

CERN-MEDICIS

An update

Thierry Stora, EN-STI



ENGINEERING
DEPARTMENT

ISOLDE

Medicis

The Scope

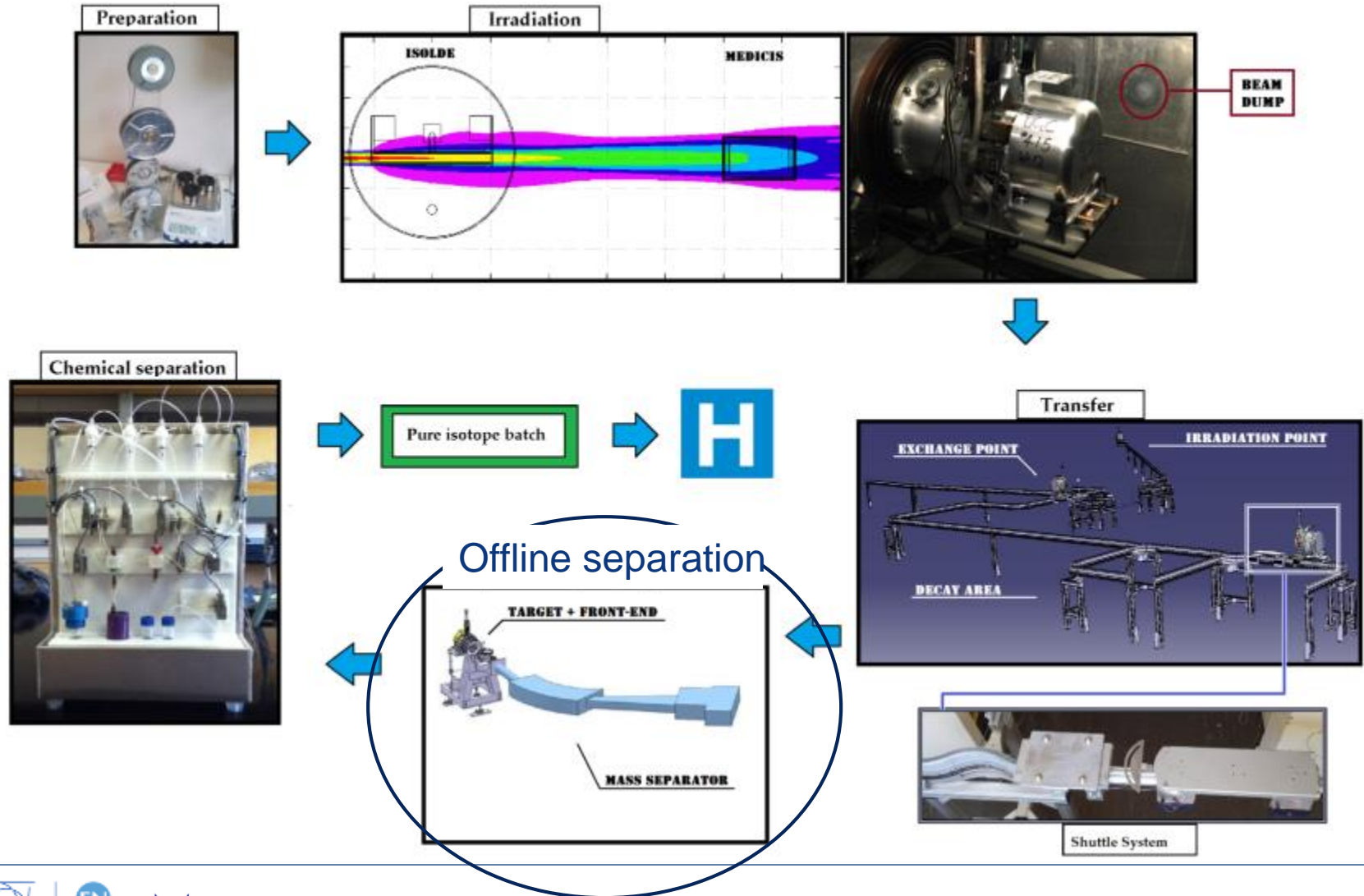
- Biomedicine
- Innovative protocols (Surgery/brachytherapy/combination)
- Innovative isotopes for imaging and treatment

Field of Application	Radiation	Chemical elements	Half lives
PET	β^+	Alkaline earth	10's min.
SPECT	γ	Halogen	Hours
TAT	α	Lanthanide	Days
Beta therapy	β^-	Transition metals	Months
Auger therapy	e^-	...	

- Studies on cells, animals (« preclinical »)
possibly extended to clinical phases
(needs upgrades, but this can be reached)

Why CERN-MEDICIS could be done *?

