA-I&T:  
Therapy  
**JNM 2023**

Fricke et al.  
 $^{161}\text{Tb}$ -DOTA-LM3  
Clinical Image  
**EJNMMI 2024**

Müller et al.  
 $^{149}\text{Tb}$ -,  $^{152}\text{Tb}$ -,  
 $^{155}\text{Tb}$ -,  $^{161}\text{Tb}$ -  
folate  
**JNM 2012**

Müller et al.  
 $^{161}\text{Tb}$ -folate &  
 $^{177}\text{Lu}$ -folate:  
Therapy  
**EJNMMI 2014**

Marie Curie  
Award



Müller et al.  
 $^{161}\text{Tb}$ -PSMA-617:  
Therapy  
**EJNMMI 2019**

Borgna et al.  
 $^{161}\text{Tb}$ -DOTA-LM3 &  
 $^{161}\text{Tb}$ -DOTATOC:  
Therapy  
**EJNMMI 2022**

Busslinger et al.  
 $^{161}\text{Tb}$ -DOTA-LM3 &  
 $^{161}\text{Tb}$ -DOTATATE  
Tolerability  
**EJNMMI 2024**



EANM'18  
Düsseldorf

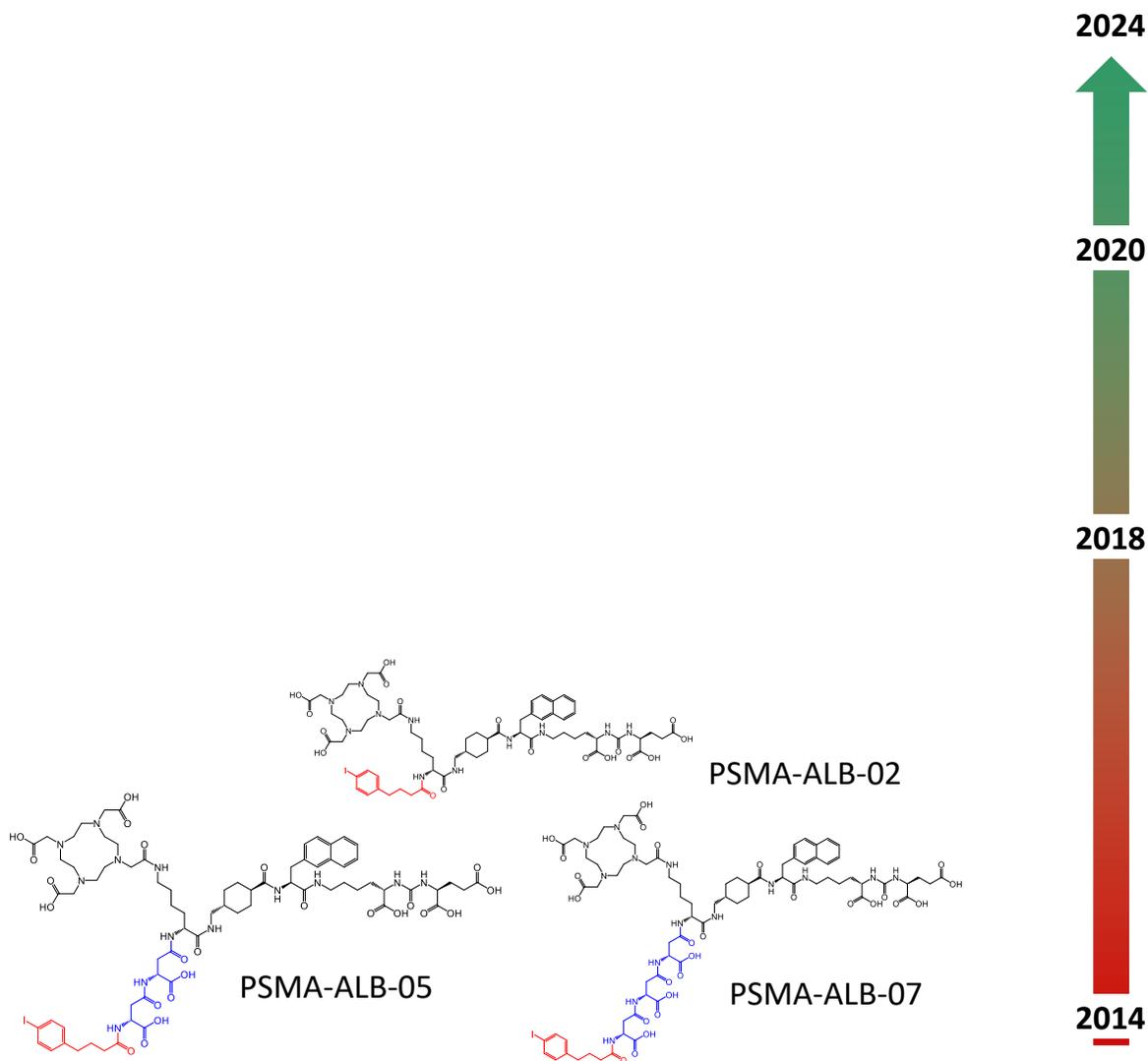


2011 2012 2014 2016 2018 2020 2022 2023 2024



**PSMA Ligands**

# Development of New PSMA Ligands



# Development of New PSMA Ligands

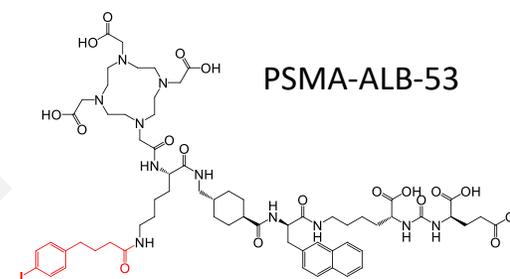
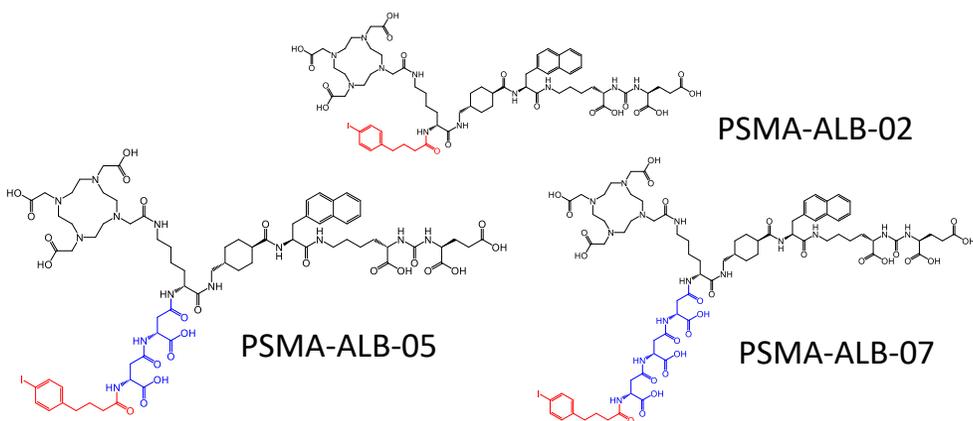


