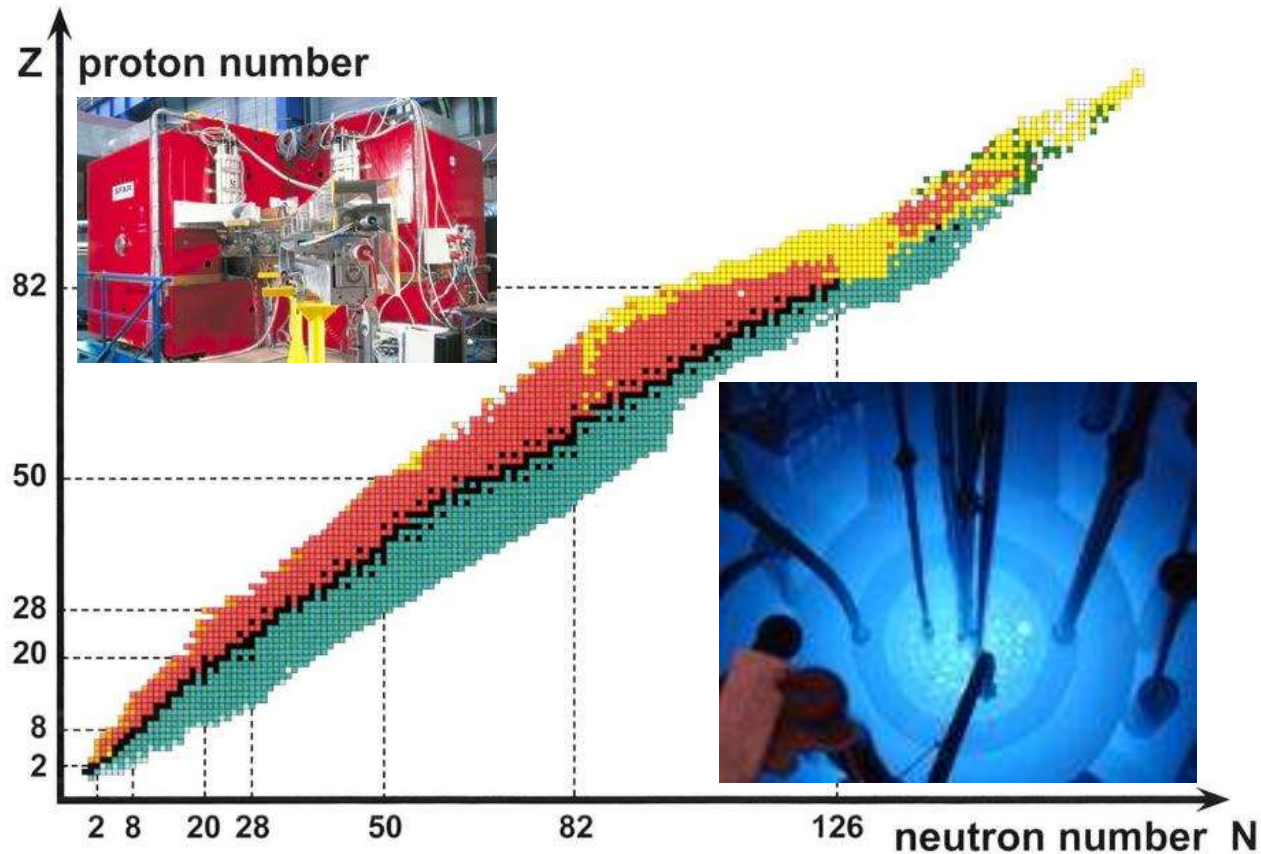
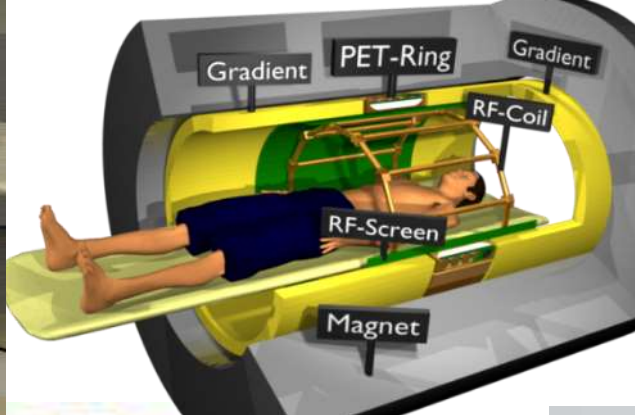


Radioisotopes: technical perspective

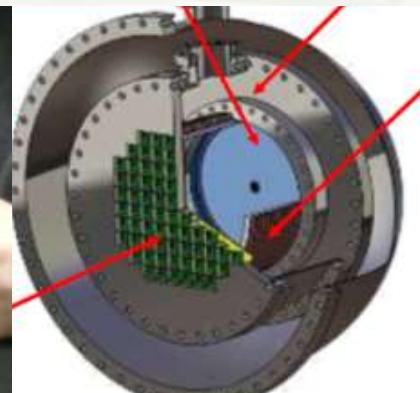
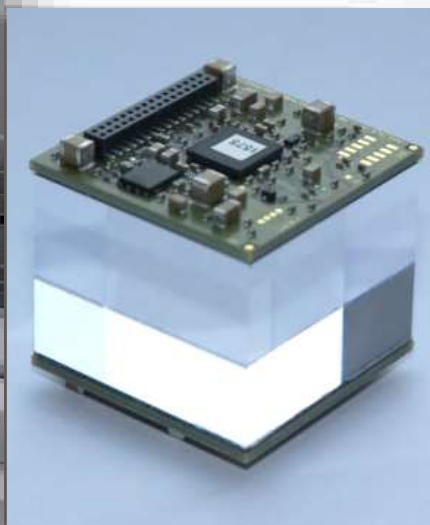
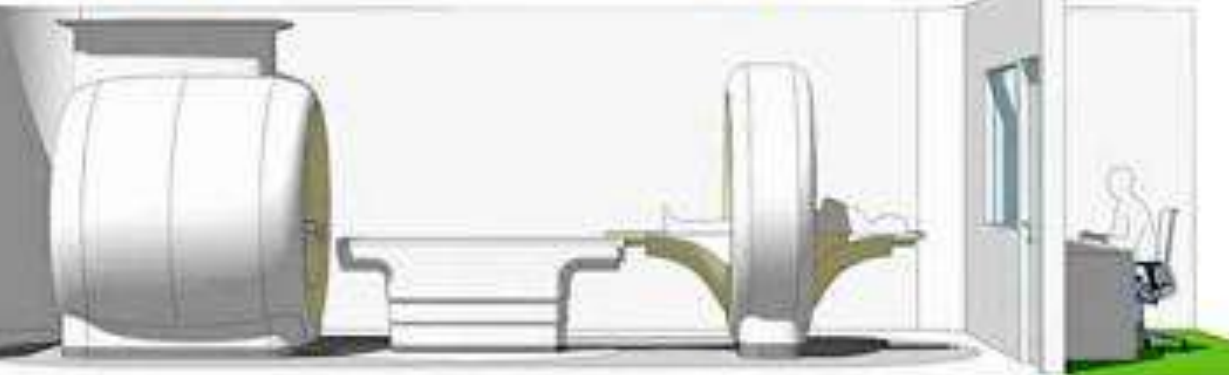
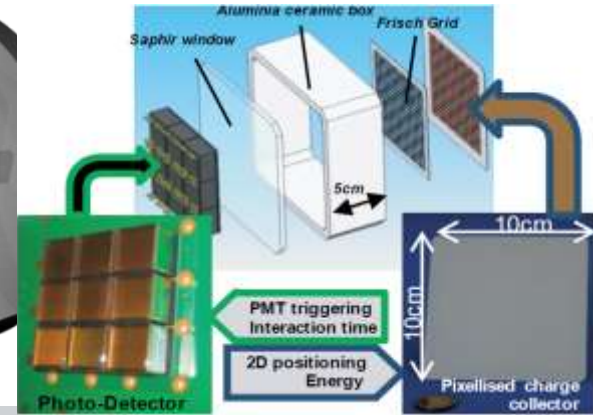