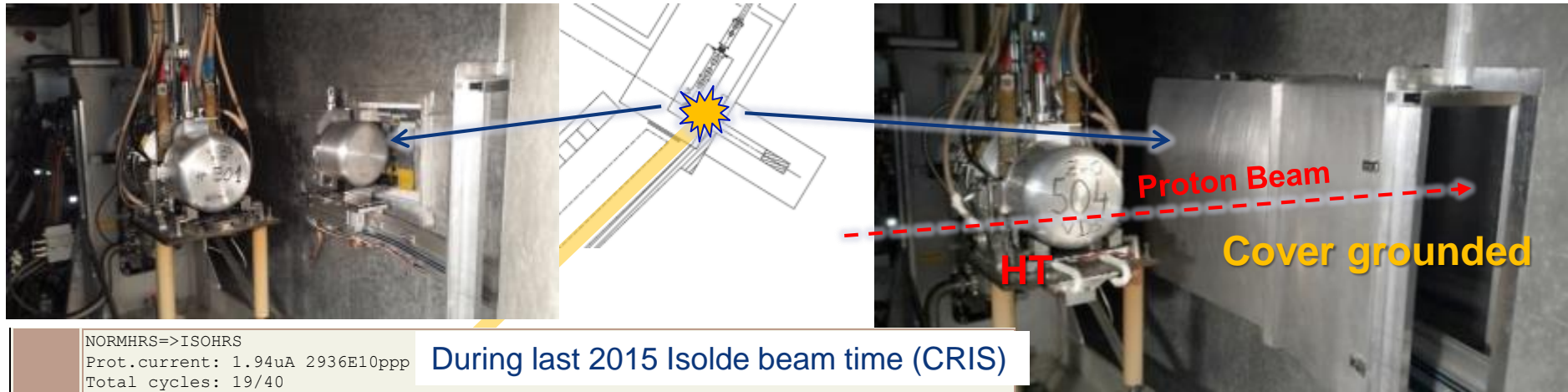
* Without an additional proton driver



Irradiation station commissioned with beam



M. Vagnoni (EN-STI fellow), et al.



NORMHRS=>ISOHRS
 Prot.current: 1.94uA 2936E10ppp
 Total cycles: 19/40
 Cycles number: 4 7 8 11 12 15 17 18 22 23 24 27 28 31 32 35 36 37 40

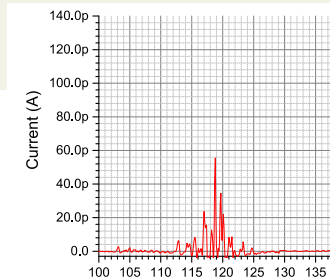
During last 2015 Isolde beam time (CRIS)

22:32

MEDICIS RCS : Target UC541 removed from the irradiation HRS dump position to its parking position on RCS.
 Integrated PoT 0.8e18.
 Will be removed with Kuka robots on Monday.
 /TS/MV



name: 20151114223532.png
 desc:



Tests with protons:
 Done successfully !



& RIB separated 1 week later on GPS

The building extension

J.L. Grenard, K. Kershaw et al.



Storage shelves
(V. Barozier & Y. Gavrikov)