2024

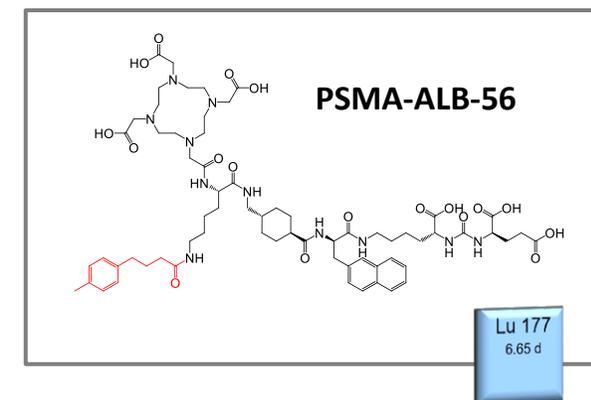
2020

2018

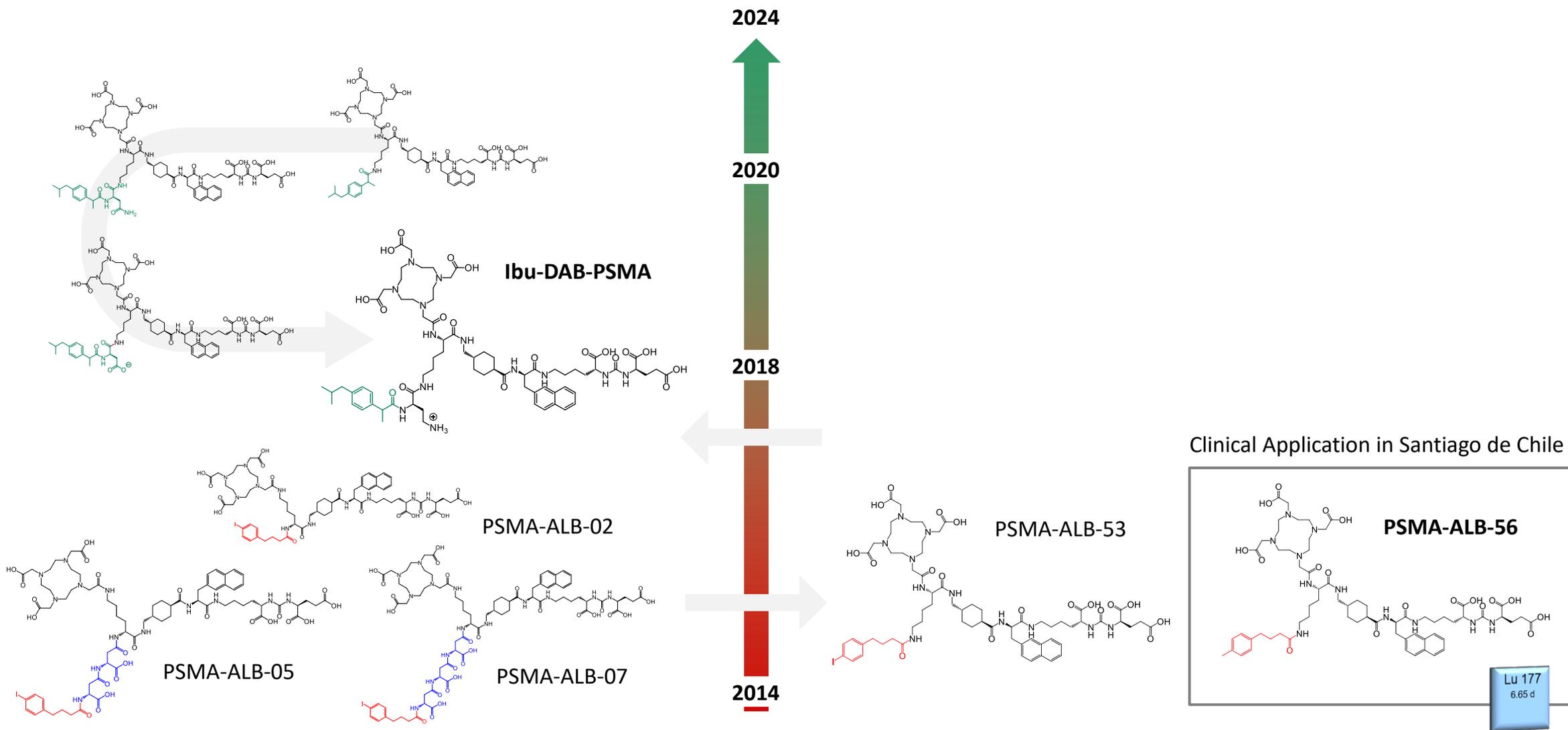
2014



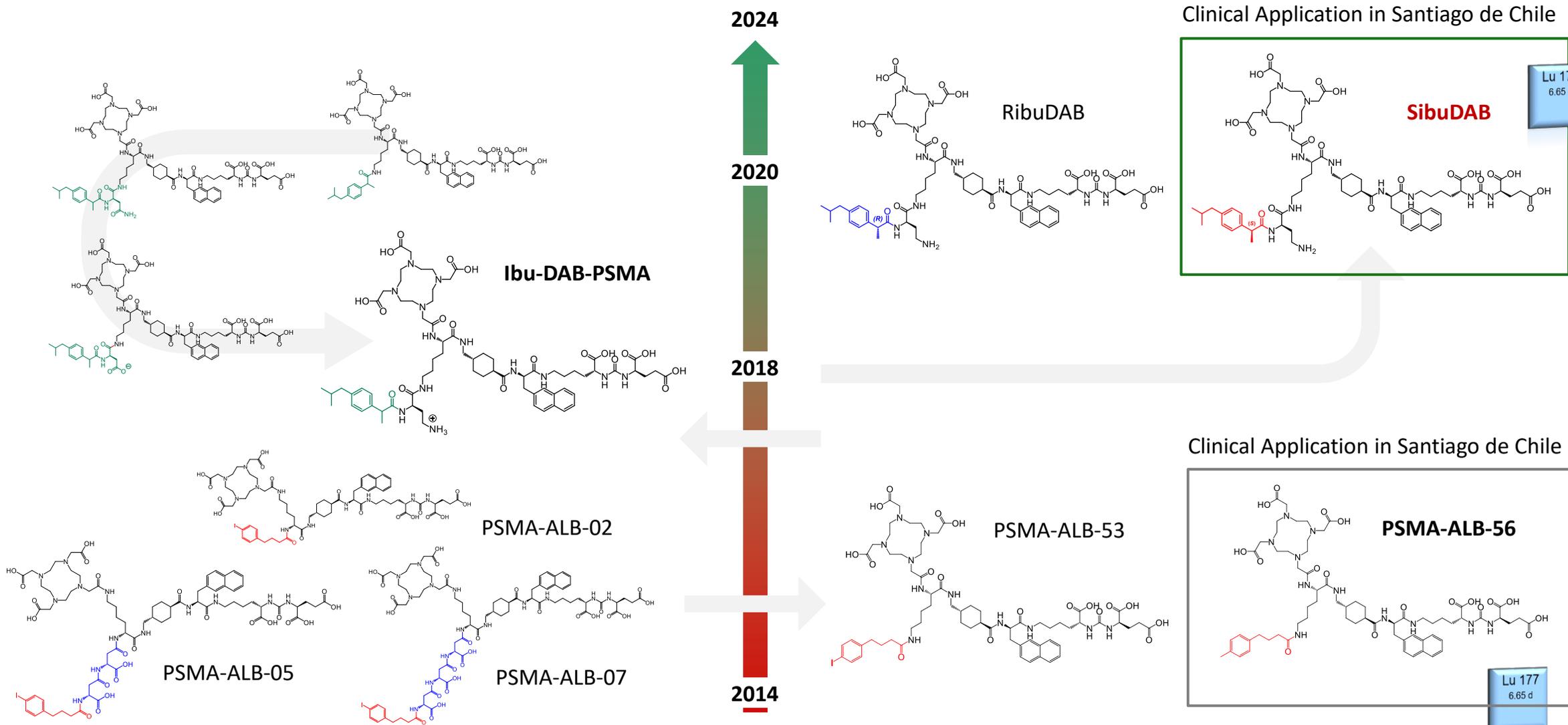
Clinical Application in Santiago de Chile



# Development of New PSMA Ligands



# Development of New PSMA Ligands



# Development of New PSMA Ligands



European Journal of Nuclear Medicine and Molecular Imaging (2022) 49:470–480  
<https://doi.org/10.1007/s00259-021-05446-5>

ORIGINAL ARTICLE



## Impact of the mouse model and molar amount of injected ligand on the tissue distribution profile of PSMA radioligands

Viviane J. Tschan<sup>1</sup> · Francesca Borgna<sup>1</sup> · Roger Schibli<sup>1,2</sup> · Cristina Müller<sup>1,2</sup>

*Theranostics* 2020, Vol. 10, Issue 4

1678



2020; 10(4): 1678-1693. doi: 10.7150/thno.40482

Research Paper

## Development of a new class of PSMA radioligands comprising ibuprofen as an albumin-binding entity

Luisa M. Deberle<sup>1,2</sup>, Martina Benešová<sup>1,2</sup>, Christoph A. Umbricht<sup>2</sup>, Francesca Borgna<sup>2</sup>, Manuel Büchler<sup>2</sup>, Konstantin Zhernosekov<sup>3</sup>, Roger Schibli<sup>1,2</sup>, Cristina Müller<sup>1,2</sup>



Article

