

Medical Isotope Production



Ulli Köster

ILL & UGA, Grenoble, France

Theranostics

=

Personalized
medicine

=

Precision
medicine

Cure

Companion
diagnostic

Therapy

Symptomatic
treatment

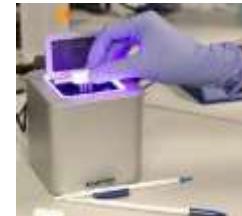
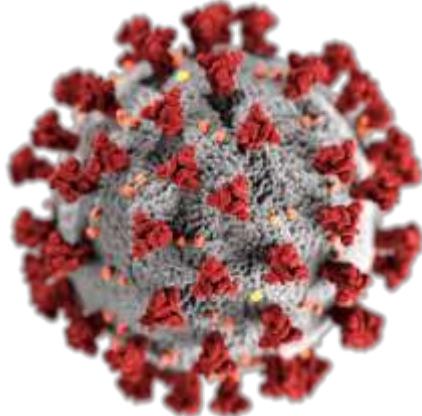
Diagnostic

Prevention

“Medical”

MD (Miracle Drug)

MD's efficacy
predictor



Theranostics

=

Personalized
medicine

=

Precision
medicine

Cure

10k

Miracle Drug *



= 0.85 G\$

Companion
diagnostic

25k

MD's efficacy
predictor



= 0.08 G\$

Therapy

25k



= 0.08 G\$

Symptomatic
treatment



Diagnostic

20M



= 1 G\$

Prevention

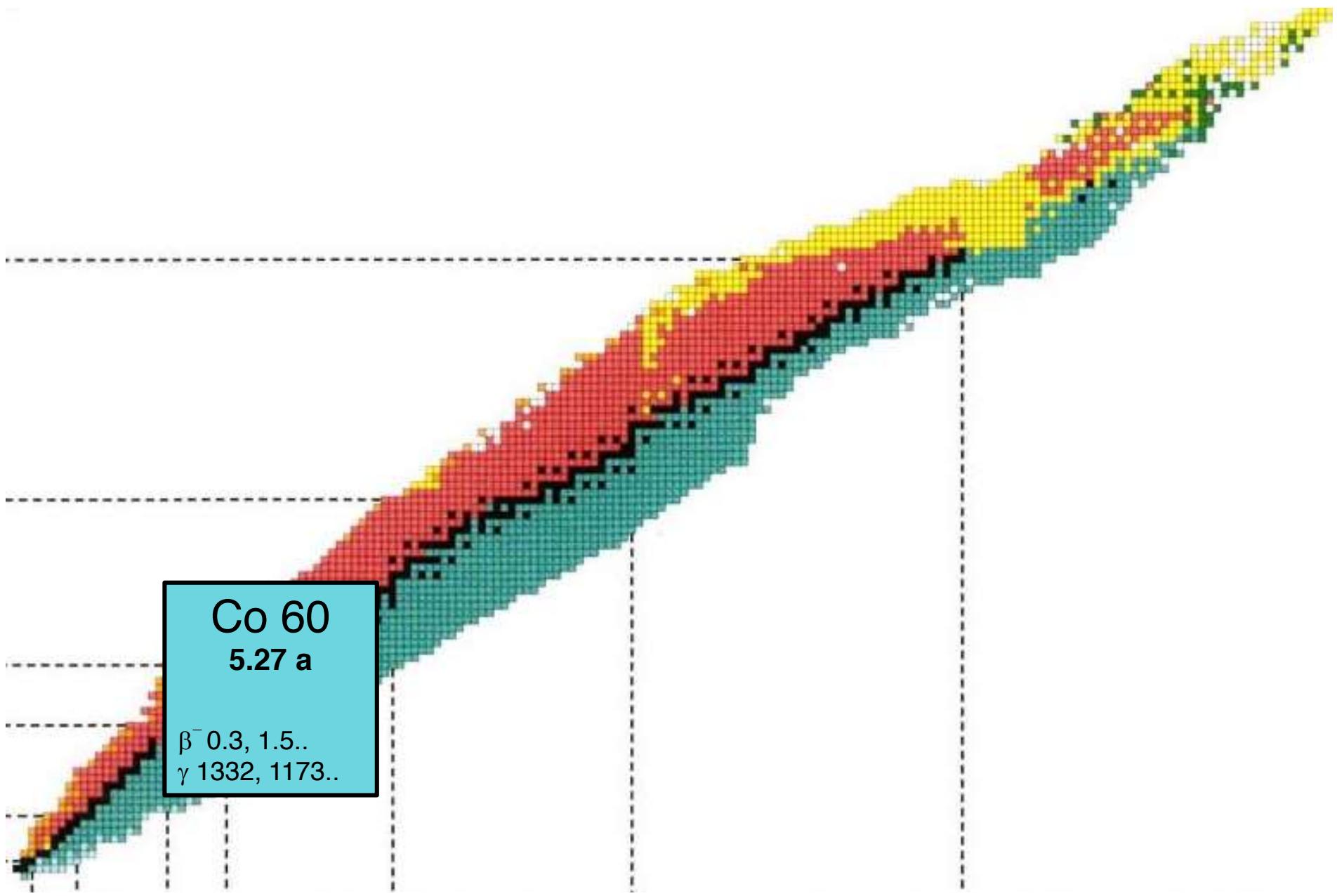
38M * 500



= 2 G\$



Radionuclides for Medicine



Radiation sterilization of medical material

- Single use consumables (surgical threads, gloves, etc.)
- Human tissues
- Implantable devices
- Pharmaceuticals

RADIATION MODALITIES



GAMMA
Exposes product to Cobalt 60 radiation

38%



ELECTRON BEAM
Exposes product to high-energy electrons

7%



X-RAY
Uses ionizing energy from electron beams

GAS MODALITIES



ETHYLENE OXIDE
Exposes product to gaseous sterilant

50%



VHP
Low temp gas process under deep vacuum

OTHER



MOIST HEAT (STEAM)
High temp, high pressure with vacuum or gravity conditioning

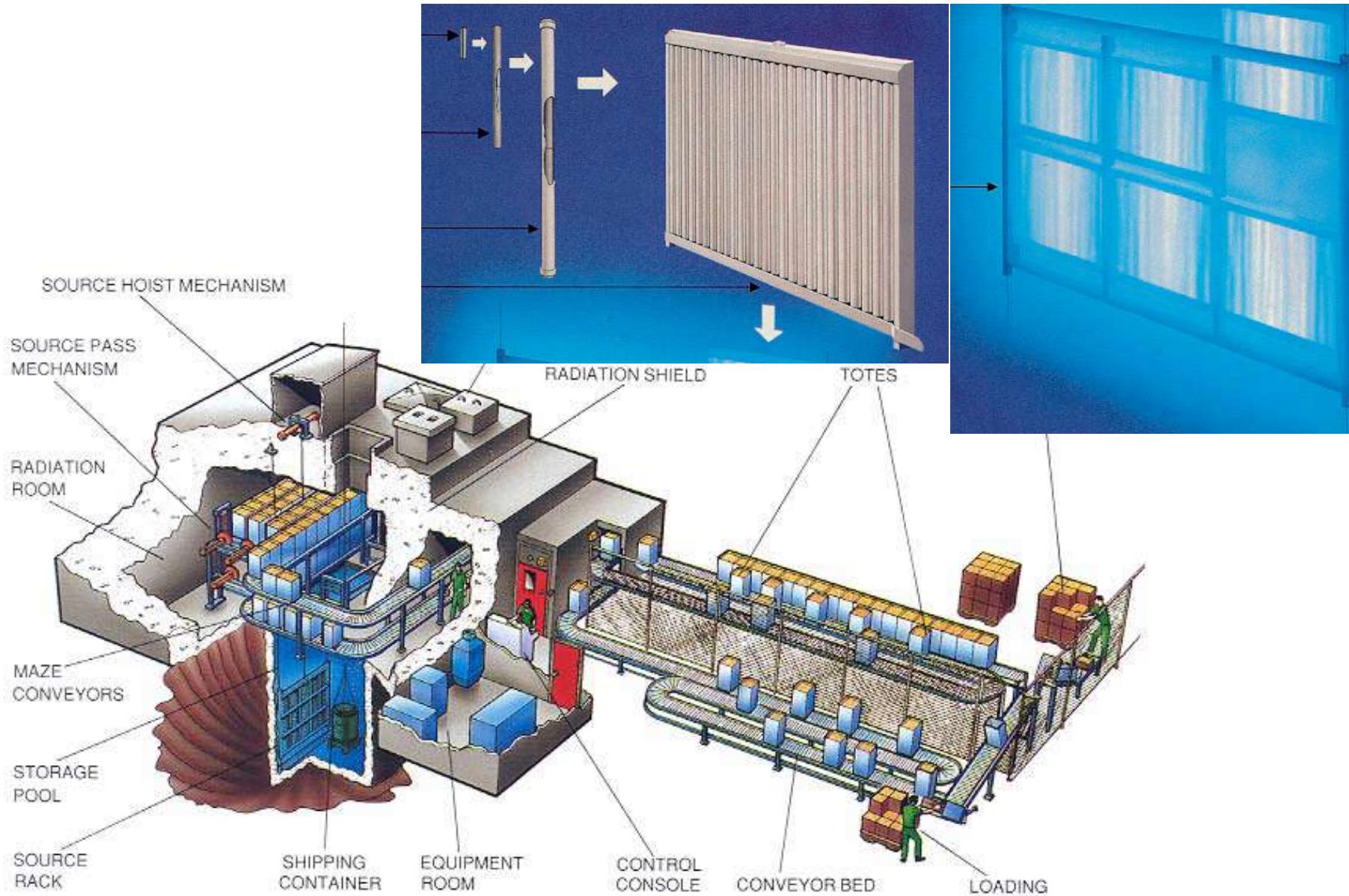


LABORATORIES
Provides microbiological and analytical testing

Market for gamma ray sterilization: $\approx 1 \text{ G\$}/\text{year}$

Total ^{60}Co activity in use: $\approx 10^{19} \text{ Bq} \approx 0.3 \text{ GCi}$

Radiation Sterilization of Medical Devices



1 MCi = 37 PBq ^{60}Co sterilizes 650 kg/hour at 25 kGy

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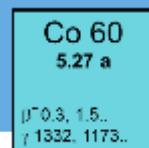
Companion
diagnostic

Therapy

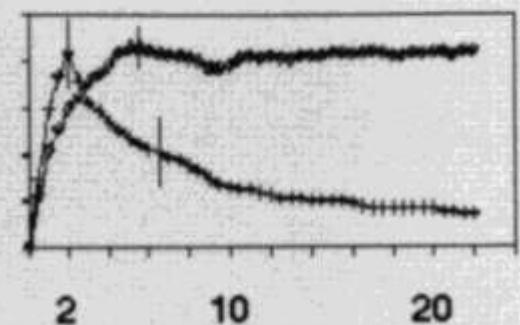
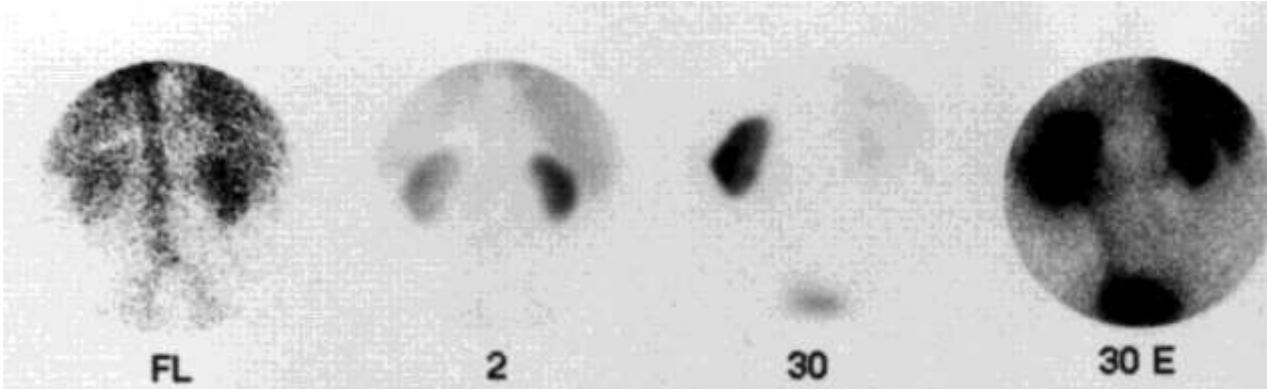
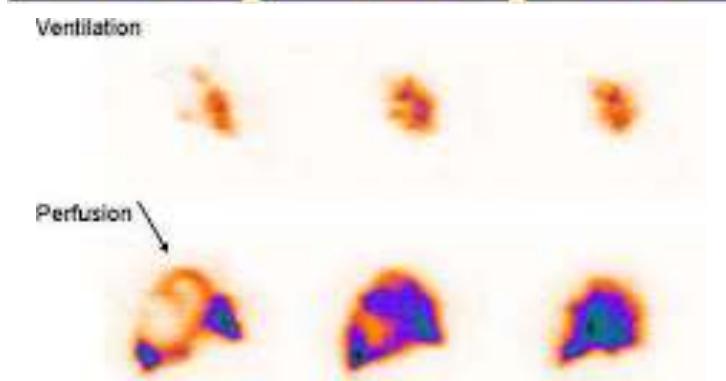
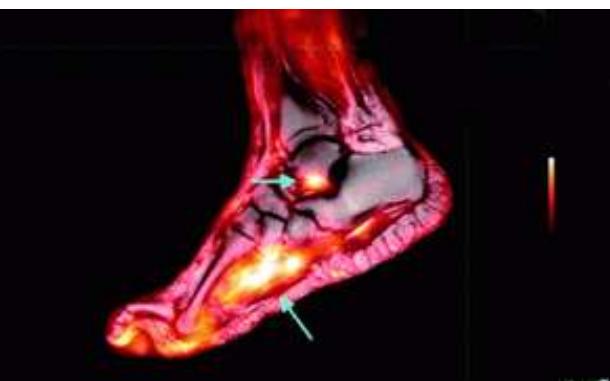
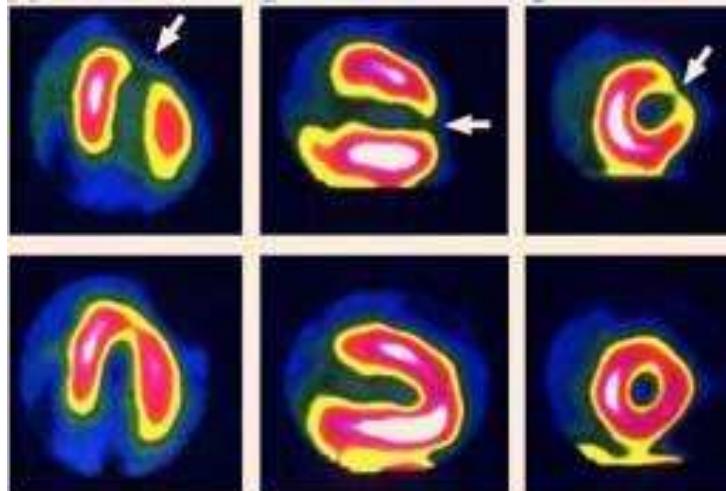
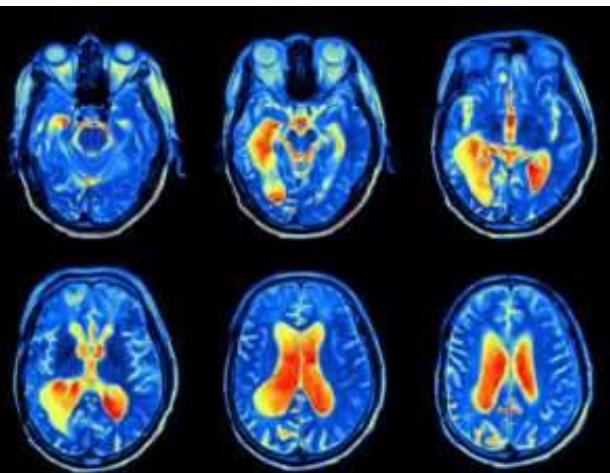
Symptomatic
treatment

Diagnostic

Prevention



Molecular imaging



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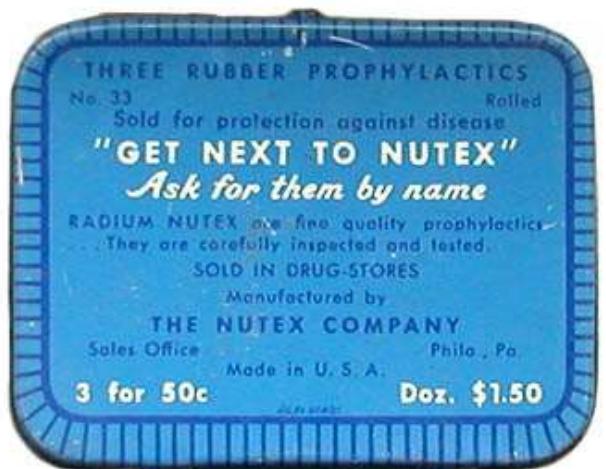
Tc 99
6.0 h
β^- 141...
γ 0.3...
σ 22.8

F 18
110 m
β^- 0.533
$\mu\alpha\gamma$

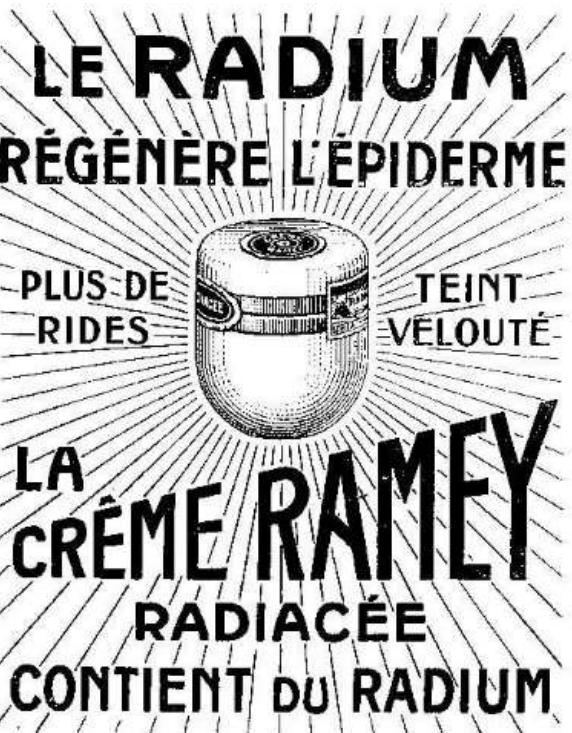
Co 60
5.27 a
β^- 0.3, 1.5...
γ 1332, 1173...