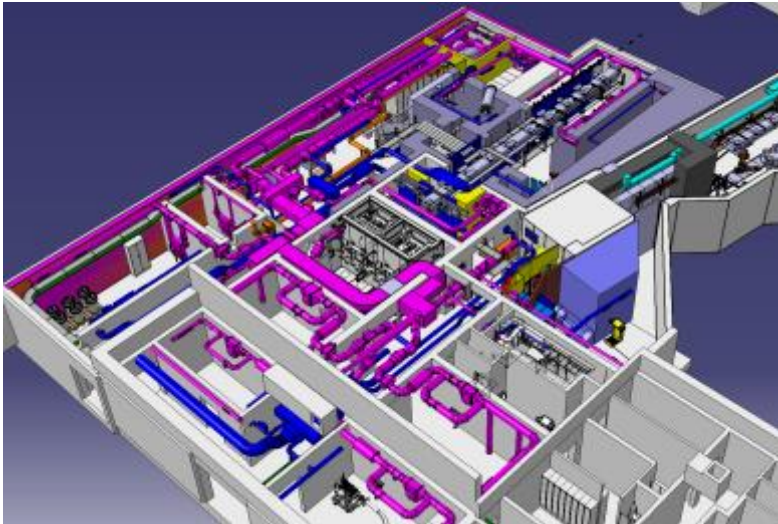
Robot hardware installed



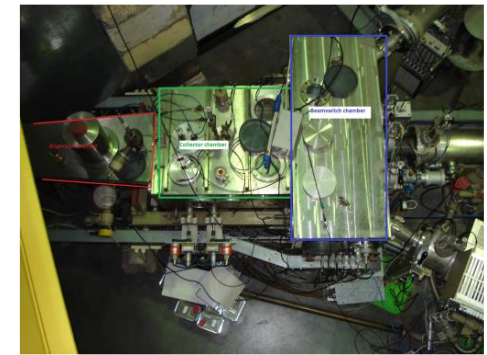
Last shielded door



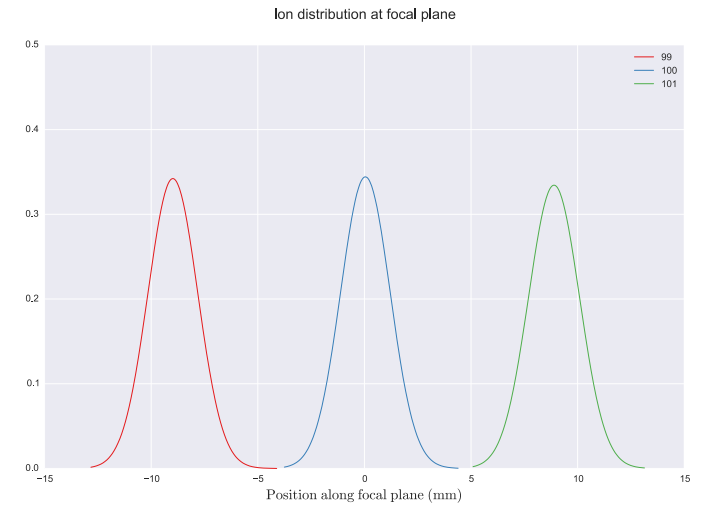
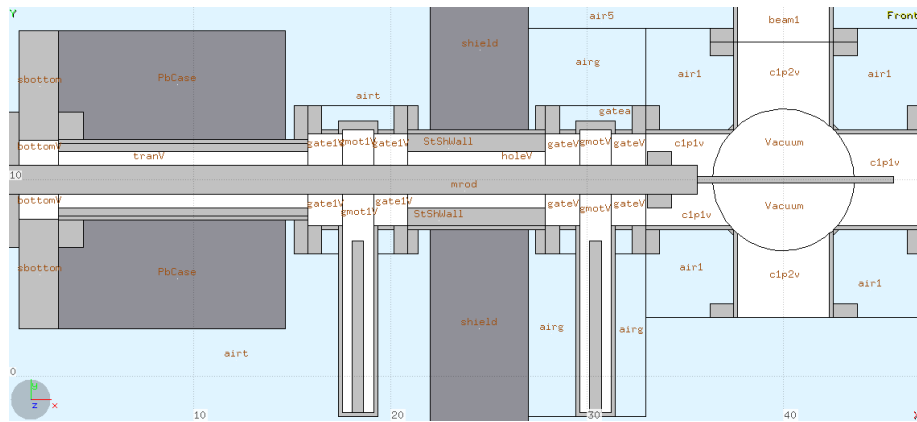
The nuclear ventilation