Cite This: *Mol. Pharmaceutics* 2018, 15, 934–946

## Albumin-Binding PSMA Ligands: Optimization of the Tissue Distribution Profile

Martina Benešová,<sup>†,‡</sup> Christoph A. Umbricht,<sup>†</sup> Roger Schibli,<sup>†,‡</sup> and Cristina Müller<sup>\*,†,‡</sup>

<sup>†</sup>Center for Radiopharmaceutical Sciences ETH-PSI-USZ, Paul Scherrer Institut, 5232 Villigen-PSI, Switzerland

<sup>‡</sup>Department of Chemistry and Applied Biosciences, ETH Zurich, 8093 Zurich, Switzerland

2024

2022

2020

2018



pubs.acs.org/molecularpharmaceutics

Article

## Preclinical Investigations to Explore the Difference between the Diastereomers [<sup>177</sup>Lu]Lu-SibuDAB and [<sup>177</sup>Lu]Lu-RibuDAB toward Prostate Cancer Therapy

Francesca Borgna,<sup>§</sup> Luisa M. Deberle,<sup>§</sup> Sarah D. Busslinger, Viviane J. Tschan, Laura M. Walde, Anna E. Becker, Roger Schibli, and Cristina Müller<sup>\*</sup>

European Journal of Nuclear Medicine and Molecular Imaging (2021) 48:893–903  
<https://doi.org/10.1007/s00259-020-05022-3>

ORIGINAL ARTICLE



## Biodistribution and dosimetry of a single dose of albumin-binding ligand [<sup>177</sup>Lu]Lu-PSMA-ALB-56 in patients with mCRPC