Discontinued radium applications



Évitez et combattez la TUBERCULOSE
par la **TUBÉRADINE**
(Antiseptique pulmonaire radioactif)



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Diagnostic



Tc 99	
6.0 h	
β^- 141...	β^- 0.3...
e	γ (90)
β	σ 22.8

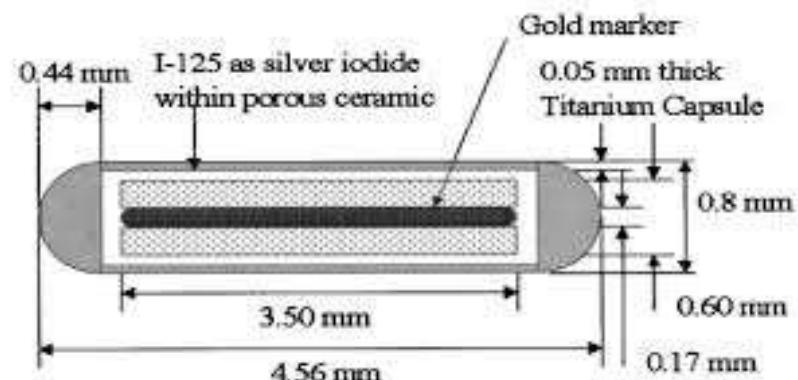
Prevention

Co 60
5.27 a
β^- 0.3, 1.5.. γ 1332, 1173..

Brachytherapy

High Dose Rate (HDR) brachytherapy
short-term insertion of ^{60}Co , ^{137}Cs ,
 ^{169}Yb or ^{192}Ir sources

Low Dose Rate (LDR) brachytherapy
long-term insertion of ^{32}P , ^{103}Pd , ^{125}I ,
 ^{131}Cs , etc. sources (“seeds”)



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Co 60
5.27 a
β^- 0.3, 1.5..
γ 1332, 1173..

Symptomatic
treatment

Tc 99
6.0 h
$2.1 \cdot 10^5$ a
I_β^- 141...
e^- 0.3...
γ (90)
σ 22.8

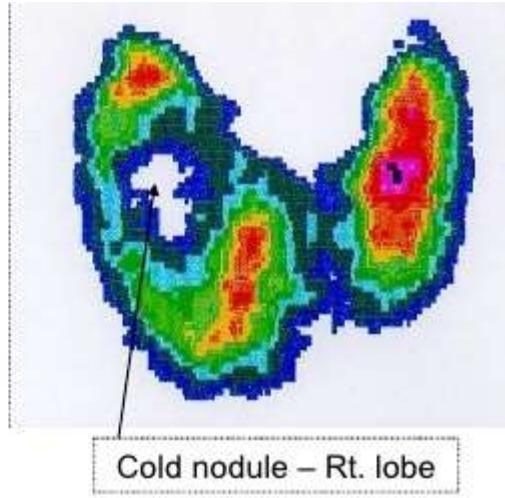
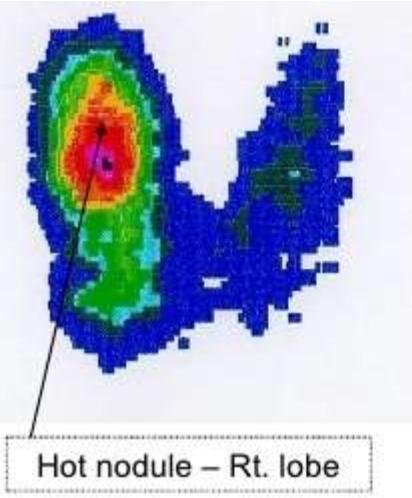
Diagnostic

F 18
110 m
β^- 0.633
μo γ

Prevention

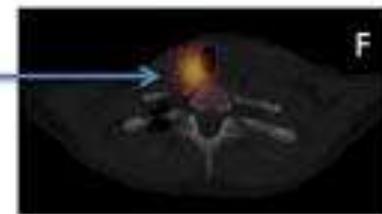
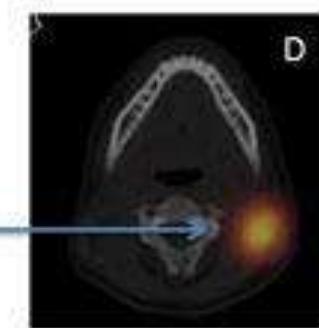
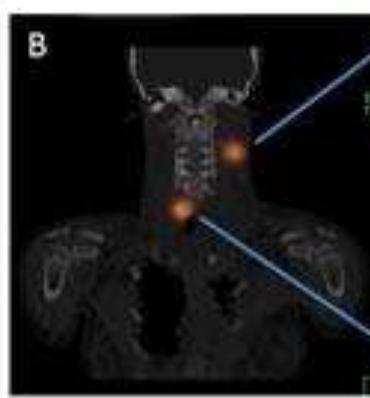
Co 60
5.27 a
β^- 0.3, 1.5..
γ 1332, 1173..

Thyroid scintigraphy and therapy



I 123
13.2 h
 ε
no β^+
 γ 159

I 131
8.0 d
 β^- 1.0, 1.8,...
 γ 364, 637...



^{123}I , ^{131}I or $^{99\text{m}}\text{TcO}_4$ for scintigraphy
 ^{131}I for therapy

(papillary) thyroid cancer has the **highest survival** of all malignant cancers !

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Co 60
5.27 a
 β^- 0.3, 1.5..
 γ 1332, 1173..

I 131
8.0 d
 β^- 1.0, 1.8,...
 γ 364, 637...

Symptomatic
treatment

Tc 99
6.0 h | $2.1 \cdot 10^5$ a
 I_f 141... | β^- 0.3...
e | γ (90)
 β | σ 22.8

Diagnostic

F 18
110 m
 β^- 0.633
 $\mu\alpha\gamma$

I 123
13.2 h
 μ
no β^+
 γ 159

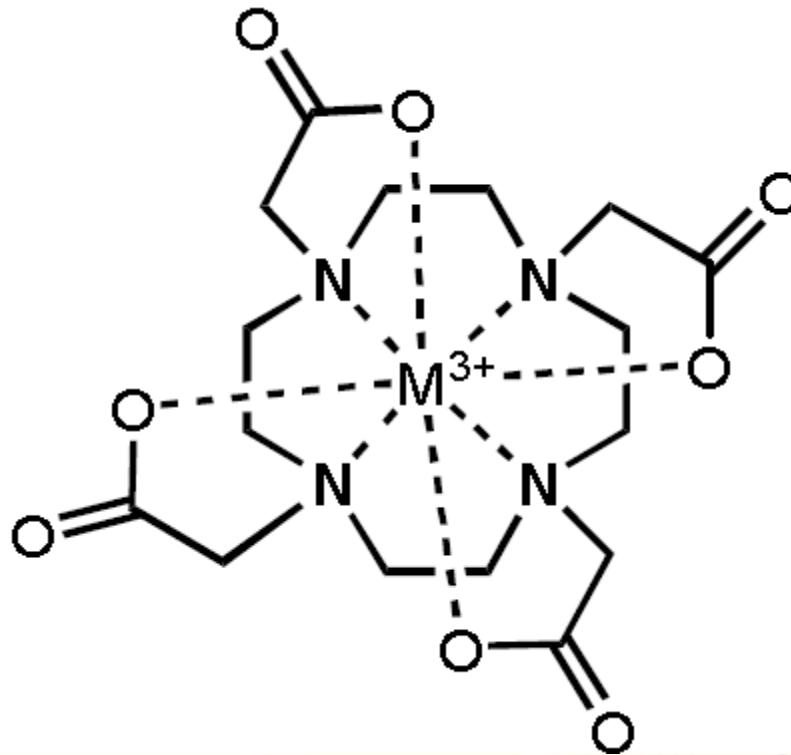
Prevention

Co 60
5.27 a
 β^- 0.3, 1.5..
 γ 1332, 1173..

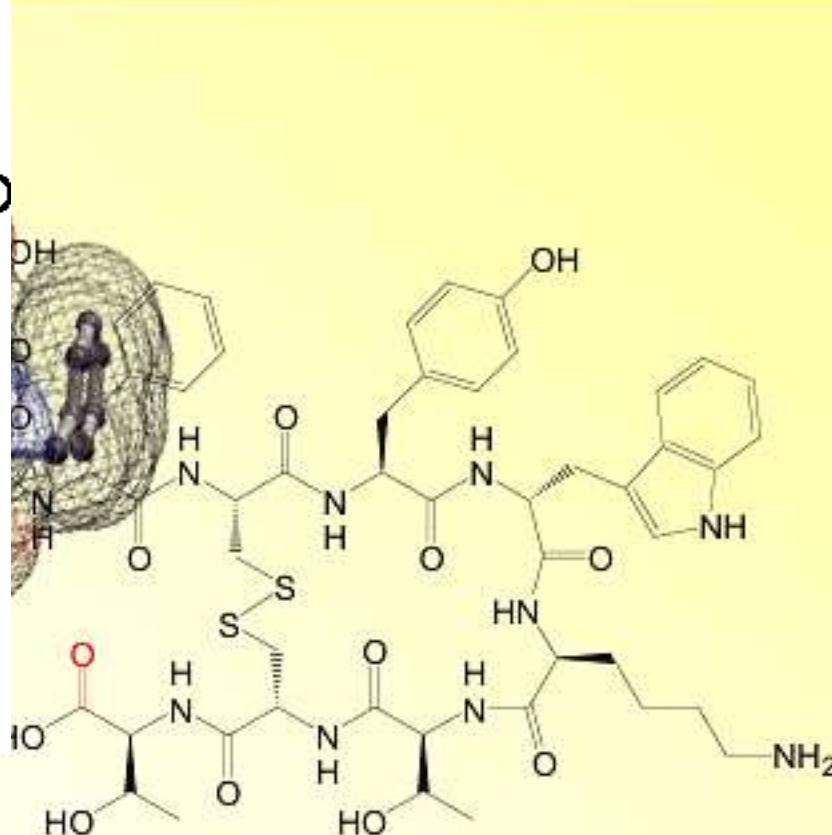
How can one treat such patients?



Structural Formula of DOTA-TOC/TATE



DOTA-TATE



1,4,7,10-tetraazacyclododecane tetraacetate

^{111}In

^{90}Y

^{67}Ga

^{177}Lu

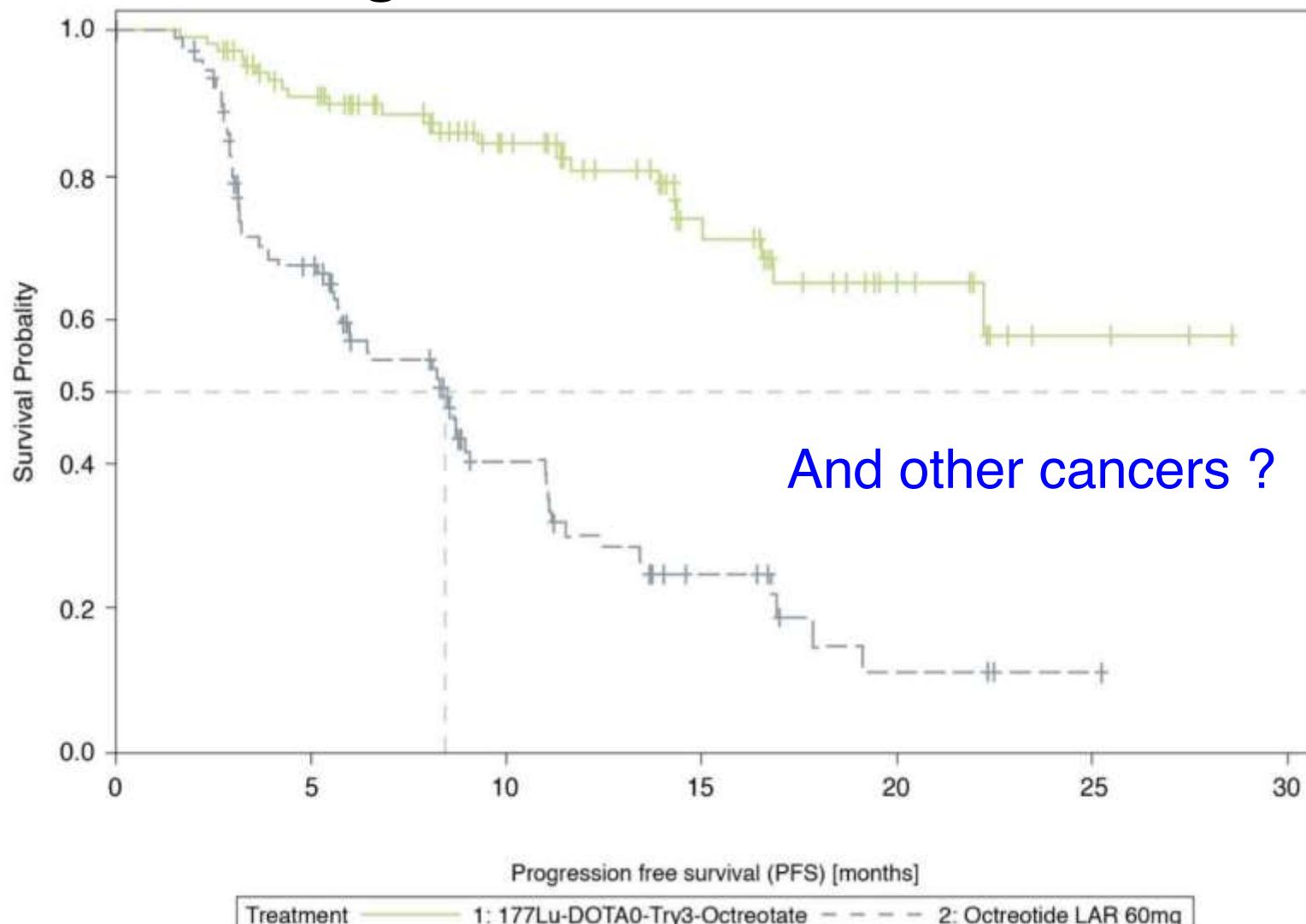
^{68}Ga

^{213}Bi

$\text{IC}_{50} (\text{Y}^{\text{III}}) = 1.6 \pm 0.4 \text{ nM}$

Helmut Maecke, EANM-2007.

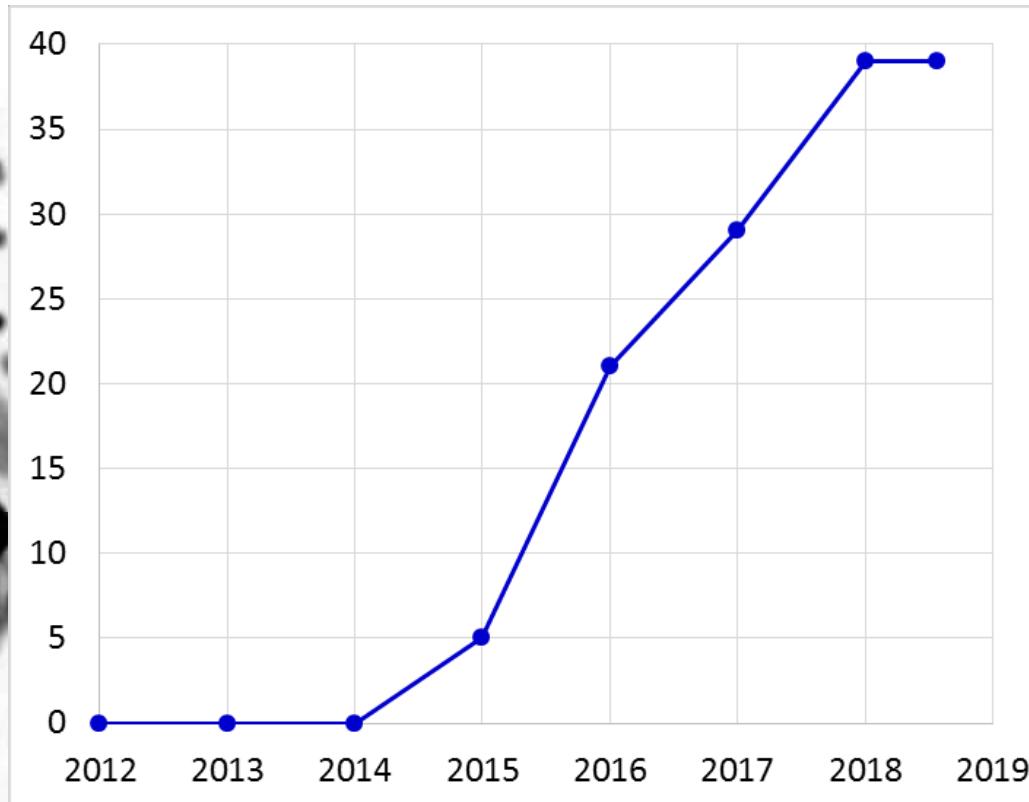
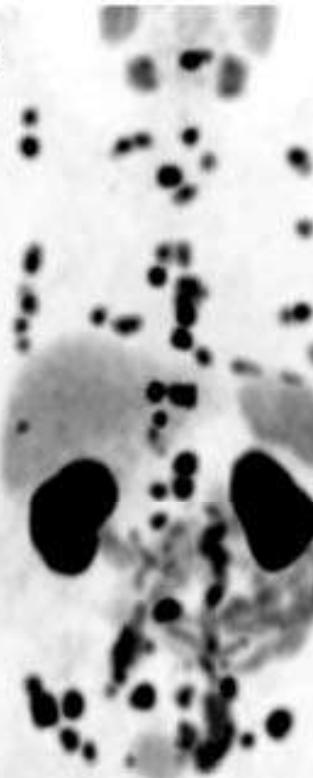
^{177}Lu -Peptide Receptor Radionuclide Therapy of midgut neuroendocrine tumors



J. Strosberg et al., N Engl J Med 2017;376:125.

^{177}Lu -radioligand therapy of advanced prostate cancer

A



C. Kratochwil et al., *Eur J Nucl Med Mol Imaging* 2015;42:987.

R.P. Baum et al., *J Nucl Med* 2016;57:1006.

