

# Accelerator-Based Boron Neutron Capture Therapy

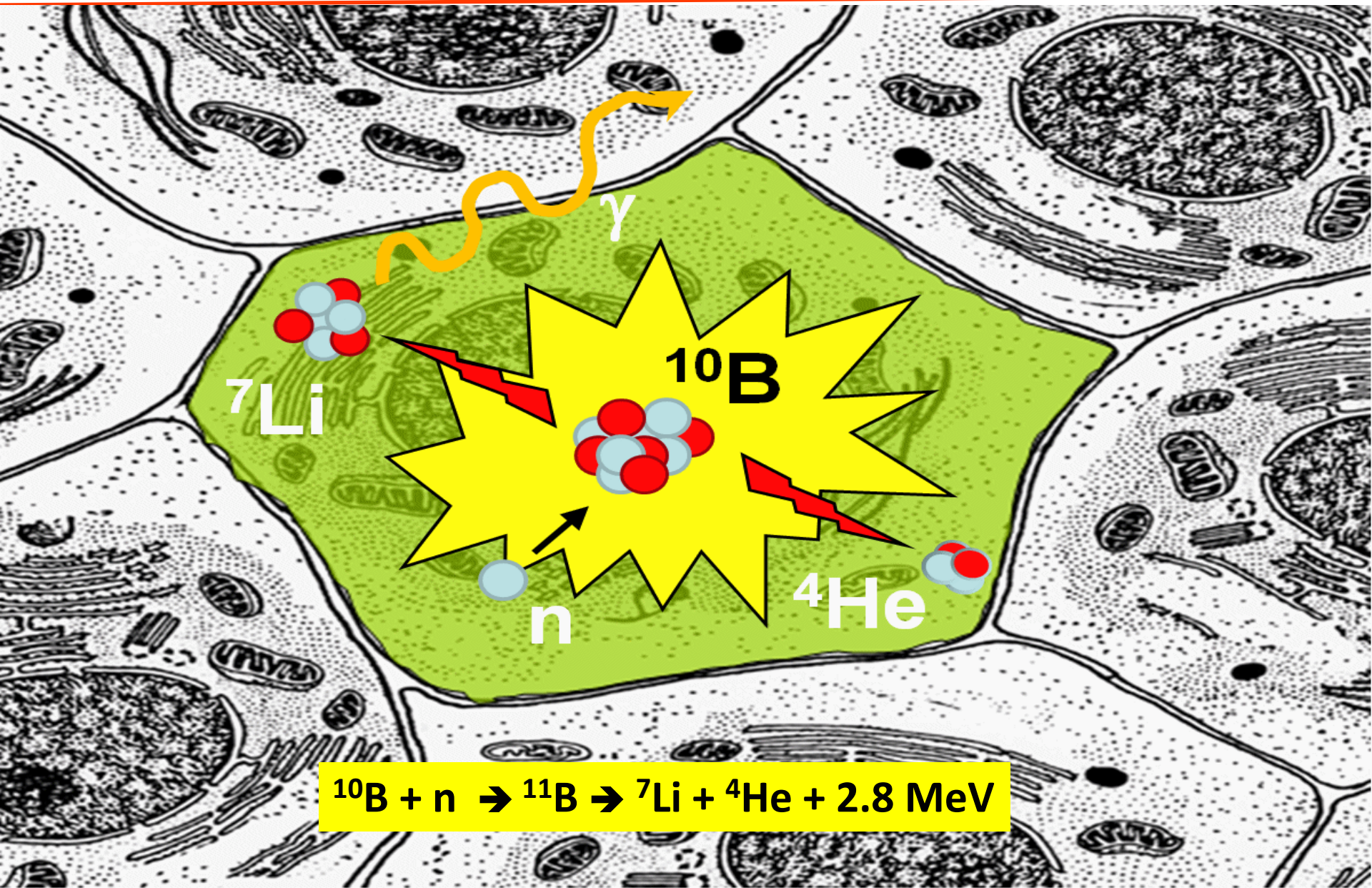
**Paolo Colautti, *INFN***

*Town meeting on IFMIF/ELAMAT Rzeszów 14-15 April 2016*

## ***Talk Topics***

- 1. BNCT rationale.**
- 2. BNCT drawbacks.**
- 3. BNCT medical results.**
  
- 4. Neutron sources from particle accelerators.**
- 5. The Neutron *Beam-Shaping Assembly*.**
- 6. Treatable tumours: *the Advantage Depth*.**
  
- 7. The AB-BNCT 4 pillars:**
  - 1. powerful accelerator;***
  - 2. safe target handling;***
  - 3. tracing the Boron carrier;***
  - 4. microdosimetric detector.***

# Rationale: BNCT is a Cellular Hadronic Therapy



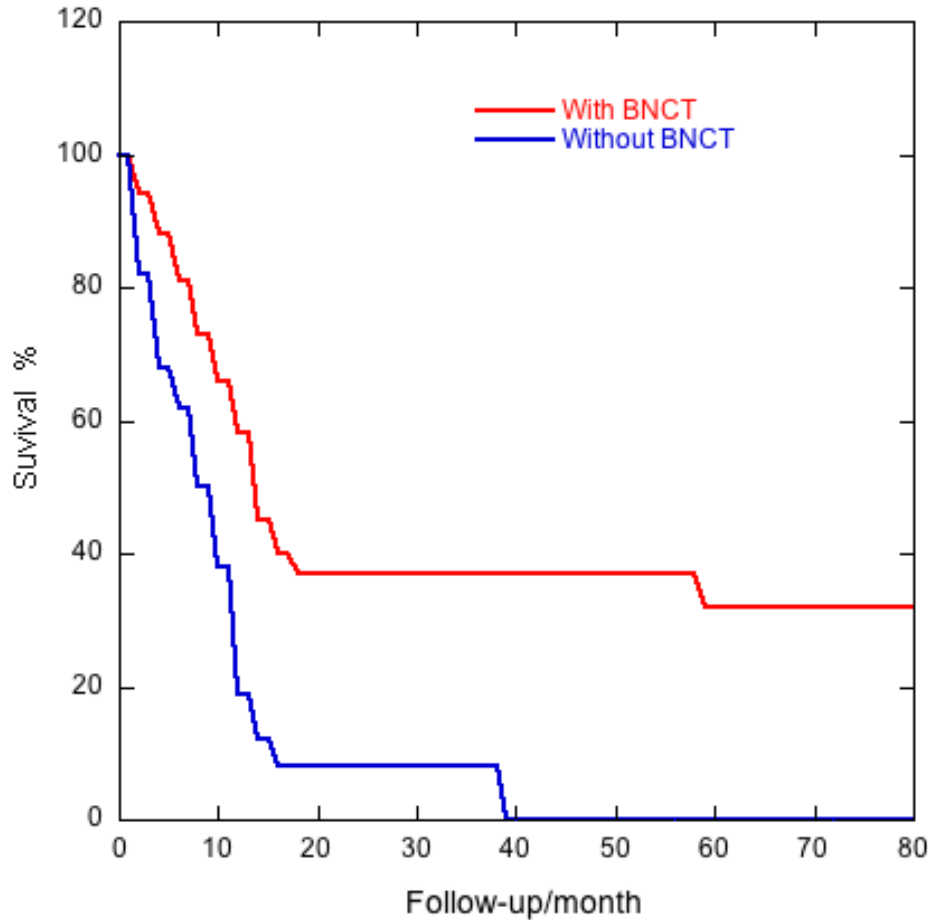
# BNCT Drawbacks: Nuclear Reactions Which Damage Also Health Cells

Reaction	Q-Value (MeV)	Cross Section barn	Threshold MeV
${}^1\text{H}(n, \gamma){}^2\text{H}$	+2.22	20.8	---
${}^{14}\text{N}(n, p){}^{14}\text{C}$	+0.63	1.6	---
${}^{14}\text{N}(n, \alpha){}^{11}\text{B}$	-0.158	0.084	0.17
${}^{40}\text{Ca}(n, \alpha){}^{37}\text{Ar}$	+1.75	0.05	---
${}^{16}\text{O}(n, \alpha){}^{13}\text{C}$	-2.16	0.008	2.36