Vasko Kramer<sup>1,2</sup> · René Fernández<sup>1</sup> · Wencke Lehnert<sup>3,4</sup> · Luis David Jiménez-Franco<sup>3</sup> · Cristian Soza-Ried<sup>1</sup> · Elisabeth Eppard<sup>2</sup> · Matias Ceballos<sup>1</sup> · Marian Meckel<sup>5</sup> · Martina Benešová<sup>6,7</sup> · Christoph A. Umbricht<sup>6</sup> · Andreas Kluge<sup>3</sup> · Roger Schibli<sup>6,7</sup> · Konstantin Zhernosekov<sup>5</sup> · Horacio Amaral<sup>1,2</sup> · Cristina Müller<sup>6,7</sup>



Article

Cite This: *Mol. Pharmaceutics* 2018, 15, 2297–2306

## Preclinical Development of Novel PSMA-Targeting Radioligands: Modulation of Albumin-Binding Properties To Improve Prostate Cancer Therapy

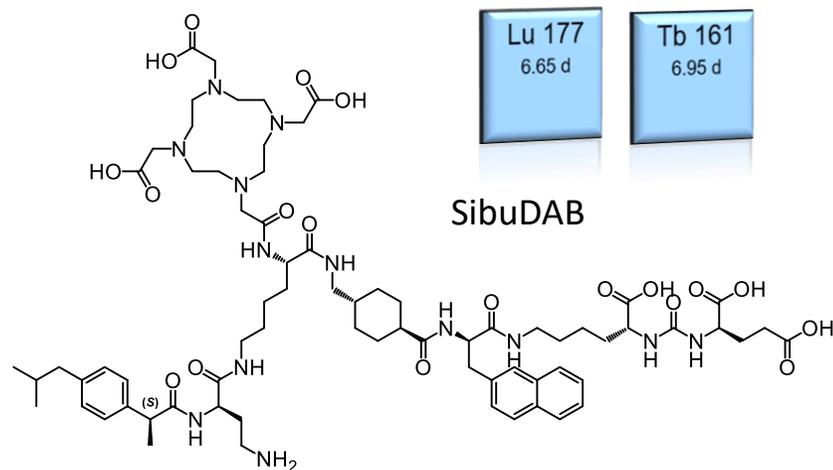
Christoph A. Umbricht,<sup>†</sup> Martina Benešová,<sup>†,‡</sup> Roger Schibli,<sup>†,‡</sup> and Cristina Müller<sup>\*,†,‡</sup>

<sup>†</sup>Center for Radiopharmaceutical Sciences ETH-PSI-USZ, Paul Scherrer Institut, 5232 Villigen-PSI, Switzerland

<sup>‡</sup>Department of Chemistry and Applied Biosciences, ETH Zurich, 8093 Zurich, Switzerland

# SibuDAB in Combination with $^{161}\text{Tb}$ and $^{177}\text{Lu}$

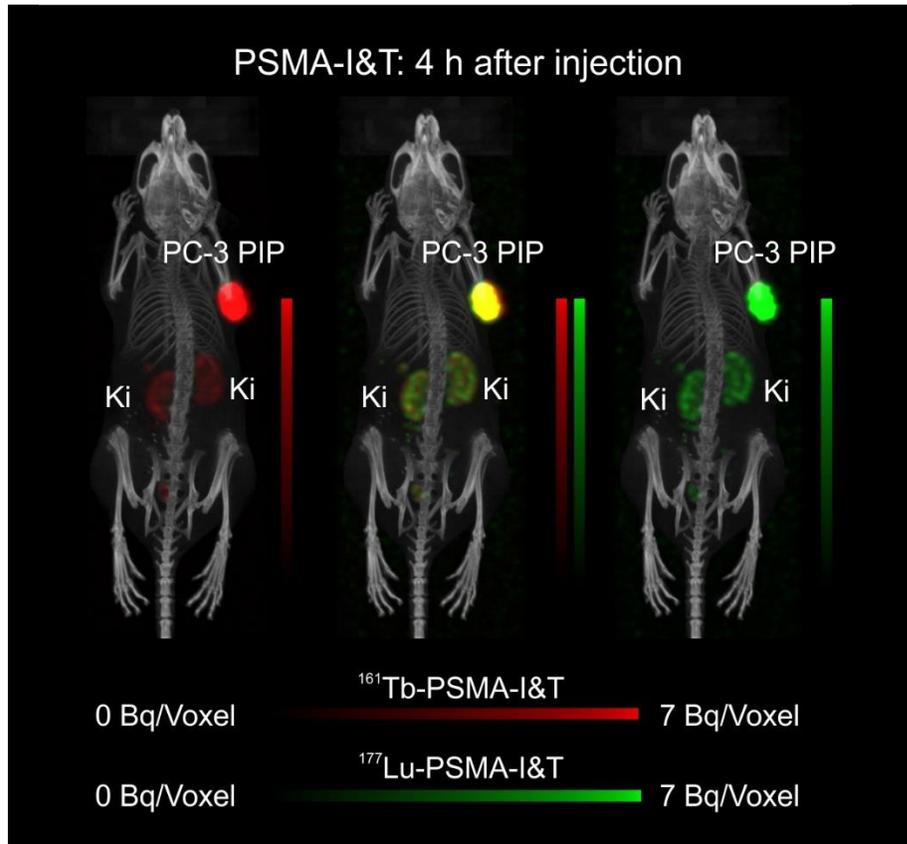
## SibuDAB



Are  $^{161}\text{Tb}$  and  $^{177}\text{Lu}$  interchangeable?

