

# Proton Computed Tomography

F. Cassol

*Atelier proton CT, CPPM, 14 November 2012*



# Contents

## The Proton Computed Tomography (pCT)

- Where?
- Why?
- How?
- Conclusion

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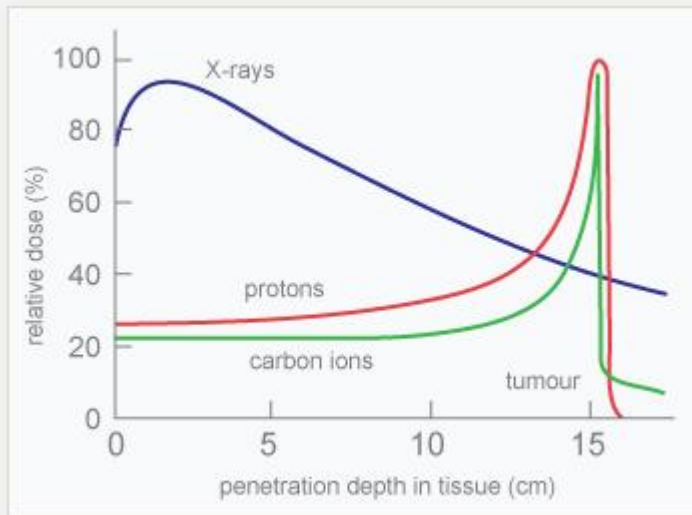
# pCT useful in proton therapy

Advance form of radiotherapy based

on the way protons lose energy in matter

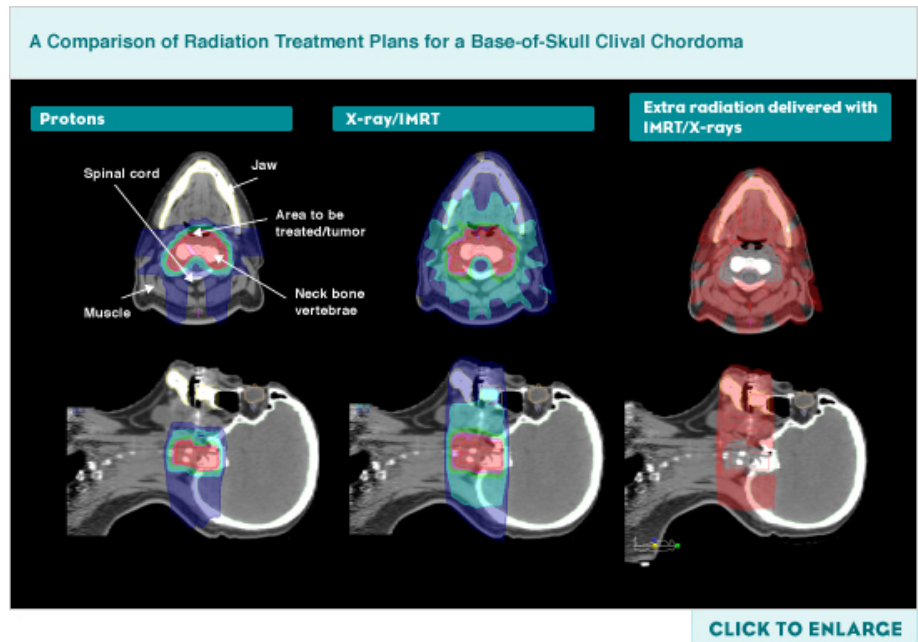
Thanks to the Bragg peak:

- Tumors can be precisely irradiated
- Close sensitive tissues can be avoided



In general:

