Detectors for online monitoring of ion therapy and imaging

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Problems raised in particle (i.e. proton and light ion) therapy

- particle range control
 - for a treatment fraction or
 - in real time
- in-vivo dosimetry
- beam monitoring (future accelerators)

33% of the voters of an AAPM poll in Aug. 2012 considered particle range uncertainties as the main obstacle for a wide application of particle therapy

ENVISION (2010-2014) + ENTERVISION (2011-2015)

Two CERN-coordinated European collaborative projects































Objectives:

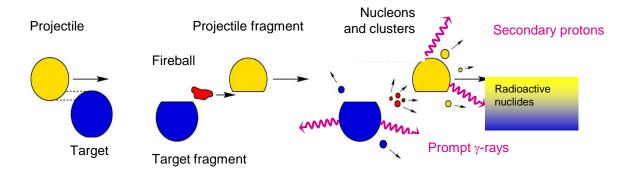
Real-time non invasive monitoring, quantitative imaging, precise determination of delivered dose. fast feedback for optimal treatment planning, real-time response to moving organs, simulation studies.

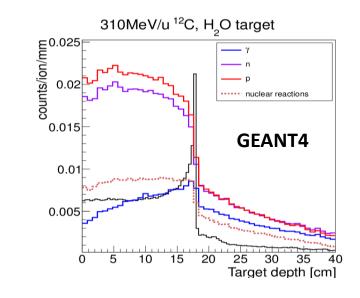
Imaging solutions: ToF-PET, Prompt radiations

Leading aspects

- Basic measurements and modelling
- Novel detectors/electronics developments
- Software (simulations, image reconstruction)
- Clinical constraints (moving organs)
- Clinical feedback and relevance
- Translationnal aspects
- Education and dissemination

Rationale: Secondary particle imaging





| Production / projectile (15 cm range) | | | |
|---|---------|--------------------|--|
| | Protons | Carbon ions | |
| Prompt-γ | 0.05 | 1 | |
| $\beta^{\scriptscriptstyle +}$ emitters | 0.05 | 0.25 | |
| Protons | - | 3 | |

| | Protons | Carbon ions |
|--------------------------|-------------------------------------|------------------------------------|
| Ions /treatment fraction | 10 ¹¹ - 10 ¹² | 10 ⁹ - 10 ¹⁰ |
| Ions/pencil spot | 10 ⁸ (distal) | 10 ⁵ - 10 ⁶ |

Particle therapy positron emission tomography - PT-PET (I)

In-beam, in-room, off-line

Beam 60 s 10 min

Situation:

- Several applications since first clinical proof at the GSI facility
- Technologically well established imaging technique
- Can be implemented on the basis of mature and high quality
 PET components delivered by industry
- Because of the low number of installations in therapy, industry is not willing to perform R&D for in-beam scanners (i.e. integration into a therapeutic beam line)

Problems:

- Degradation of dose-activity-correlation by metabolism increasing from in-beam via in-room to off-line PT-PET
- Low counting statistics
- Missing real time capability

Particle therapy positron emission tomography - PT-PET (II)

Research needed:

for

- Very fast detectors and acquisition systems of
 - system, not detector time resolution well below 100 ps (FWHM)
 - direct time-of-flight reconstruction and real time in-beam PT-PET

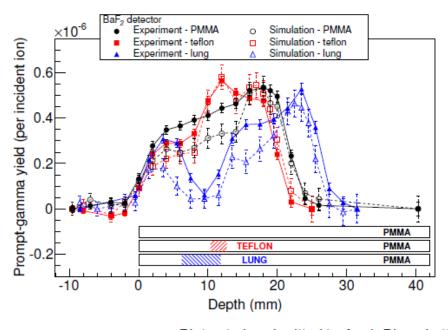
Explicitely, no PT-PET research is needed on:

- PET detectors with comparable properties as those being commercially available
- Improving a factor of 2 or 3 in time resolution, we need a factor of >10
- Organic scintillators, RPC and other low atomic number, low density detectors

Either there is to expect a breakthrough, or the PT-PET community should buy their detectors (tomographs) from companies

Prompt gamma imaging

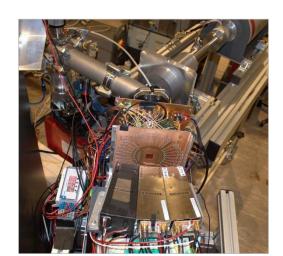
- Collimated gamma cameras
 - reach real size prototypes
 - millimetric resolution on range variation
 - Pencil beam scale for protons
 - Energy slice scale for carbon ions
 - TOF: better sensitivity (lower background)
 - Mandatory for C ions



Pinto et al. submitted to Appl. Phys. Lett.