The Mass separator



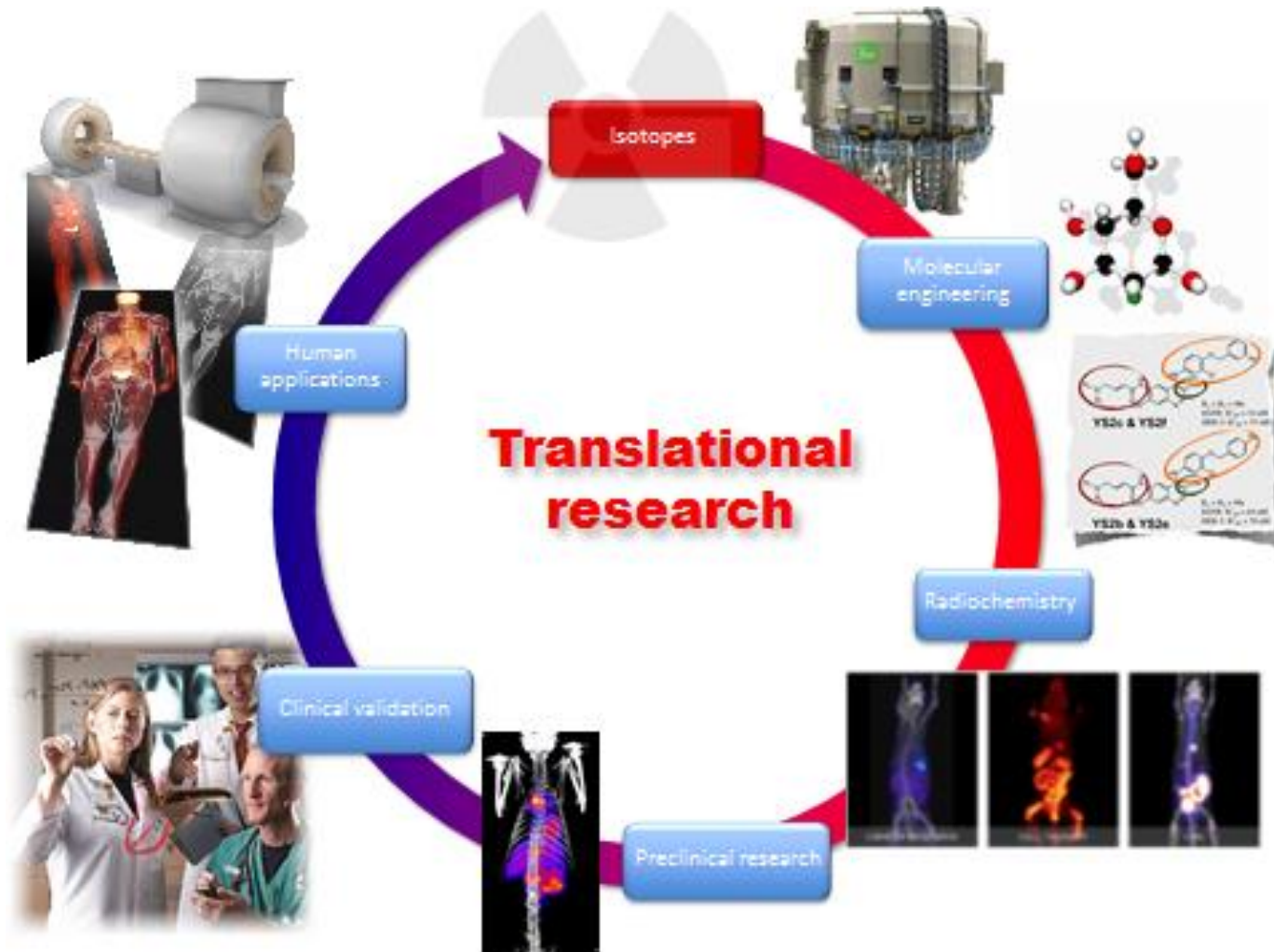
LISOL dipole ready for shipment
In Louvain La Neuve



Separator in CERN-MEDICIS
Configuration (Y. Martinez)

Conceptual design – collection chamber (A. Brown)

The scientific case



Courtesy prof. Ratib, in the context of the CERN-MEDICIS project

Collaboration with JRC-ITU

Intracavity injection +resection of Glioblastoma

Targeted alpha-radionuclide therapy of functionally critically located gliomas with ^{213}Bi -DOTA-[Thi⁸,Met(O₂)¹¹]-substance P: a pilot trial

D. Cordier • F. Forrer • F. Bruchertseifer •
A. Morgenstern • C. Apostolidis • S. Good •
J. Müller-Brand • H. Mäcke • J. C. Reubi • A. Merlo



JOINT RESEARCH CENTRE

The European Commission's in-house science service

European Commission > JRC Science Hub > News & events > JRC News > CERN and the JRC to scale up production of alpha-emitters against cancer

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23
SEP
2015

CERN and the JRC to scale up production of alpha-emitters against cancer

A novel, accelerator-driven method could produce nuclides for targeted alpha therapy of cancer in practically unlimited amounts, overcoming current obstacles for its wider use due to a limited production of alpha-emitters. The JRC and the Conseil Européen pour la Recherche Nucléaire (CERN) have embarked to explore the potential of the jointly proposed method.