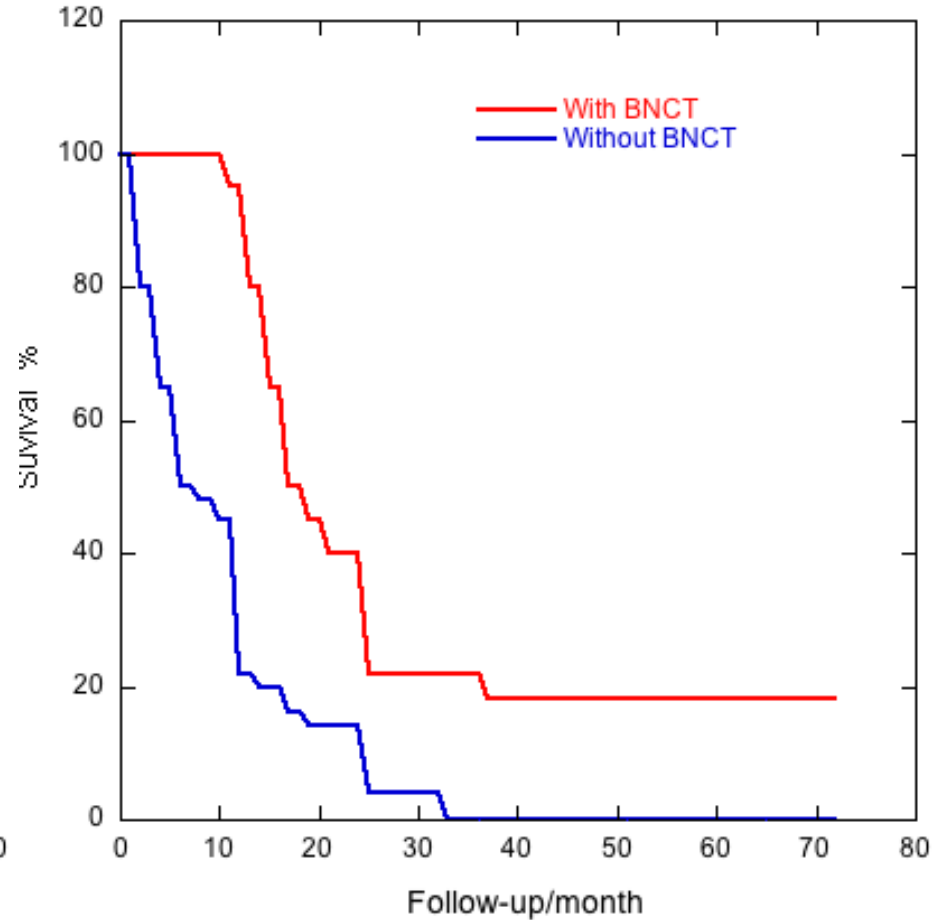
${}^{10}\text{B}+n \rightarrow {}^7\text{Li}+{}^4\text{He}$	<b>2.79</b>	<b>3837</b>	---
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# BNCT Medical Results: Kaplan-Meier Plots

## Head and Neck Cancer



## Glioblastoma

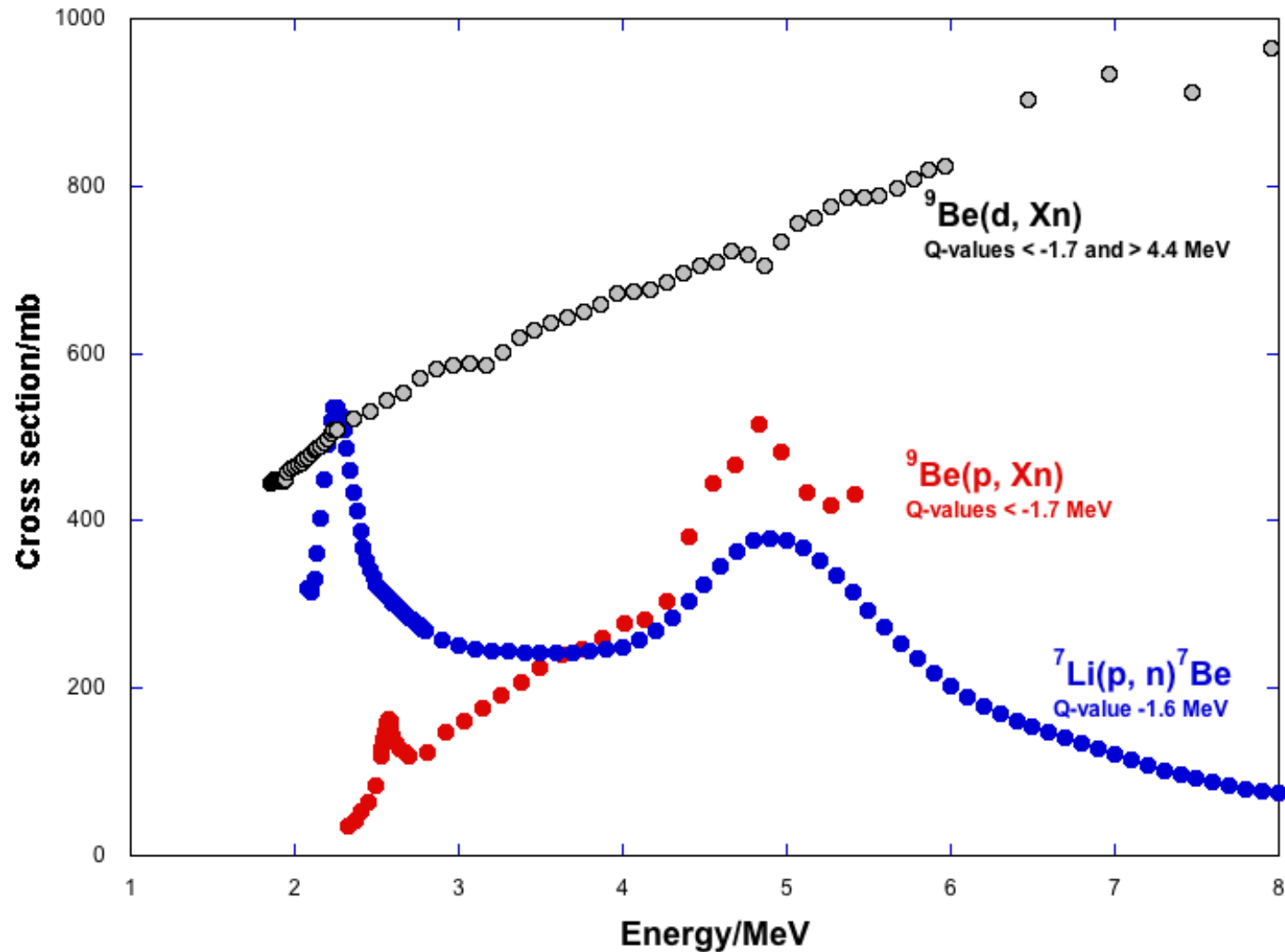


Nuclear Physics for Medicine. NuPECC Report, edit A.Bracco et al. 2014, [http://www.nupecc.org/npmed/npmed2014\\_hires.pdf](http://www.nupecc.org/npmed/npmed2014_hires.pdf)

- **Treatments with thermal neutrons (skin melanomas, explanted liver, brain-glioblastoma intraoperative irradiation) were successful.**
- **Treatments with epithermal neutrons are only partially successful.**

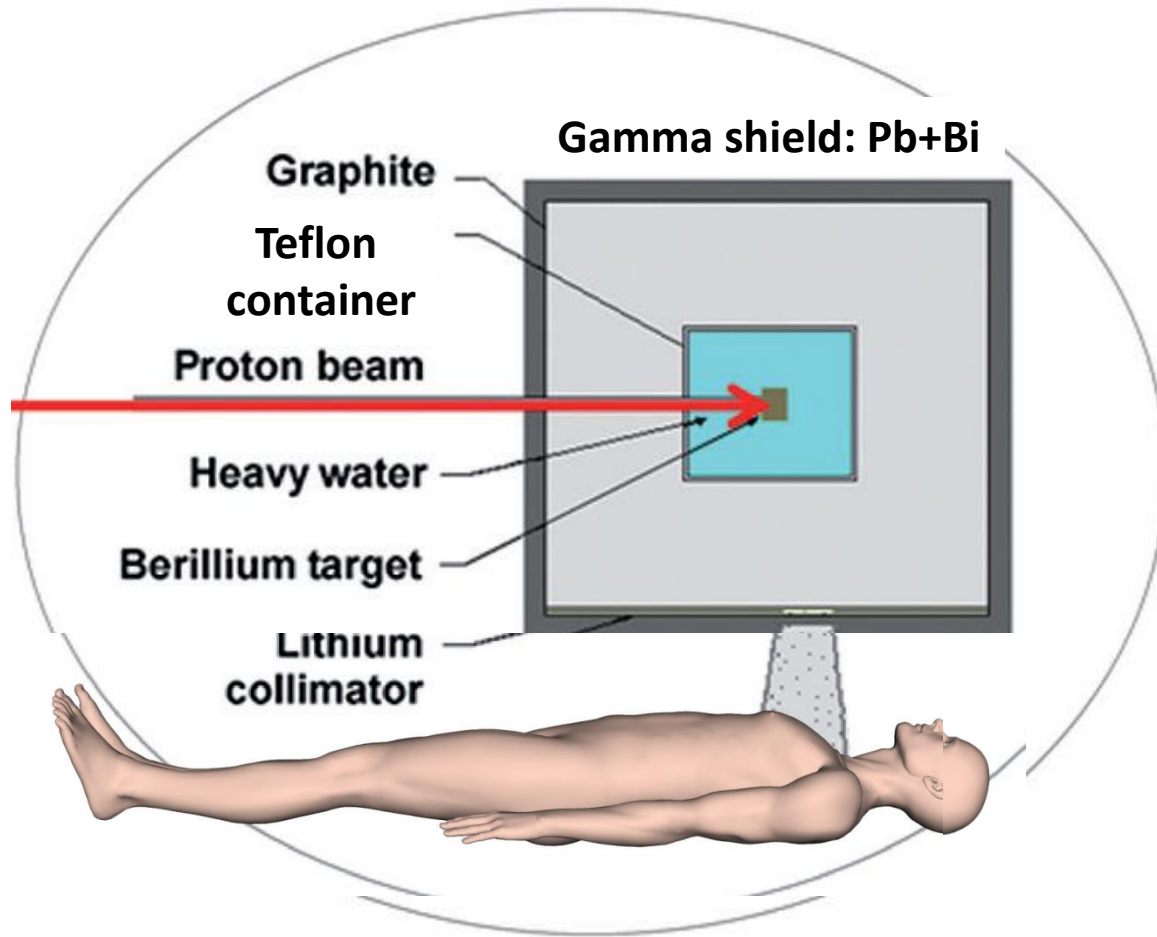
- **So far, all BNCT treatments have used nuclear research reactors.**
- **Unlikely nuclear reactors could be installed inside a hospital.**
- **The use of a dedicated particle accelerator, rather than a research reactor, will implement a real clinical BNCT.**

