

Readiness of CERN-MEDICIS for... 2016 or 2017

Thierry Stora, EN-STI

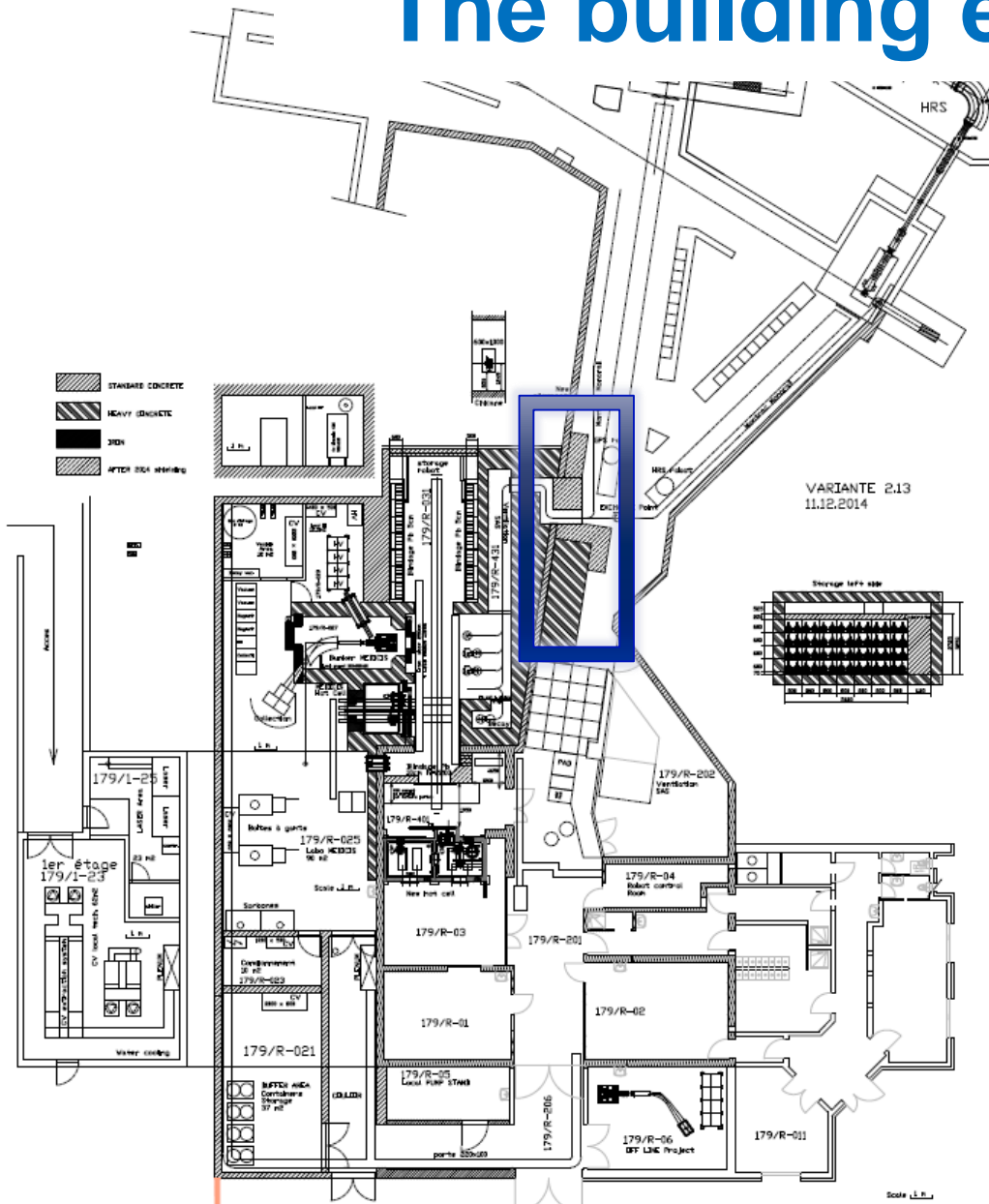


ENGINEERING
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ISOLDE

MEDICIS

The building extension