The method for production of



Current radiotherapy against cancer mostly uses beta-emitters as medical isotopes

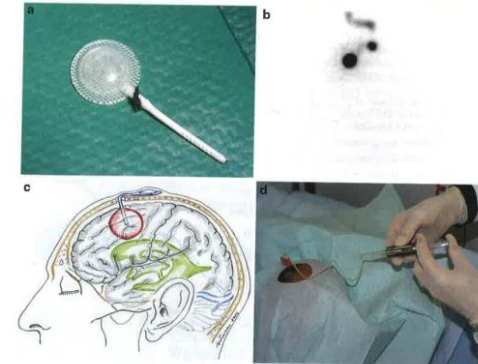
© Alex Tihonov, Fotolia.com

Related Topics

Medical applications of radionuclides and targeted alpha therapy
Public health

JRC Institutes

ITU



12th SWISS EXPERIMENTAL SURGERY SYMPOSIUM

New Radio Isotopes for Diagnosis & Treatment

In Pre-clinical and Clinical Research

Organized by the HUG and the University of Geneva

With the participation of:

L. Buhler, Ph. Morel, B.H. Walpoth

Co-organized with CERN, CHUV, EPFL, ISREC

With the participation of:

D. Hanahan, J. Prior, O. Ratib, T. Stora

Friday, 15 January 2016

08h30 – 17h00

EUROPEAN ORGANIZATION FOR NUCLEAR RESEARCH

Letter of Intent to the ISOLDE and Neutron Time-of-Flight Committee

Radium and Francium beam tests to produce $^{225}\text{Ac}/^{213}\text{Bi}$ generators at CERN-MEDICIS

14 October 2015

F. Bruchertseifer¹, A. Morgenstern¹, Y. Martinez^{2,3}, T. Cocolios², T. Stora³ and the CERN-MEDICIS collaboration

¹ JRC-ITU, Karlsruhe, Germany

² KULeuven, Leuven, Belgium

³ CERN-MEDICIS project, CERN, Switzerland

Outreach



1st yearly Grace-MEDICIS collaboration/public lecture took place on 15th October 2014

The 2nd one is in preparation:
Prof. W. Weber, Memorial Sloan Kettering Cancer Center New York



Prof Doug Hanahan

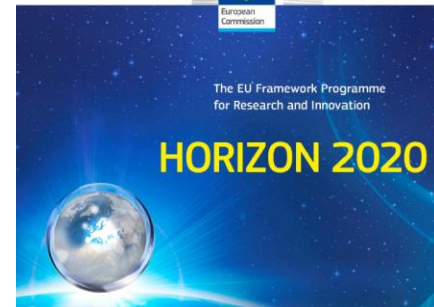
Director ISREC Lausanne

AACR's Lifetime Achievement Award

Tentative planning

Phase	Action	Date
PHASE I	Commissioning: without beam (*)	2016
PHASE II	Commissioning with beam and light targets to gain operational experience	2017
PHASE II B	Isotope production with light targets	Mid 2017
PHASE III	Extending to heavy targets up to Tantalum	End 2017
PHASE IV	Collection of short lived alpha emitters (e.g. ^{149}Tb)	2018
PHASE IV B	Operation with lasers	2018
PHASE V	Operation with uranium targets/possible proton beam upgrade	2019