# Neutron Sources Exploitable for AB-BNCT

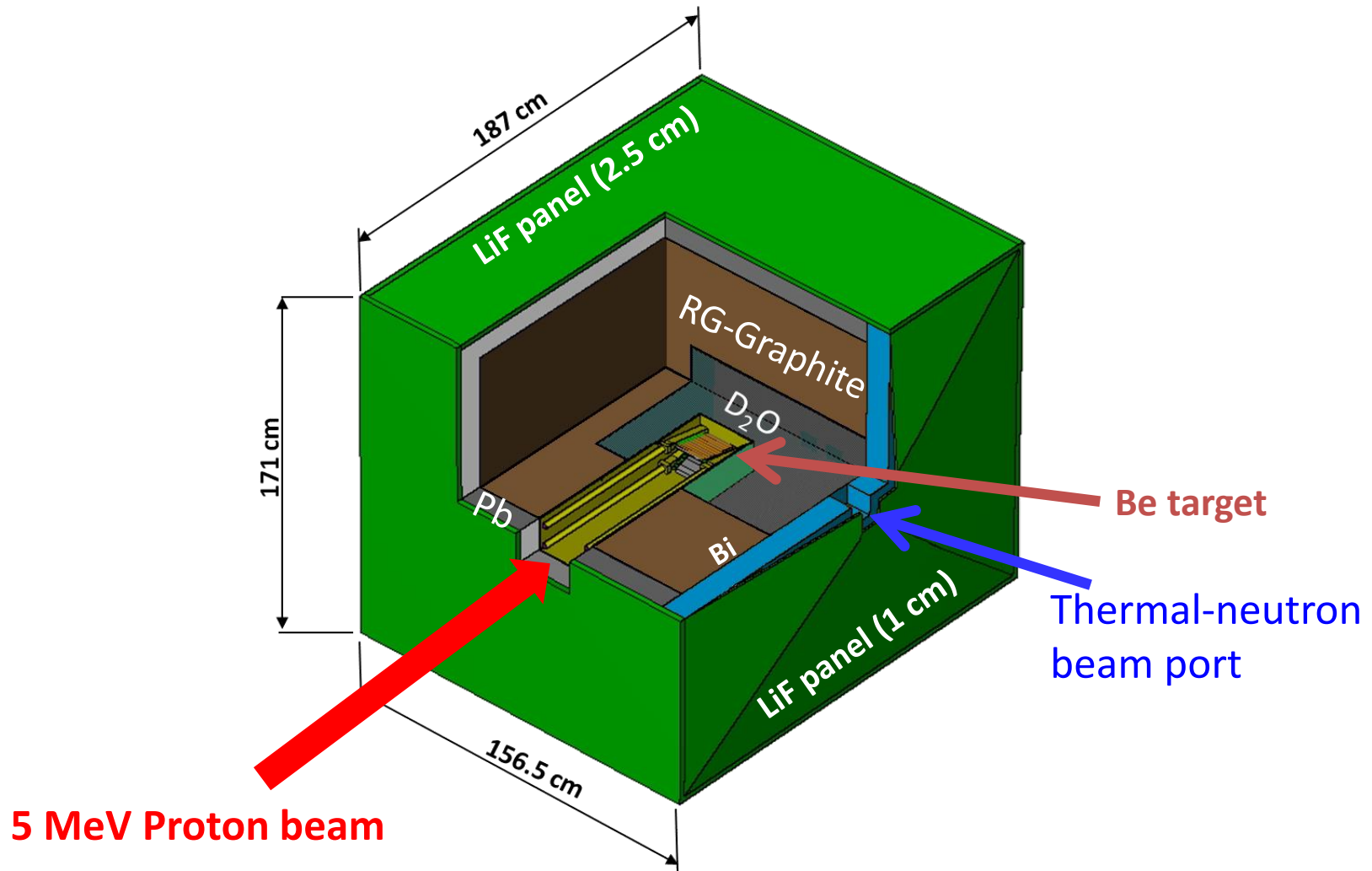




# The Beam Shaping Assembly to Slow Down and Shaping the Neutron Beam



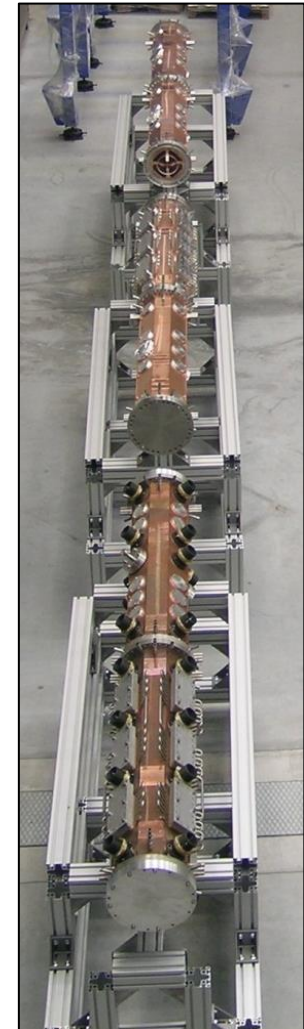
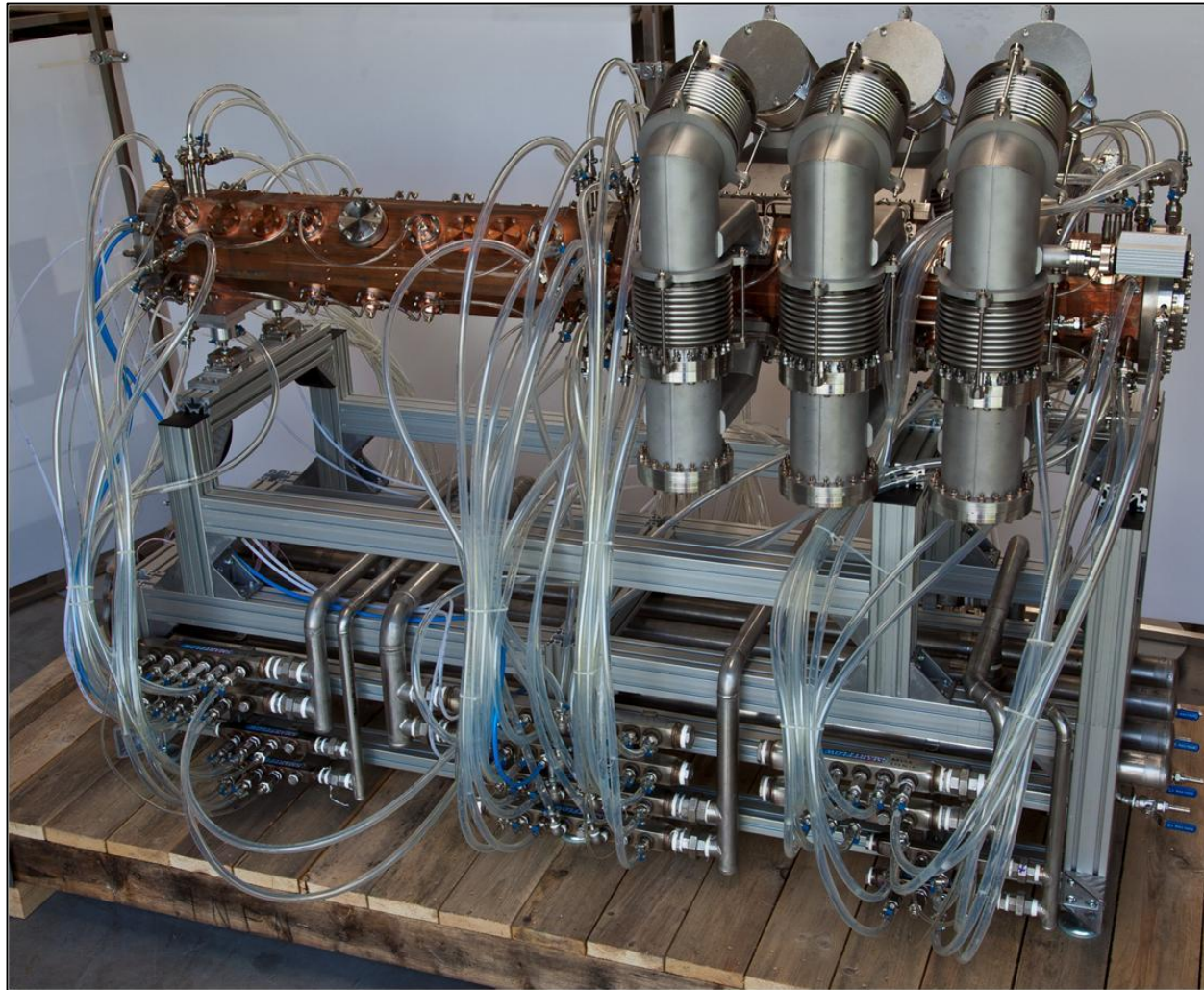
# The LNL Beam-Shaping Assembly