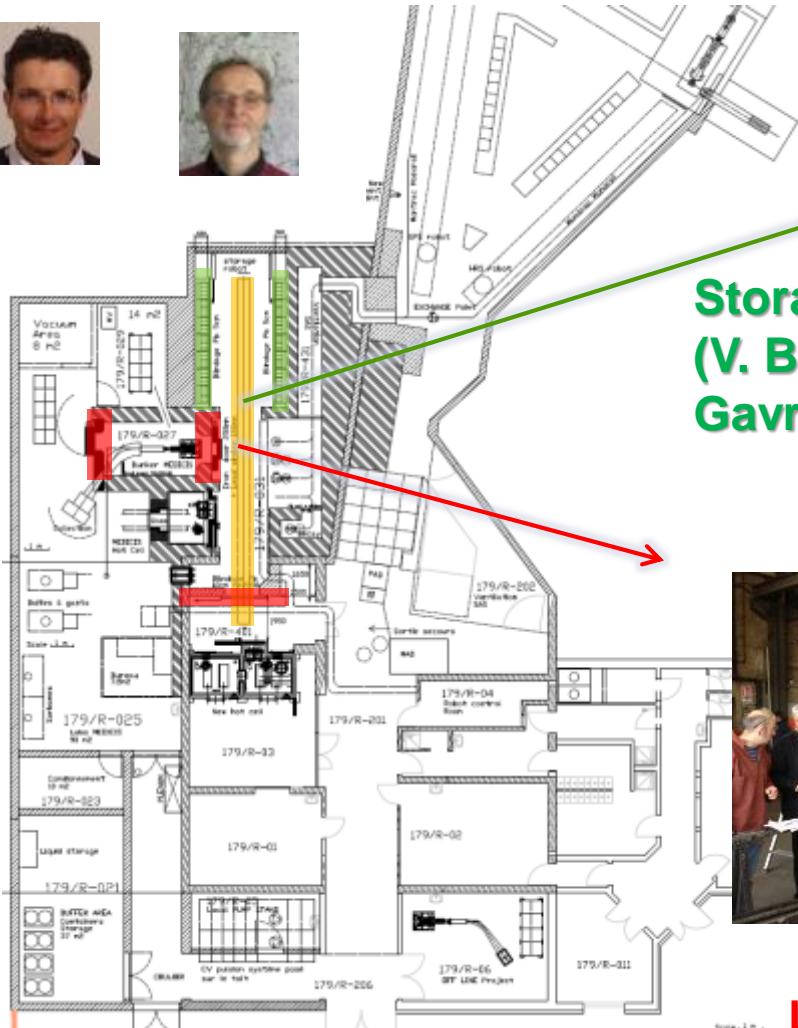


Temporary shielding removed
And final shielding/access under way



The building extension

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



Storage shelves
(V. Barozier & Y. Gavrikov)