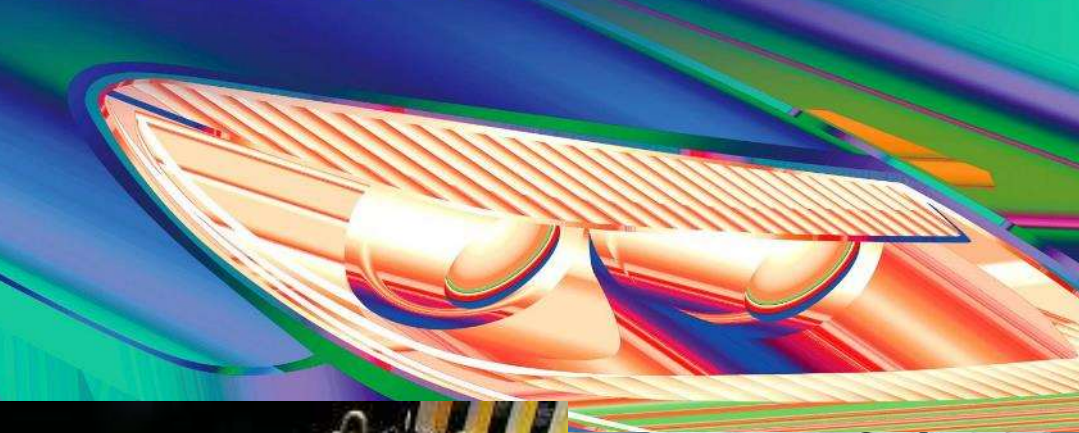
IRM



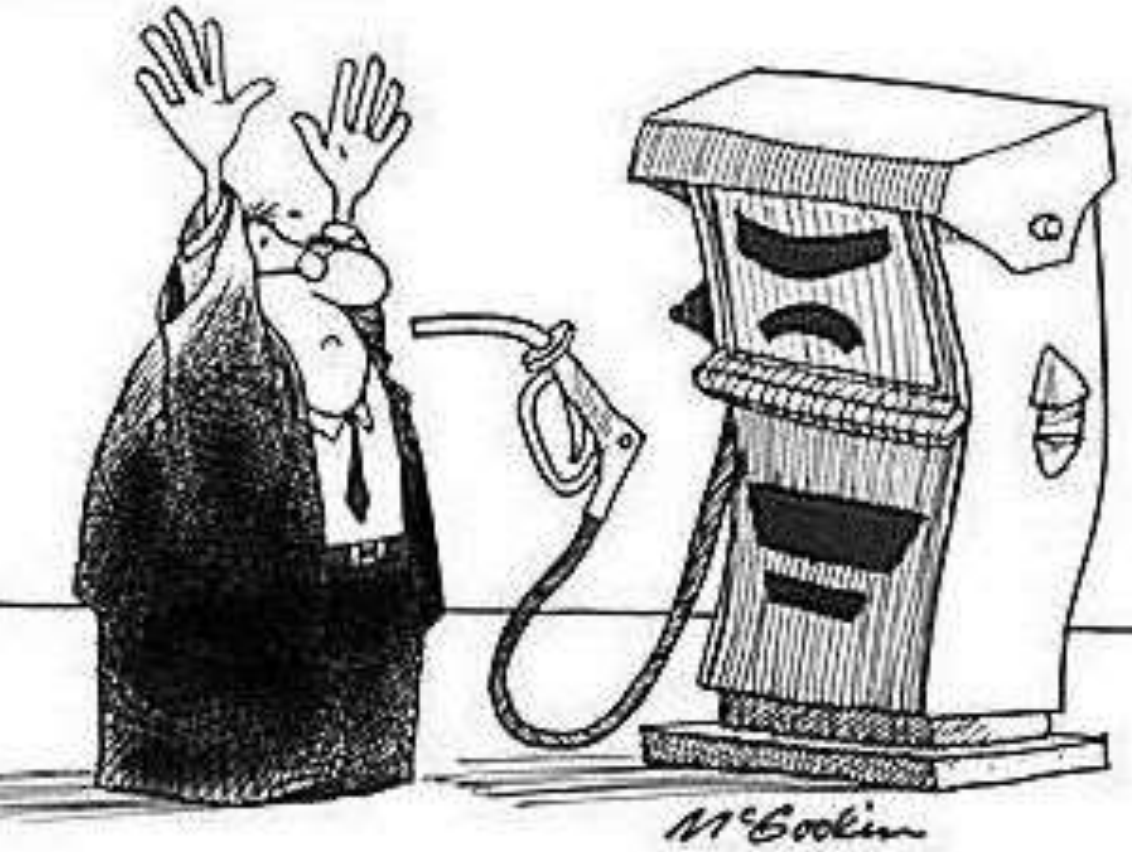
PET



85th International Motor Show & accessories 5–15 March 2015, Geneva



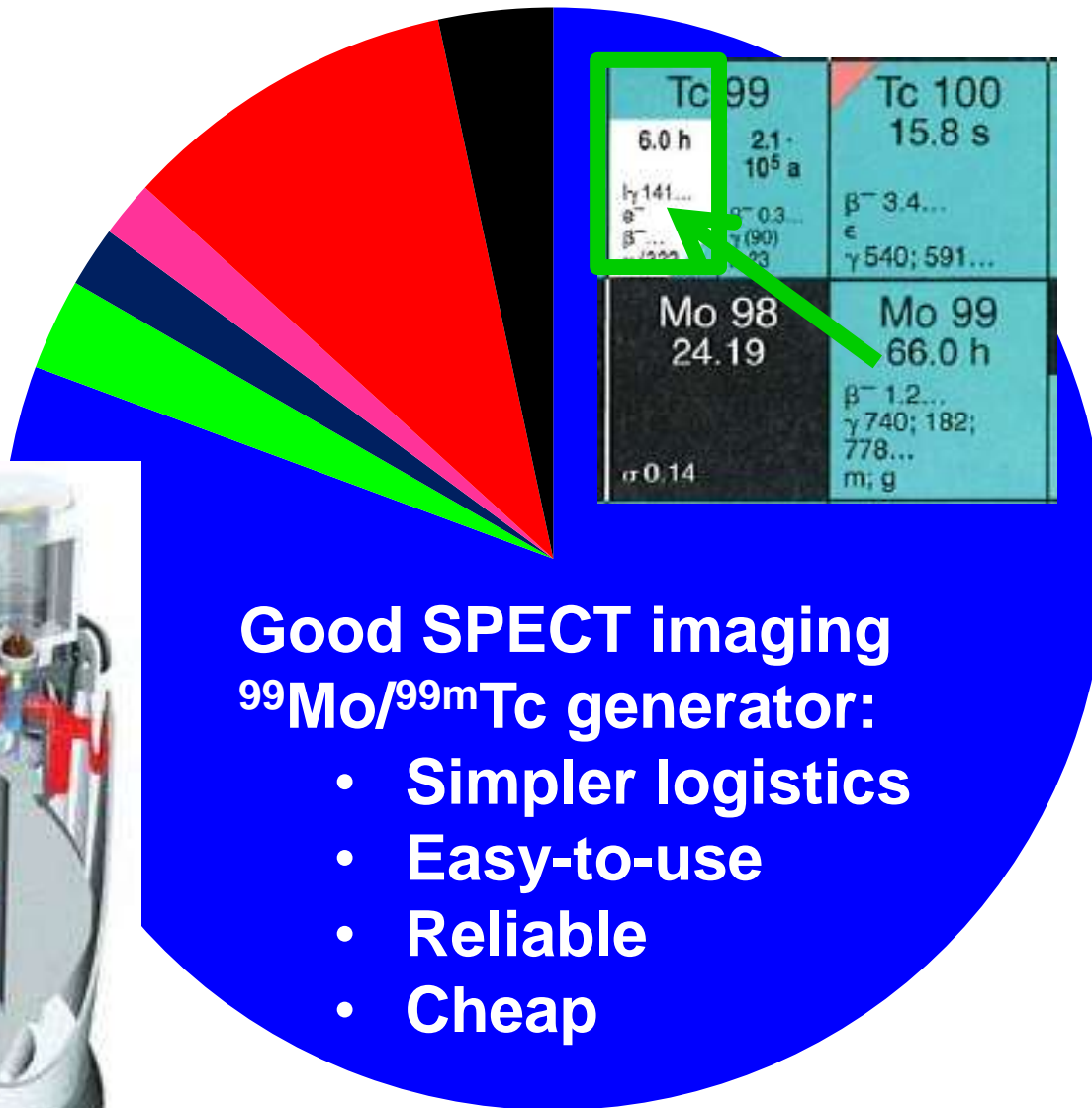
Don't forget the fuel!