* Preferable but may be hard to achieve

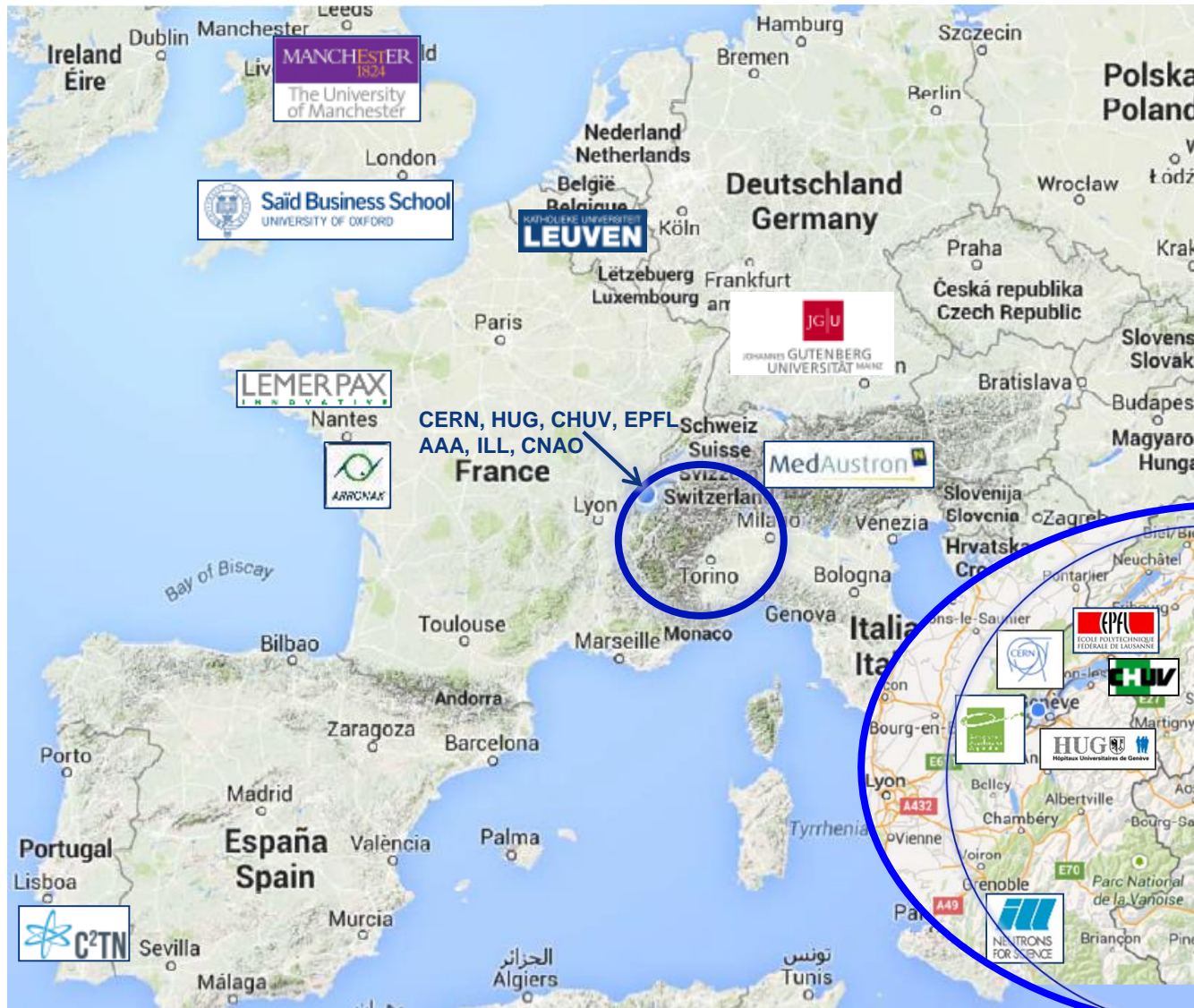


MEDICIS-PROMED

« MEDICIS-Produced radioisotope beams for medicine »

www.cern.ch/medicis-promed

The intersectorial distributed network



The intersectorial “regional” network

Overview of the Research Network

>7/15 young researchers have been hired : recruitment → Dec 2015

MEDICIS_PROMED training network

"Timely

Coordination Dr. T. Stora, CERN Medical coordination : PhD, MD J. Prior, CHUV

Innovations" WP3 : theranostic pharmaceuticals/surgery for new ovarian cancer personalized treatment

Terbium isotope theranostic pairs
Biological targets for ovarian cancers

AAA (FR) lead- radiopharmaceuticals - ESR6
IST (PT)/dna targetting - ESR8
CERN MEDICIS (EU)/molecular break-up - ESR1
HUG (CH)/surgery - ESRCH3
CHUV(CH)/preclinical tests - ESRCH2

"Timely
Innovations"

WP 1 : mass separation of new medical isotopes

WP 2 : Pet aided 11C hadrontherapy

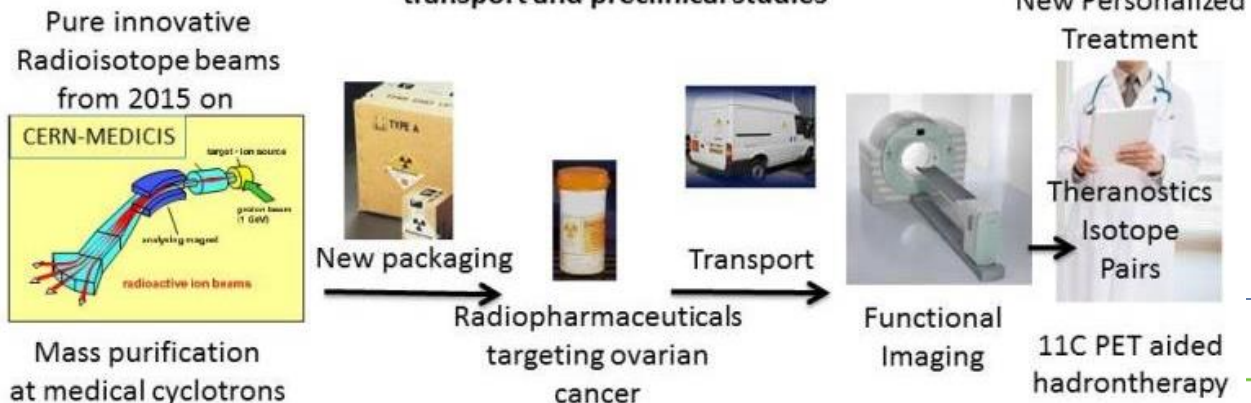
"Timely
Innovations"