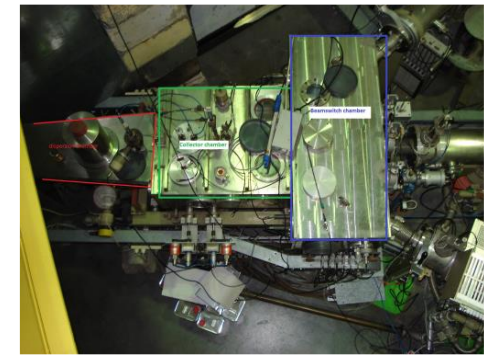
Robot hardware
installed



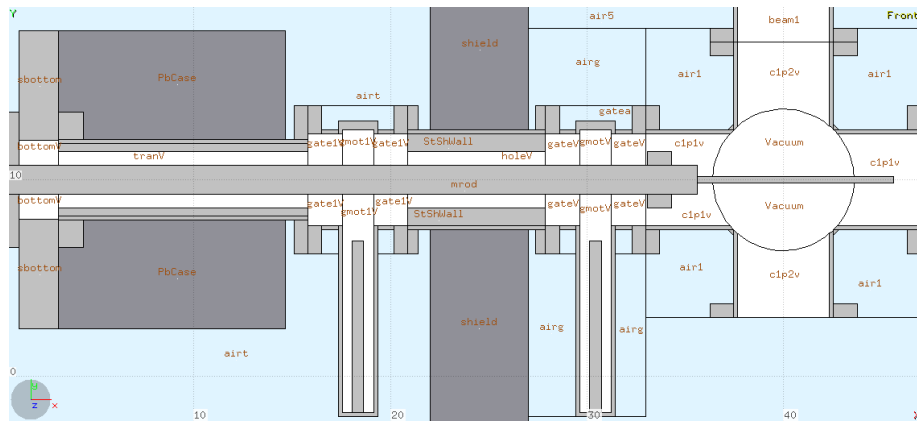
Last shielded door



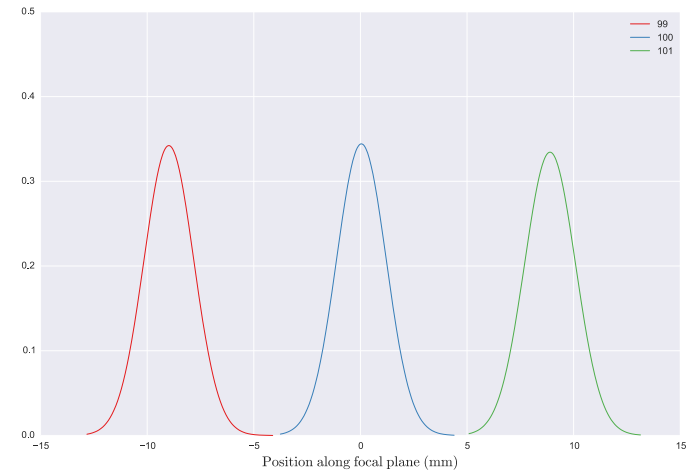
The Mass separator



LISOL dipole ready for shipment
In Louvain La Neuve



Ion distribution at focal plane



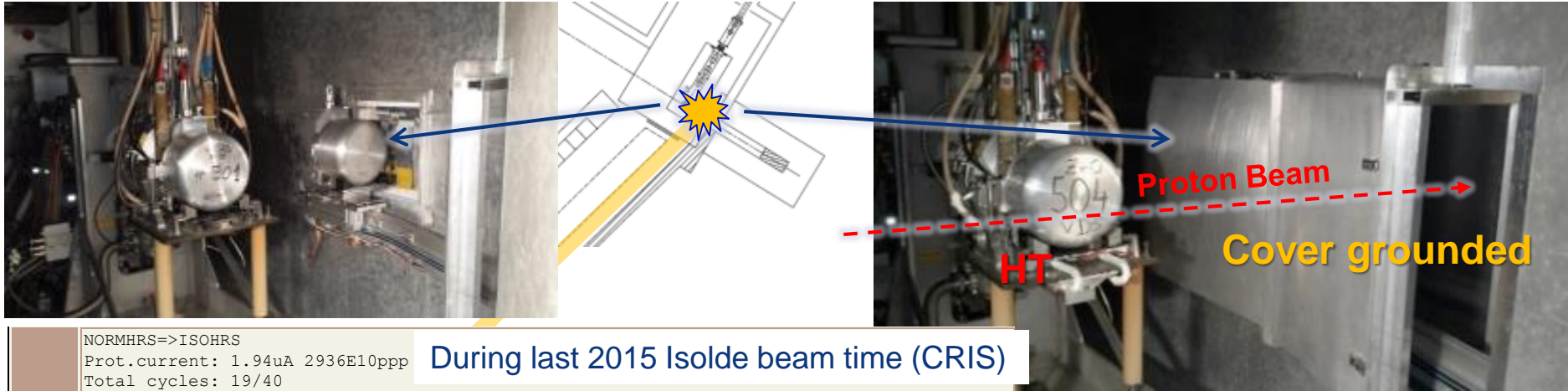
Separator in CERN-MEDICIS
Configuration (Y. Martinez)