Dose outside the target volume is reduced of a factor 2-5 compared to photons



# Proton therapy

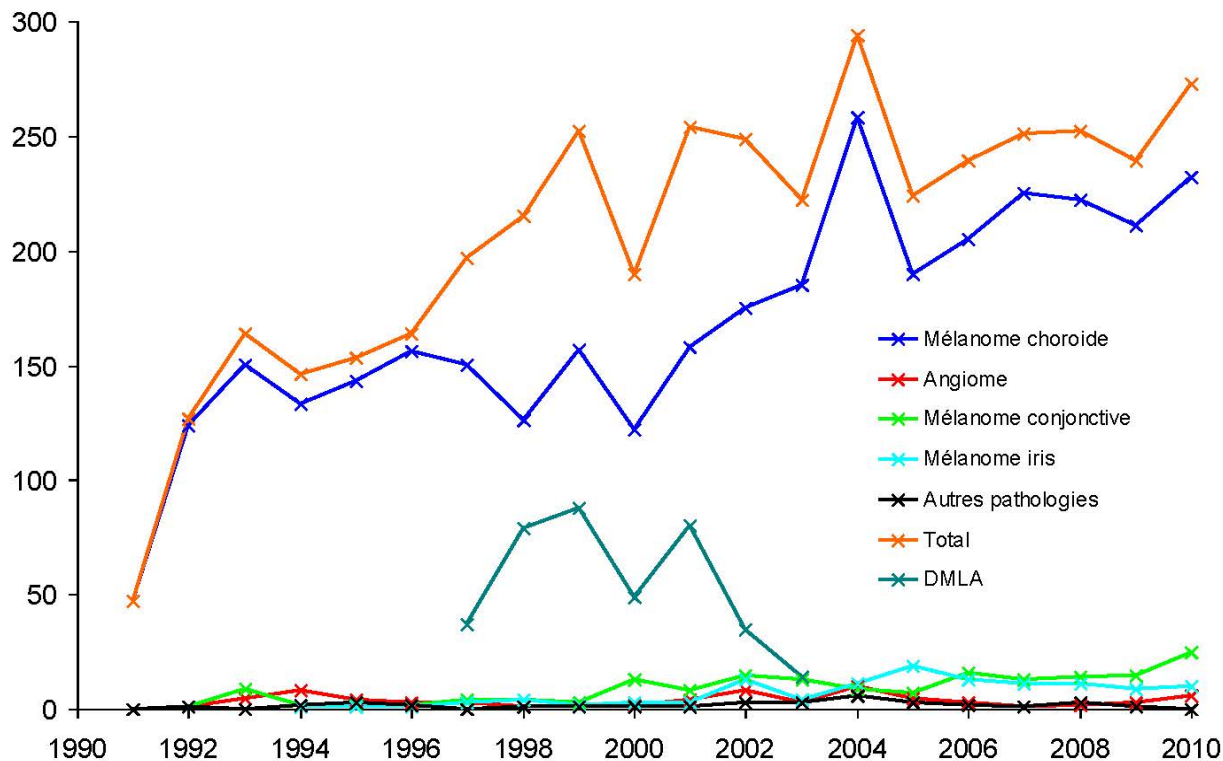
## Good results overall for:

- Cancers that need **high doses**: eye, skull base, spinal tumor
- Cases where **other tissues** must be **preserved**: pediatric tumors
- For several cases the usefulness of PT is still a controversy
  
- Several centers from the '90-'00:
  - Usa, Japan, Germany, France, Italy, Russia
- In France: Nice (1991) and Orsay (1991)