# The AB-BNCT 4 Pillars

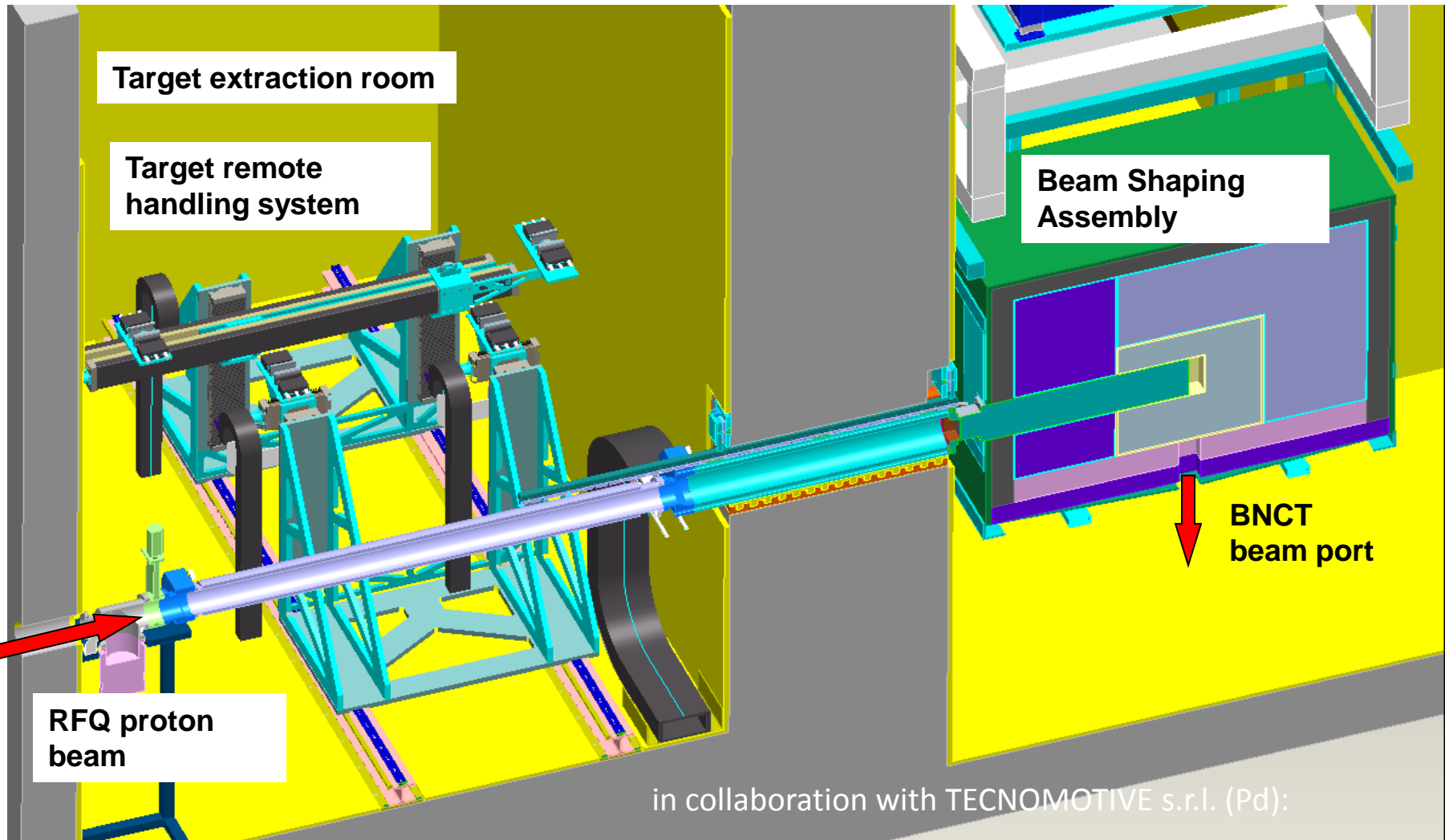
## 1. Powerful accelerator, the LNL RFQ accelerator



5 MeV 30 mA proton beam  $\implies 10^{14} \text{ s}^{-1}$  neutron production rate

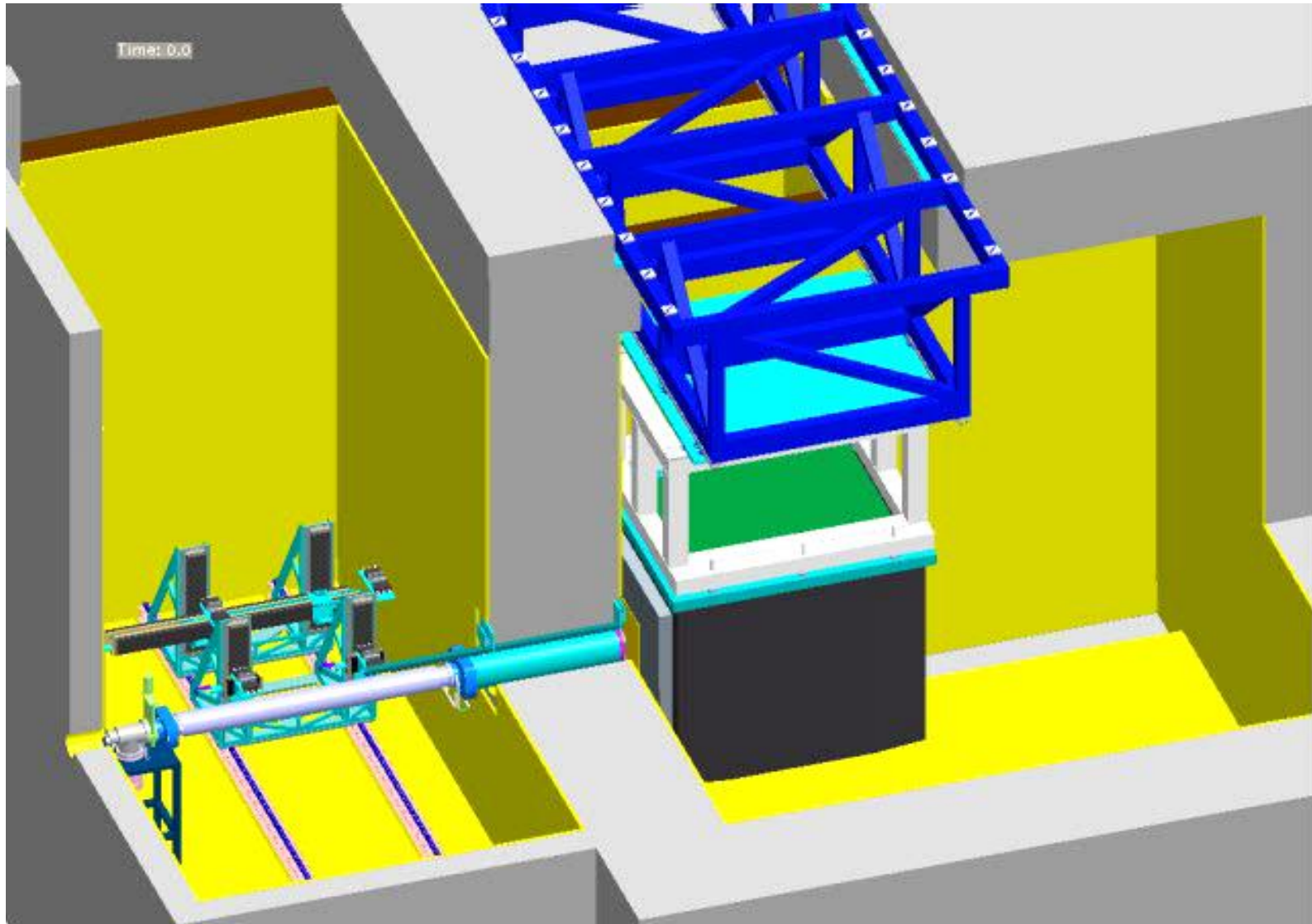
# The AB-BNCT 4 Pillars

## 2. Safe target handling, the LNL design



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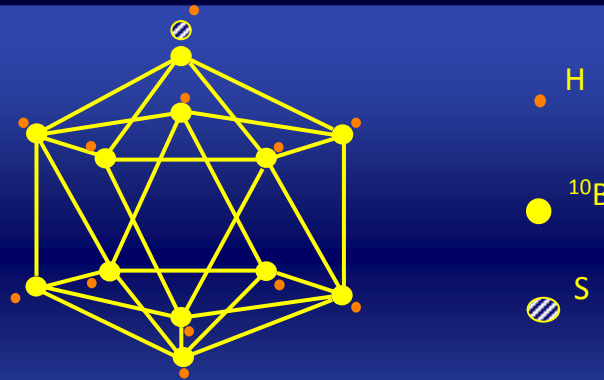
# The 4 AB-BNCT Pillars