JOGU (DE) lead - laser purification - ESR5
UNI MANCHESTER (UK)/adv material- ESR4
CERN MEDICIS (EU)/ production safety - ESR2
Lemer-Pax (FR) /transport - ESR10
IST (PT)/nanofibers - ESR7

CNAO (IT) lead - 11C hadrontherapy - ESR9
KUL (BE) - mass sep 11C - ESR11
CERN MEDICIS (EU) - 11C acceler. - ESR3
HUG (CH) - imaging tests - ESRCH1
EPFL (CH) - biochemical synthesis - ESRCH4
Medauston (AT) - hadrontherapy

Medauston
animal models

MEDICIS-PROMED: Innovative treatments based on radioactive ion beam production, transport and preclinical studies



Training : Events and models

Kick-off week – CERN (EU) 1st half feb 2016, and ICTR-PHE 2016

General training 1 – Manchester (UK)

Workshop on functional multimodal SPECT/PET imaging – Lausanne/Geneva (CH)

Specialized training 2 – Leuven (BE)

Summer school 1 at CNAO – Pavia (IT).

Summer school 2 at C2TN-IST – Lisbon (PT)

K. Novoselov, Graphene Institute – Physics Nobel Prize 2010 – Scientific Innovation and Advanced Materials

U. Koester, ILL- chairman of the NuPECC working group for *Nuclear Physics for Medicine-Radioisotope production*– Production of medical radioisotopes

P. Van Duppen, KUL – Adv ERC – Radioactive Ion Beams and Lasers

S. Buono, AAA – Radiopharmaceuticals marketing and Entrepreneurship

G. Coukos, CHUV – Adv. ERC – Immunotherapy and cancer treatment

P. Lecoq, CERN – Adv ERC – Detectors and Medical imaging

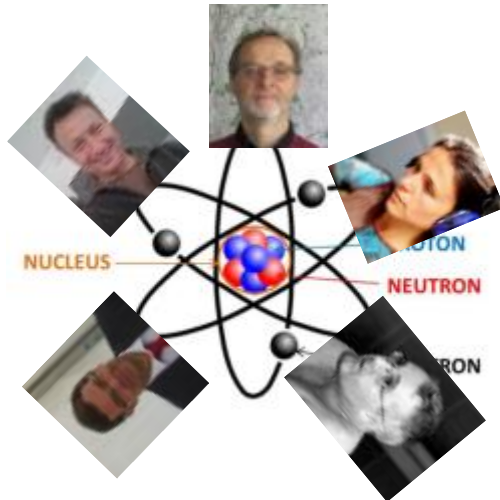
K. Noda-san – NIRS – PET-aided hadron therapy with carbon ions

Program cohesion : Oxford University Said Business School (ECTS, PhD)





ENGINEERING DEPARTMENT



And many others

Thank you, questions , comments ?