Take home messages

1. Radioisotopes are the “fuel” for nuclear medicine

Cumulative use of diagnostic isotopes in Europe



■ 99mTc

■ 201Tl

■ 131I

■ 123I

■ PET

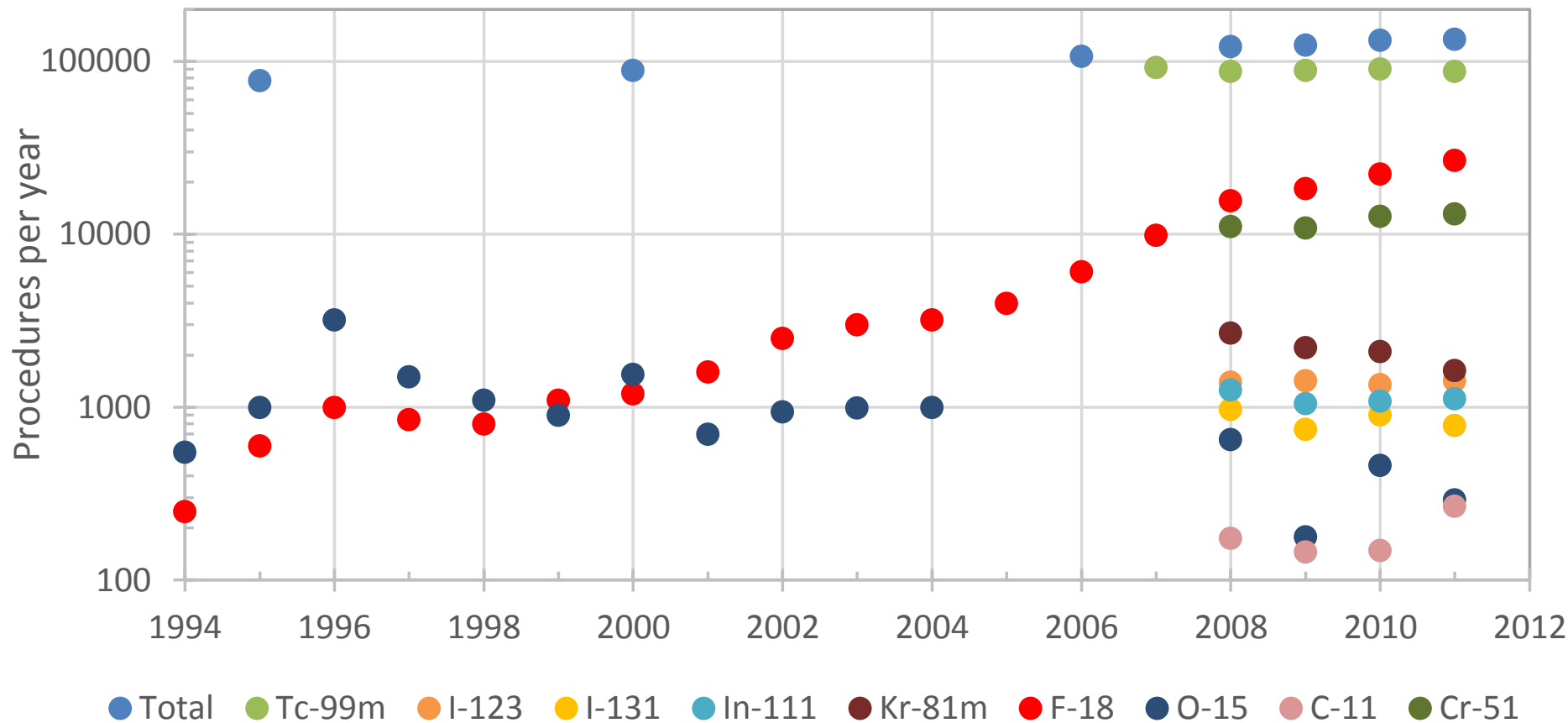
■ Other

Good SPECT imaging
⁹⁹Mo/^{99m}Tc generator:

- Simpler logistics
- Easy-to-use
- Reliable
- Cheap



Diagnostic procedures in Denmark



The State Institute of Radiation Protection Denmark.

2011 breakdown:

69% SPECT/scintigraphy - 21% PET - 10% Cr-51 EDTA (kidney)

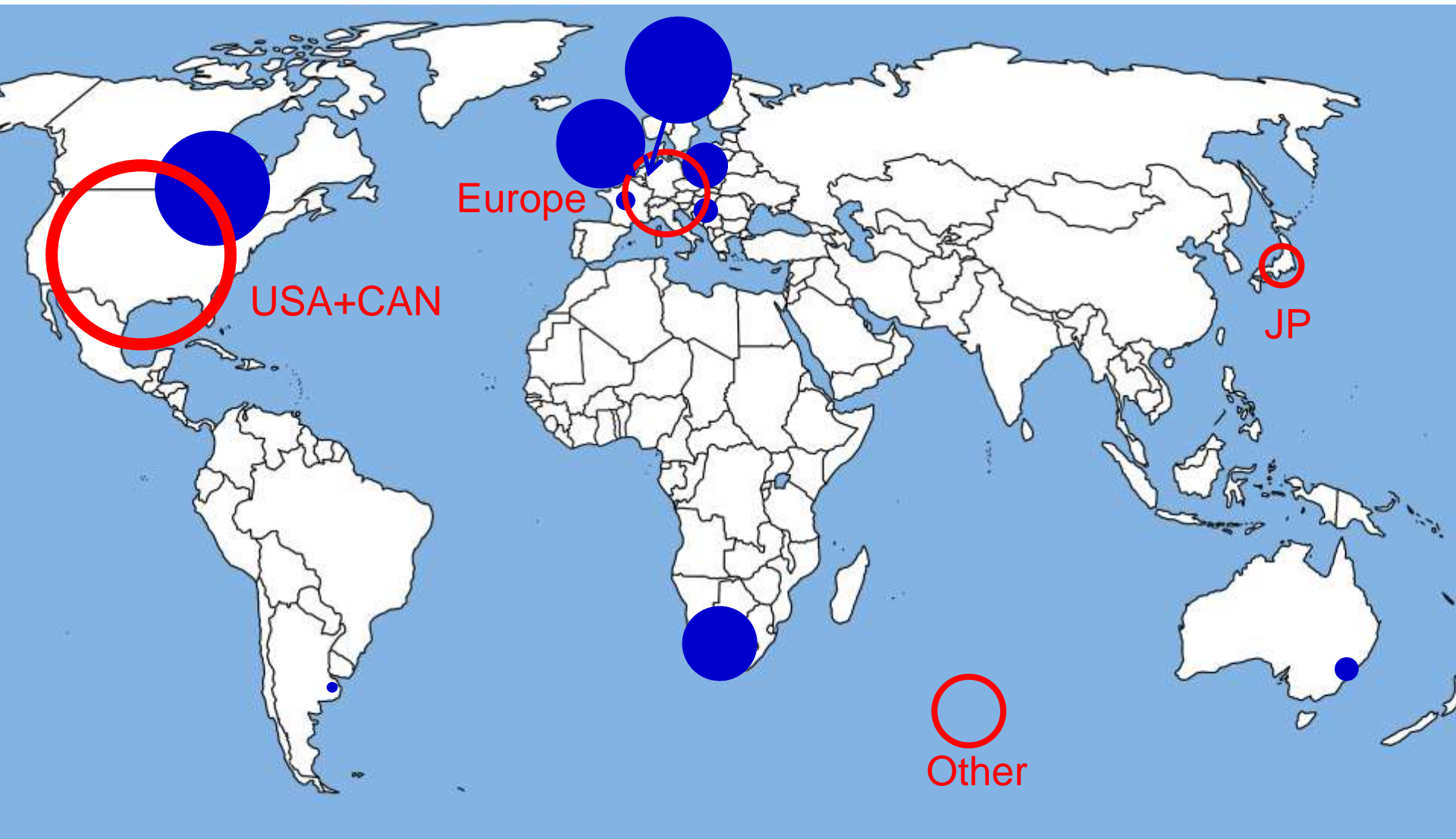
PET: 93% FDG - 4% other F-18 tracers - 3% other isotopes