## 3. Selective Boron Carrier, Actual Commercial Compounds.



Boronophenylalanine: BPA

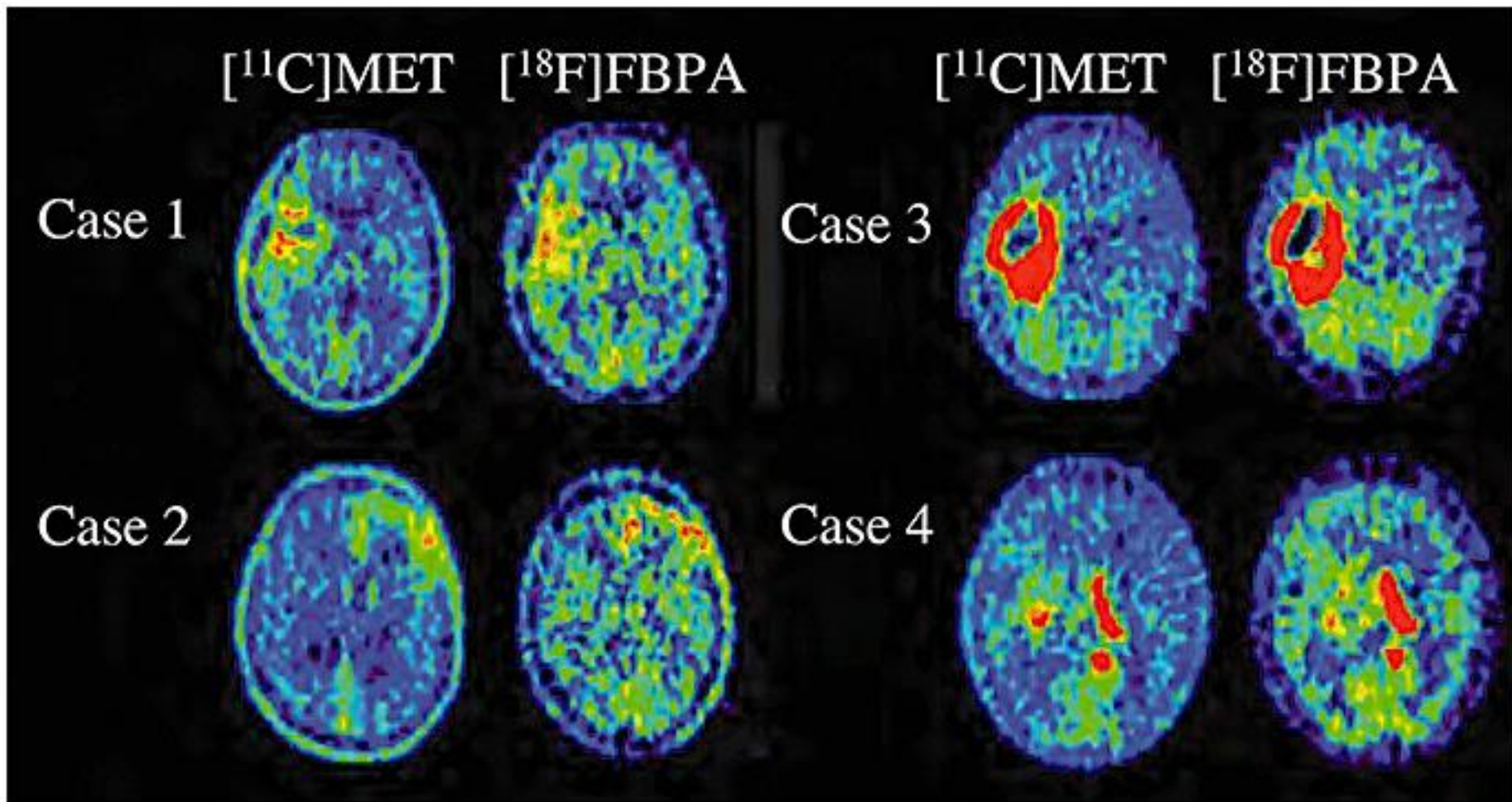
$Na^+$



Sodium Borocaptate: BSH

# The 4 AB-BNCT Pillars

## 3. Tracing Boron carrier, fluorinated BPA and PET .

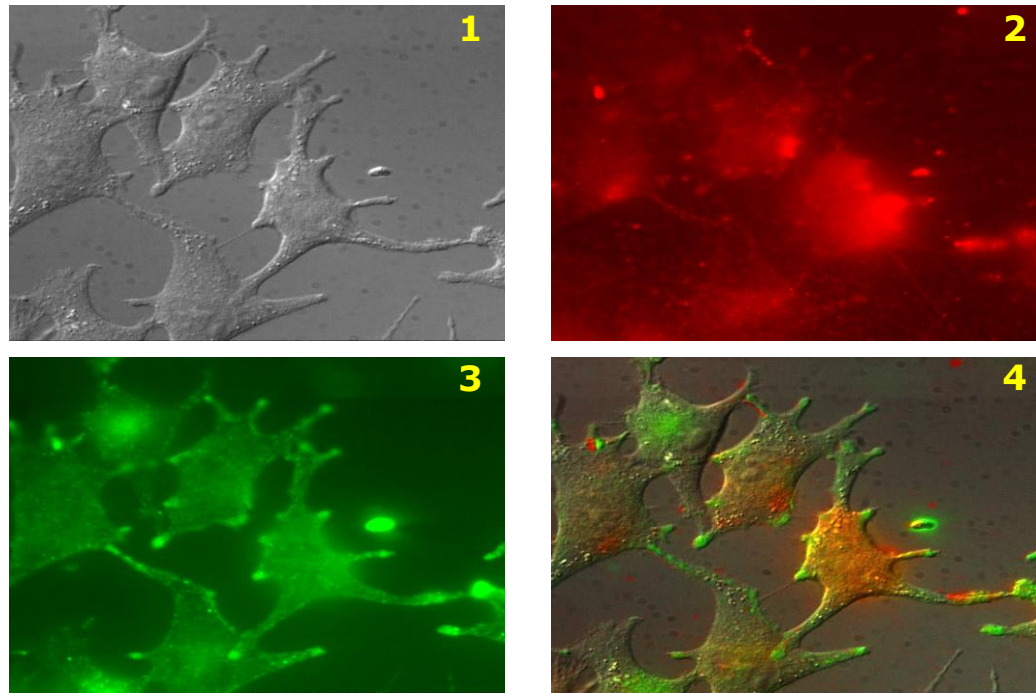




# The 4 AB-BNCT Pillars

## 3. Tracing Boron carrier, Boronophthalocyanine and fluorescence induced by a laser beam

**B<sub>4</sub>Pc: the selective uptake can be simply traced by fluorescence**

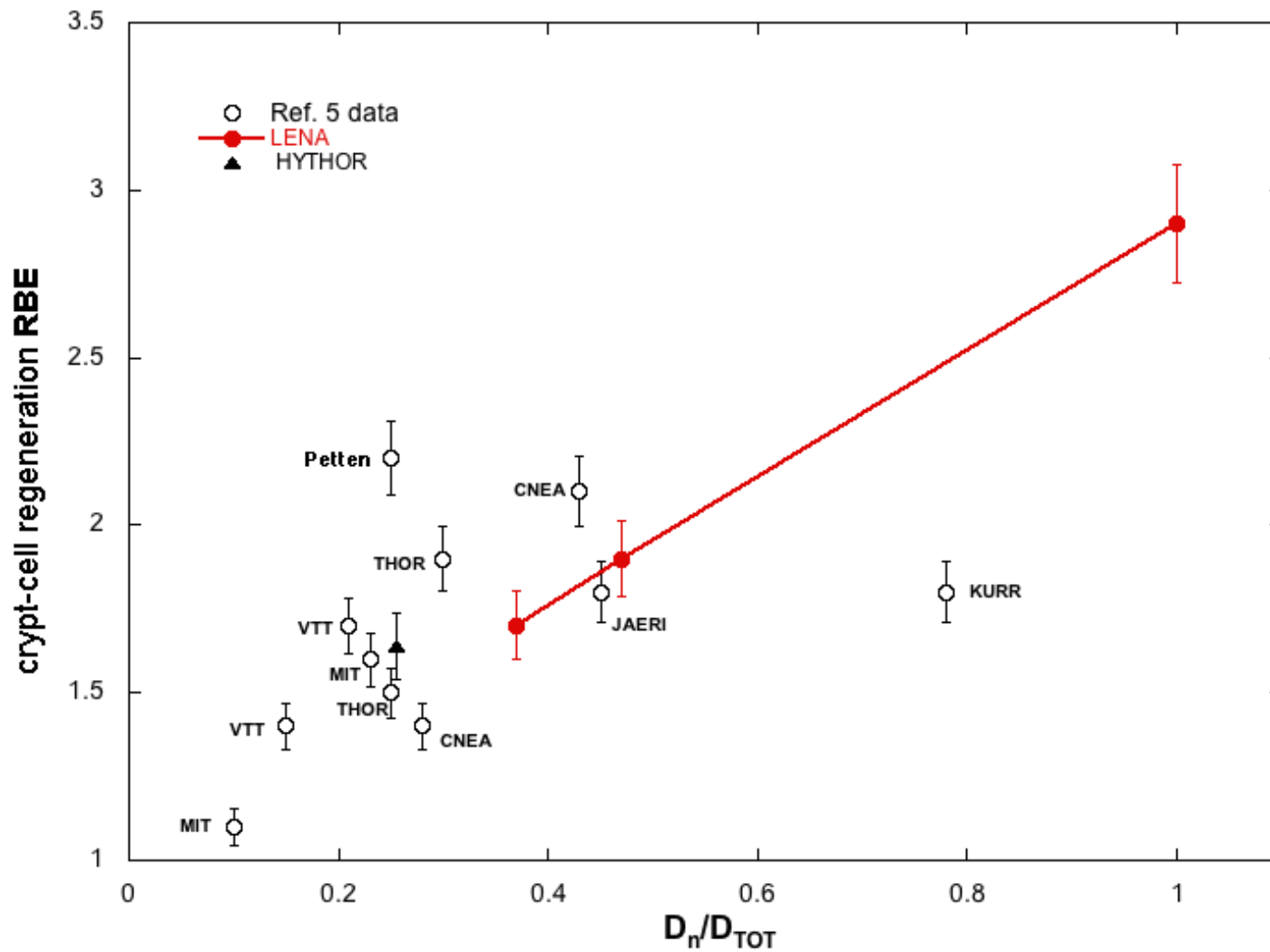


**Fluorescence micrographs of cells after 24 h incubation with 7  $\mu$ M DOPC liposome-incorporated B<sub>4</sub>Pc 1 bright field image, 2 fluorescence of phthalocyanine, 3 fluorescence of endosomal probe Lucifer Yellow, 4 overlay of images 2 and 3**

E.Friso et al., Photochem.Photobiol. Sci. 5, 39-50, 2006.