# Equal Biodistribution of $^{161}\text{Tb}$ - and $^{177}\text{Lu}$ -based PSMA Ligands

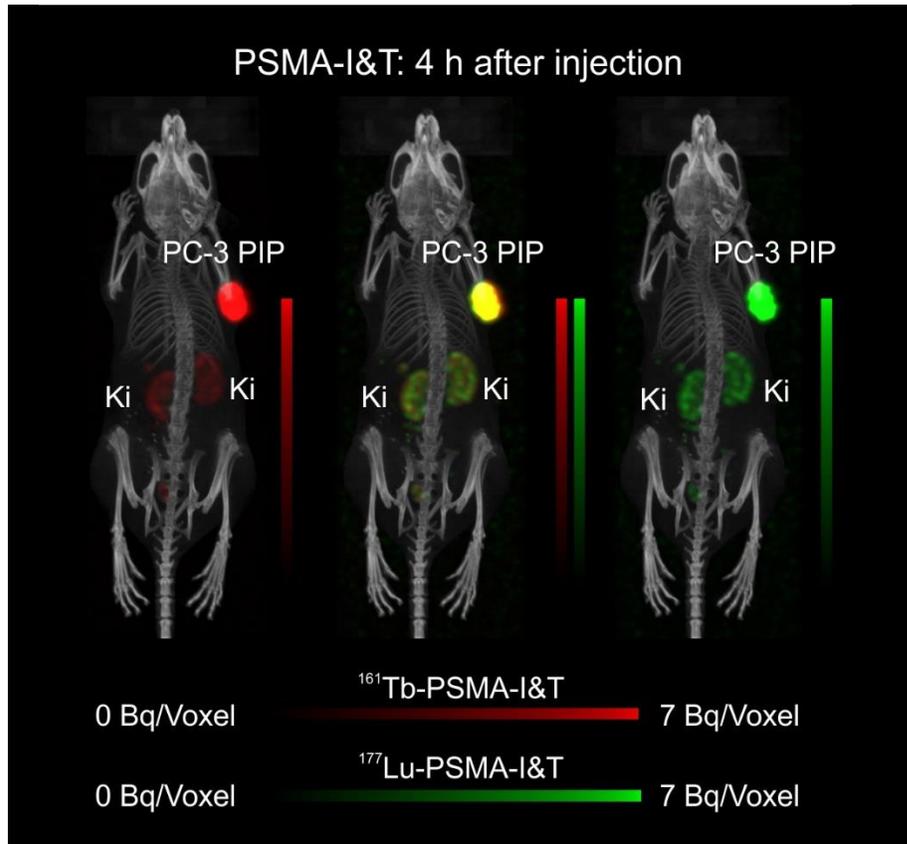
## PSMA-I&T



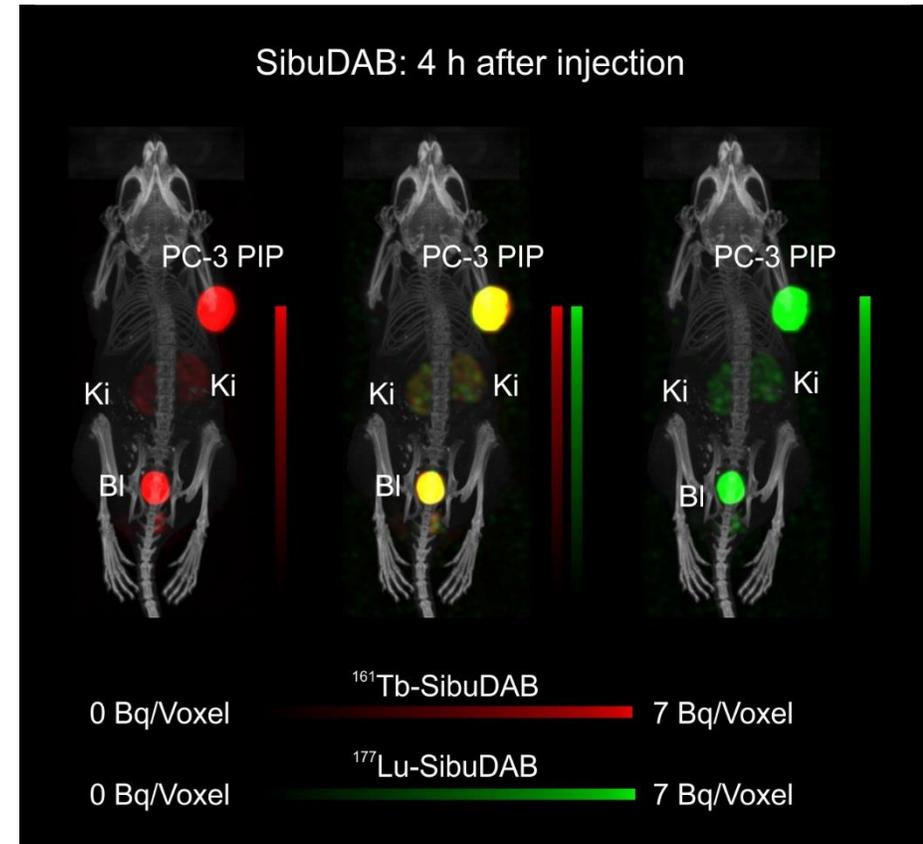
Are  $^{161}\text{Tb}$  and  $^{177}\text{Lu}$  interchangeable?