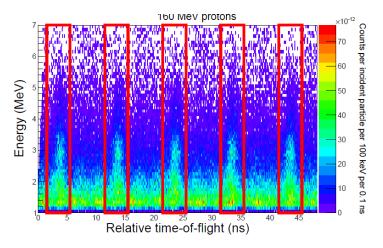
- Compton cameras
 - Small size prototypes
 - Further developments are ongoing
 - Fortuitous coincidence issue

Dresden Compton camera



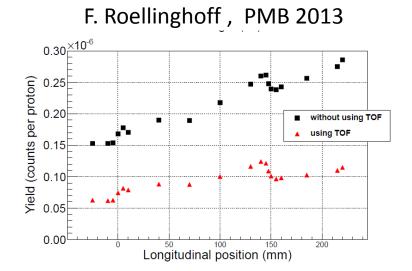
Prompt gamma imaging

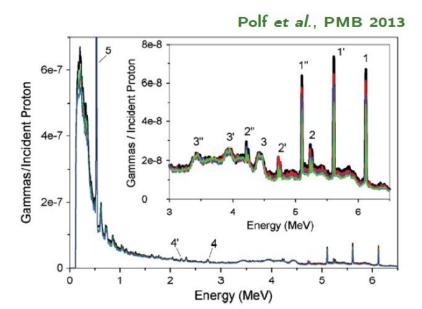
 Beam time structure issue IBA C230, 160 MeV protons



Synchrotrons: beam pulses ~20 ns Ion per ion difficult at 108 pps

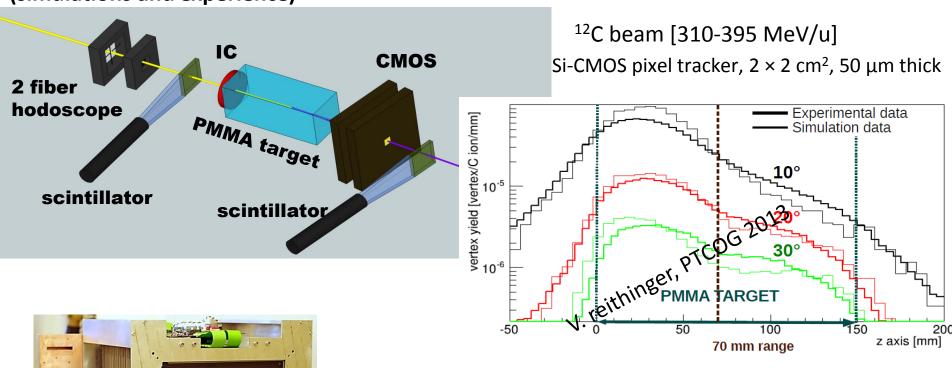
Energy issue
 Chemical composition information
 Localization?





Secondary proton imaging for range verification

Millimetric resolution of range variations in homogenous targets at pencil beam scale (simulations and experience)



TERA: Large Acceptance tracker (GEM) + proton range (plastic scintillator stack) : PRR30 under test 30 × 30 cm², high rate (~ 10⁶ s⁻¹)

Patient dependent calibration (multiple scattering)

Other methods of particle range assessment

Acoustic signals

- Several experimental attempts since 1979
- Potential for clinical application unclear
- → Proof of principle at therapeutic beams with different time characteristics

Radiation induced MRI visible signal

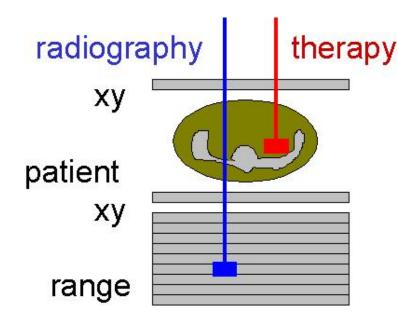
- MR visible signals shown for
 - vertebral bone marrow
 - liver (contrast agent based)
- Time dependence of MR signal appearance under investigation
- → Feasibility of transfer to other organs and tissues
- → Identification of dose sensitive MRI contrast agents (vision)