C. Kratochwil et al., *J Nucl Med* 2016;57:1170.

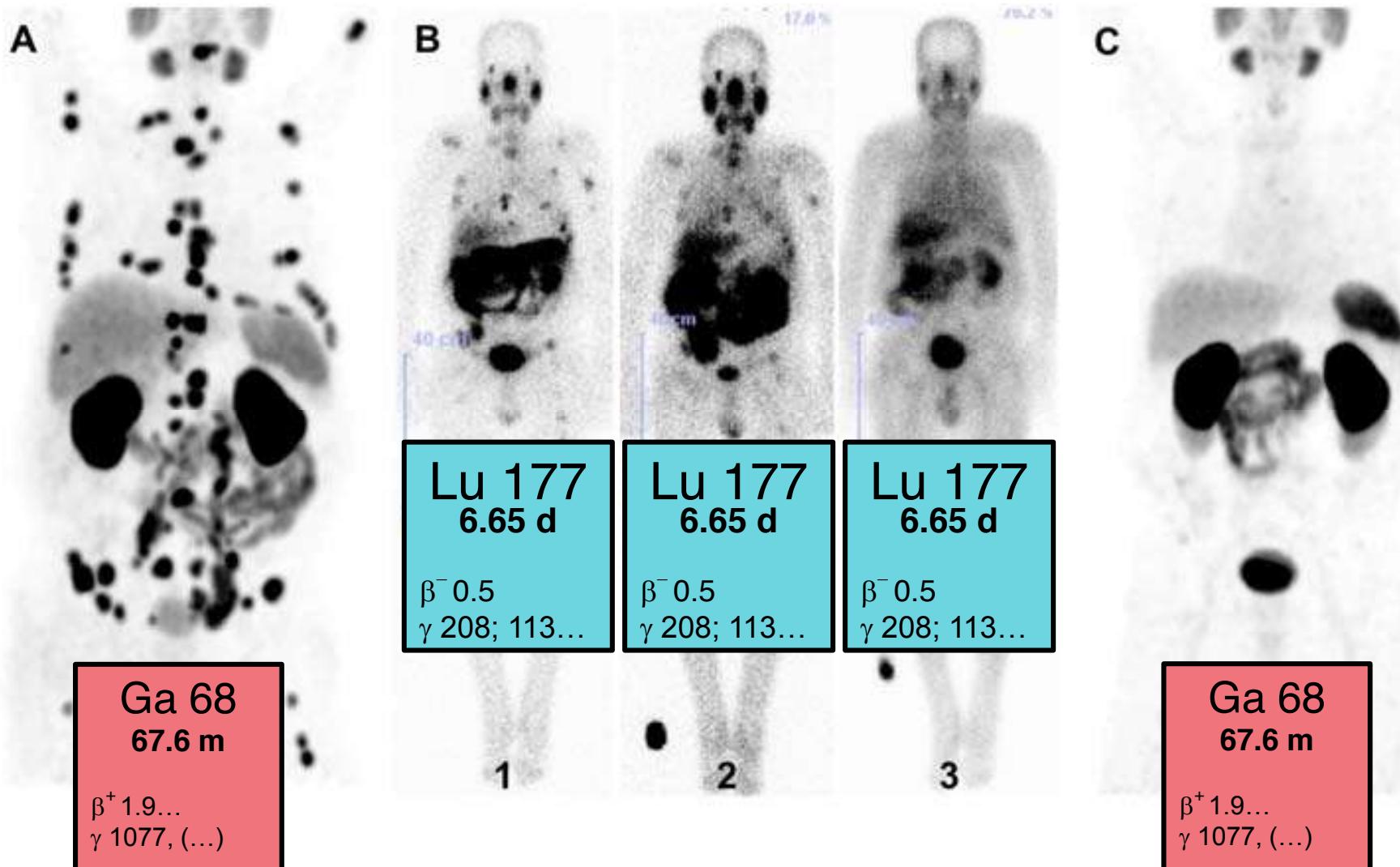
K. Rahbar et al., *J Nucl Med* 2017;58:85.

M.S. Hofman et al., *Lancet Oncol* 2018;19:825.

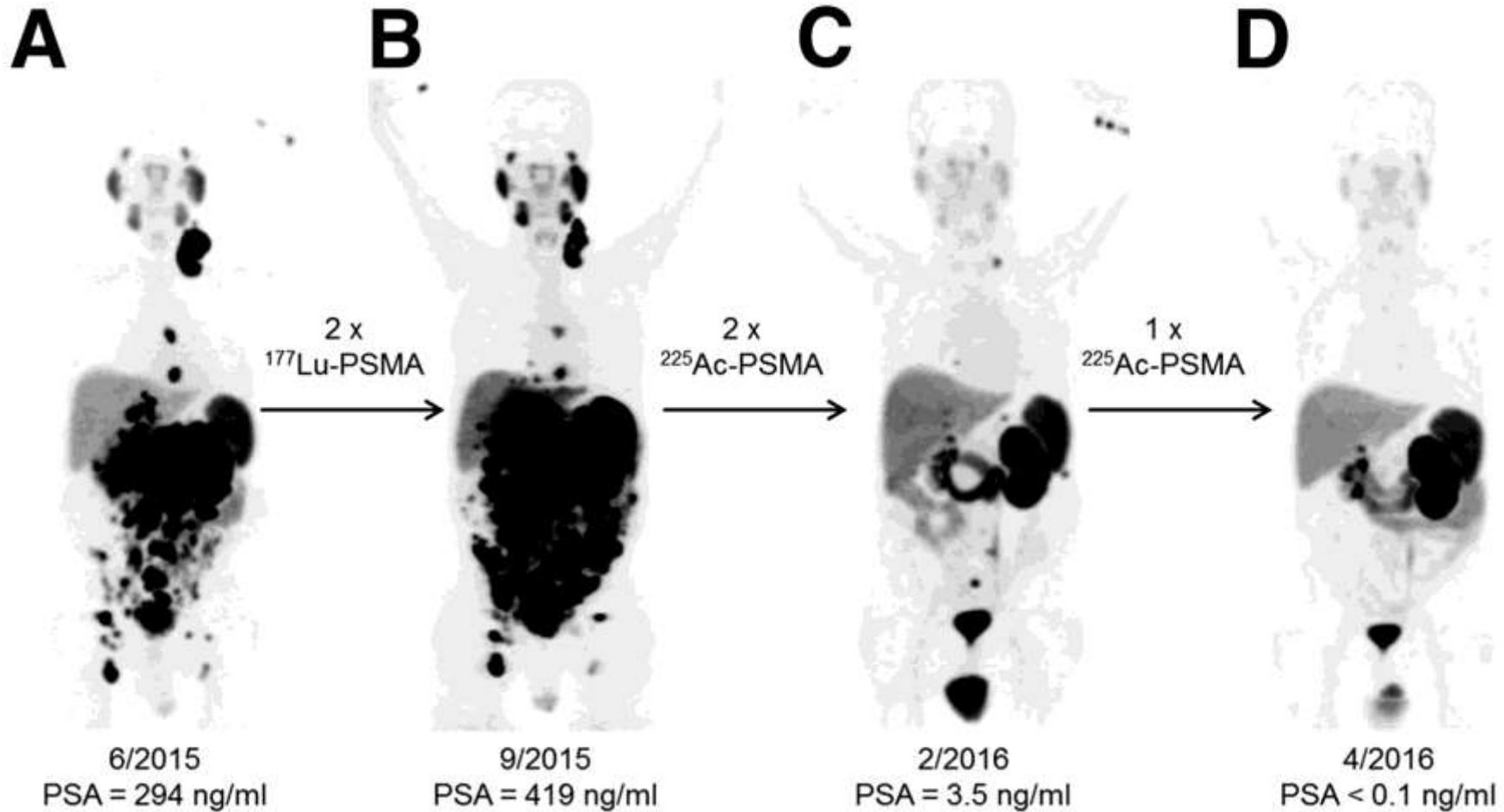
M.M. Heck et al., *Eur Urol* 2019;75:920.

T.W. Barber et al., *J Nucl Med* 2019; 60:955.

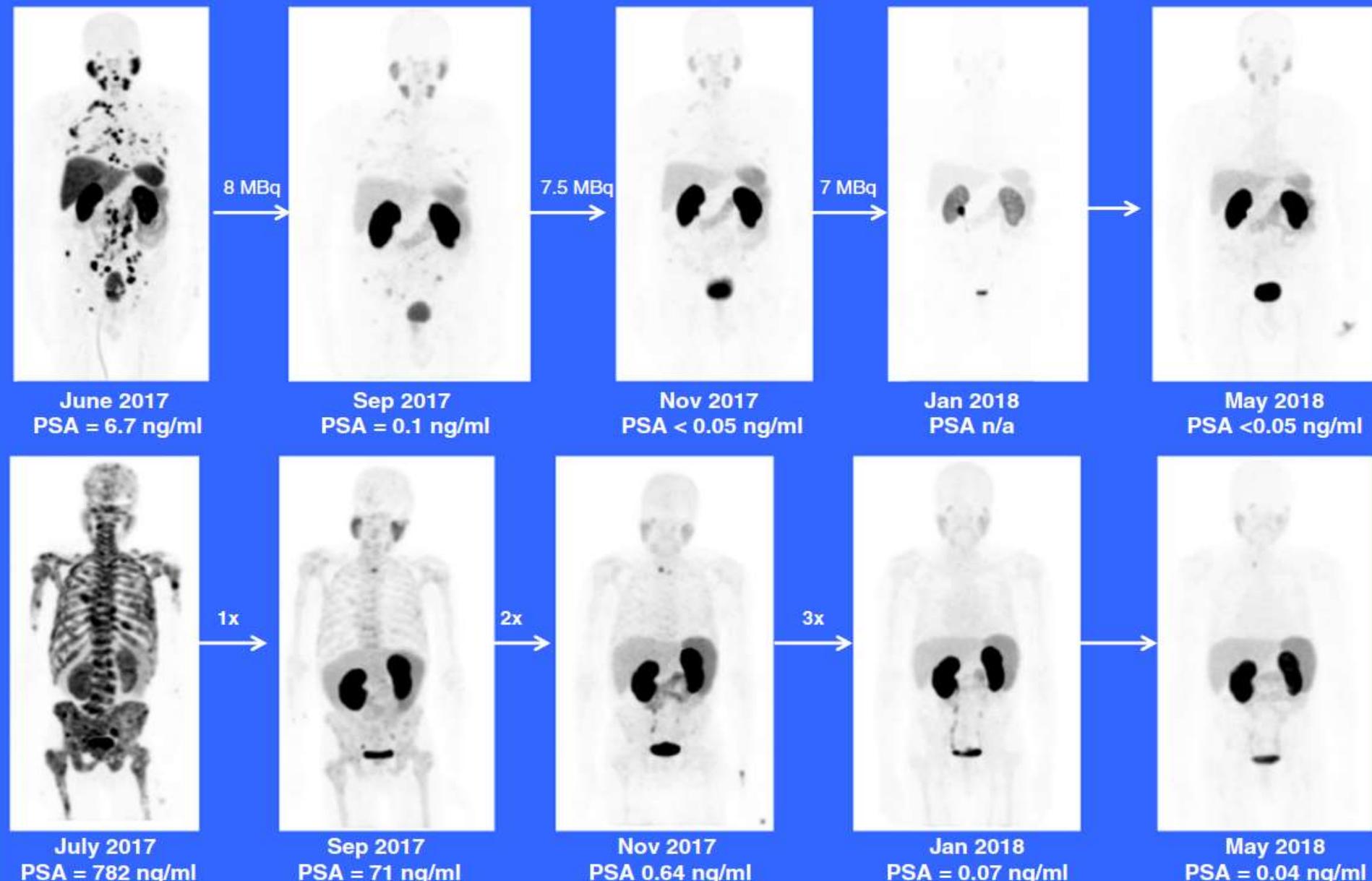
^{177}Lu -radioligand therapy of advanced prostate cancer



Targeted therapy with ^{225}Ac



Targeted therapy with ^{225}Ac



Theranostics

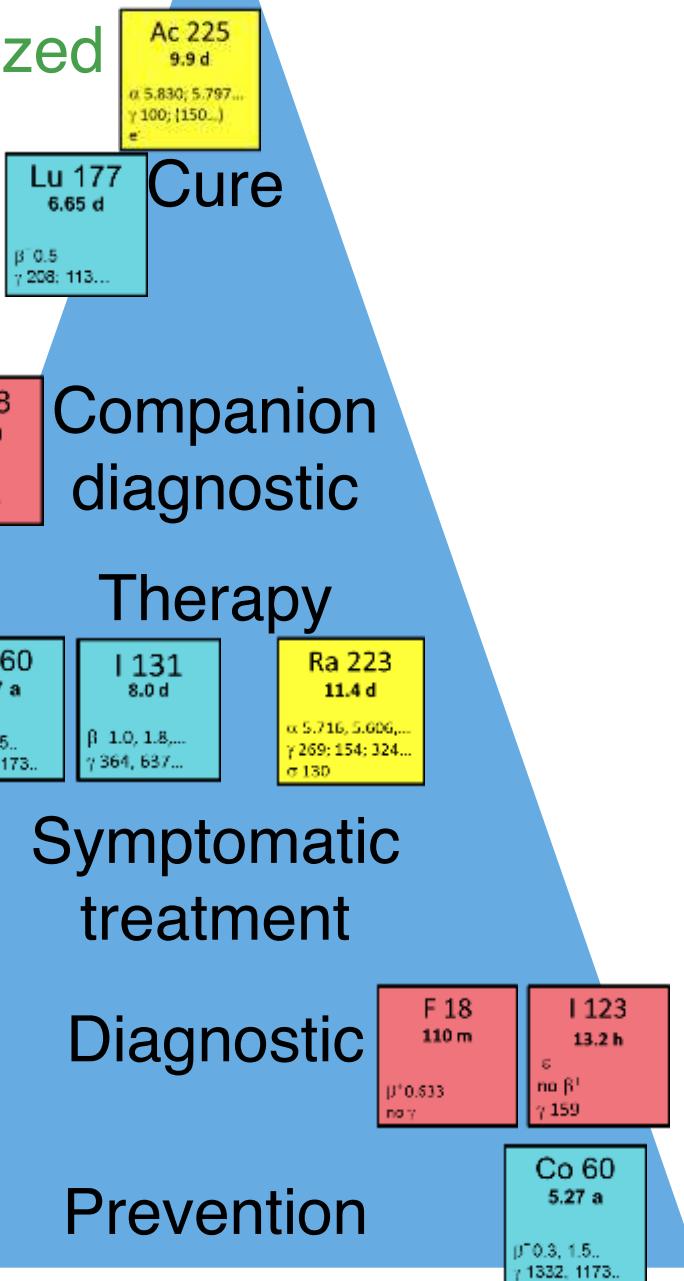
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Personalized
medicine