# Equal Biodistribution of $^{161}\text{Tb}$ - and $^{177}\text{Lu}$ -based PSMA Ligands

## PSMA-I&T



## SibuDAB

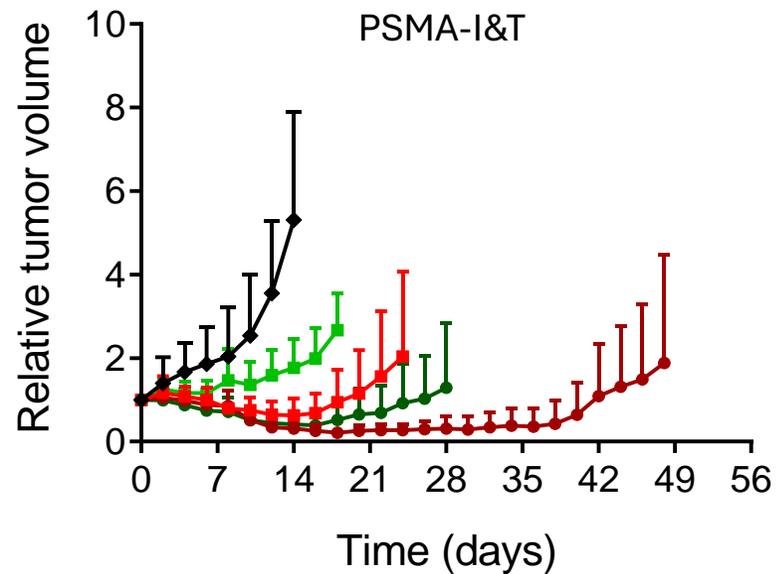


# PSMA-I&T and SibuDAB: Tumor Growth Curves



## Tumor growth curves

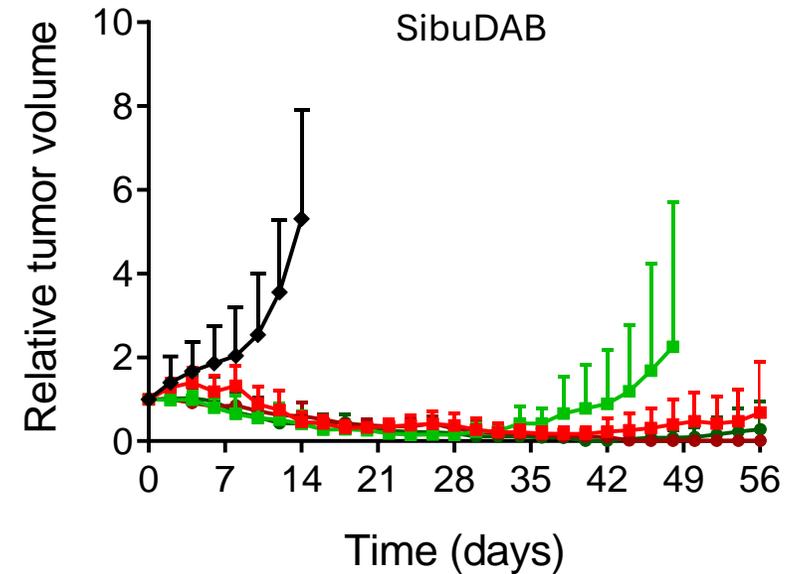
(PC-3 PIP tumor-bearing nude mice)



- Vehicle (control)
- <sup>161</sup>Tb-PSMA-I&T (5 MBq)
- <sup>161</sup>Tb-PSMA-I&T (10 MBq)
- <sup>177</sup>Lu-PSMA-I&T (5 MBq)
- <sup>177</sup>Lu-PSMA-I&T (10 MBq)

## Tumor growth curves

(PC-3 PIP tumor-bearing nude mice)



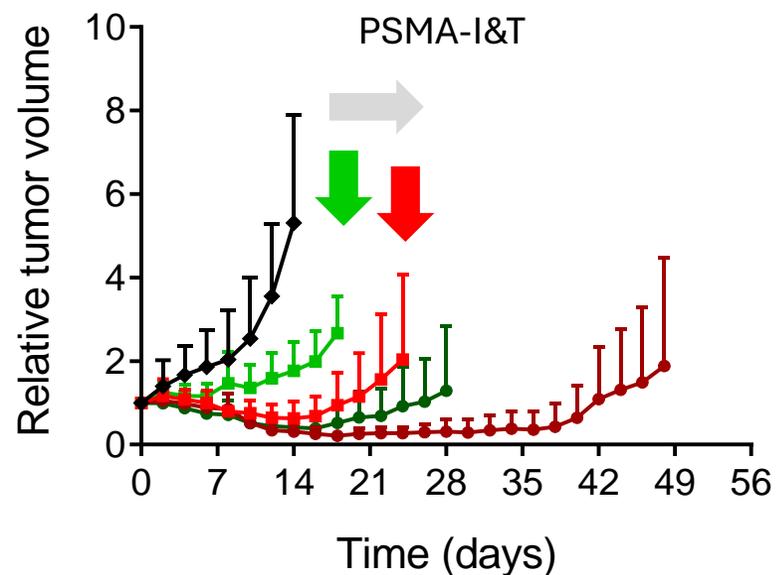
- Vehicle (control)
- <sup>161</sup>Tb-SibuDAB (5 MBq)
- <sup>161</sup>Tb-SibuDAB (10 MBq)
- <sup>177</sup>Lu-SibuDAB (5 MBq)
- <sup>177</sup>Lu-SibuDAB (10 MBq)

# PSMA-I&T and SibuDAB: $^{161}\text{Tb}$ versus $^{177}\text{Lu}$



## Tumor growth curves

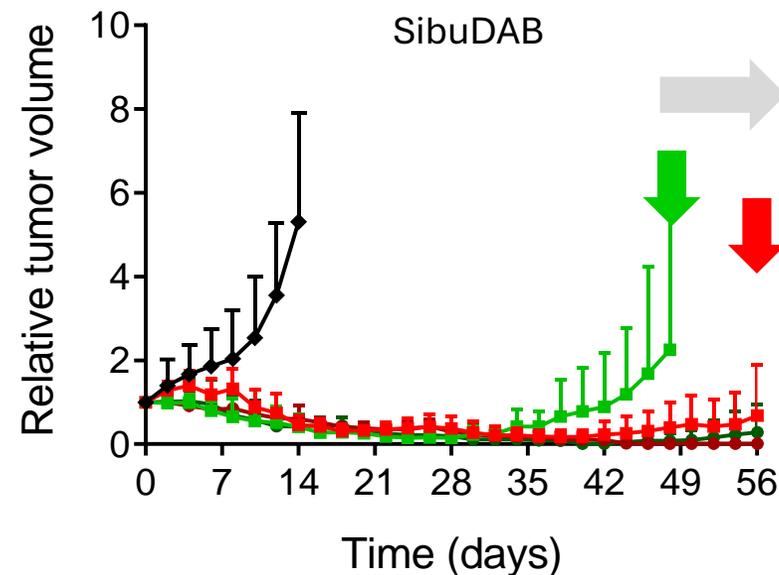
(PC-3 PIP tumor-bearing nude mice)



- Vehicle (control)
- $^{161}\text{Tb}$ -PSMA-I&T (5 MBq)
- $^{161}\text{Tb}$ -PSMA-I&T (10 MBq)
- $^{177}\text{Lu}$ -PSMA-I&T (5 MBq)
- $^{177}\text{Lu}$ -PSMA-I&T (10 MBq)

## Tumor growth curves

(PC-3 PIP tumor-bearing nude mice)