Production: 76% reactor – 24% cyclotron

Take home messages

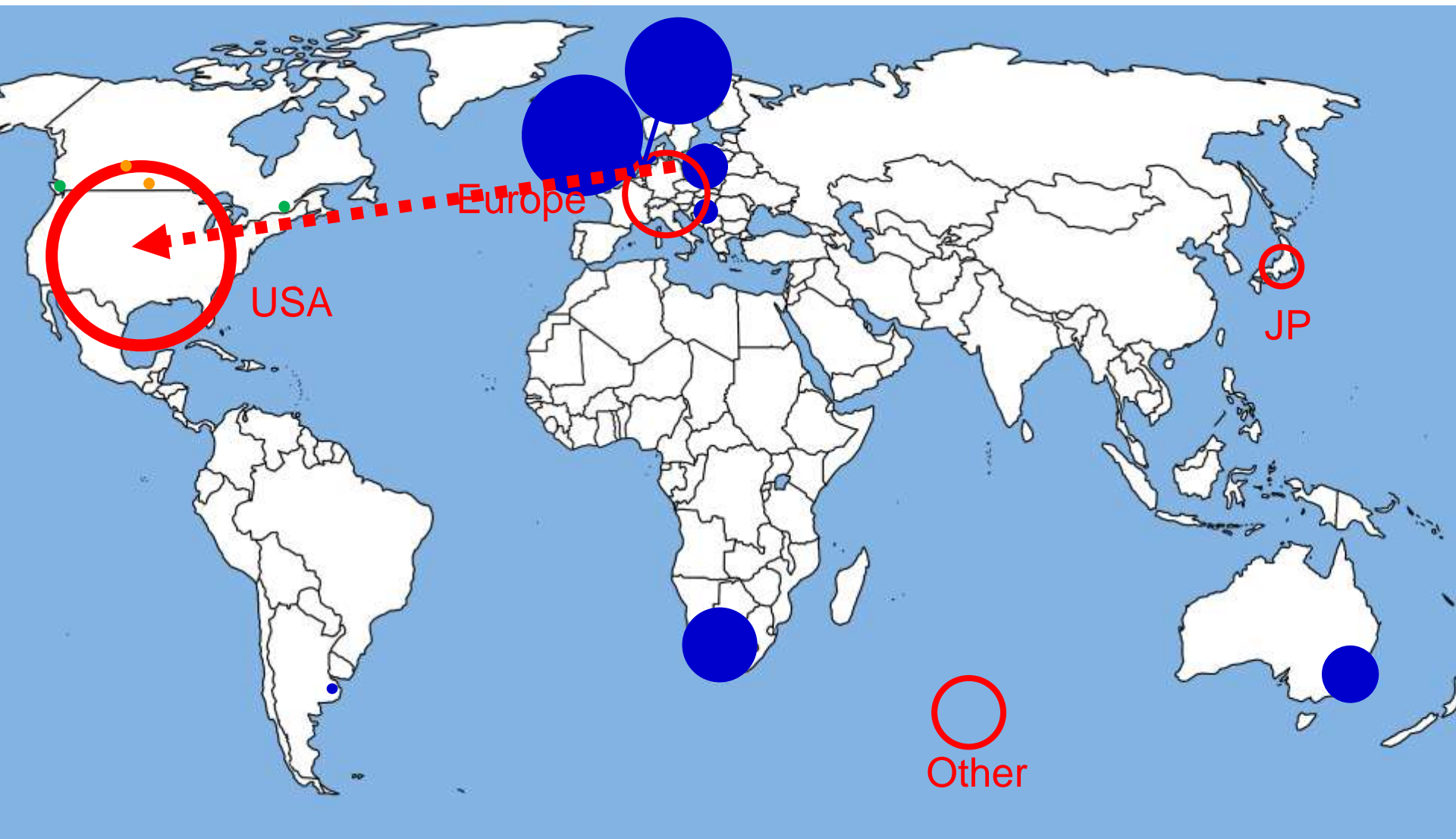
1. Radioisotopes are the “fuel” for nuclear medicine
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2014: ^{99}Mo production capacity and demand



Circle diameter proportional to annual reactor capacity (blue) and demand (red).

End 2016: ^{99}Mo production capacity and demand



Circle diameter proportional to annual production capacity (blue: reactor; green: cyclotron; orange: LINAC) and demand (red).

Isotope shortage means a healthcare crisis

European hospitals cope with Mo-99 supply crisis

EUROPE

MOLYBDENUM SUPPLY

L'inquiétante pénurie d'isotopes pour l'imagerie

Engpässe in der Tumormedizin

Medical isotope shortage reaches crisis level

Krebsärzten gehen die Diagnosemittel aus

Médecine nucléaire : il faut prolonger le réacteur Osiris

We Need to Expand Medical Isotope Production!

ANALYSIS | The made-in-Canada isotope shortage facing medical scans

Desperately Seeking Moly

Isotopes médicaux - Crise mondiale à l'horizon

Aucune solution n'existe pour résoudre le problème d'approvisionnement

Isotope shortage to get worse with closing of more reactors

L'OCDE s'inquiète des risques de pénurie d'isotopes médicaux

Mangel an medizinisch verwendbaren Isotopen

Mo-99 crisis

Szintigraphien fallen aus, für Februar droht der Notstand

Niowave broke ground last month on a \$79 million isotope production facility that looks out on the runways of Capital Region International. The company expects to move in by the end of the year, to make its first medical isotopes early next year, to be producing them commercially by early 2016. It expects to add 90 jobs to its 70-member staff in the process.

UBC SCIENTISTS HELP AVERT A NUCLEAR MEDICINE MELTDOWN

January 8, 2015

Moly 99 reactor using Sandia design could lead to U.S. supply of isotope to track disease

Commodities | Sat Feb 7, 2015 4:11am IST

Canada seeks to avoid medical isotope shortage, extends nuclear reactor

Isotope breakthrough may stave off shortage concerns

MARCH 12, 2015 BY PAUL MAYNE



Coquí Pharma completes design of medical isotope facility

Lab confirms new commercial method for producing medical isotope (Argonne)

June 15, 2015

09 April 2015

NorthStar Medical Radioisotopes ready to begin Mo-99 production

Michael Walter

Aug 31, 2015

It Takes Two: GE Healthcare and SHINE team up to solve longstanding radiopharmaceutical supply concerns in medical imaging

Successful generation of Tc-99m is a supply chain advancement that can help ensure patient access to critical medical imaging scans.

CHALFONT ST. GILES, UK - 9 November 2015 - Technetium-99m (Tc-99m) is used in more than 40





A STEVEN SPIELBERG FILM

tom hanks
saving isotope Tc-99m

edward burns matt damon tom sizemore

the mission is a nuclide.