=

Precision
medicine

Radionuclides



Isotopes for targeted alpha therapy

12 s	Ac 213 0.80 s	Ac 214 8.2 s	Ac 215 0.17 s	Ac 216 0.44 ms	Ac 217 0.76 μ s	Ac 218 1.1 μ s	Ac 219 11.8 μ s	Ac 220 26 ms	Ac 221 52 ms	Ac 222 63 s	Ac 223 2.10 m	Ac 224 2.9 h	Ac 225 10.0 d	Ac 226 29 h		
	α ; 7.36	α ; 7.215; 7.061; β^- ; 5.39; 244...; γ (395...)	α ; 7.600; 7.211...; β^- ; 5.83; 854...; γ (395...)	α ; 9.029; 9.105...; β^- ; 4.95; 435...; γ 83; 854; 771...;	α ; 9.60...; β^- ; 4.85; 382...; γ 10.54...;	α ; 9.85	α ; 9.205 g	α ; 9.884	α ; 7.86; 7.81; β^- ; 7.68...; γ 134...;	α ; 7.65; 7.44; β^- ; 7.36...;	α ; 6.647; 6.662...; β^- ; 6.58...; γ (99; 191; 84...);	α ; 6.142; β^- ; 6.06...; γ 216; 132...;	α ; 5.830; 5.793...; β^- ; 5.732...; C 14...; γ 100; 150...; 198; 93...; e-	β^- ; 0.9... 1.1...; γ 0.534...; 1230; 158...; 264; 186...;		
11 s	Ra 212 13.0 s	Ra 213 2.1 ms	Ra 214 2.46 s	Ra 215 1.67 ms	Ra 216 2.0 ms	Ra 217 0.18 μ s	Ra 218 1.6 μ s	Ra 219 25.6 μ s	Ra 220 10 ms	Ra 220 23 ms	Ra 221 28 s	Ra 222 38 s	Ra 222 11.4 d	Ra 224 3.66 d	Ra 225 14.8 d	
	α ; 7.88...; β^- ; 7...; γ 5.899...; γ (635...)	α ; 5.56...; β^- ; 5.524...; γ 6.371...; 6.321...; 6.321...; γ 8.498...; γ 11.10...; γ 21...; 10...;	α ; 7.137; 5.505...; β^- ; 6.0...; γ (642...);	α ; 8.700; 7.879...; β^- ; 8.34...; γ 834; 540...;	α ; 8.68...; β^- ; 8.47...; γ 8.348...;	α ; 8.99	α ; 8.99 g	α ; 7.679; 7.389...; β^- ; 7.310; 214...; γ 5.95...;	α ; 7.48...; β^- ; 7.465...;	α ; 6.613; 6.761...; β^- ; 6.58...; γ 149; 93; 174...; C 14...;	α ; 6.559; 6.237...; β^- ; 6.58...; γ (329; 473...); C 14...;	α ; 6.142...; β^- ; 6.06...; γ 216; 132...;	α ; 5.830; 5.793...; β^- ; 5.732...; C 14...; γ 100; 150...; 198; 93...; e-	β^- ; 0.9... 1.1...; γ 0.534...; 1230; 158...; 264; 186...;		
10 m	Fr 211 3.10 m	Fr 212 20.0 m	Fr 213 34.6 s	Fr 214 3.35 ms	Fr 215 5.0 ms	Fr 216 0.09 μ s	Fr 217 0.70 μ s	Fr 218 16 μ s	Fr 218 22 ms	Fr 218 1.0 ms	Fr 219 21 ms	Fr 220 27.4 s	Fr 221 4.9 m	Fr 222 14.2 m	Fr 223 21.8 m	Fr 224 3.3 m
	α ; 0.535...; β^- ; 0...; γ 540; 918...; 281...;	α ; 6.162; 6.304...; β^- ; 6.406; 6.310...; γ 1274; 227; 1180...;	α ; 6.775	α ; 8.47...; β^- ; 8.347...; γ 8.336...;	α ; 9.36	α ; 9.01	α ; 8.315	α ; 7.679...; β^- ; 7.580...; γ 7.580...;	α ; 7.48...; β^- ; 7.312...; γ (352; 517...); C 14...;	α ; 6.68; 6.63...; β^- ; 6.58...;	α ; 6.341; 6.126...; β^- ; 6.218; (101; 411...); C 14...;	α ; 6.341; 6.126...; β^- ; 6.218; (101; 411...); C 14...;	α ; 6.288...; β^- ; (550); γ 180; 150...;	α ; 5.34...; β^- ; 5.21; 211...; 242...;	α ; 5.285 d	β^- ; 2.8... 2.8...; γ 216; 152...; 337; 1341...;
09 m	Rn 210 2.4 h	Rn 211 14.8 h	Rn 212 24 m	Rn 213 19.5 ms	Rn 214 65 ms	Rn 214 0.76 ms	Rn 215 2.3 μ s	Rn 216 45 μ s	Rn 217 0.54 ms	Rn 218 35 ms	Rn 218 3.9 s	Rn 220 55.5 s	Rn 221 25 m	Rn 222 3.825 d	Rn 223 23.2 m	
	α ; 0.40...; β^- ; 0...; γ 458; (571...); 648; 73...;	α ; 5.783; 5.851...; β^- ; 6.74; 1083...; γ 779...; 0...;	α ; 6.264...; β^- ; 5.40...;	α ; 8.088; 7.252...; β^- ; 8.54...;	α ; 8.68...; β^- ; 8.442...; γ 8.30...;	α ; 9.36	α ; 8.67	α ; 8.05	α ; 7.740...;	α ; 7.133...; β^- ; (809); γ 402...;	α ; 6.553...; β^- ; 6.553...;	α ; 6.288...; β^- ; (550); γ 180; 150...;	α ; 5.48948...; β^- ; 5.407; 5.788...; γ 5778...;	α ; 5.10...; β^- ; 0.74	β^- ; 0.8... 1.1...; γ 593; 417...; 636; 655...;	
08 h	At 209 5.4 h	At 210 8.3 h	At 211 7.22 h	At 212 119 ms	At 212 214 ms	At 213 0.11 μ s	At 214 0.76 μ s	At 214 0.09 μ s	At 215 0.1 ms	At 216 0.3 ms	At 217 32.3 ms	At 218 ~ 2 s	At 219 0.9 m	At 220 3.71 m	At 221 2.3 m	At 222 54 s
	α ; 5.647...; β^- ; 5.524...; γ 5422; 5.861...; 548; 782...; 780...;	α ; 5.867...; β^- ; 5.718...; γ (687...);	α ; 6.211...; β^- ; 6.111...; γ 6.081...;	α ; 7.84...; β^- ; 7.68...; γ 7.63...;	α ; 9.08	α ; 8.782...; β^- ; 8.677...; γ (779);	α ; 8.819...; β^- ; 8.719...; γ (405);	α ; 8.026...; β^- ; 7.908...; γ 7.816...;	α ; 7.069...; β^- ; 7.004...; γ (259; 334...);	α ; 6.694; 6.683...; β^- ; 6.683...;	α ; 6.27	β^- ?	β^- ; 5.493...; γ 341; 290...; 422...;	β^- ?	β^- ?	
07 d	Po 208 2.898 a	Po 209 102 d	Po 210 138.38 d	Po 211 0.016 s	Po 212 45.1 s	Po 212 17.1 ms	Po 213 4.2 μ s	Po 214 164 μ s	Po 215 1.78 μ s	Po 216 1.78 s	Po 217 0.15 s	Po 218 1.53 s	Po 218 3.05 m	Po 219 >300 ns	Po 220 >300 ns	
	α ; 5.1152...; β^- ; 5.1152...; γ (292; 571...); 0...;	α ; 5.90458...; β^- ; 5.803...; γ < 0.0005...;	α ; 6.205...; β^- ; 6.104...; γ 6.04...;	α ; 7.85...; β^- ; 7.78...; γ 7.73...;	α ; 9.02...; β^- ; 8.92...; γ 8.87...;	α ; 8.376...; β^- ; 8.276...; γ (779);	α ; 7.889...; β^- ; 7.802...; γ (690); 298...;	α ; 7.802...; β^- ; 7.702...; γ (690); 298...;	α ; 7.069...; β^- ; 7.004...; γ (259; 334...);	α ; 6.7783...; β^- ; (805);	α ; 6.543	α ; 6.0024...; β^- ?	β^- ?	β^- ?		
06 d	Bi 207 31.55 a	Bi 208 3.68 \cdot 10 ⁻⁵ a	Bi 209 100	Bi 210 1.0 \cdot 10 ⁻⁸ s	Bi 210 5.013 d	Bi 211 2.17 s	Bi 212 25 m	Bi 213 45.59 m	Bi 214 19.9 m	Bi 215 38.9 s	Bi 216 7.7 m	Bi 216 38 m	Bi 217 98.5 s	Bi 218 33 s		
	α ; 5.16...; β^- ; 5.16...; γ 5770; 1064...; 1770...;	α ; 5.95; 261...;	α ; 6.011...; β^- ; 6.011...;	α ; 7.85...; β^- ; 7.75...; γ 7.70...;	α ; 9.02...; β^- ; 8.92...; γ 8.87...;	α ; 8.376...; β^- ; 8.276...; γ (779);	α ; 7.889...; β^- ; 7.802...; γ (690); 298...;	α ; 7.802...; β^- ; 7.702...; γ (690); 298...;	α ; 7.069...; β^- ; 7.004...; γ (259; 334...);	α ; 6.7783...; β^- ; (805);	α ; 6.543	α ; 6.0024...; β^- ?	β^- ?			
05 a	Pb 206 24.1	Pb 207 22.1	Pb 208 52.4	Pb 209 3.253 h	Pb 210 22.3 a	Pb 211 36.1 m	Pb 212 10.64 h	Pb 213 10.2 m	Pb 214 26.8 m							
	α ; 0.027	α ; 0.61	α ; 0.00023	β^- ; 0.6	β^- ; no γ	α ; 0.02... 0.06... β^- ; 0.71... 0.79... γ 3.72... δ 0.5	α ; 0.74...; β^- ; 0.84...; γ 4.05...; δ 0.47...	α ; 0.3... 0.6... β^- ; 0.405... 0.602... γ 4.27...;	α ; 0.7...; β^- ; 0.7...; γ 4.05...; δ 0.47...	α ; 0.7...;	α ; 0.7... 1.0... β^- ; 0.759... 295... γ 342...					
04 a	Tl 205 70.48	Tl 206 3.7 m	Tl 207 1.33 s	Tl 208 3.053 m	Tl 209 2.16 m	Tl 210 1.30 m	Tl 211 >300 ns	Tl 212 >300 ns								
	α ; 0.11	β^- ; 0.0006...; γ 0.0006...; 453...; 238...; 295...; 1021...;	β^- ; 0.13...; γ 0.03...; 381...; 1088...;	β^- ; 1.0... 2.4...; γ 2615; 583...; 511; 880; 277...;	β^- ; 1.8...; γ 1567; 465...; 117...;	β^- ; 1.9... 2.3...; γ 800; 208...;	β^- ?	β^- ?								

136

134

132

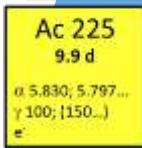
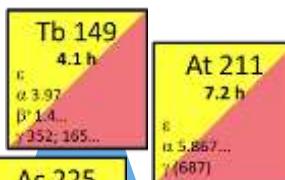
Theranostics

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medicine

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Precision
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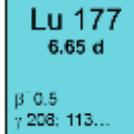
Ac 225

9.9 d

α 5.830; 5.797...

γ 100; (150...)

ϵ ...



Lu 177

6.65 d

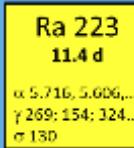
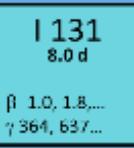
β^- 0.5...

γ 208; 113...

Cure

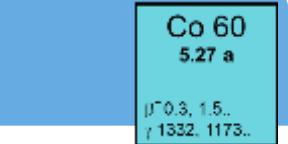
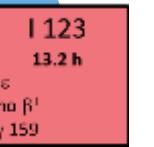
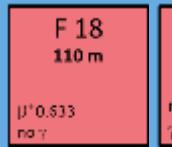
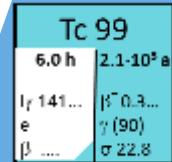
Companion
diagnostic

Therapy



Symptomatic
treatment

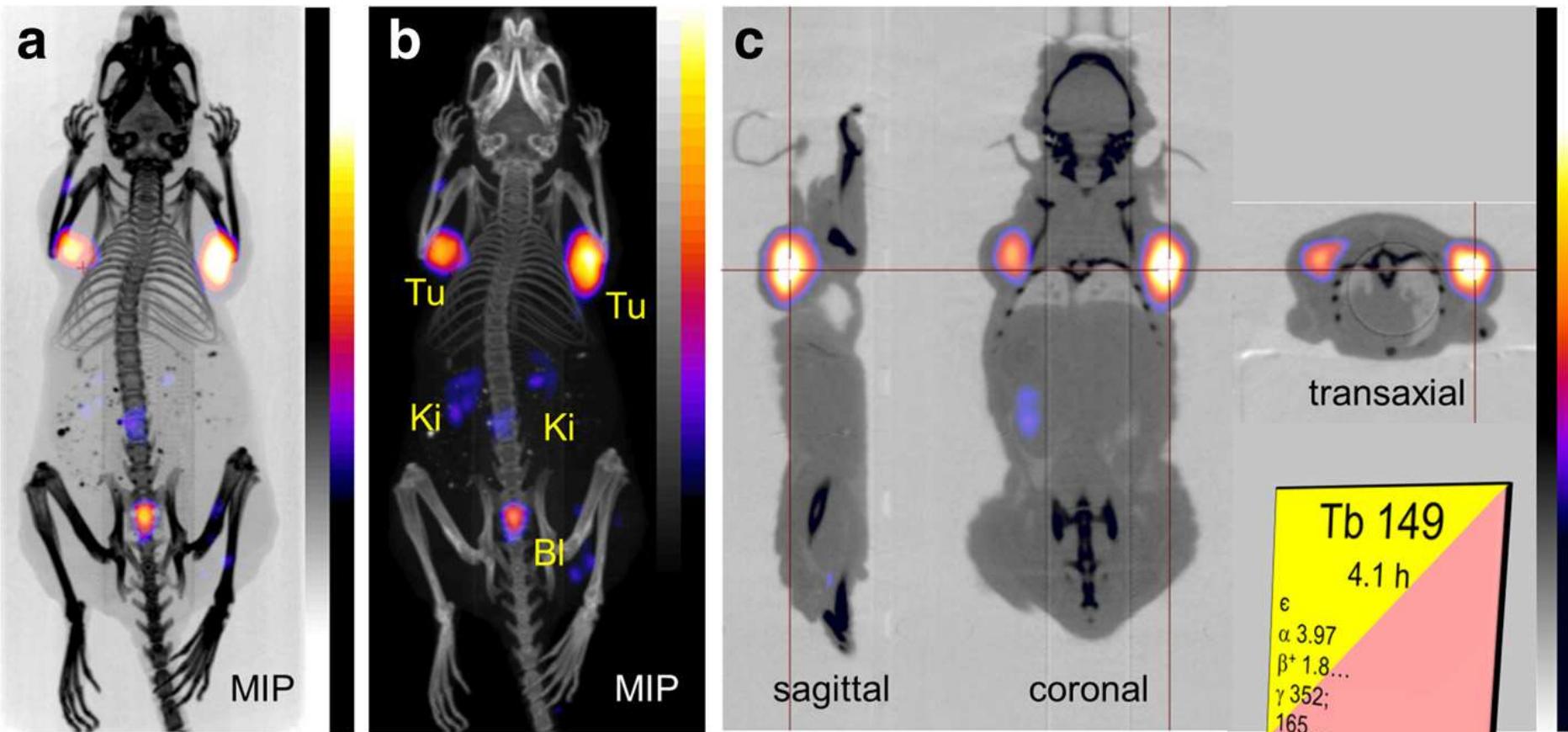
Diagnostic



Prevention

Radionuclides

Alpha-PET with ^{149}Tb



7 MBq $^{149}\text{Tb-DOTANOC}$ 2 h p.i.



PAUL SCHERRER INSTITUT



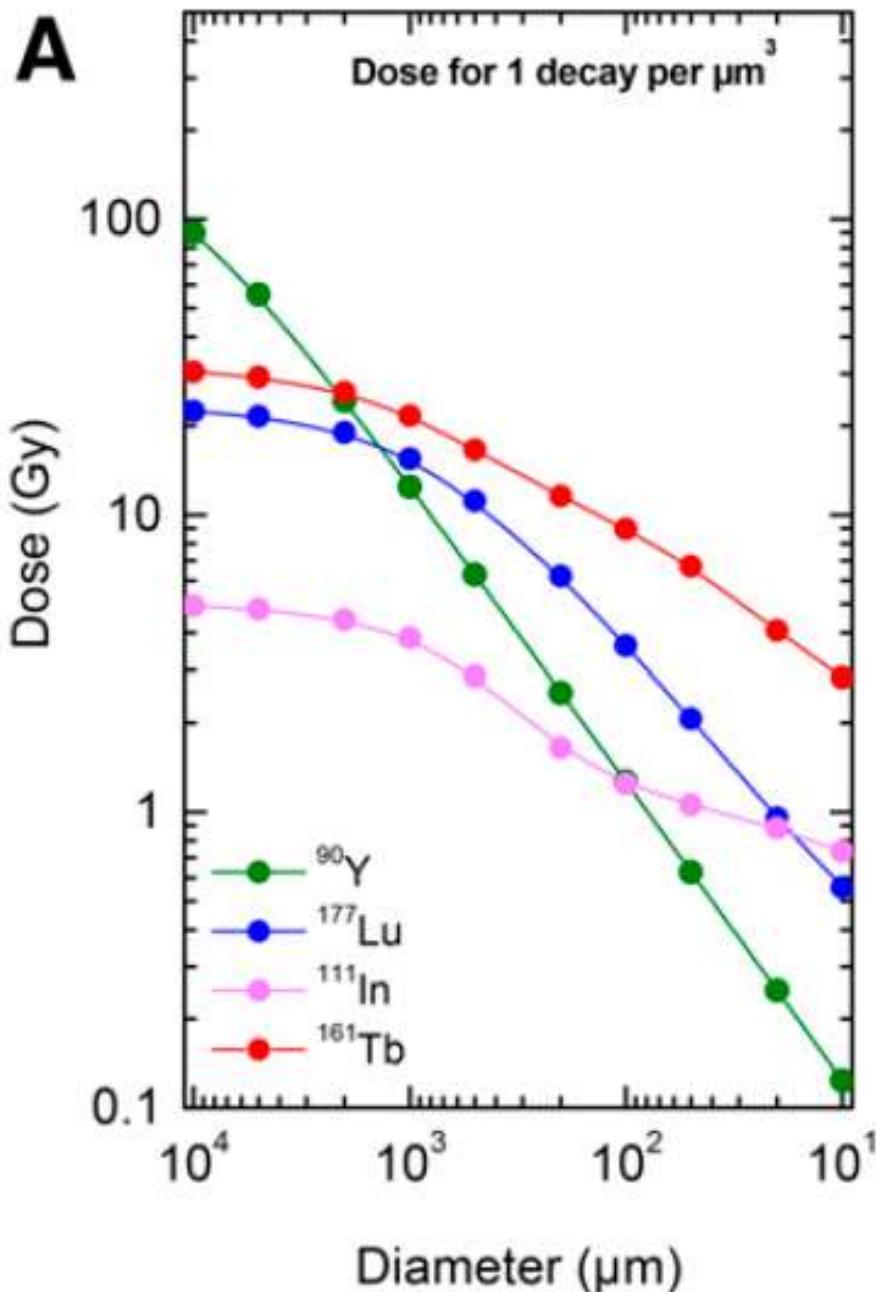
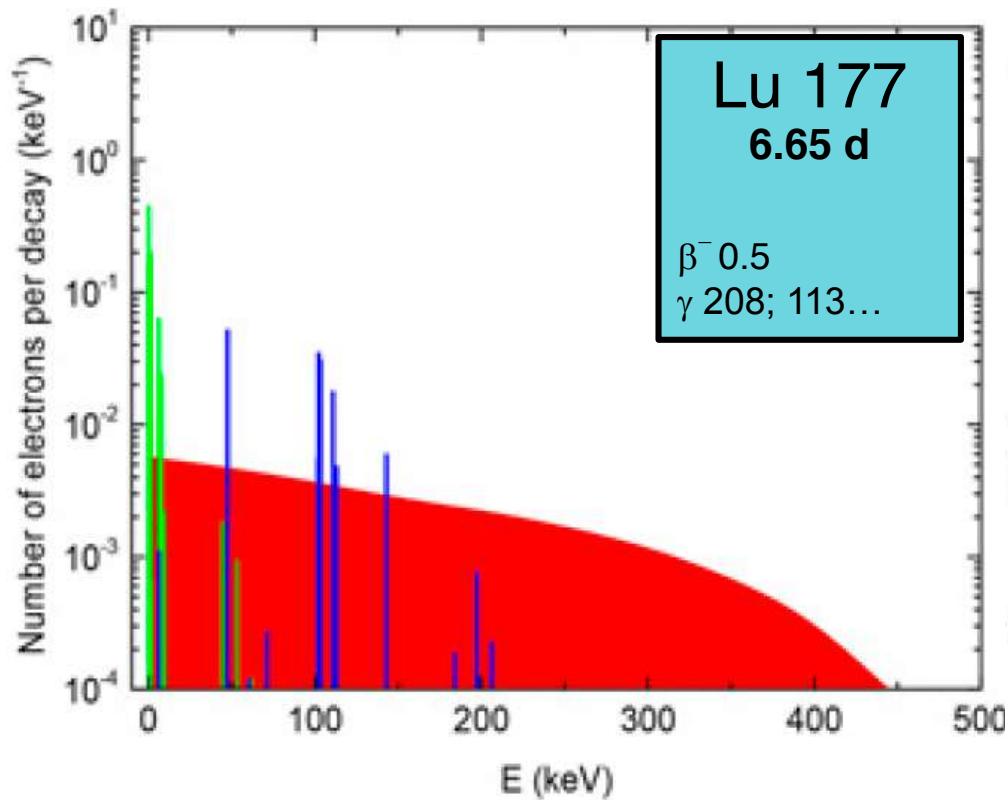
ETH zürich