# The 4 AB-BNCT Pillars

## 4. Microdosimetric detector, the RBE case



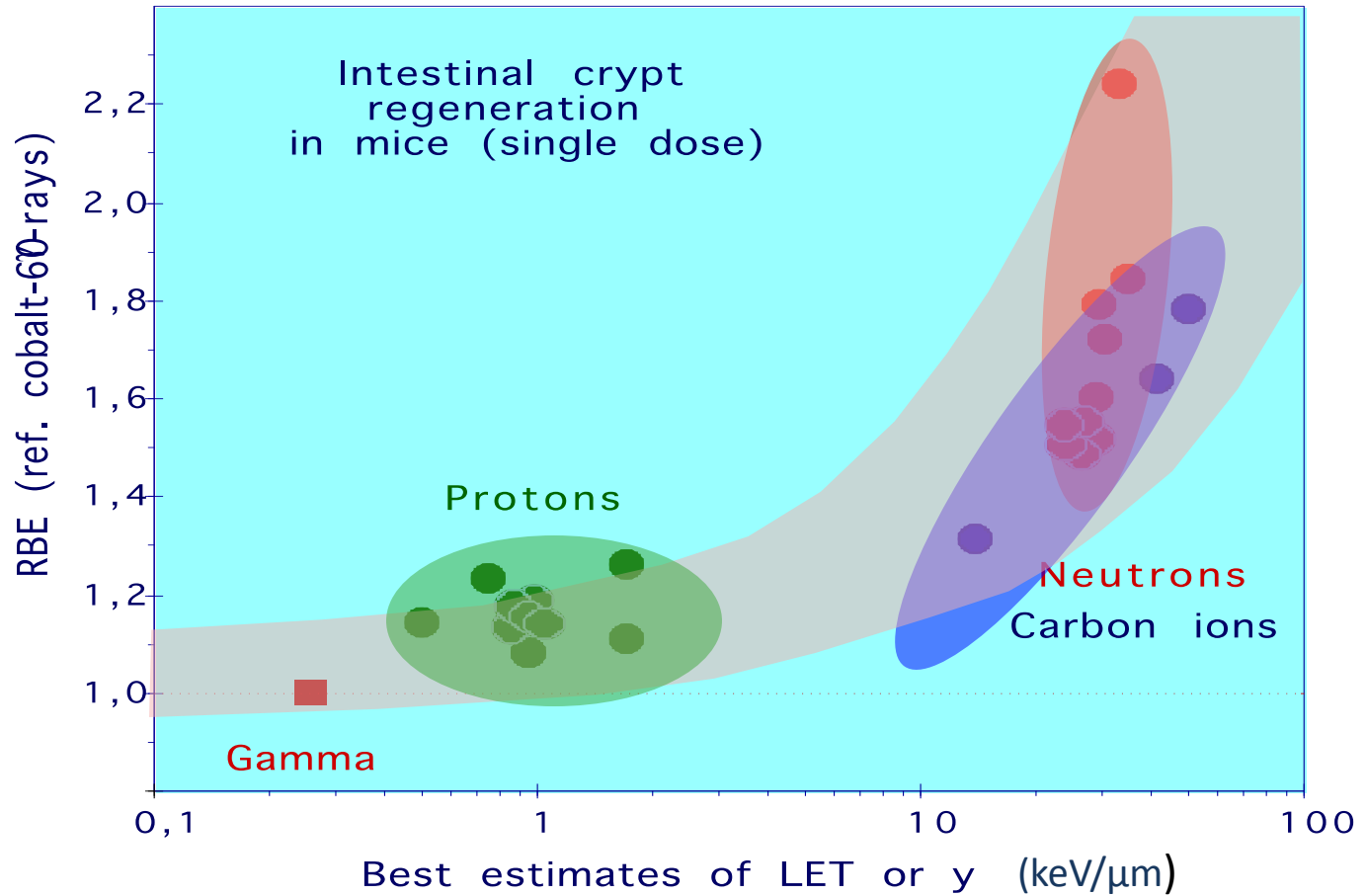
J.Guellette et al. Proceedings ICNCT 12,81-84, 2006

P.Colautti et al., ARI 88, 147-152, 2014

# The 4 AB-BNCT Pillars

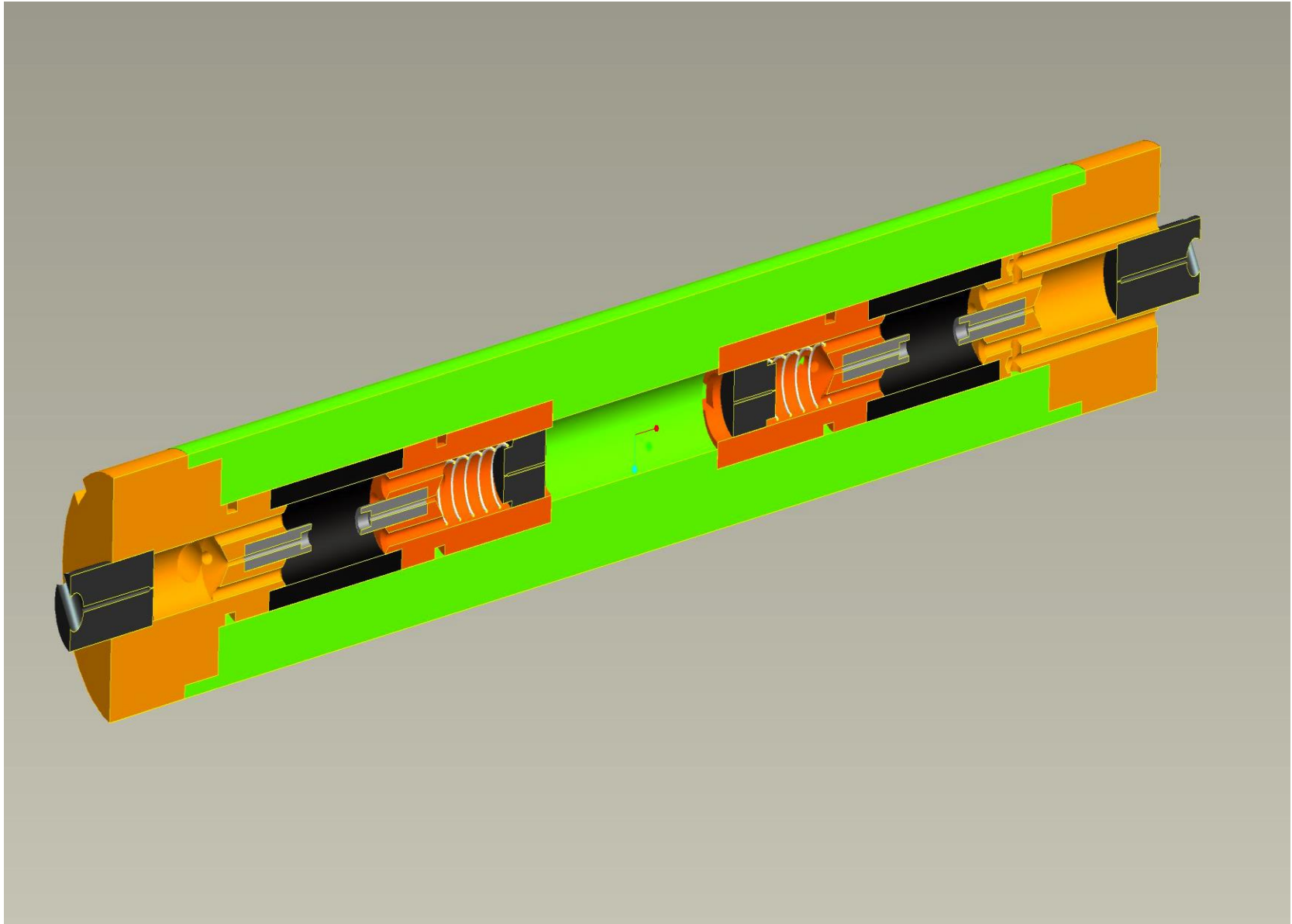
## 4. Microdosimetric detector, the RBE case