DREAMWORKS PICTURES AND PARAMOUNT PICTURES PRESENT AN AMBLIN ENTERTAINMENT PRODUCTION IN ASSOCIATION WITH MUTUAL FILM COMPANY TOM HANKS
"SAVING PRIVATE RYAN" EDWARD BURNS MATT DAMON TOM SIZEMORE COSTUME DESIGNER BONNIE CURTIS AND ALLISON LYON SEGAN MUSIC BY JOHN WILLIAMS EDITOR JOANNA JOHNSTON
EXECUTIVE PRODUCERS MICHAEL KAHN AND TOM SANDERS DIRECTOR OF PHOTOGRAPHY JANUSZ KAMINSKI A.S.C. PRODUCED BY STEVEN SPIELBERG & IAN BRYCE AND MARK GORDON & GARY LEVINSON
SCREENPLAY BY ROBERT RODAT **July 24** DIRECTED BY STEVEN SPIELBERG
DREAMWORKS PICTURES

Take home messages

1. Radioisotopes are the “fuel” for nuclear medicine
2. $^{99\text{m}}\text{Tc}$ and ^{18}F are the main fuel, alternative fuels catching up
3. $^{99\text{m}}\text{Tc}$ is an industrial/economic challenge, not a scientific one

THE 11TH FILM BY QUENTIN TARANTINO

KILL TC



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4. Opportunity to switch to alternative fuels (PET)

Facilities producing ^{82}Sr

LANL, USA – 100 MeV, 200 μA

BNL, USA – 200 MeV, 100 μA

INR, Russia – 160 MeV, 120 μA

TRIUMF, Canada – 110 MeV, 70 μA

iThemba, South Africa – 66 MeV, 250 μA

ARRONAX, France – 70 MeV, < 750 μA

SPES, Italy – 70 MeV, < 1000 μA

Zevacor, USA – 70 MeV, < 750 μA



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The rising star for therapy



Production of non-carrier-added ^{177}Lu

1. Enrichment of stable isotope ($> 99.8\%$ ^{176}Yb)
2. Irradiation in high flux reactor
3. Radiochemical separation of n.c.a. ^{177}Lu

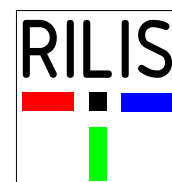
Lu 177 6.71 d β^- 0.5... γ 208; 113... g σ 1000	
Yb 176 12.76 if 3.1 IFA, α $< 1\text{E-}6$	Yb 177 1.9 h β^- 1.4... γ 150; 1080; 122; 1241... g



Resonant Laser Ionization for Isotope Separation



ISOLDE



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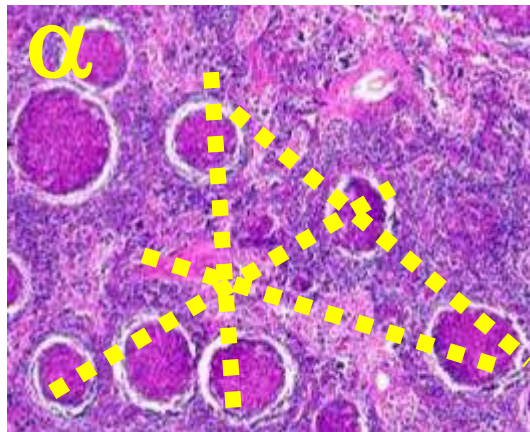
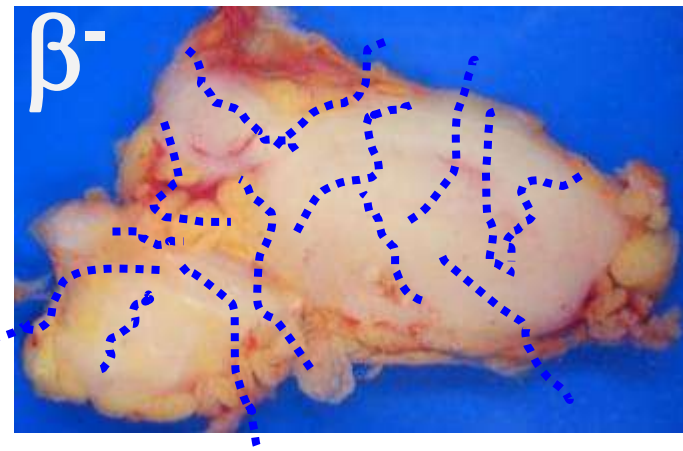
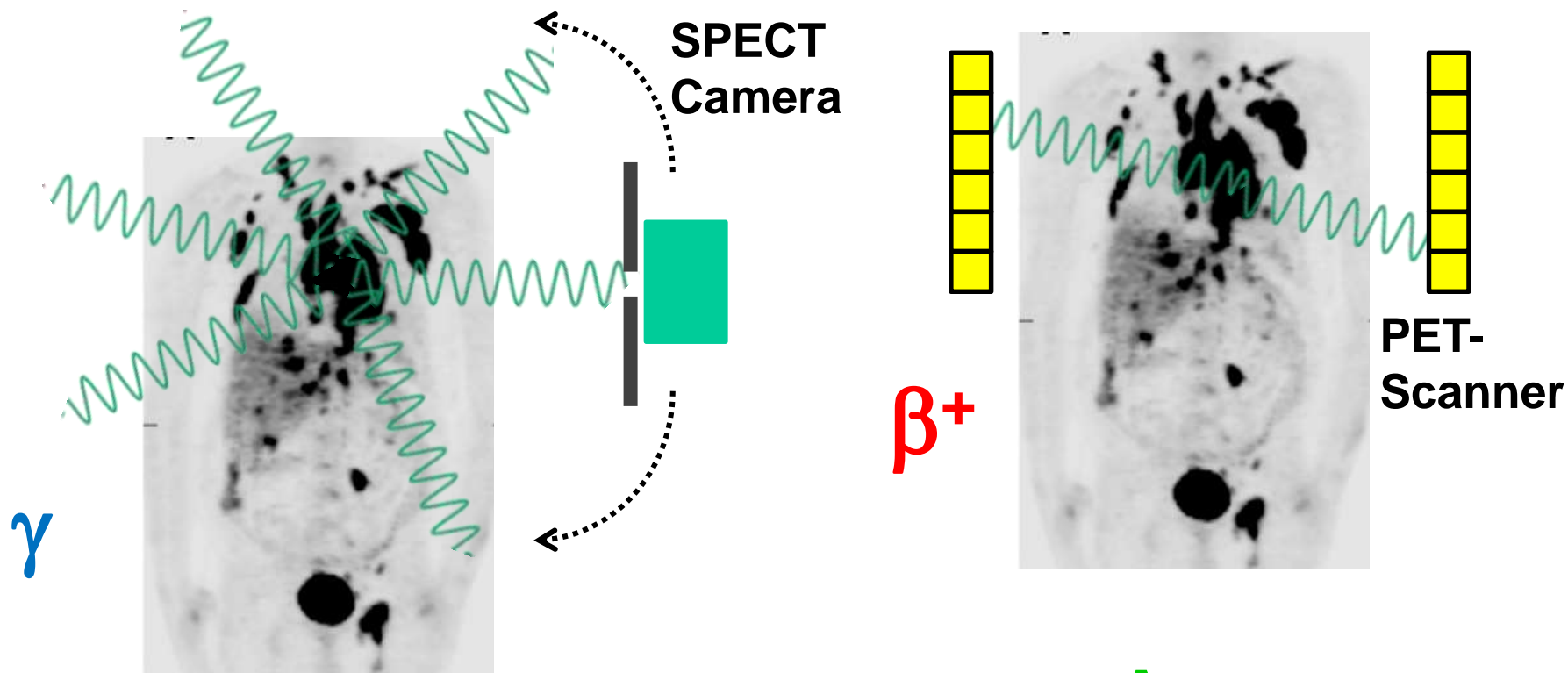
H																	He	
Li	Be											B	C	N	O	F	Ne	
Na	Mg											Al	Si	P	S	Cl	Ar	
K	Ca	Sc	Ti	V	Cr	Mn	Fe	Co	Ni	Cu	Zn	Ga	Ge	As	Se	Br	Kr	
Rb	Sr	Y	Zr	Nb	Mo	Tc	Ru	Rh	Pd	Ag	Cd	In	Sn	Sb	Te	I	Xe	
Cs	Ba			Hf	Ta	W	Re	Os	Ir	Pt	Au	Hg	Tl	Pb	Bi	Po	At	Rn
Fr	Ra			Rf	Db	Sg	Bh	Hs	Mt	Ds	Rg	Cn	Uut	Uuq	Uup	Uuh	Uus	Uuo
		La	Ce	Pr	Nd	Pm	Sm	Eu	Gd	Tb	Dy	Ho	Er	Tm	Yb	Lu		
		Ac	Th	Pa	U	Np	Pu	Am	Cm	Bk	Cf	Es	Fm	Md	No	Lr		