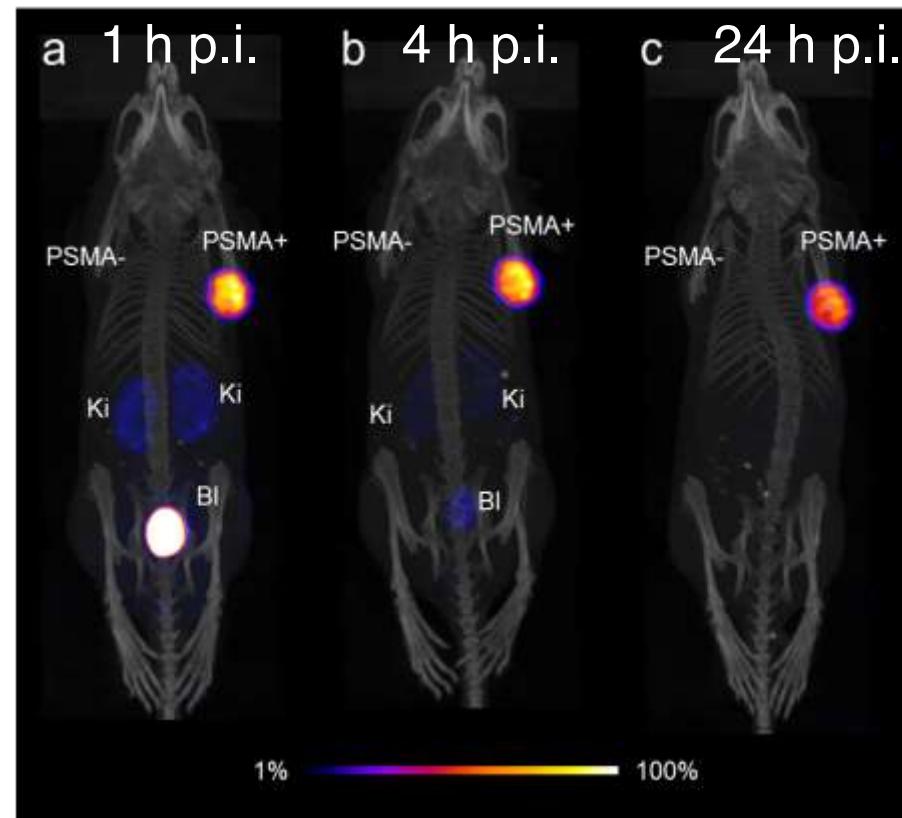
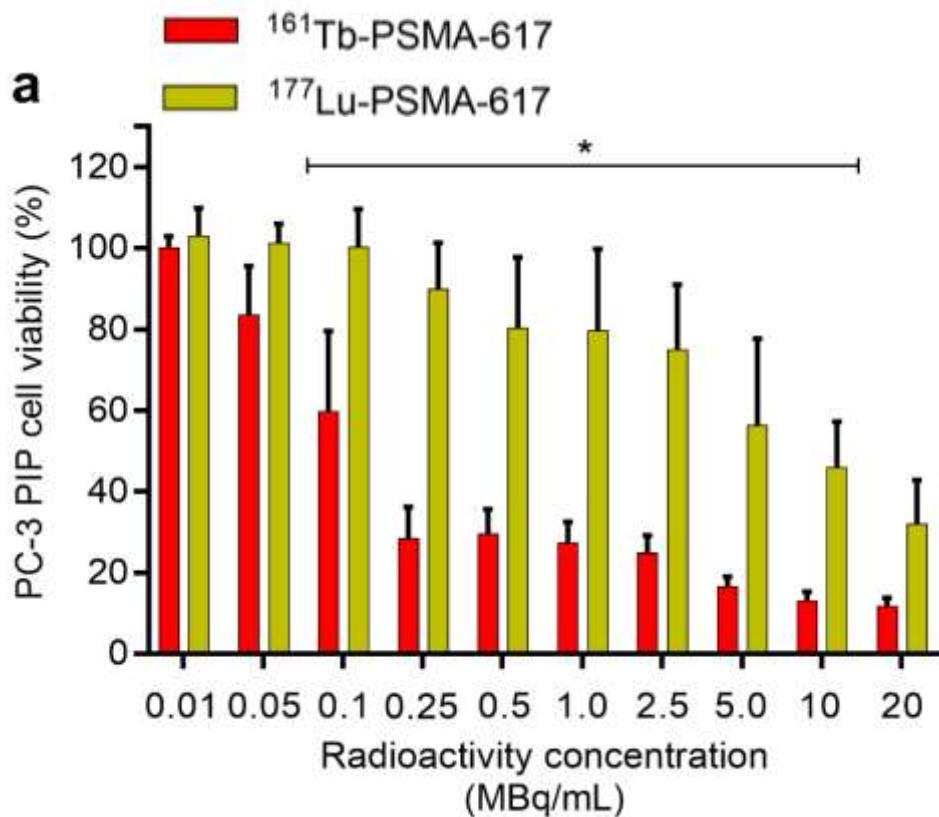
u^b

UNIVERSITÄT
BERN

^{161}Tb versus ^{177}Lu



Terbium-161 for PSMA-targeted radionuclide therapy of prostate cancer



C. Müller et al., Eur J Nucl Med Mol Imaging 2019;46:1919.

Theranostics

=

Personalized
medicine

=

Precision
medicine

Cure

Companion
diagnostic

Therapy

Ga 68
67.6 m

β^+ 1.9...

γ 1077, (...)

Lu 177
6.65 d

β 0.5

γ 208, 113...

Tb 149
4.1 h

α 3.97

β^+ 1.4...

γ (352; 105...)

Ac 225
9.9 d

α 5.830; 5.797...

γ 100; (150,...)

Tb 161
6.9 d

β^+ 0.5; 0.6

γ 26; 49; 75...

Lu 177
6.65 d

β 0.5

γ 208, 113...

Symptomatic
treatment

Diagnostic

Tc 99
6.0 h
 β^- 141...
 γ 141...
 ϵ
 β

Prevention

Radionuclides

Co 60
5.27 a

β^- 0.3, 1.5...
 γ 1332, 1173...

I 131
8.0 d

β^- 1.0, 1.8...
 γ 364, 637...

Ra 223
11.4 d

α 5.716, 5.606...
 γ 269; 154; 124...
 σ 130

F 18
110 m

Tc 99
6.0 h
 β^- 141...
 γ 141...
 ϵ
 β

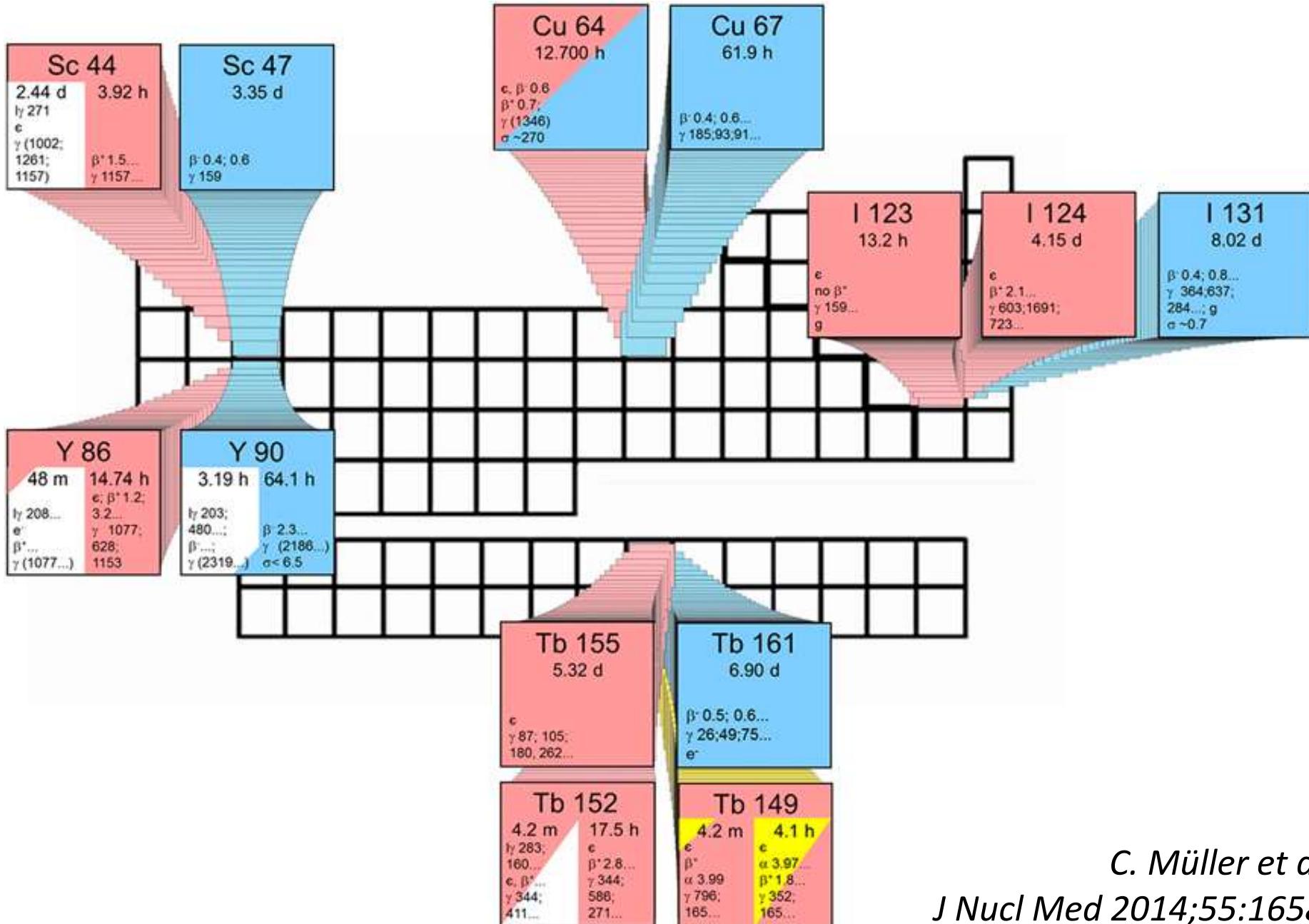
I 123
13.2 h

β^- no β^+
 γ 159

Co 60
5.27 a

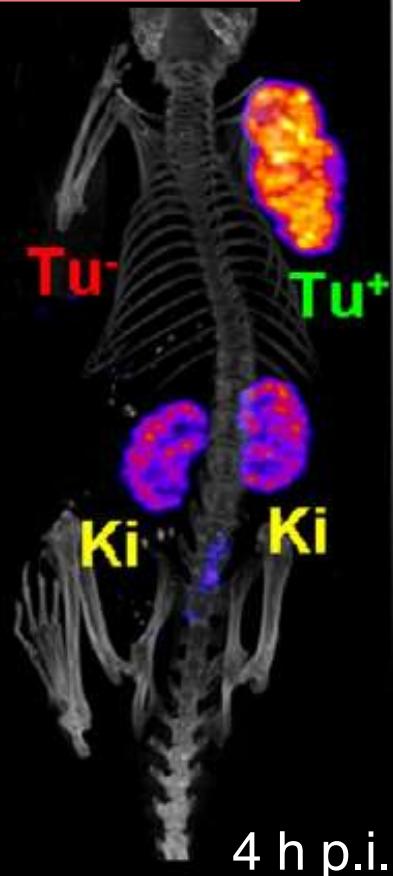
β^- 0.3, 1.5...
 γ 1332, 1173...

Matched pairs for theranostics



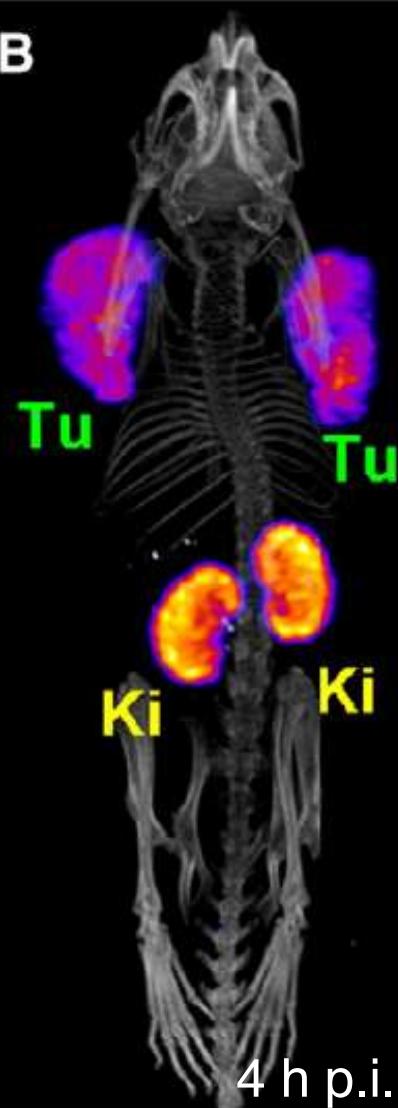
Tb 155
5.3 d

ε
 γ 87; 105; 180...

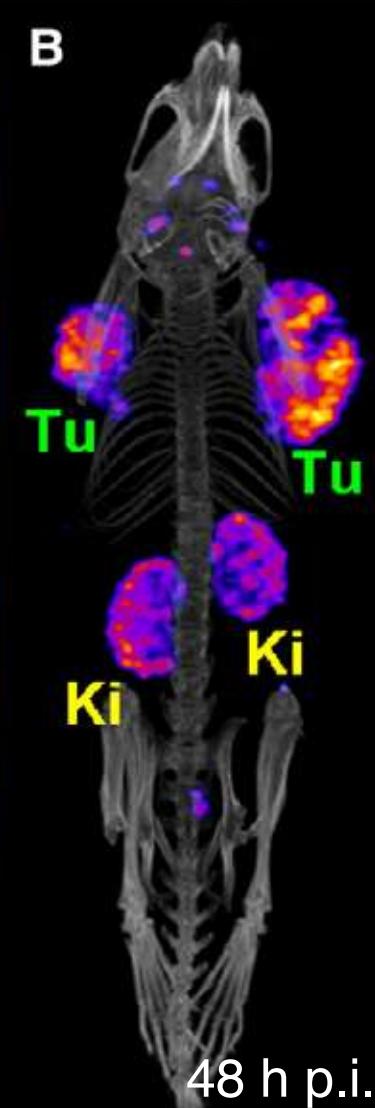


155Tb for SPECT

B



B

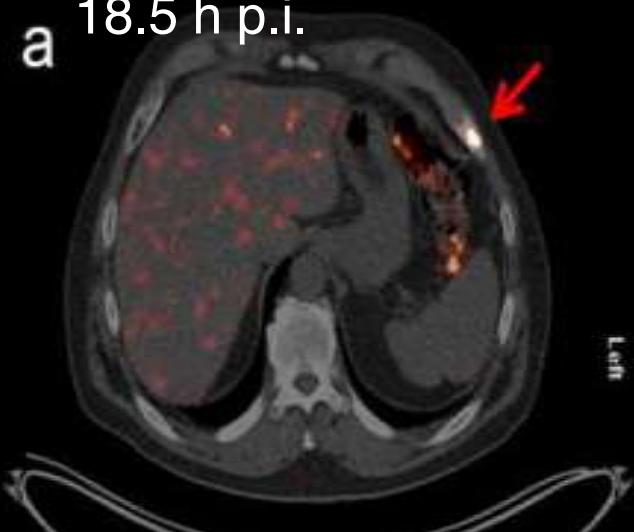


72 h p.i.

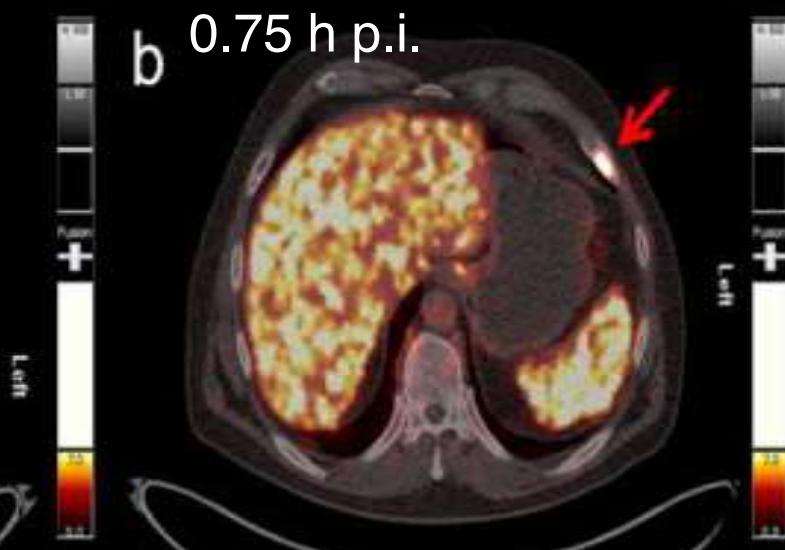
chCE7
SKOV-3ip tumor

First-in-human study with ^{152}Tb -PSMA-617

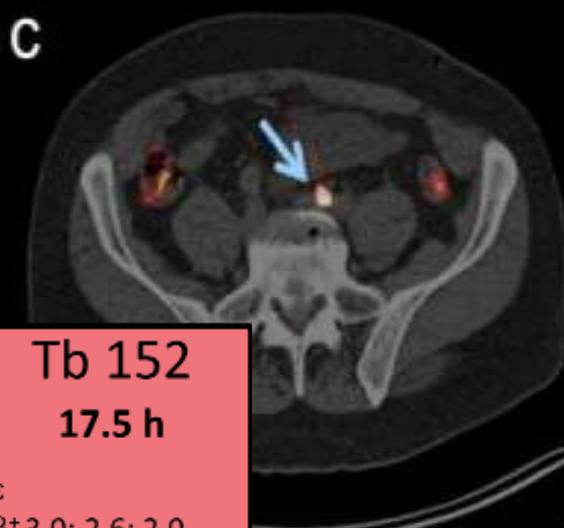
a 18.5 h p.i.



b 0.75 h p.i.