# RBE of BNCT components



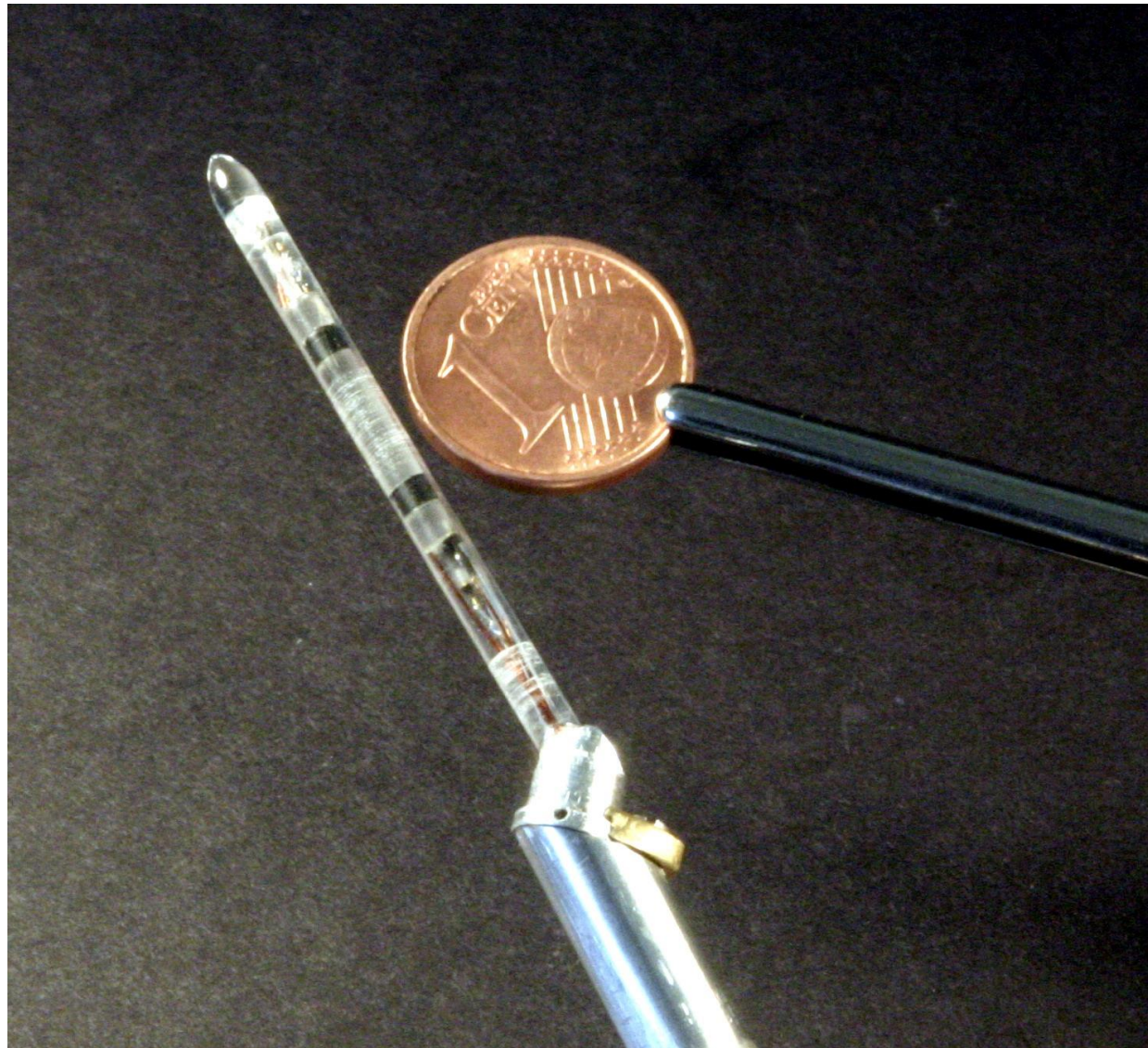
# The 4 AB-BNCT Pillars

## 4. Microdosimetric detector, the LNL twin TEPC



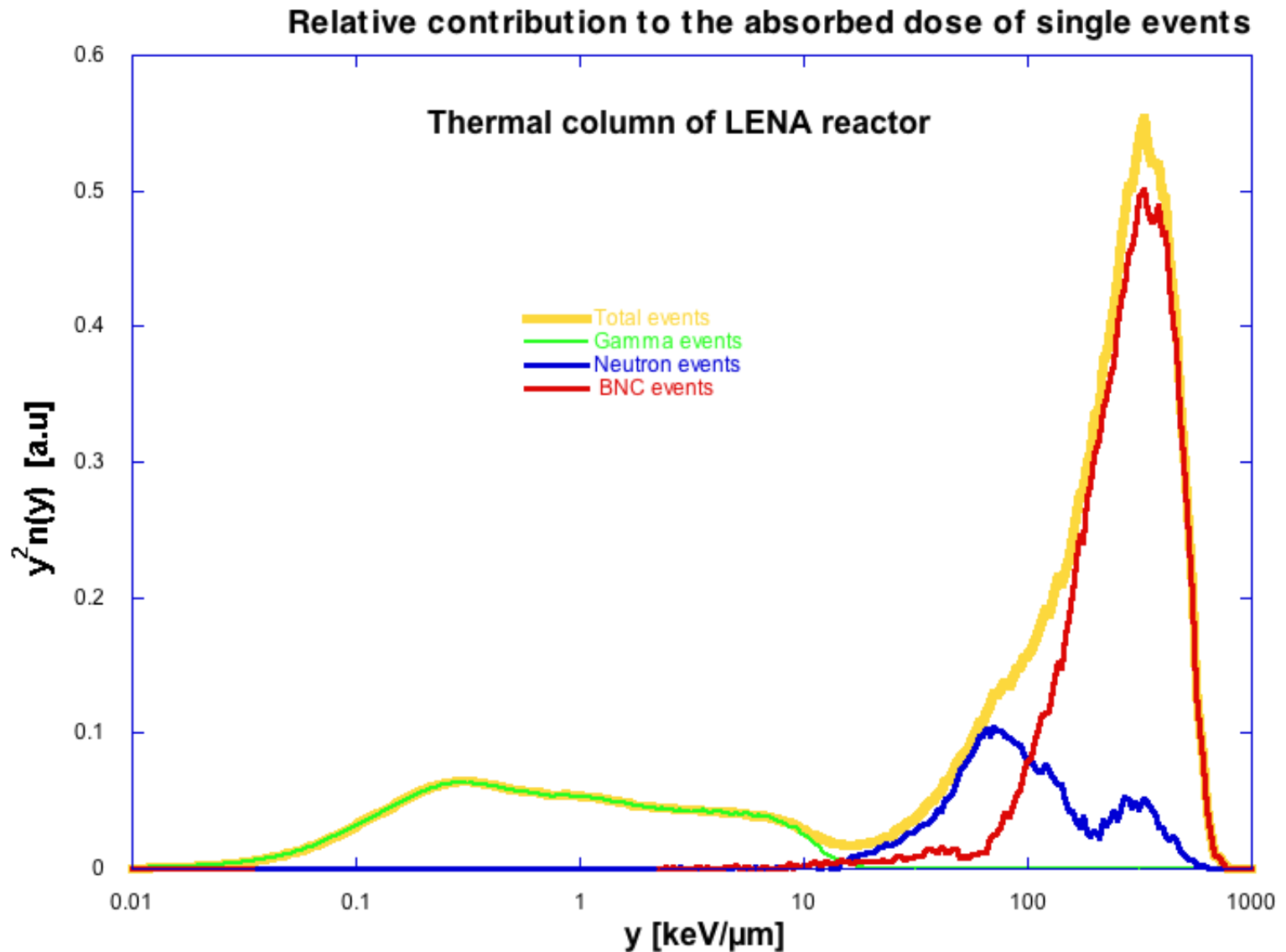
# The 4 AB-BNCT Pillars

## 4. Microdosimetric detector, the LNL twin TEPC



# The 4 AB-BNCT Pillars

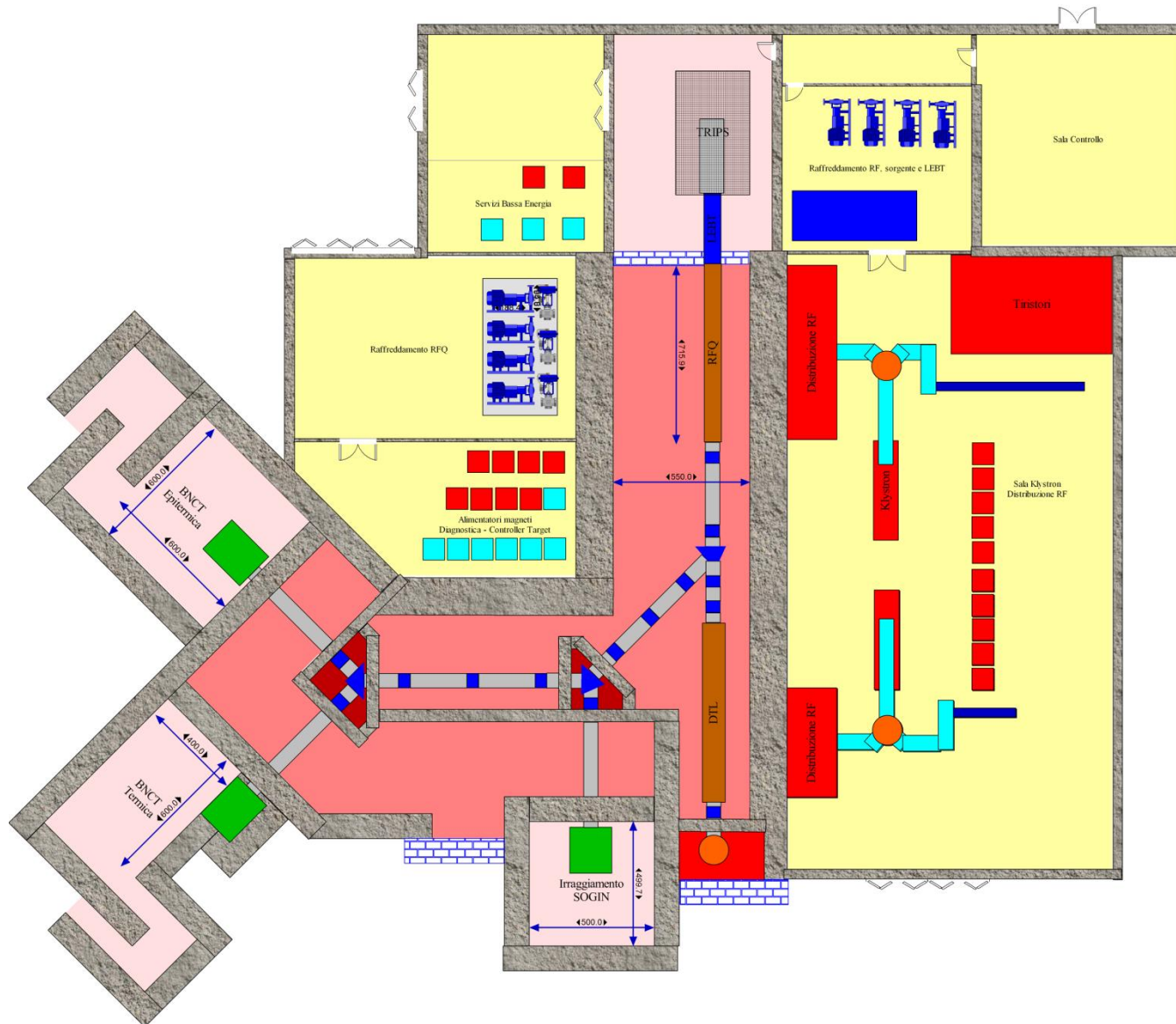
## 4. Microdosimetric detector, the LENA reactor BNCT microdosimetric spectrum