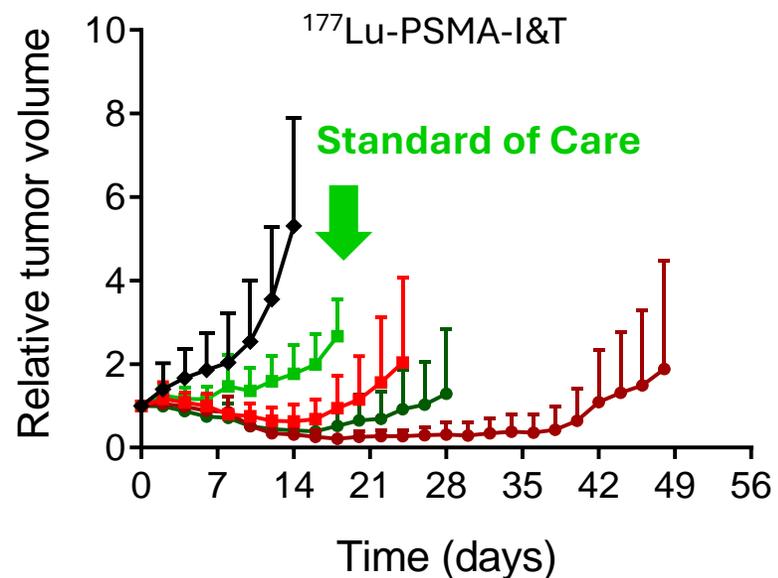
- Vehicle (control)
- $^{161}\text{Tb}$ -SibuDAB (5 MBq)
- $^{161}\text{Tb}$ -SibuDAB (10 MBq)
- $^{177}\text{Lu}$ -SibuDAB (5 MBq)
- $^{177}\text{Lu}$ -SibuDAB (10 MBq)

# $^{177}\text{Lu}$ -PSMA-I&T (Standard) versus $^{161}\text{Tb}$ -SibuDAB



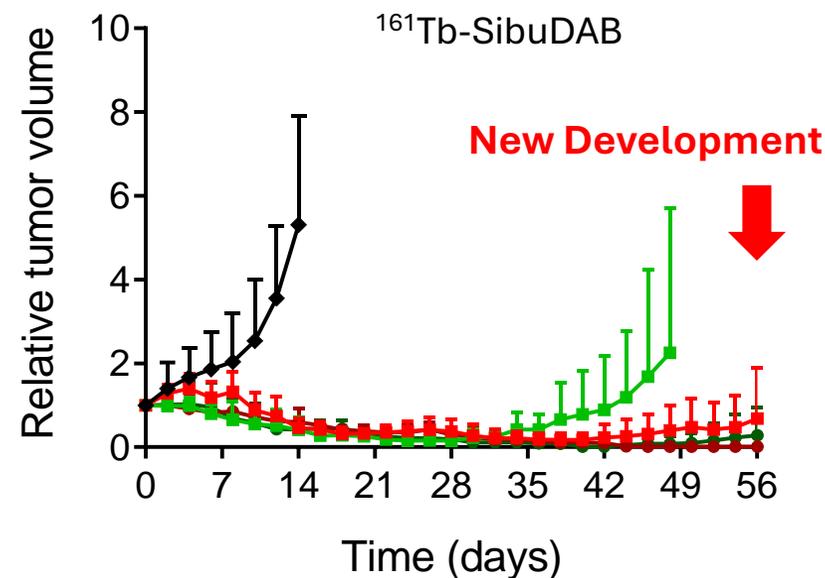
## Tumor growth curves

(PC-3 PIP tumor-bearing nude mice)



## Tumor growth curves

(PC-3 PIP tumor-bearing nude mice)

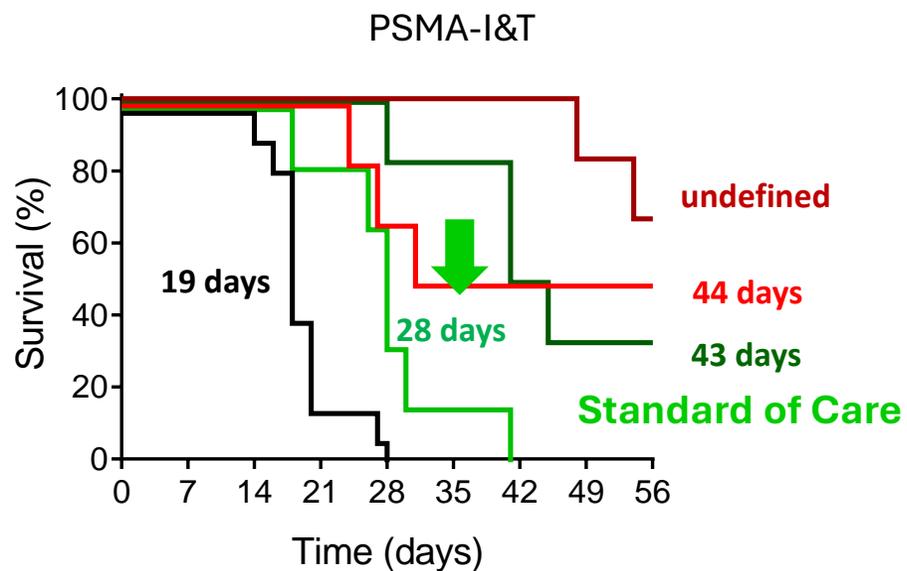


# $^{177}\text{Lu}$ -PSMA-I&T (Standard) versus $^{161}\text{Tb}$ -SibuDAB



## Survival curves

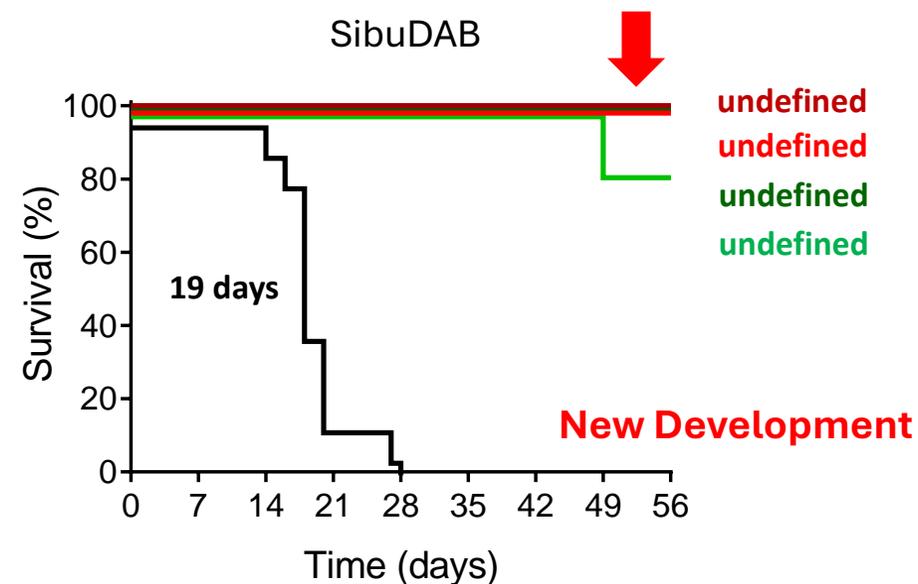
(PC-3 PIP tumor-bearing nude mice)