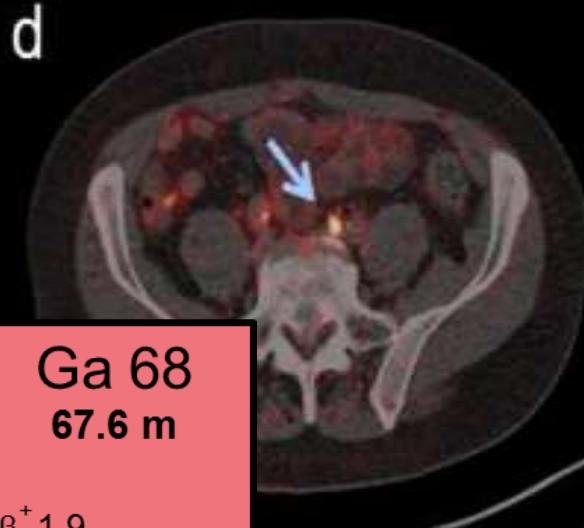
c



Tb 152
17.5 h

β^+ 3.0; 2.6; 2.0...
 γ 344; 271; 586...

d



Ga 68
67.6 m

β^+ 1.9...
 γ 1077, (...)



Zentrale Bad Berka

PAUL SCHERRER INSTITUT



ETH zürich



ENSAR²

Terbium: the Swiss knife for nuclear medicine

Tb 155

5.3 d

ε

γ 87; 105; 180...

e^-

Tb 152

17.5 h

ε

β^+ 3.0; 2.6; 2.0...

γ 344; 271; 586...

Tb 161

6.9 d

β^- 0.5; 0.6

γ 26; 49; 75...

e^-

Tb 149

4.1 h

ε

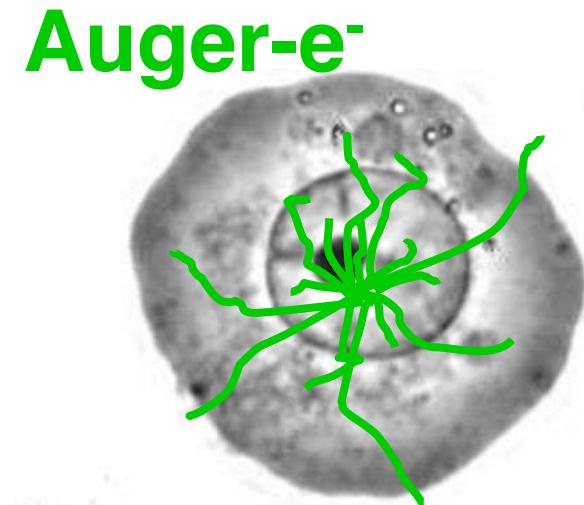
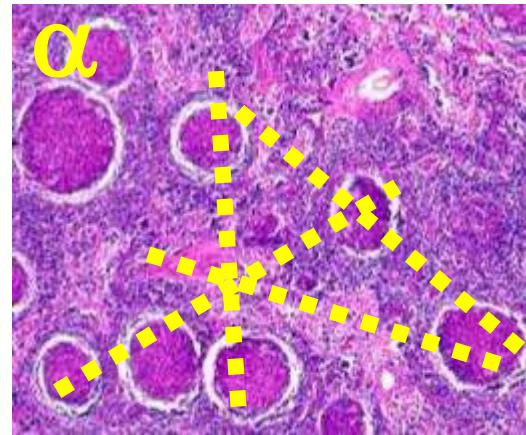
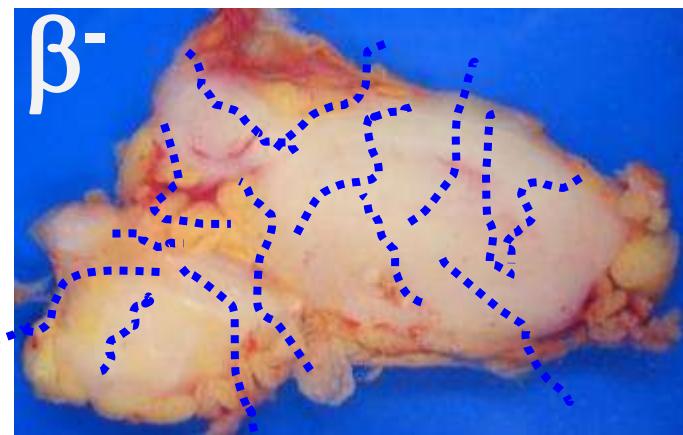
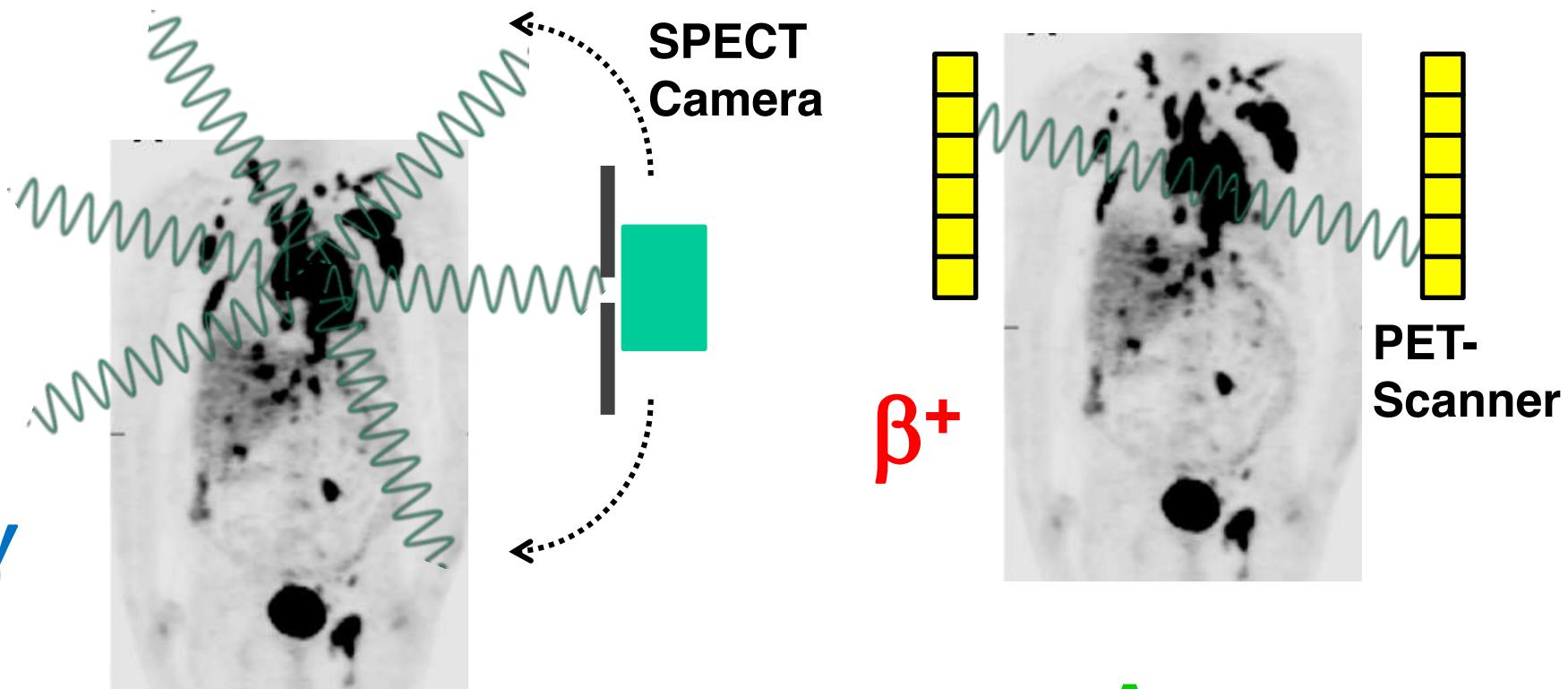
α 3.97

β^+ 1.4...

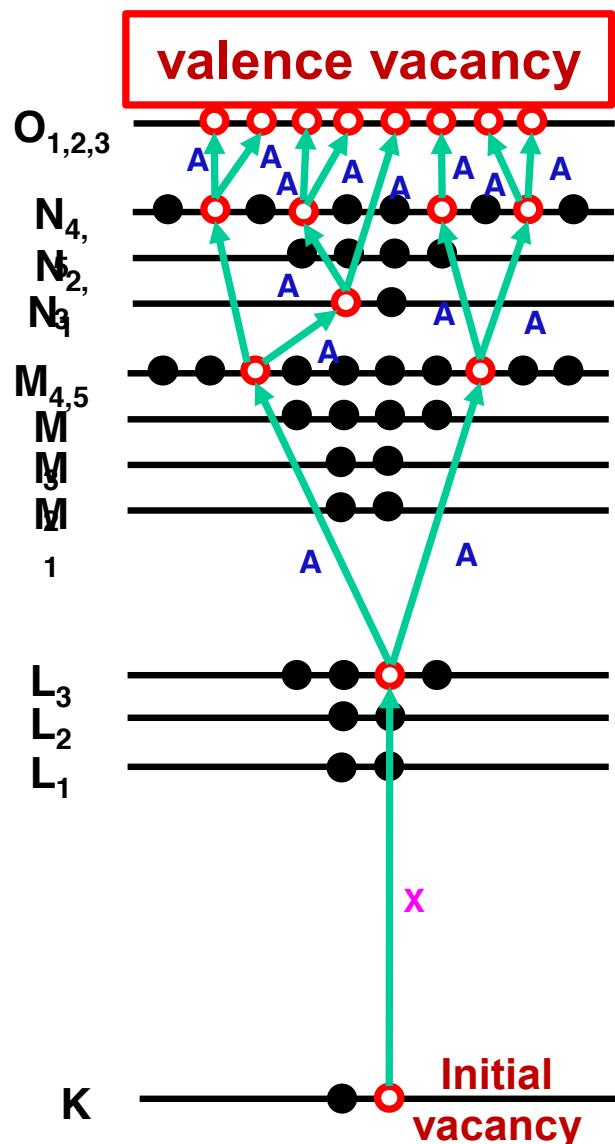
γ 352; 165...



The Nuclear Medicine Alphabet



BrlccEmis: Simulation of Auger electron spectra

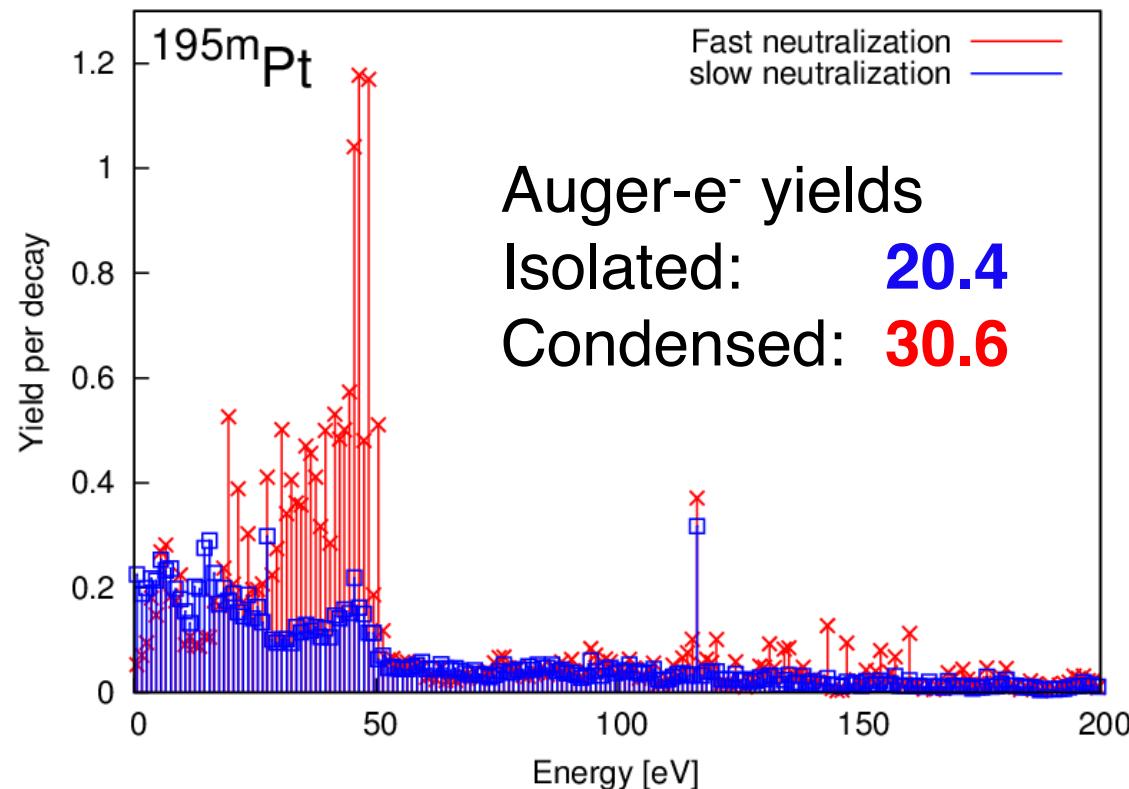


Two approaches:

1. No neutralization during Auger cascade (**slow**) isolated atom

2. Valence vacancy immediately neutralized (**fast**) condensed phase

Pt 195
4.0 d
ly 99;
130...
e ⁻



=

=

Radionuclides

Tb 149
4.1 h
 α 3.97
 β^+ 1.4...
 γ 352; 165...

At 211
7.2 h
 α 5.867...
 γ (687)

Ac 225
9.9 d
 α 5.830; 5.797...
 γ 100; 150...

Er 165
10.3 h
 α no
 β^- no
 γ 5867...

Pt 195
4.0 d stable
 α 99;
 β^- 130...
 e^-

Lu 177
6.65 d
 β^- 0.5
 γ 208; 113...

Tb 161
6.9 d
 β^- 0.5; 0.6
 γ 26; 49; 75...
 e^-

Sc 47
3.35 d
 β^- 0.4; 0.6
 γ 159

Cu 67
2.6 d
 β^- 0.4; 0.6
 γ 185; 93; 91...

Ga 68
67.6 m
 β^+ 1.9...
 γ 1077, (...)

Tb 152
17.5 h
 α
(β^- 3.0; 2.6; 2.0...
 γ 344; 271; 586...)

Tb 155
5.3 d
 α
(β^- 87; 105; 180...
 e^-)

Sc 43
3.9 h
 β^- 1.2...
 γ 373...

Cu 64
12.7 h
 α
(β^- 0.6; $\beta^+ D.7$
 γ (1346))

Sc 44
4.0 h
 β^- 1.5...
 γ 1157...

Co 60
5.27 a
 β^- 0.3, 1.5...
 γ 1332, 1173...

I 131
8.0 d
 β^- 1.0, 1.8...
 γ 364, 637...

Ra 223
11.4 d
 α 5.716, 5.606,...
 γ 269; 154; 324...
 σ 130

Symptomatic
treatment

Tc 99
6.0 h
2.1-10³ Bq
 β^- 141...
 e^-
 β^- 0.3...
 γ (90)
 σ 22.8

Diagnostic

F 18
110 m
 β^- 0.633
 α
 γ

I 123
13.2 h
 α
 β^-
 γ 159

Prevention

Co 60
5.27 a
 β^- 0.3, 1.5...
 γ 1332, 1173...

Prospects of targeted radionuclide therapies ?

2014	Algeta	⇒ Bayer	2.4 G\$
2018	AAA	⇒ Novartis	3.9 G\$
2018	Sirtex	⇒ CGE/CDH	1.4 G\$
2019	Endocyte	⇒ Novartis	2.1 G\$
2019	Blue Earth	⇒ Bracco	0.5 G\$
2019	BTG	⇒ Boston Scientific	4.2 G\$
Sum over 5 years			14.5 G\$