Some yield estimates

Medical application	Isotope half-life	Parent isotope beam	Target - Ion source	ISOLDE [†]		RIB ϵ_{ext}^{**} (%)	CERN-MEDICIS [†]		CERN-MEDICIS 2GeV 6μA		Comments
				In-target			In-target Activity ^{EOB} (Bq)	Extracted Activity ^{EOB} (Bq)	Possible gain ϵ_{ext} (%)	In-target Activity ^{EOB} /Extracted Activity ^{EOB} (Bq)	
				Production rate (pps)	Activity ^{EOB} (Bq)						
3-therapy/ CT/dosimetry	²¹³ Bi 45.6m	²²⁵ Ac	UCX-Re	1.5E9*	7.2E8	²²¹ Fr 10	2.8E8	2.8E7	50	8.4E8 4.2E8	Only mass separation
β therapy	²¹² Bi 60.6m	²²⁴ Ac	UCX-Re	1.5E9*	1.4E9	²²⁰ Fr 10	1.7E9	1.7E8	50	5.1E9 2.5E9	Only mass separation
β therapy	¹⁷⁷ Lu 6.7d	¹⁷⁷ Lu RILIS/VD	Ta-Re/ Re-VD5	3.3E9	7.4E8	¹⁷⁷ Lu 1	6.4E8	6.4E6	20	8.3E8 1.7E8	Chemical purification
α therapy	¹⁶⁶ Yb 56.7h	¹⁶⁶ Yb	Ta-Re	1.4E10	5.4E10	¹⁶⁶ Yb 5	4.1E10	2.1E9	20	5.4E10 1.1E10	Chemical purification
β therapy	¹⁶⁶ Ho 25.8h	¹⁶⁶ Ho	Ta-Re	1.4E7	1.2E7	¹⁶⁶ Ho 5	9.6E6	4.8E5	20	2.9E7 6.0E6	Chemical purification
α therapy	¹⁶¹ Tb 6.9d	¹⁶¹ Tb	UCX-Re	2.1E7	2.7E7	¹⁶¹ Tb 5	1.9E7	9.5E5	20	2.7E7 5.4E6	Chemical purification
3-therapy	¹⁵⁶ Tb 5.35d	¹⁵⁶ Tb	Ta-Re	2.5E8	8.9E7	¹⁵⁶ Tb 1	5.5E7	5.5E5	20	6.3E7 1.3E7	Chemical purification
SPECT	¹⁵⁵ Tb 5.33d	¹⁵⁵ Dy/ Tb	Ta-Re	3.2E9/ 7.4E8	7.9E9	¹⁵⁵ Dy 1	5.3E9	5.3E7	20	3.4E9 6.8E8	RILIS Dy
3 therapy	¹⁵³ Sm 46.8h	¹⁵³ Sm	UCX-Re	1.5E8	2.2E9	¹⁵³ Sm 5	2.8E9	1.4E8	20	5.2E9 1.0E9	Chemical purification
PET/CT	¹⁵² Tb 17.5h	¹⁵² Dy/ Tb	Ta-Re	1.3E10/ 3.3E9	5.6E10	¹⁵² Dy 1	3.7E10	3.7E8	20	1.1E11 2.2E10	RILIS Dy
α therapy	¹⁴⁹ Tb 4.1h	¹⁴⁹ Tb	Ta-Re	1.1E10	6.0E10	¹⁴⁹ Tb 1	3.8E10	3.8E8	20	1.2E11 2.4E10	Chemical purification

^{40}Pr -PET/ ger therapy	^{140}Nd 3.4d	^{140}Nd	Ta-Re	1.8E9	2.0E10	^{140}Nd 5	1.2E10	6.0E8	20	2.0E10	4.0E9	Chemical purification
- therapy	^{89}Sr 50.5d	^{89}Sr	UCX-Re	1.2E10	2.3E9	^{89}Sr 5	2.0E9	1.0E8	20	2.7E9	5.4E8	Only mass searation
PET	^{82}Sr 25.5d	^{82}Sr	UCX-Re	3.6E10	4.6E9	^{82}Sr 5	1.7E9	8.5E7	20	2.0E9	4.0E8	Only mass separation
- therapy	^{77}As 38.8h	^{77}As	UCX- VD5	5.7E9	1.1E10	^{77}As 5	5.8E9	2.9E8	20	9.4E9	1.4E9	Chemical purification
PET	^{74}As 17.8d	^{74}As	Y_2O_3 -VD5	6.5E9	1.2E9	^{74}As 5	3.8E8	1.9E7	20	4.5E8	9.0E7	Chemical purif
PET	^{72}As 26.0d	^{72}As	Y_2O_3 -VD5	1.6E10	2.8E10	^{72}As 5	9.1E9	4.6E8	20	1.5E10	3.0E9	Chemical purification
PET	^{71}As 65.3h	^{71}As	Y_2O_3 -VD5	1.8E10	1.8E10	^{71}As 5	5.9E9	3.0E8	20	8.0E9	1.6E9	Chemical purification
β therapy	^{67}Cu 61.9h	^{67}Cu	UCX-Re	2.7E9	3.4E9	^{67}Cu 7	1.5E9	1.1E8	20	2.7E9	5.4E8	Chemical purification
PET	^{64}Cu 12.7h	^{64}Cu	Y_2O_3 -VD5	1.1E10	2.3E10	^{64}Cu 5	7.1E9	3.6E8	20	2.1E10	3.6E9	Chemical purification
γ , dosimetry	^{61}Cu 3.3h	^{61}Cu	Y_2O_3 -VD5	7.7E9	1.7E10	^{61}Cu 5	5.1E9	2.6E8	20	2.1E10	4.0E9	Only mass separation
β therapy	^{47}Sc 3.4d	^{47}Sc	Ti	6.4E10	5.0E10	^{47}Sc 5	4.2E10	2.1E9	20	5.9E10	1.2E10	Evaporation
PET	^{44}Sc 4.0h	^{44}Sc	Ti	4.4E10	6.6E10	^{44}Sc 6.4	5.7E10	2.9E9	20	1.6E11	3.2E10	Evaporation
PET	^{11}C 20.3m	^{11}CO	NaF-LiF- VD5 ⁰	-	-	- 15	-	1.4E9	-	-	4.2E9	Only mass separation