In-room scanners with molecular imaging capabilities

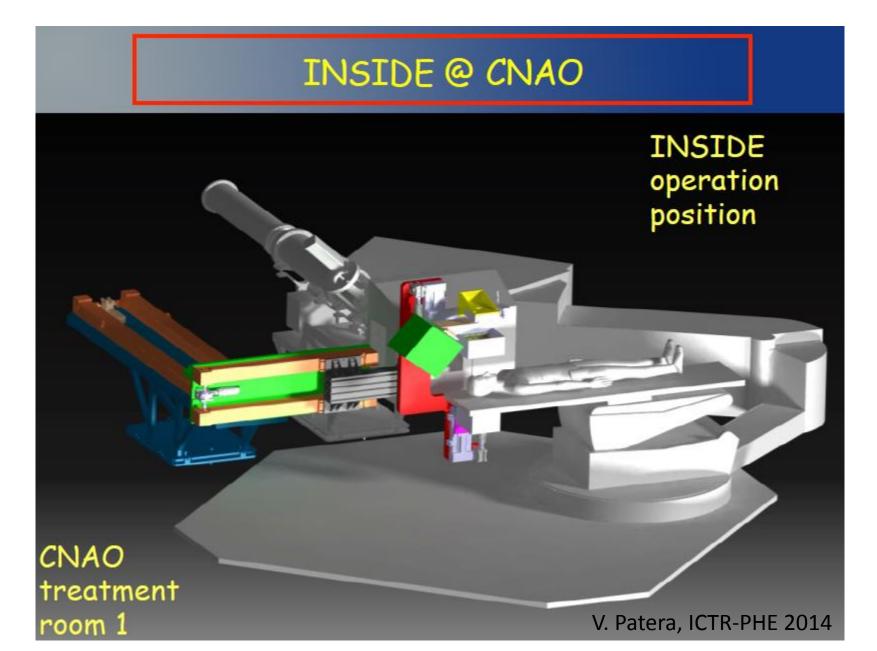
→ Visualisation of biomarkers for biologically-adapted radiotherapy

Particle radiography

- Direct measurement of dE/dx
 - TPS verification in-beam
- Several prototypes or projects in Europe (PSI, TERA, Firenze, HIT-Munich...)
- Requires high energy protons or light ions
- Lower dose than X-rays?
- Modelling and experiments are neccesary



Multimodal solutions



Novel in-beam monitors

- Next generation accelerators (laser, DWA, synchrocyclotron)
 - High instantaneous dose rate
 - Mixed fields of different ions
 - Variable LET
- Faster beam scanning
 - Fast readout
- Therapeutic and transmission/absorption verification
 - Large dynamic range
- → New solutions should be envisaged (GEM, diamond...)
 Radiation hardness, stability, cost...

Measurement of nuclear data

- Range verification by means of secondary radiations requires scalable knowledge of production cross sections
- In-vivo dosimetry requires precise calculations from treatment planning
 - Monte-Carlo simulations based on measured or modelled cross sections
 - Feasibility of yield based simulation proven for PT-PET
- → Measurements at LEIR for more accurate Monte Carlo simulations:
- PT-PET: β +-activity yield as a function of: $(A, Z)_{\text{target}}$, $E_{\text{projectile}}$, Range
- PGI: γ -ray emission density as function of: $(A, Z)_{\text{target}}$, $E_{\text{projectile}}$, Range, E_{γ}
- IVI: p,d,t,He production yields as function of: $(A, Z)_{target}$, $E_{projectile}$, Range, E, angle
- Low statistical error (< 1%)
- Dedicated apparatus at a flexible beam of high availability

Requirements for LEIR experiments

Close to clinical beams – and beyond

- Beam intensities: 1pA to 10nA (clinical dose rates)
- Beam energies: above 250 MeV protons?
- Beam time structure:
 - Cyclotron like (1ns pulse every 10-1000 ns)
 - Synchrotron like (20-50 ns pulse microstructure)
- Pencil beam scanning and passive beam delivery?
- Robotic patient positioning system

In the more general field of imaging

- Novel PET technology
- Sub-10 ps electronics
- Digital DAQ based on modern FPGAs for high granularity high connectivity system
- Computing networks for parametric images strategy and biomarkers mining
- New Imaging methodology (e.g., Photoacustic, Cerenkov,..)
- Hybrid imaging methodology
- Very fast detectors for synchrotron radiation biological applications, e.g., dynamic unfolding of proteins and protein crystallography
- and more

Simulations and image processing

It is absolutely mandatory that software and hardware scientists work together.

There is no real/effective detector for medical imaging if there has not been a simulation/design, construction and testing, data acquisition and reconstruction of the images.

CERN has the unique opportunity to provide an environment where these two communities have been for a long time and still are working strictly and productively together, e.g. ATLAS, CMS.

(A. Del Guerra)

Questions

- Should online control adapt to the treatment modality, or should the treatment adapt to the control?
- Which strategy (strategies) to be envisaged?
 - CERN driven research?
 - International collaboration on a single device?
 - Open platform to external users?