Bayer HealthCare



NOVARTIS



Boston
Scientific
Advancing science for life™



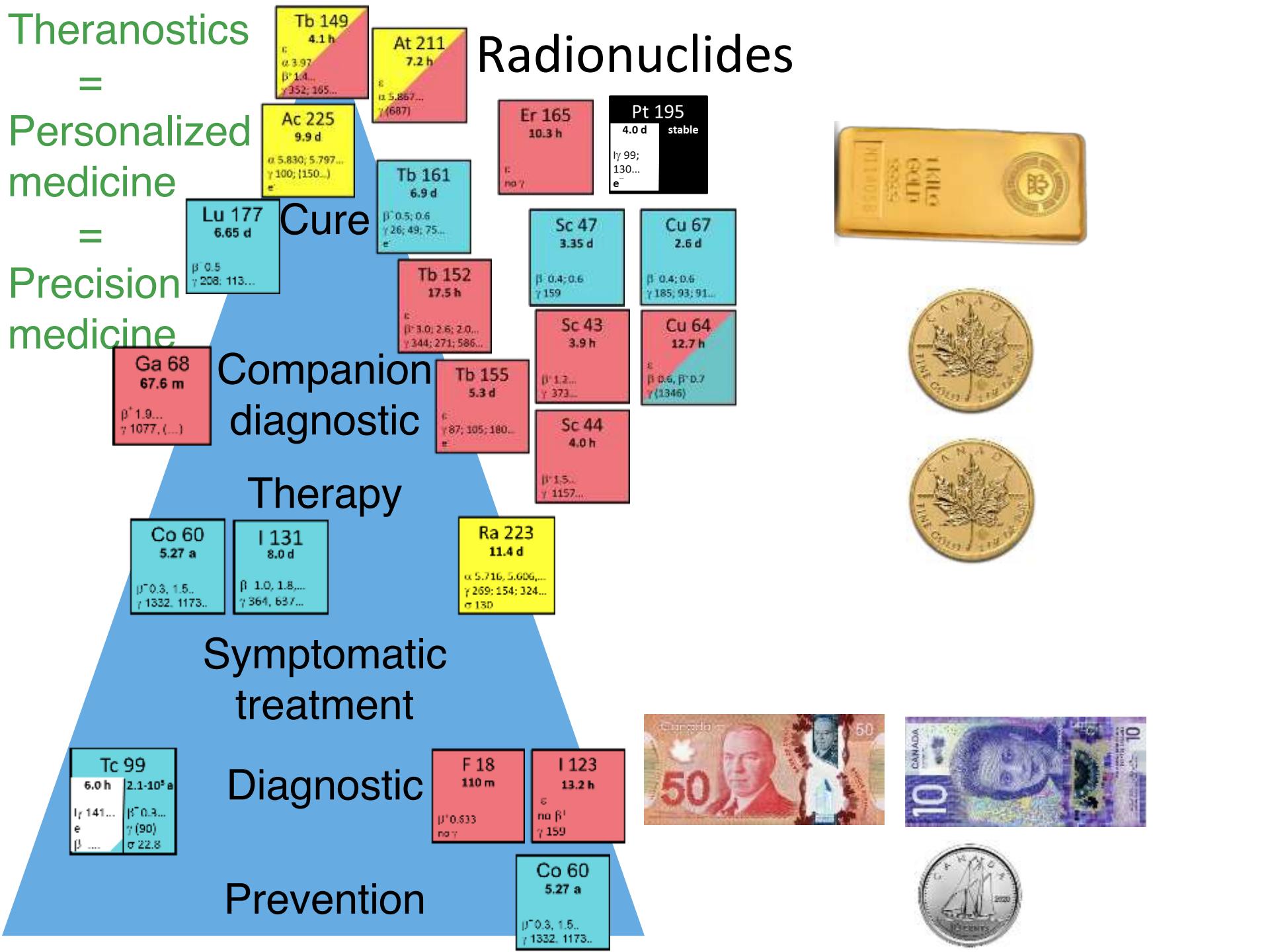
远大健康
CGE HEALTHCARE

SANOFI

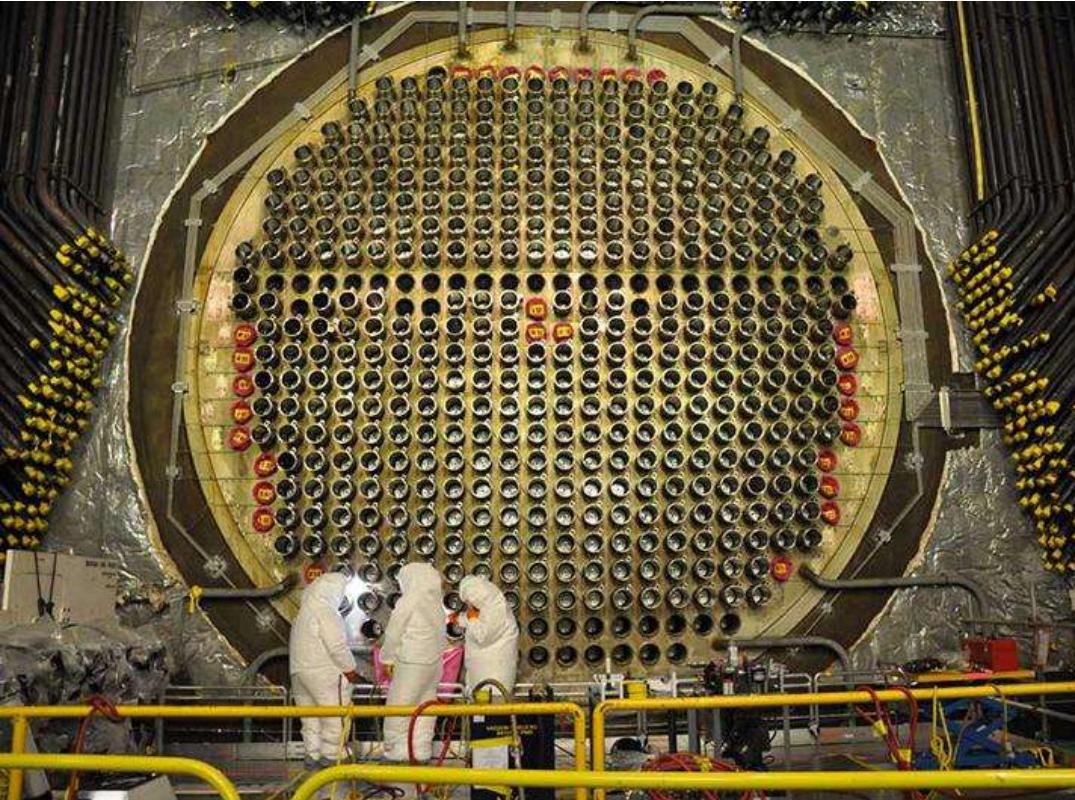


oranomed





^{60}Co production in CANDU reactors



$\Phi \approx 1\text{E}14 \text{ cm}^{-2}\text{s}^{-1}$ over many m^2

“particle current” $\approx 16 \text{ p}\mu\text{A}/\text{cm}^2$

Total “particle current” $\approx 40 \text{ A}$
easily usable $\gg 100 \text{ mA}$



Different ways to ^{60}Co (as a theoretical case study)

Cu 58 3.20 s β^+ 7.5... γ 1454, 1448 40...	Cu 59 82 s β^+ 3.8... γ 1302, 878 339, 465...	Cu 60 23 m β^+ 2.0, 3.9... γ 1332, 1792 826...	Cu 61 3.4 h β^+ 1.2... γ 283, 656, 67 1186...	Cu 62 9.74 m β^+ 2.9... γ (1173...)	Cu 63 69.15 σ 4	Cu 64 12.7004 h β^- 0.6, β^+ 0.7 γ (1346) σ -270	Cu 65 30.85 σ 2.17
Ni 57 36 h β^+ 0.8... γ 1378, 1920 127...	Ni 58 68.077 σ 4.6 $\sigma_{n,\alpha} < 3E-5$	Ni 59 7.5·10 ⁴ a β, β^+ ... no γ σ 77.7, $\sigma_{n,\alpha}$ 14 $\sigma_{n,p}$ 2, σ_{abs} 92	Ni 60 26.223 σ 2.9	Ni 61 1.1399 σ 2.5 $\sigma_{n,\alpha}$ 5	Ni 62 3.6346 σ 1.5	Ni 63 100 a β^- 0.07 no γ σ 20	Ni 64 0.9255 σ 1.6
Co 56 77.236 d β, β^+ 1.5... γ 847, 1238 2598, 1144 1038...	Co 57 271.80 d σ 122, 136, 14	Co 58 8.94 h β, β^+ 0.5, 1.3 γ 811 σ 140000 σ 1900	Co 59 100 σ 20.7 + 16.5	Co 60 10.5 m β, β^+ 0.5, 1.3 γ 1332... σ 58	Co 61 1.65 h β, β^+ 1.2... γ 67, 909...	Co 62 14.0 m β^- 2.9... γ 1173 1163 2003...	Co 63 1.5 m β^- 4.1... γ 1173 1163 2302 1129...
Fe 55 2.73 a σ no γ σ 13 $\sigma_{n,\alpha}$ 0.01	Fe 56 91.754 σ 2.8	Fe 57 2.119 σ 1.4	Fe 58 0.282 σ 1.3	Fe 59 44.494 d β^- 0.5, 1.6... γ 1099, 1292... σ < 10	Fe 60 2.62·10 ⁶ a β^- 0.2 m	Fe 61 6.0 m β^- 2.6, 2.8... γ 1205, 1027 298...	Fe 62 68 s β^- 2.5 γ 506 g

$^{59}\text{Co}(n,\gamma)^{60}\text{Co}$: 1 n \approx 1 atom ^{60}Co

“160 mA” = 1 EBq ^{60}Co

$^{59}\text{Co}(d,p)^{60}\text{Co}$: 40 MeV d \approx 1.2E-3 ^{60}Co

25000x SPIRAL2 = 1 EBq ^{60}Co

$^{nat}\text{Ni}(p,*)^{60}\text{Co}$: 30 MeV p < 1E-5 ^{60}Co

70000x TR30 = 1 EBq ^{60}Co equiv.

$^{nat}\text{Fe}(p,*)^{56}\text{Co}$: 30 MeV p \approx 1.2E-3 ^{56}Co

100000x main cyc. = 1 EBq ^{60}Co

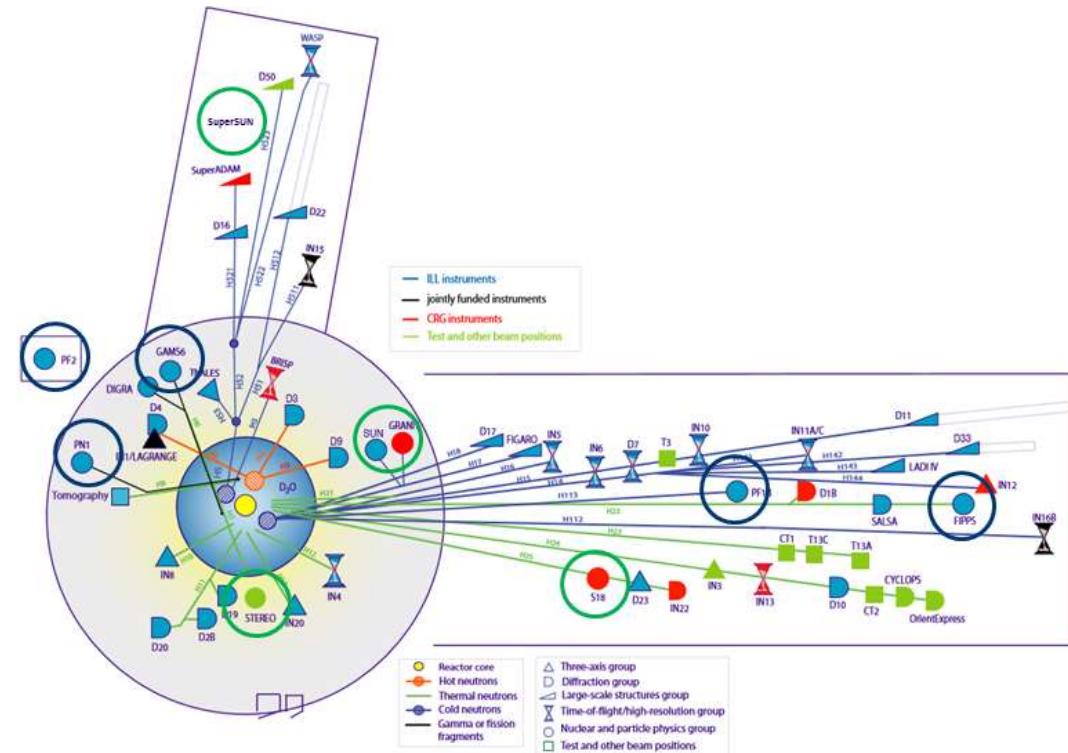
$^{nat}\text{Cu}(p,*)^{60}\text{Co}$: 500 MeV p \approx 0.015 ^{60}Co

$^{nat}\text{Ni}(\gamma,*)^{60}\text{Co}$: 30 MeV e⁻ < 1E-6 ^{60}Co

^{177}Lu : The “gold standard” for radionuclide therapy

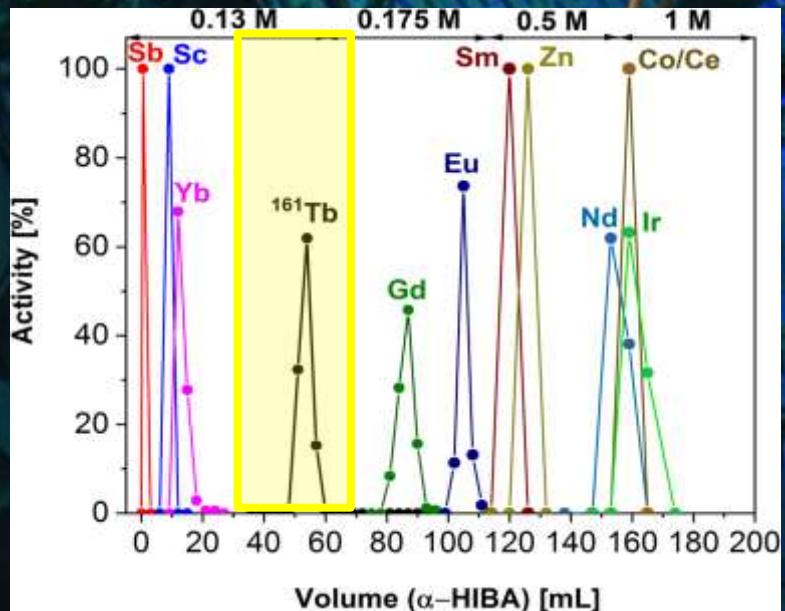


Institut Laue-Langevin 2018:
≈ 1600 scientific users at ILL
≈ 4000 patients get ^{177}Lu from ILL



Production of non-carrier-added ^{161}Tb

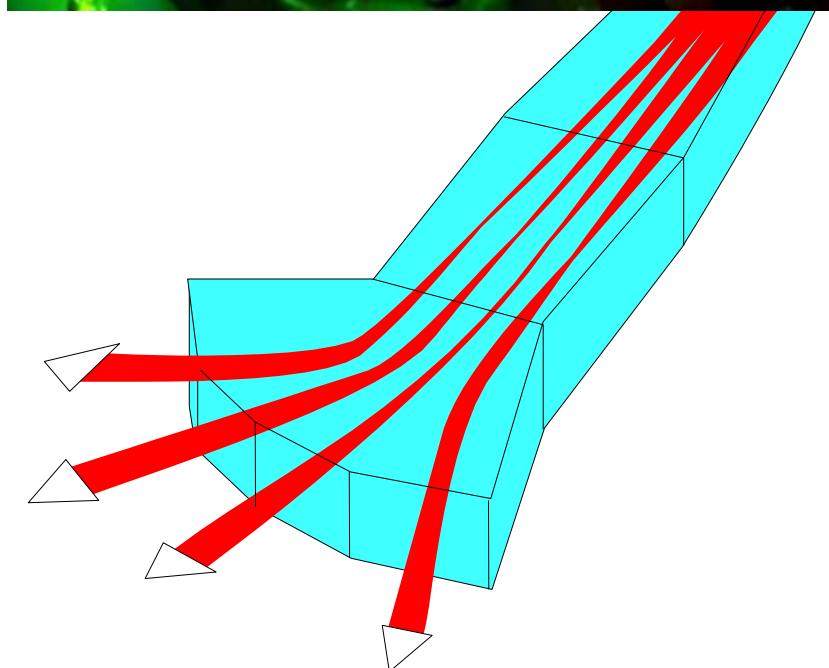
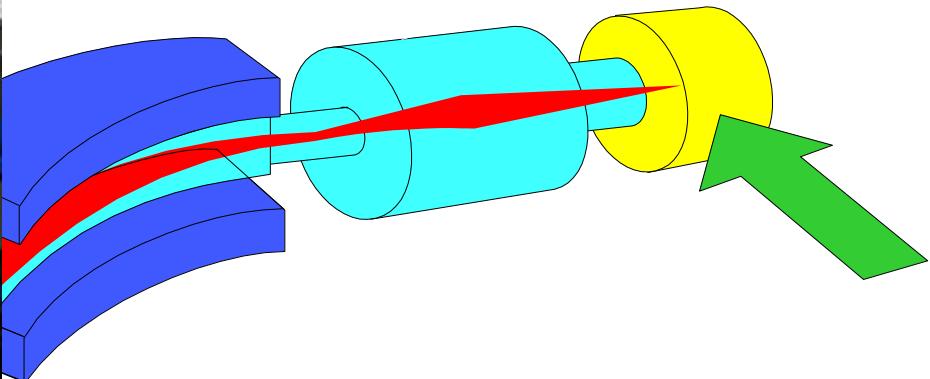
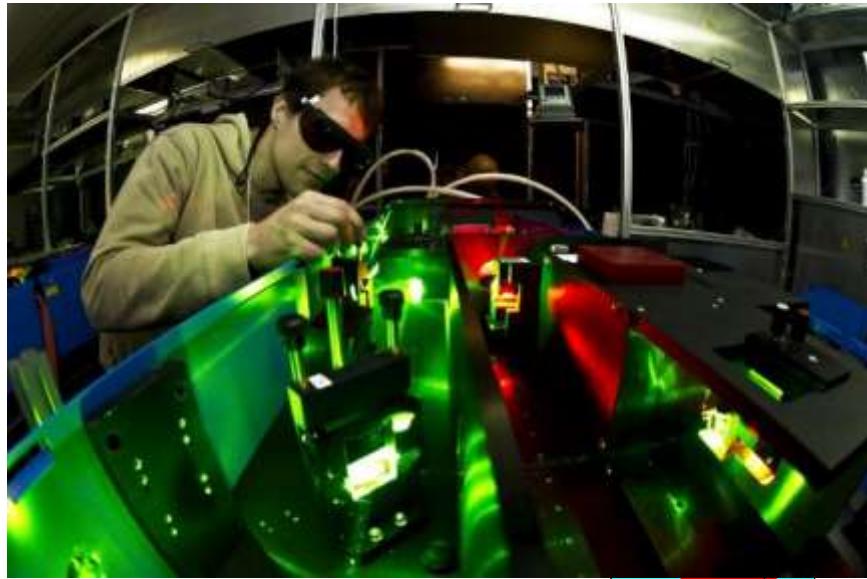
Dy 160 2.329	Dy 161 18.889	Dy 162 25.475	Dy 163 24.896	Dy 164 28.260
σ_{60} $\sigma_{n,\alpha} < 0.0003$	σ_{600} $\sigma_{n,\alpha} < 1E-6$	σ_{170}	σ_{120} $\sigma_{n,\alpha} < 2E-5$	$\sigma_{1610} + 1040$
Tb 159 100	Tb 160 72.3 d $\beta^- 0.6; 1.7\dots$ $\gamma 879; 299;$ $966\dots$ $\sigma 570$	Tb 161 6.90 d $\beta^- 0.5; 0\dots$ $\gamma 26; 49; 5\dots$ e^-	Tb 162 7.76 m $\beta^- 1.4; 2.4\dots$ $\gamma 260; 808;$ $888\dots$	Tb 163 19.5 m $\beta^- 0.8; 1.3\dots$ $\gamma 351; 390;$ $494\dots$
Gd 158 24.84	Gd 159 18.48 h $\beta^- 1.0\dots$ $\gamma 364; 58\dots$	Gd 160 21.86 $\sigma 1.5$	Gd 161 3.66 m $\beta^- 1.6; 1.7\dots$ $\gamma 361; 315;$ $102\dots$ $\sigma 20000$	Gd 162 8.2 m $\beta^- 1.0\dots$ $\gamma 442; 403\dots$



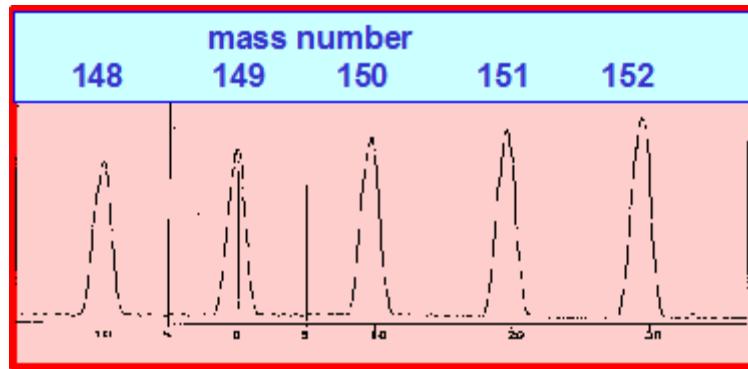
Irradiation at ILL & Necsa, chemical separation at PSI