Feasible Dye schemes tested Ti:Sa schemes tested Dye and Ti:Sa schemes tested

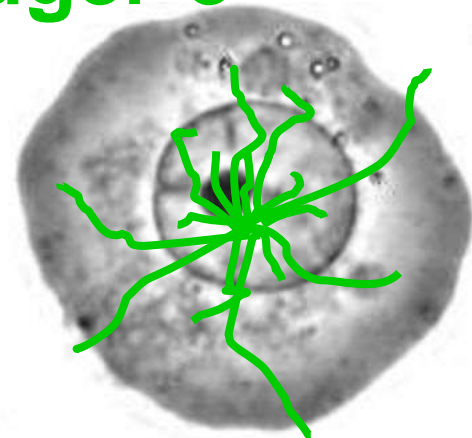
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The Nuclear Medicine Alphabet

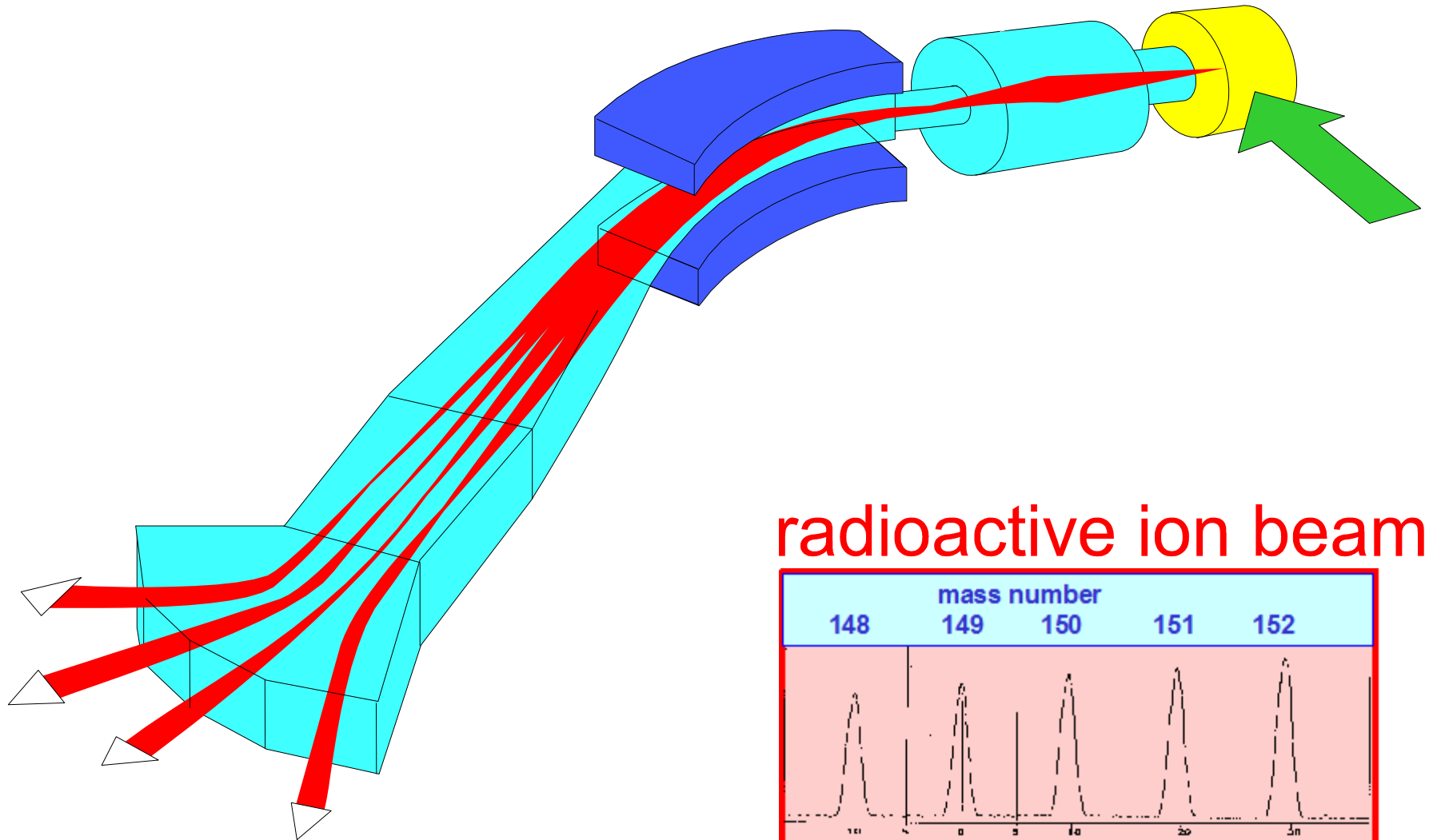


Auger- e^-

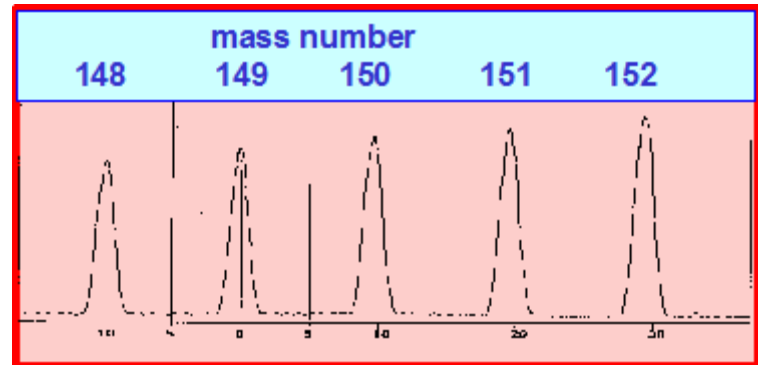


Radio-nuclide	Half-life	Daughters	Half-life	Cumulative α /decay	E_{α} mean (MeV)	Range (μm)
Tb-149	4.1 h			0.17	3.97	25
<i>Pb-212</i>	<i>10.6 h</i>	Bi-212 Po-212	1.01 h 0.3 μs	1	7.74	65
Bi-212	1.01 h	Po-212	0.3 μs	1	7.74	65
<i>Bi-213</i>	<i>0.76 h</i>	Po-213	4 μs	1	8.34	75
At-211	7.2 h	Po-211	0.5 s	1	6.78	55
Ra-223	11.4 d	Rn-219 Po-215 <i>Pb-211</i> Bi-211	4 s 1.8 ms <i>0.6 h</i> 130 s	4	6.59	>50
Ra-224	3.66 d	Rn-220 Po-216 <i>Pb-212</i> Bi-212	56 s 0.15 s <i>10.6 h</i> 1.01 h	4	6.62	>50
Ac-225	10.0 d	Fr-221 At-217 <i>Bi-213</i> Po-213	294 s 32 ms <i>0.76 h</i> 4 μs	4	6.88	>50