Conceptual design – collection chamber (A. Brown)

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

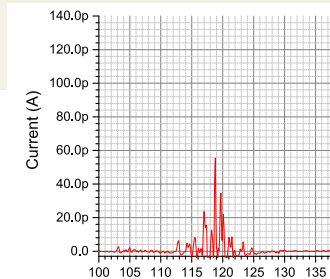
During last 2015 Isolde beam time (CRIS)

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

Facility completion and start-up

- The Medical applications at CERN have been slightly restructured.
- A review is under preparation to assess the readiness to start (the staged) operation of CERN-MEDICIS, chaired by R. Saban.

| 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

The activities and collaboration

- The CERN-MEDICIS scientific case seems to be solid, eg recent LUTHATERA® radiopharmaceutical approval (using on ^{177}Lu)

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

¹JRC-ITU, Karlsruhe, Germany

²KULeuven, Leuven, Belgium

³CERN-MEDICIS project, CERN, Switzerland

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

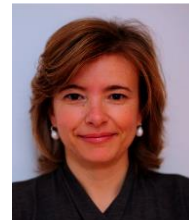
Friday, 15 January 2016

08h30 – 17h00

Outreach & Collaboration



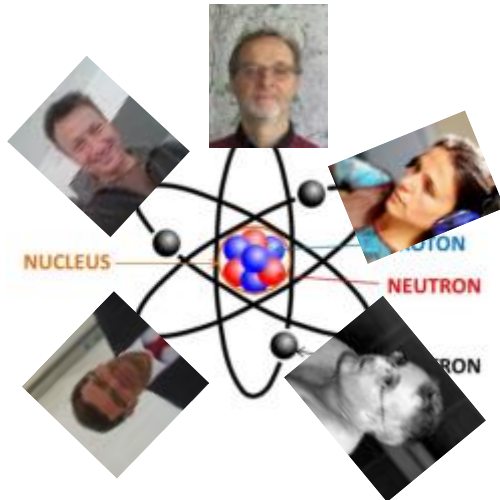
2nd yearly Grace-MEDICIS lecture :
Prof. W. Weber, Memorial Sloan Kettering Cancer
Center New York



C. Ferrari started with us
Yesterday to help on the
administration



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And many others

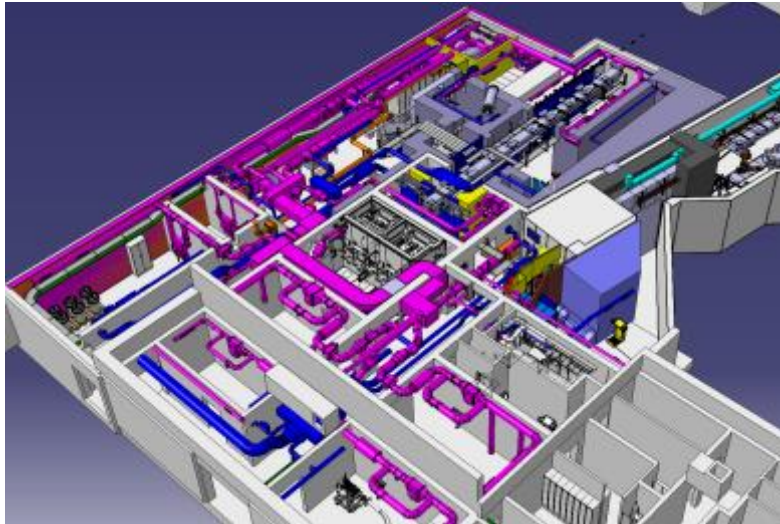
Thank you, questions , comments ?