N. Gracheva et al., EJNMMI Radiopharm Chem 2019;4:12.

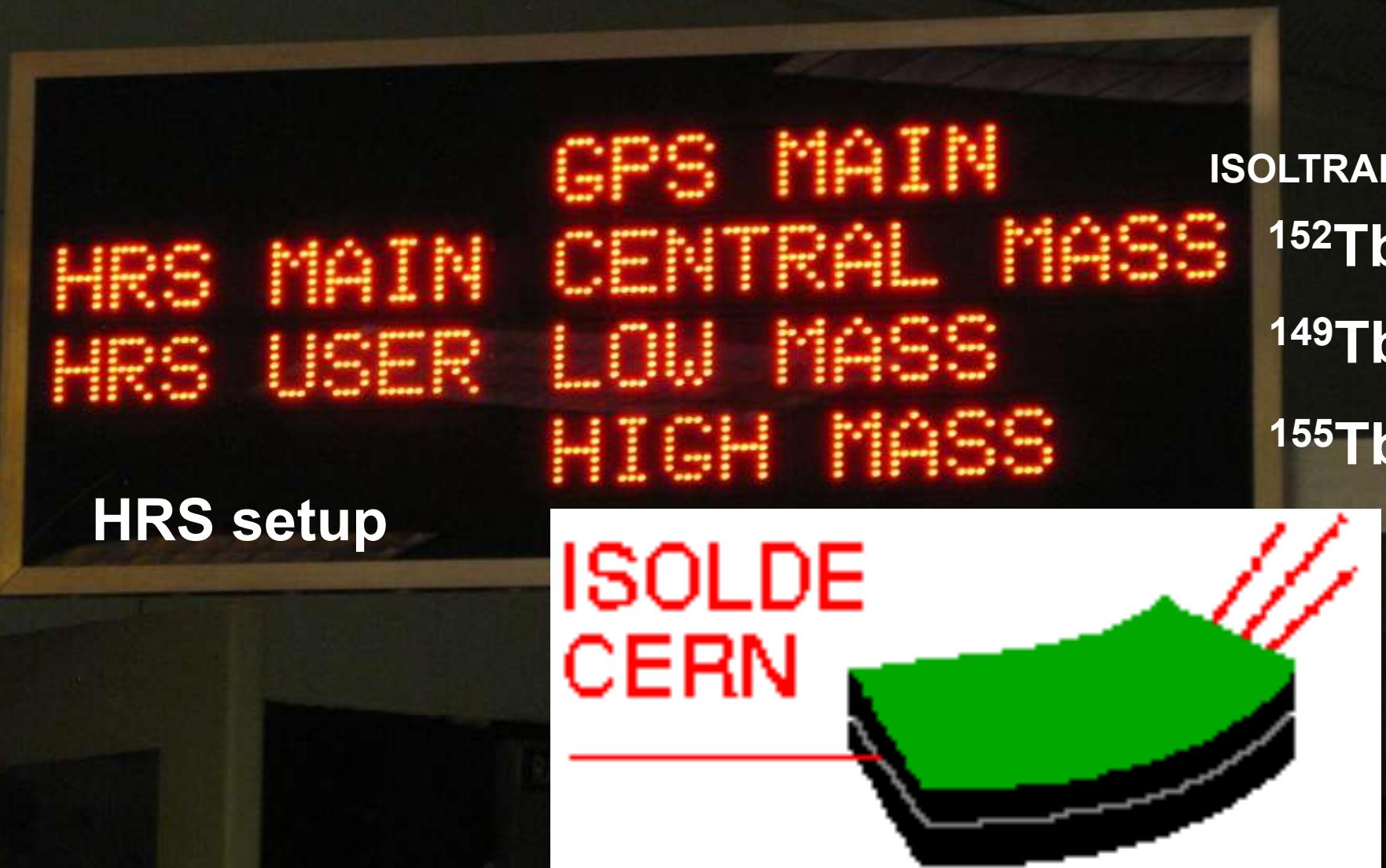
Production of ^{149}Tb , ^{152}Tb and ^{155}Tb at ISOLDE



radioactive ion beams



Efficient parallel operation



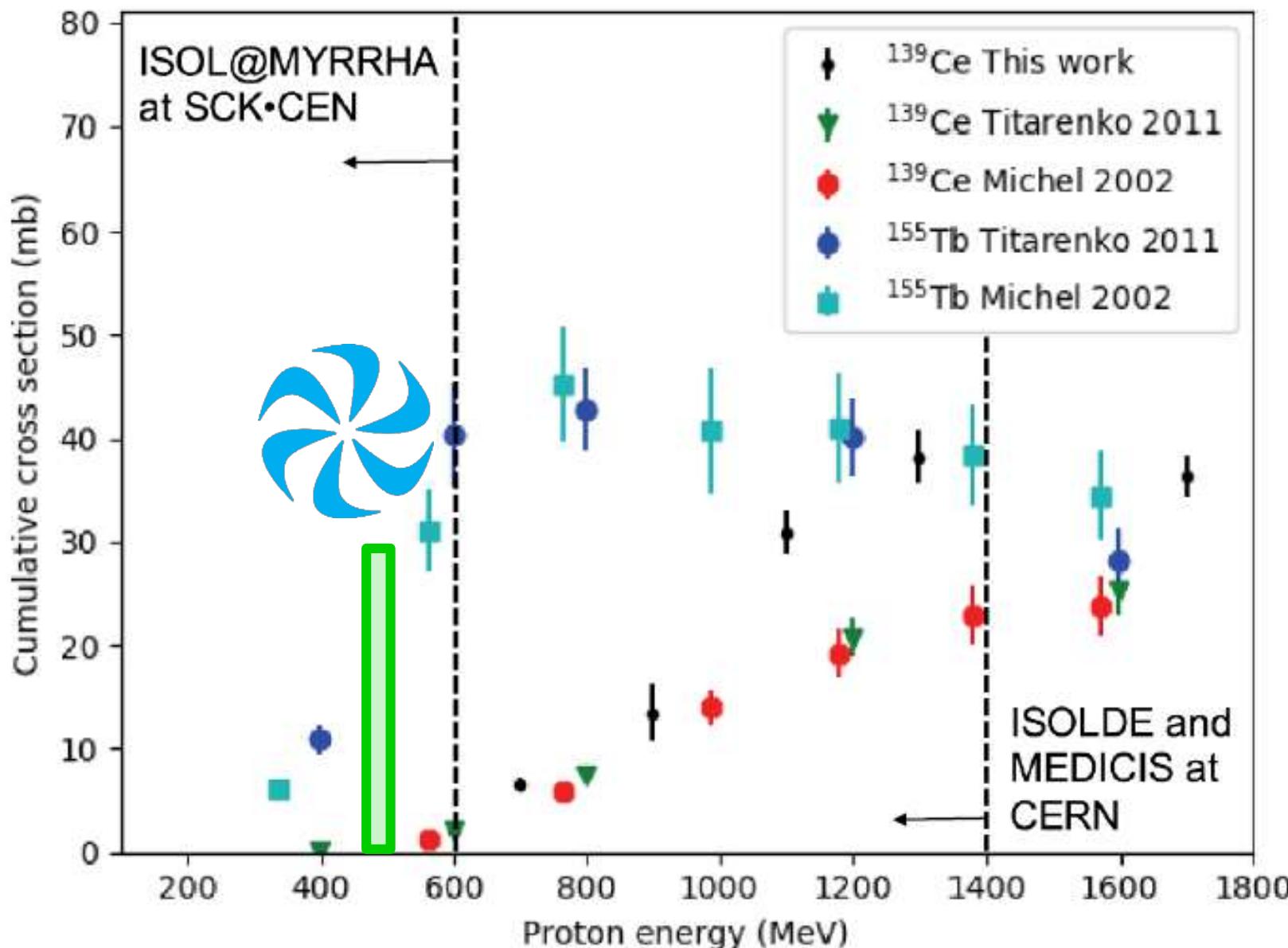


Medicis

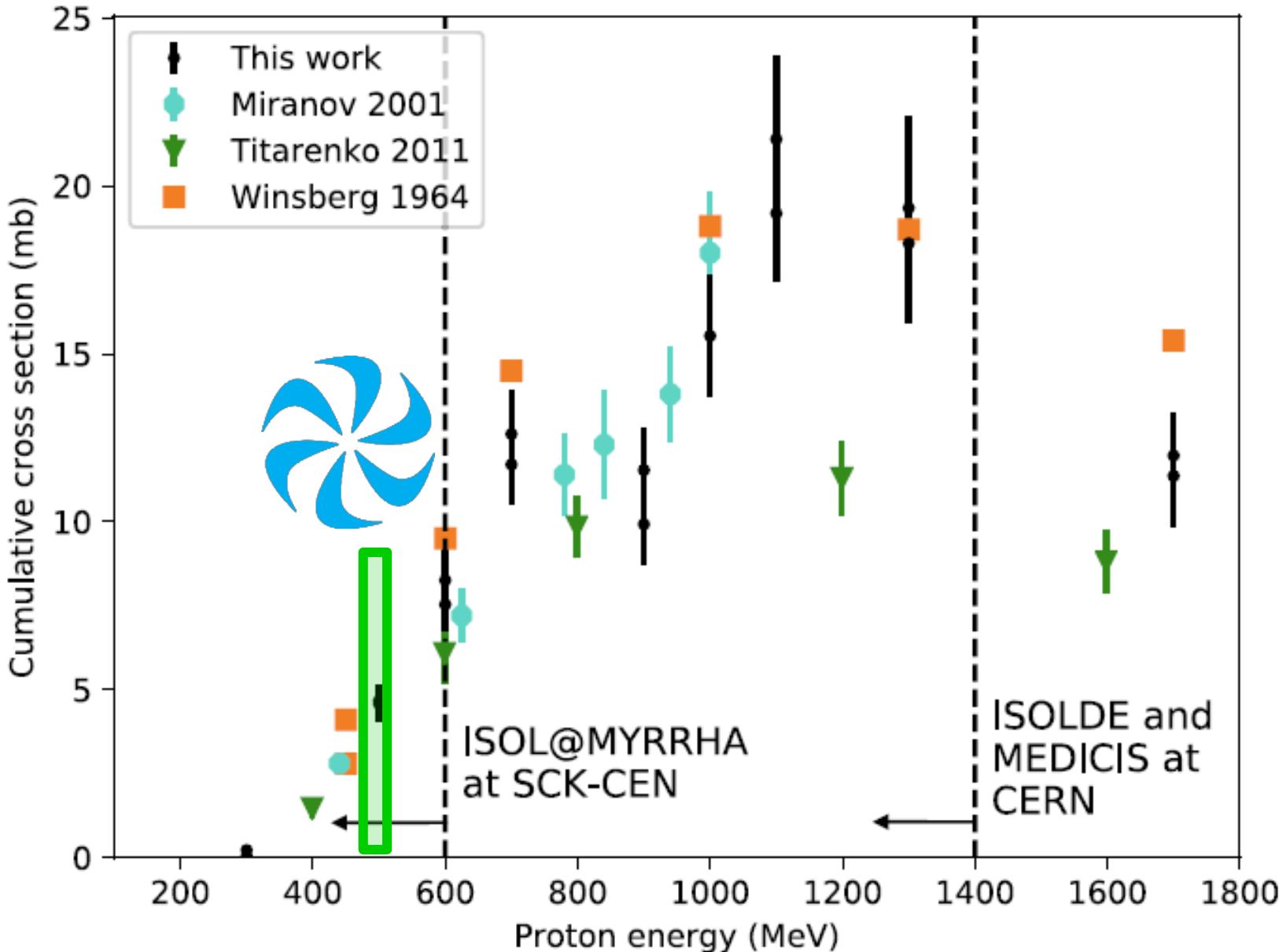
a very useful beam dump !



Spallation cross-sections: ^{155}Tb vs. ^{139}Ce



Spallation cross-sections: ^{149}Tb vs. ^{133}Ce



A great model: the US DOE Isotope Program

NIDC: National Isotope Development Center +

https://www.isotopes.gov 110% ⌂ ⌂ ⌂

NIDC NATIONAL ISOTOPE DEVELOPMENT CENTER

the government source of isotopes for science, medicine, security, & other applications

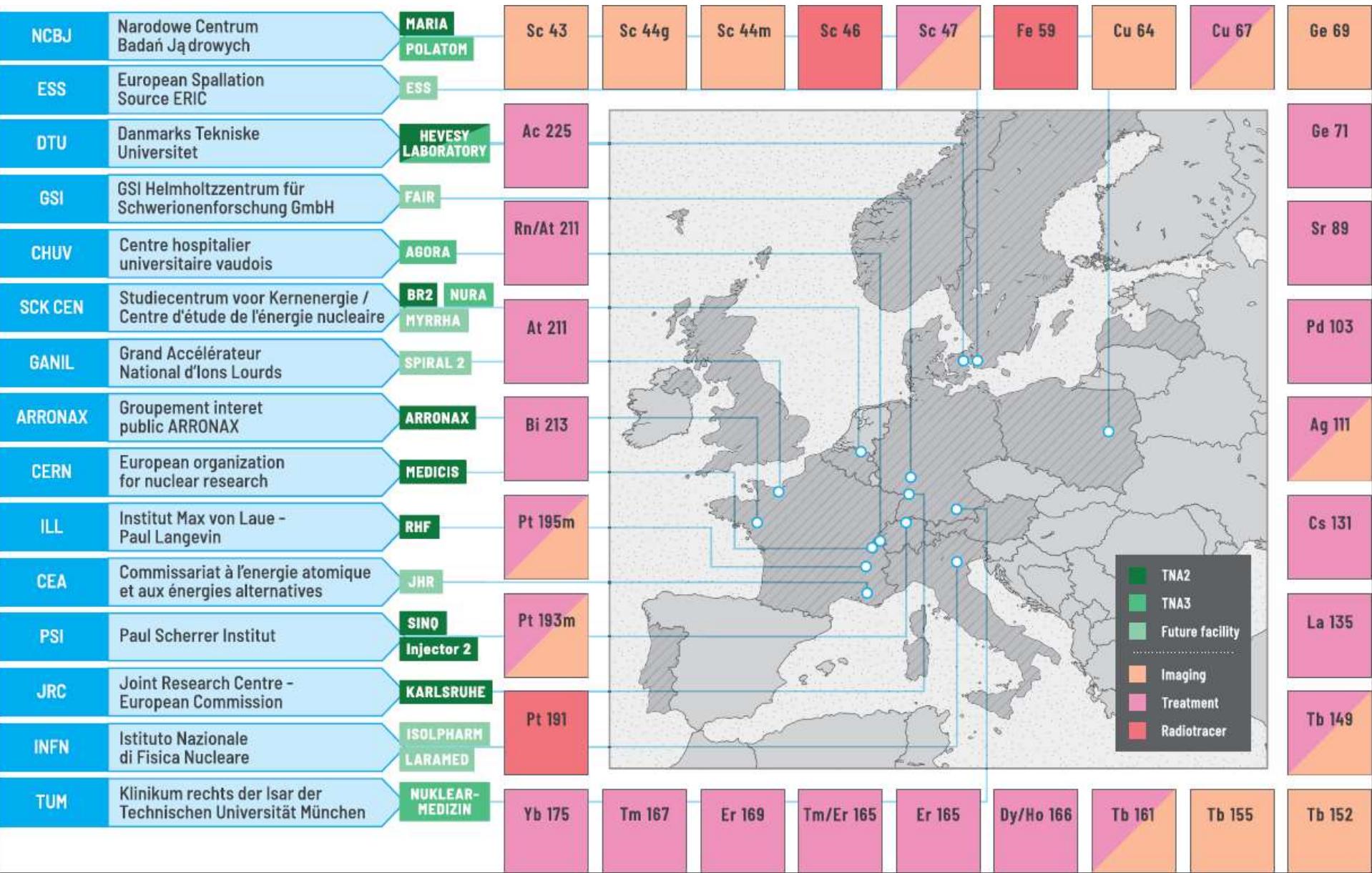
U.S. DEPARTMENT OF ENERGY Office of Science

Product Catalog News Education & Outreach Isotope Production About NIDC

Welcome to the NIDC!

The **National Isotope Development Center (NIDC)** interfaces with the isotope user community and manages the coordination of isotope production across the facilities and business operations involved in the production, sale, and distribution of isotopes. A virtual center, the NIDC is funded by the [U.S. Department of Energy Isotope Program](#) within the [Office of Nuclear Physics](#) in the [Office of Science](#).

Proposed EU project PRISMAP: PRoduction of high purity Isotopes by mass Separation for Medical Application



Summary and Outlook

- Targeted Radionuclide Therapies (TRT) are transforming nuclear medicine
- Bright future for Theranostics, in particular in oncology
- NP research infrastructures can play an important role for supply of EMERGING and UNCONVENTIONAL radionuclides ($^{149/152/155/161}\text{Tb}$, ^{211}At , ^{67}Cu , $^{193\text{m}/195\text{m}}\text{Pt}$, ...)
- but medical applications using these radionuclides must be developed in parallel
- Research infrastructures are NOT competitive in production of “industrial” radionuclides
- PRISMAP: a European Isotope Development Center !

Specific ideas/wishes for TRIUMF



1. Logic choices for ISAC:

- Tb isotopes and other n-deficient lanthanides (Auger)
- ^{225}Ac , $^{211}\text{Rn}/^{211}\text{At}$
- regular harvesting of R&D isotopes
- needs shielded collection position(s)!

2. Prospects for ARIEL

- $^{226}\text{Ra}(\gamma, \text{n})^{225}\text{Ra}(\beta^-)^{225}\text{Ac}$, $\text{Zn}(\gamma, \text{p})^{67}\text{Cu}..$

3. Alpha beams (30-40 MeV)

- for production of ^{211}At , ^{43}Sc , $^{117\text{m}}\text{Sn}$, $^{195\text{m}}\text{Pt}$, $^{193\text{m}}\text{Pt}$, etc.
- NO ^4He beams in Canada: cyclotron or LINAC prototype?

4. Synergies with laser spec., hyperfine interactions

- fundamental spectroscopy of At-molecules
- TDPAC for chelator/labelling development (cf. β -NMR)