- Vehicle (control)
- $^{161}\text{Tb}$ -PSMA-I&T (5 MBq)
- $^{161}\text{Tb}$ -PSMA-I&T (10 MBq)
- $^{177}\text{Lu}$ -PSMA-I&T (5 MBq)
- $^{177}\text{Lu}$ -PSMA-I&T (10 MBq)

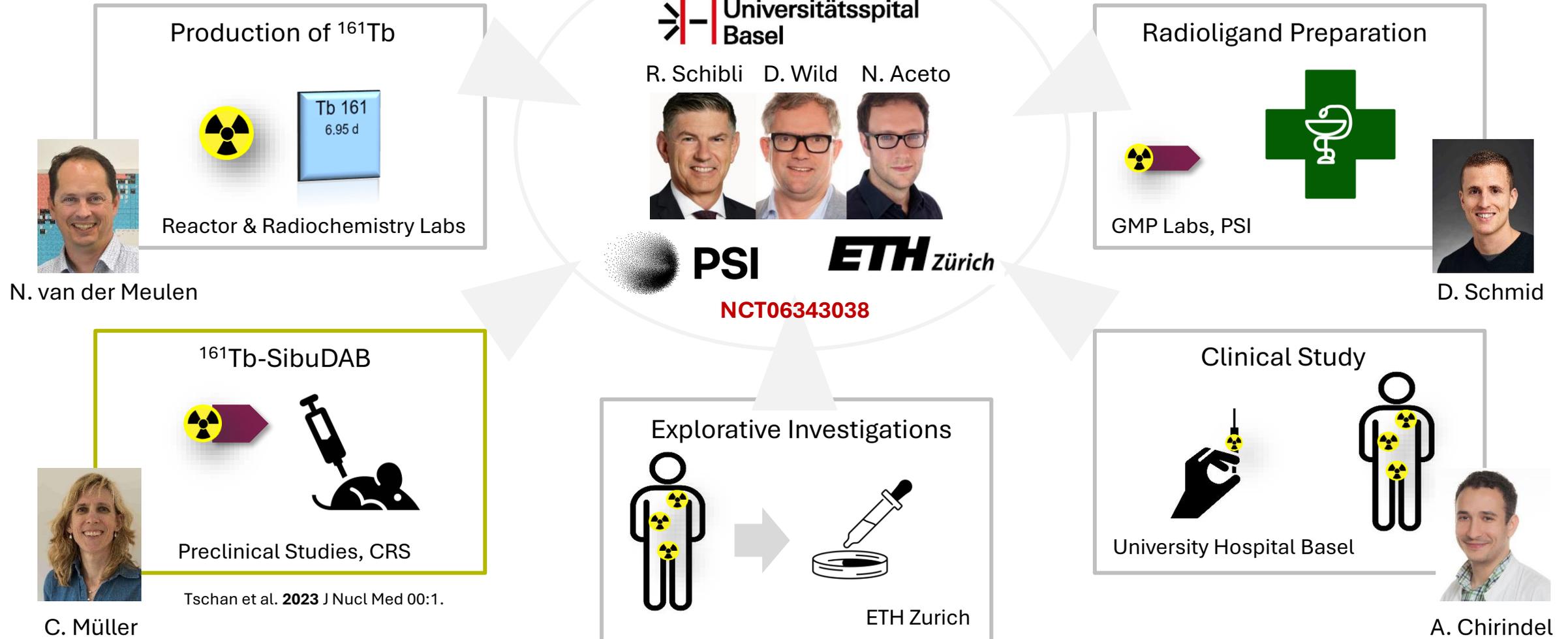
## Survival curves

(PC-3 PIP tumor-bearing nude mice)



- Vehicle (control)
- $^{161}\text{Tb}$ -SibuDAB (5 MBq)
- $^{161}\text{Tb}$ -SibuDAB (10 MBq)
- $^{177}\text{Lu}$ -SibuDAB (5 MBq)
- $^{177}\text{Lu}$ -SibuDAB (10 MBq)

# «PROGNOSTICS» Clinical Translation of $^{161}\text{Tb}$ -SibuDAB



# Acknowledgment



«Nuclide Chemistry» Group



«Radionuclide Development» Group



krebsforschung schweiz  
recherche suisse contre le cancer  
ricerca svizzera contro il cancro  
swiss cancer research



# Acknowledgment



## **Center for Radiopharm. Sciences (PSI)**

Everybody who contributed



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## **Laboratory of Radiochemistry (PSI)**

Everybody who contributed



## **Zentralklinik Bad Berka, Germany**

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## **Basel University Hospital, Switzerland**

Prof. D. Wild & Team



## **Necsa, Pelindaba, South Africa**

Dr. J. R. Zeevaart & Team



## **University of Gothenburg, Sweden**

Prof. P. Bernhardt & Team

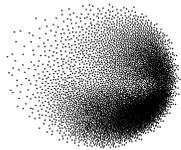


<https://www.psi.ch/en/zrw/nuclide-chemistry>

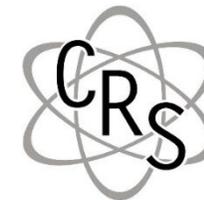


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recherche suisse contre le cancer  
ricerca svizzera contro il cancro  
**swiss cancer research**





# PSI Thank you for your Attention!



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Cristina Müller



Roger Schibli

Nick van der Meulen