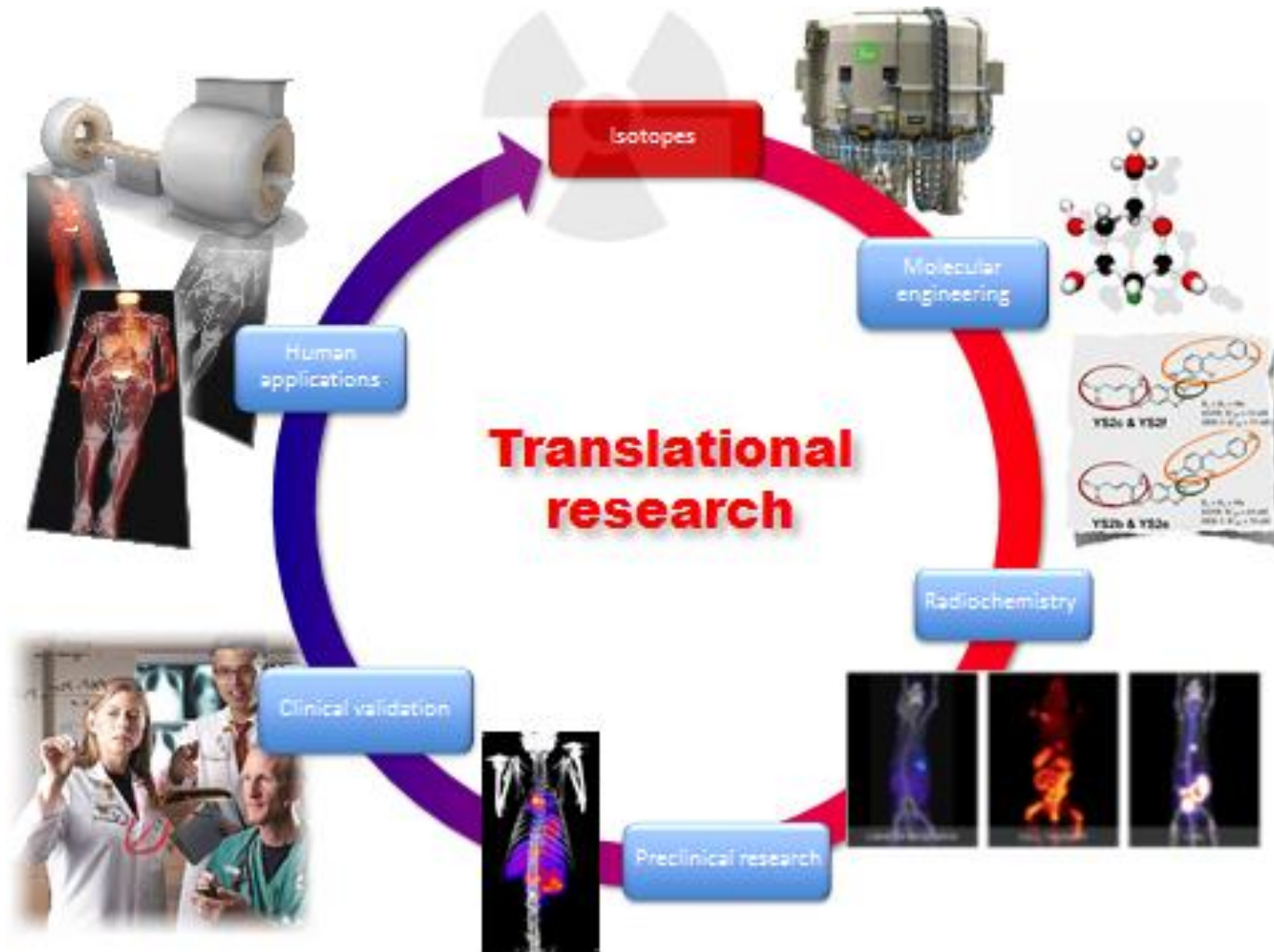
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Reserve