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



radioactive ion beams

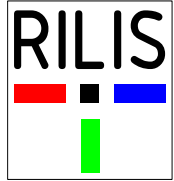


Challenges of Astatine

ASTATINE 70mg

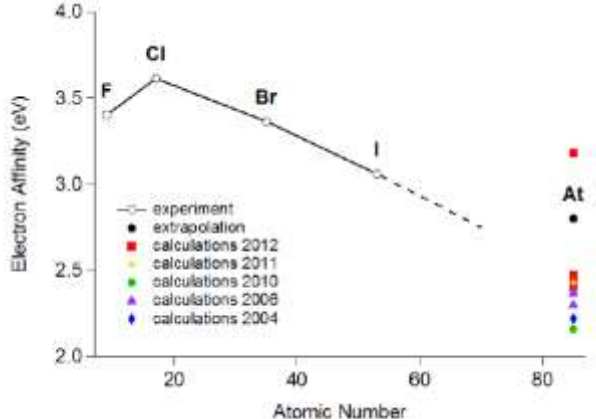


1 x per Planet (Apply to crust) ²¹⁸At

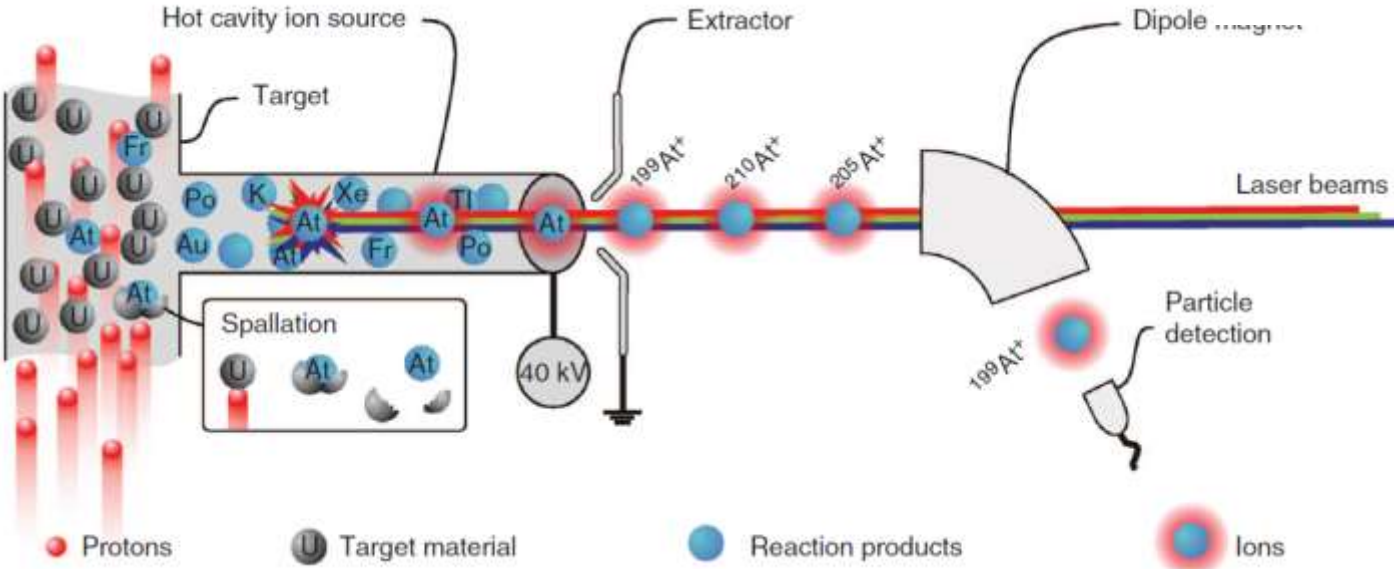


Measurement of the first ionization potential of astatine by laser ionization spectroscopy

S. Rothe^{1,2}, A.N. Andreyev^{3,4,5,6}, S. Antalic⁷, A. Borschevsky^{8,9}, L. Capponi^{4,5}, T.E. Cocolios¹, H. De Witte¹⁰,