# CONCLUSIONS

1. BNCT is a tumour radiation therapy useful when a cellular therapy is recommended. It needs high neutron-fluence rates, so far provided only by nuclear research reactors.
1. New neutron sources based on particle accelerators could give a boost to BNCT studies. Nine AB-BNCT projects are actually running in the world (Italy, Russia, UK, Japan, Israel, Argentina).
2. A poor  $^{10}\text{B}$  carrier specificity limits the BNCT therapeutic advantage. Knowledge of the  $^{10}\text{B}$  carrier metabolic distribution in the patient improves the BNCT therapeutic advantage.
3. The radiation field complexity and the poor knowledge of the radiation field RBE limit the BNCT therapeutic advantage. Experimental microdosimetry performed with TEPCs or other microdosimetric detectors could improve the BNCT therapeutic advantage.

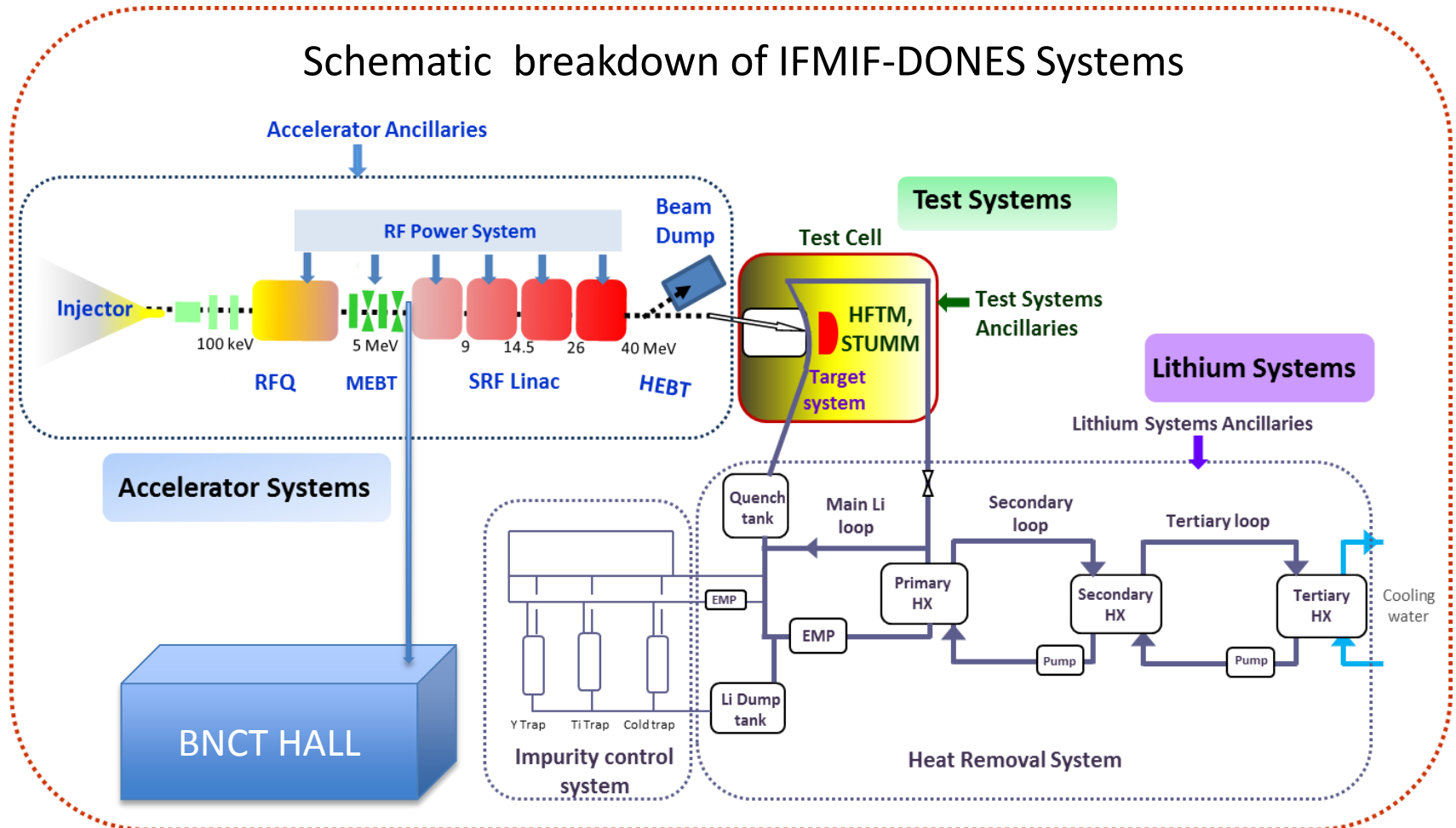
# The LNL AB-BNCT Proposed Centre





# The IFMIF AB-BNCT IFMIF Possible Draft

## Schematic breakdown of IFMIF-DONES Systems



**Site, Buildings & Plant Systems**

- Layout & Site Infrastructures
- Buildings
- Plant Systems (I and II)
- Remote Handling

Central Control Systems and Integrated Instrumentation

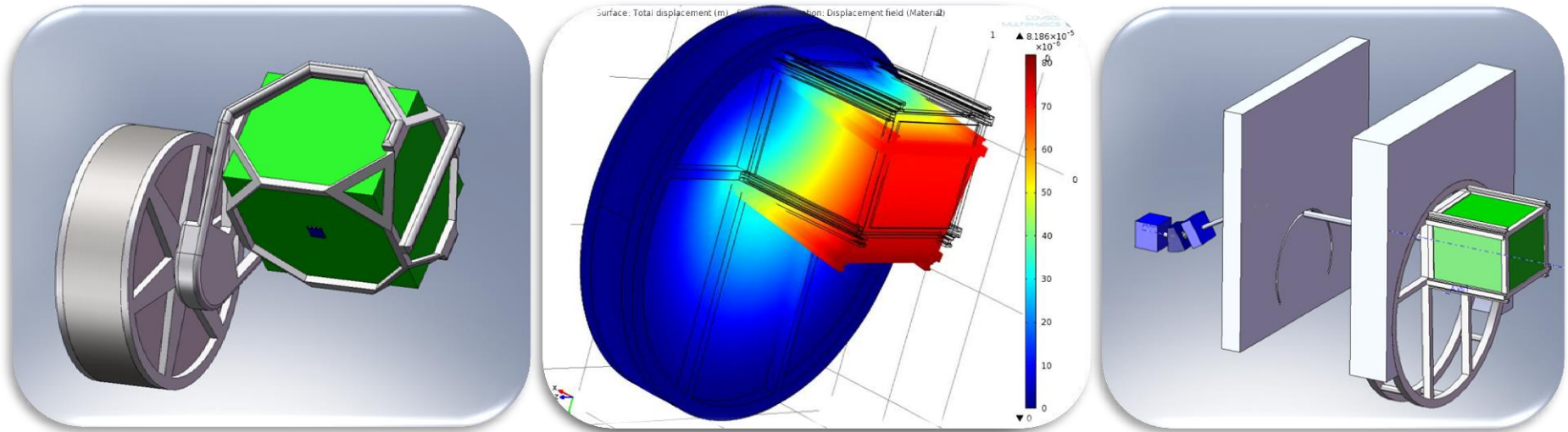
P.Colautti IFMIF/ELAMAT Rzeszów 14-15 April 2016



# BNCT Microdosimetric Spectra

## Sviluppo di gantry rotante (2)

- Da simulazioni preliminari, il punto meno critico di ancoraggio del moderatore è la parete di calcestruzzo intorno al tubo di passaggio del fascio.
- L'ancoraggio del moderatore alla parete di calcestruzzo in questo punto impone la rotazione di un cilindro di calcestruzzo di 5.5 m di diametro e 1 m di spessore.
- Il peso di tale ciambella è circa 80 t.



- Il peso complessivo degli elementi da muovere è maggiore di 110 tonnellate.
- Il costo di tale sistema rischia di essere eccessivo > 8 M€

# The third pillar: experimental microdosimetric spectrum