The nuclear ventilation



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

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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



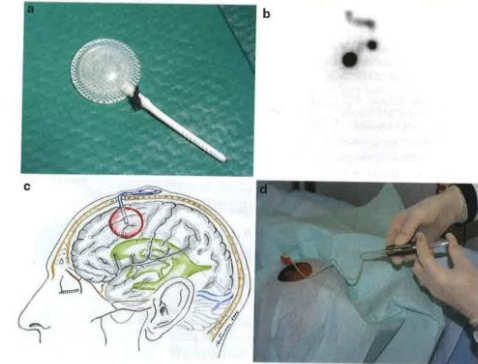
Current radiotherapy against cancer mostly uses beta-emitters as medical isotopes
© Alex Tihonov, Fotolia.com

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

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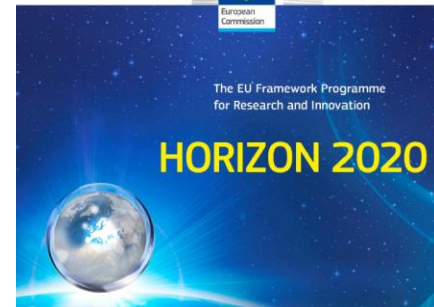
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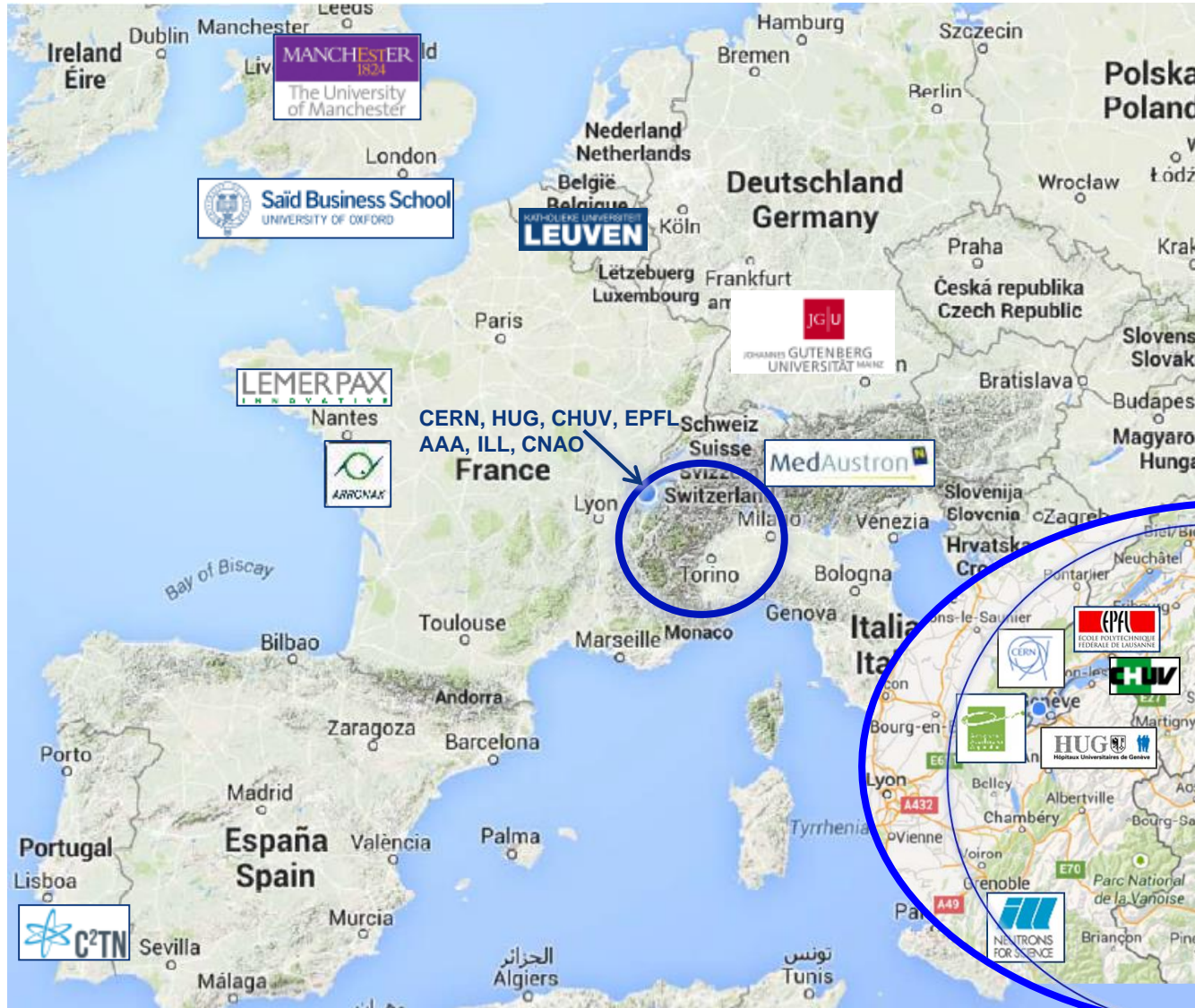