Dag Hanstorp et al.
INTC-I-148

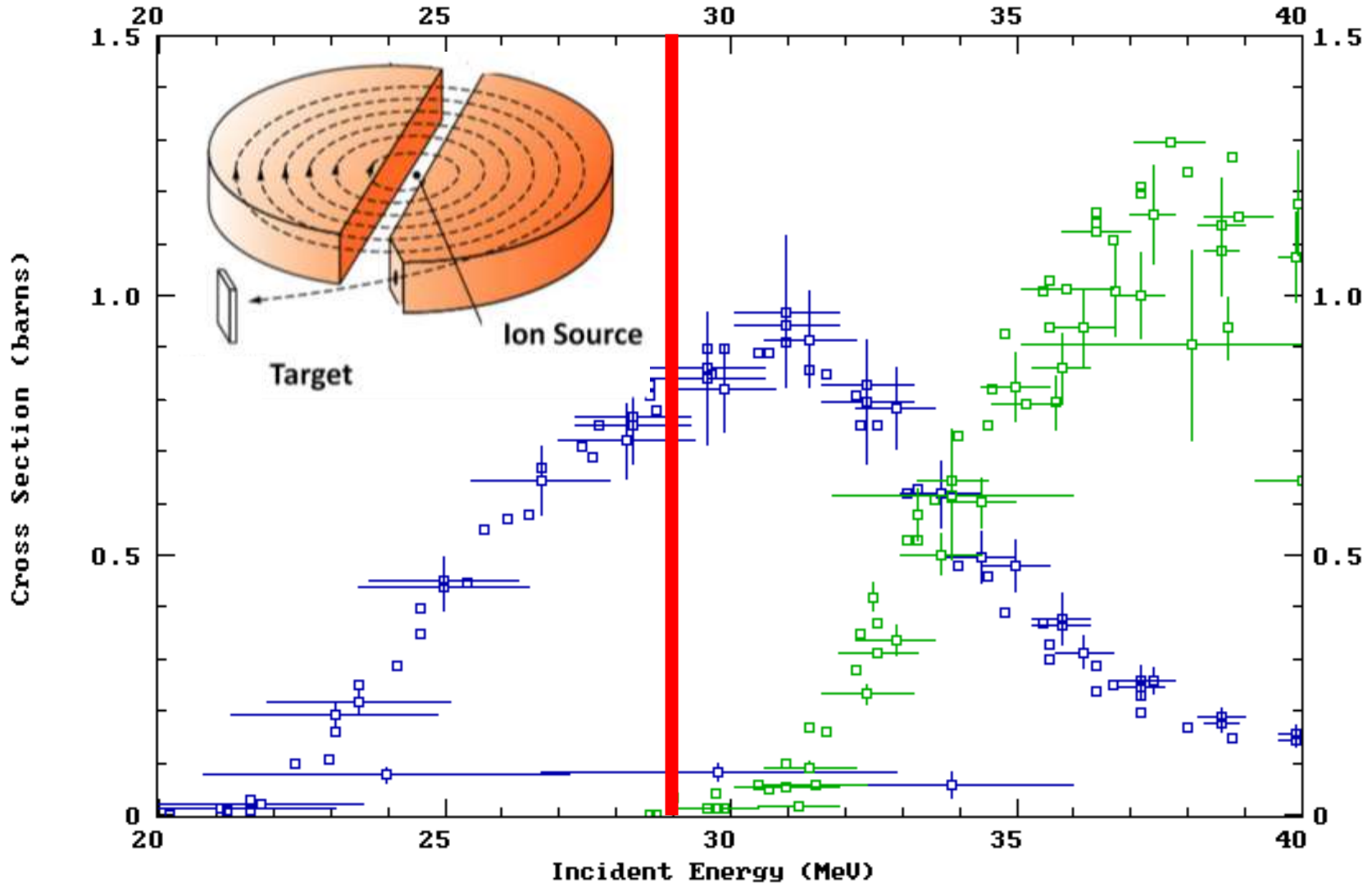


IP(At) = 9.317510(8) eV

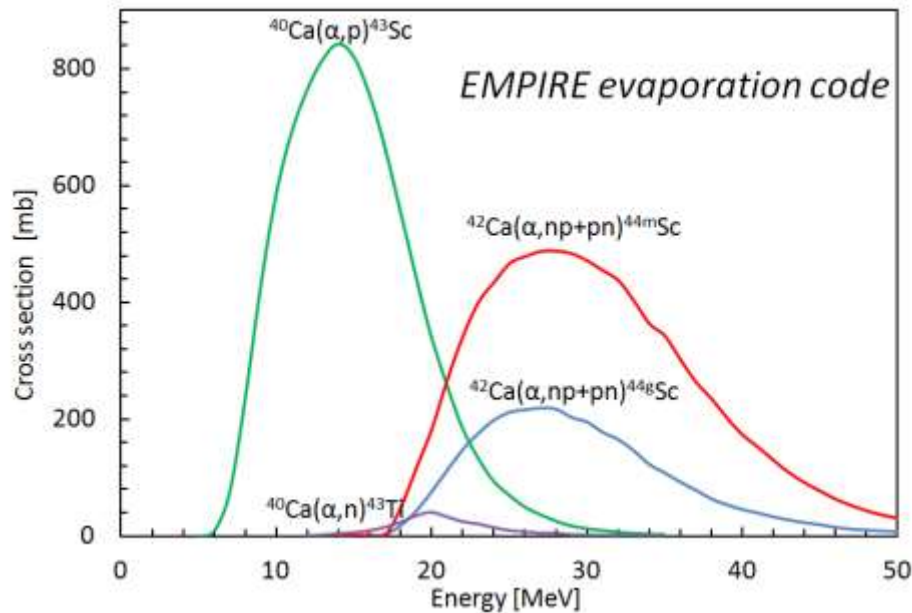
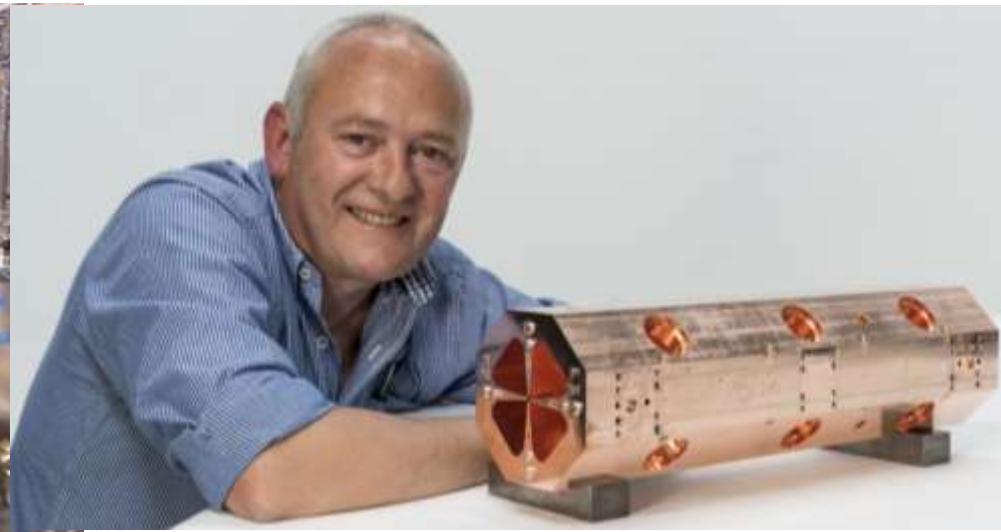
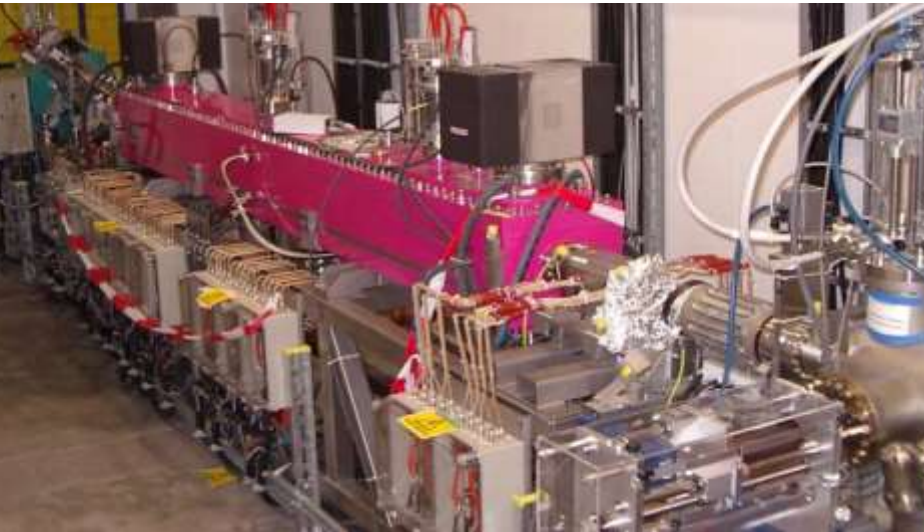
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3. ^{99m}Tc is an industrial/economic challenge, not a scientific one
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5. Compact (<10 MeV) or high energy (>100 MeV) accelerators
6. Efficient enrichment of rare isotopes
7. Provide high quality alpha emitters (^{149}Tb , ^{211}At , ^{225}Ac):
mass sep. at ISOLDE/MEDICIS & radiochemically purified

^{211}At production

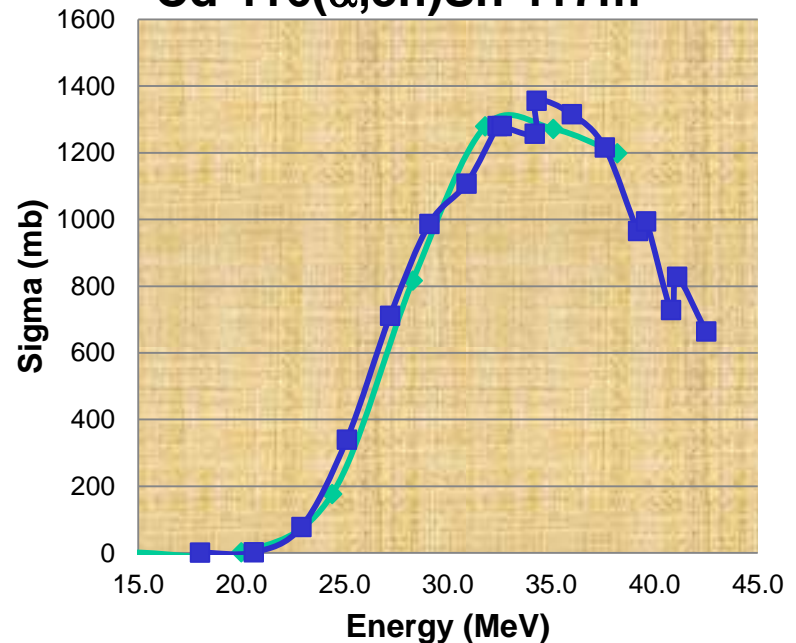


7.2-11 MeV/u light ion LINAC A/q=2



J. Jastrzębski. ICTR-PHE2016

Cd-116($\alpha, 3n$)Sn-117m



N. Stevenson. ICTR-PHE2016

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mass sep. at ISOLDE/MEDICIS & radiochemically purified
8. Great potential for linear accelerator $A/q=2$, 7.2-11 MeV/nucl.


Which medical isotopes will we need in 2030 ?

MEMORANDUM

DATE: December 4, 1958

Today >30 million clinical applications per year !

TO: Addressees Below

FROM: Daniel M. Schaeffer, Head 
BNL Patent Office

SUBJECT: P-701 and P-702 - PREPARATION OF
CARRIER-FREE MOLYBDENUM AND OF
TECHNETIUM FROM FISSION PRODUCTS

The New York Patent Group has carefully studied the information available relative to the above-identified item. The AEC does not at present desire to prepare a patent application on this item for the following reason:

"The method of producing carrier-free molybdenum-99 from fission products is disclosed in U. S. Patent Application S.N. 732,108, Green, Powell, Samos & Tucker (BNL Pat No. 58-17). It is noted that molybdenum-99 may be separated from its radioactive daughter, technetium-99, by absorption of a solution of molybdenum-99 on alumina and subsequent elution of its daughter with .1 nitric acid. While this method is probably novel, it appears that the product will probably be used mostly for experimental purposes in the laboratory. On this basis, no further patent action is believed warranted."

believe that this attitude is significant. We are not aware of a potential market for technetium-99 great enough to encourage one to undertake the risk of patenting in hopes of successful and rewarding licensing. We would recommend against filing on the Tucker, Greene and Murrenhoff separation process."

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Radioisotope production is outside of ILL’s normal sphere of activity, but we have a moral imperative to do this work.

It’s a great example of how a publicly funded facility can have a totally unexpected and unpredictable payoff for society.

Prof. Andrew Harrison (former ILL Director)