MEDICIS-PROMED

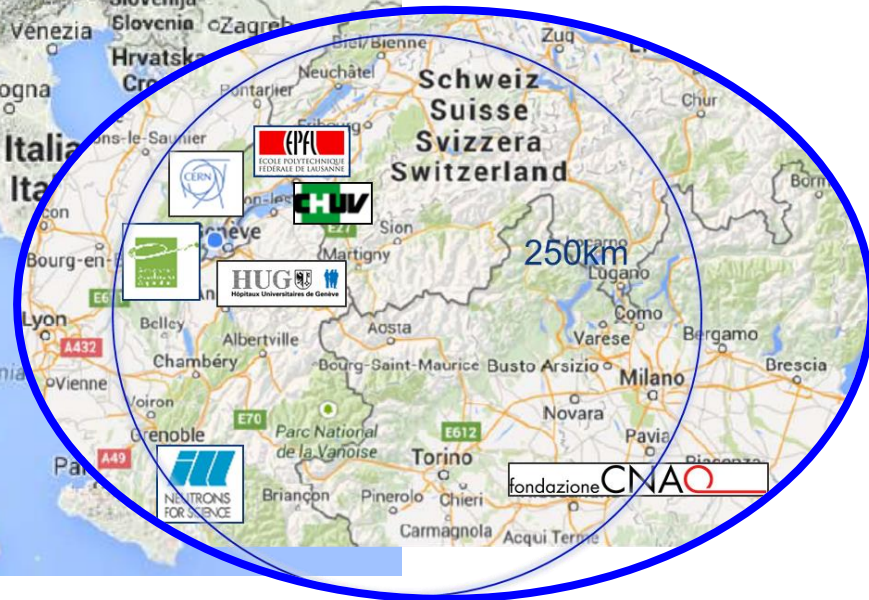
« MEDICIS-Produced radioisotope beams for medicine »

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)



Some yield estimates

| Medical application | Isotope half-life | Parent isotope beam | Target - Ion source | ISOLDE [†] | | RIB ϵ_{ext}^{**} (%) | CERN-MEDICIS [†] | | CERN-MEDICIS 2GeV 6 μ A | | 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) | | | | | | | |
| α -therapy/ T/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 |
| ger 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 |
| uger 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 |
| α -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 |
| α 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 |