# Centre Antoine Lacassagne Nice

Types de pathologies traitées **France HADRON**

Lyon, 28-29 Novembre 2017



IRSN

INSTITUT  
CLAUDIUS REGAUD

CAL  
Nice

Centre ETOILE  
Département de Cardiologie

Archade  
HÔPITAL PASTEUR - NICE

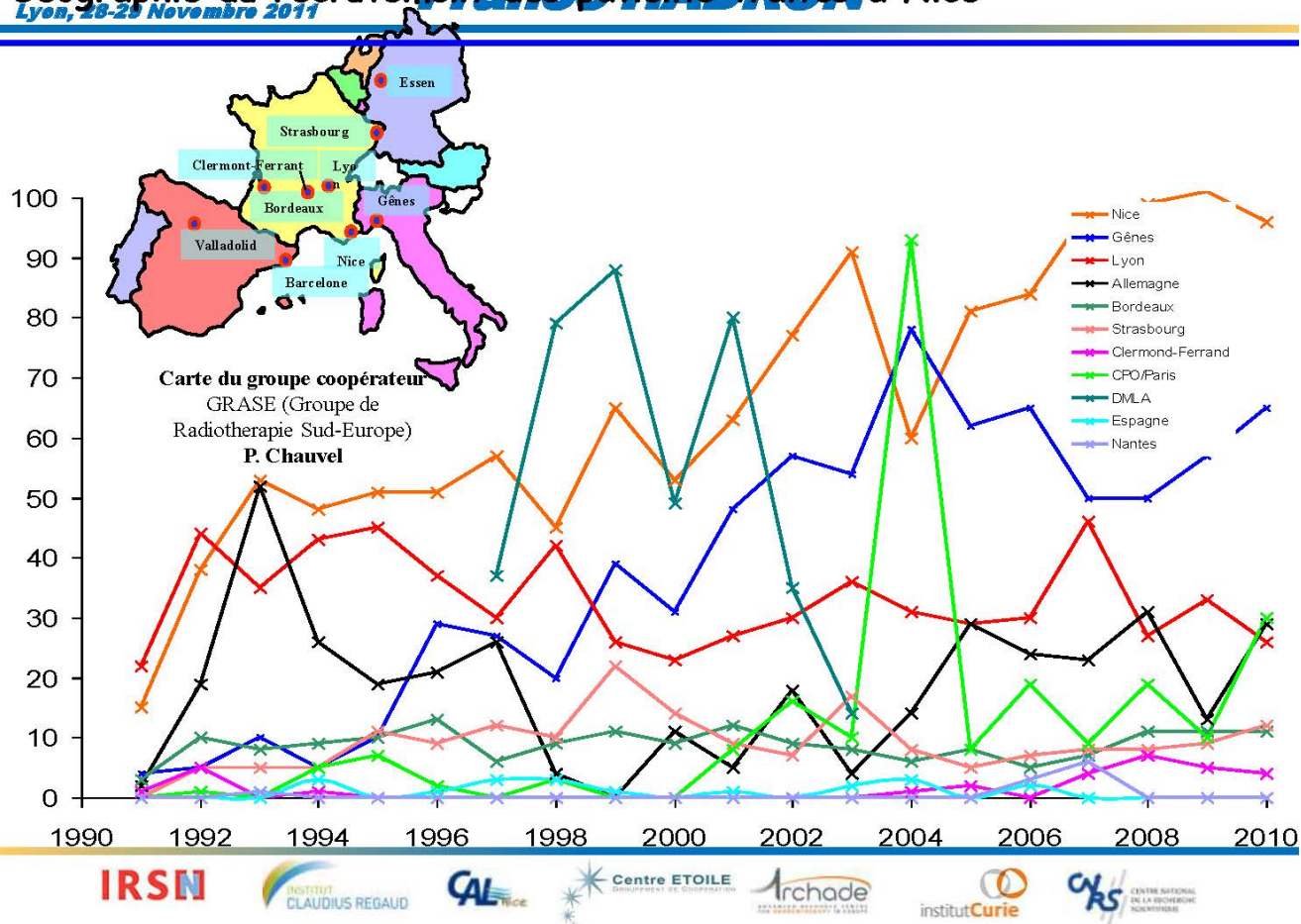
institut Curie

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CENTRE NATIONAL  
DE LA RECHERCHE  
SCIENTIFIQUE

# Centre Antoine Lacassagne Nice

## Géographie du recrutement des patients traités à Nice

Lyon, 28-29 Novembre 2011



Patient Statistics (for the facilities in operation end of 2011):

| WHERE        |                             | PARTICLE | FIRST PATIENT | PATIENT TOTAL | DATE OF TOTAL |                     |
|--------------|-----------------------------|----------|---------------|---------------|---------------|---------------------|
| Canada       | Vancouver (TRIUMF)          | p        | 1995          | 161           | Dec-11        | ocular tumors only  |
| China        | Wanjie (WPTC)               | p        | 2004          | 1078          | Dec-11        | no patients in 2011 |
| China        | Lanzhou                     | C ion    | 2006          | 159           | Dec-11        |                     |
| England      | Clatterbridge               | p        | 1989          | 2151          | Dec-11        | ocular tumors only  |
| France       | Nice (CAL)                  | p        | 1991          | 4417          | Dec-11        | ocular tumors       |
| France       | Orsay (CPO)                 | p        | 1991          | 5634          | Dec-11        | 4540 ocular tumors  |
| Germany      | Berlin (HMI)                | p        | 1998          | 1859          | Dec-11        | ocular tumors only  |
| Germany      | Munich (RPTC)               | p        | 2009          | 895           | Dec-11        |                     |
| Germany      | HIT, Heidelberg             | C ion    | 2010          | 568           | Dec-11        |                     |
| Germany      | HIT, Heidelberg             | p        | 2010          | 94            | Dec-11        |                     |
| Italy        | Catania (INFN-LNS)          | p        | 2002          | 290           | Dec-11        | ocular tumors only  |
| Italy        | Pavia (CNAO)                | C ion    | 2011          | 5             | Dec-11        |                     |
| Japan        | Chiba (HIMAC)               | C ion    | 1994          | 6569          | Dec-11        | 11 with scanning    |
| Japan        | Kashiwa (NCC)               | p        | 1998          | 870           | Dec-11        | estimated           |
| Japan        | Hyogo (HIBMC)               | p        | 2001          | 3198          | Dec-11        |                     |
| Japan        | Hyogo (HIBMC)               | C ion    | 2002          | 1271          | Dec-11        |                     |
| Japan        | Tsukuba (PMRC, 2)           | p        | 2001          | 2166          | Dec-11        |                     |
| Japan        | Shizuoka                    | p        | 2003          | 1175          | Dec-11        |                     |
| Japan        | Koriyama-City               | p        | 2008          | 1378          | Dec-11        |                     |
| Japan        | Gunma                       | C ion    | 2010          | 271           | Dec-11        |                     |
| Japan        | Ibusuki (MMRI)              | p        | 2011          | 180           | Dec-11        |                     |
| Korea        | Ilsan, Seoul                | p        | 2007          | 810           | Dec-11        |                     |
| Poland       | Krakow                      | p        | 2011          | 11            | Dec-11        | ocular tumors only  |
| Russia       | Moscow (ITEP)               | p        | 1969          | 4300          | Dec-11        | estimated           |
| Russia       | St. Petersburg              | p        | 1975          | 1372          | Dec-11        |                     |
| Russia       | Dubna (JINR, 2)             | p        | 1999          | 828           | Dec-11        |                     |
| South Africa | iThemba LABS                | p        | 1993          | 521           | Dec-11        |                     |
| Sweden       | Uppsala (2)                 | p        | 1989          | 1185          | Dec-11        |                     |
| Switzerland  | Villigen PSI, incl OPTIS2   | p        | 1996          | 1107          | Dec-11        | 277 ocular tumors   |
| USA, CA.     | UCSF - CNL                  | p        | 1994          | 1391          | Dec-11        | ocular tumors only  |
| USA, CA.     | Loma Linda (LLUMC)          | p        | 1990          | 16000         | Dec-11        | estimated           |
| USA, IN.     | Bloomington (IU Health PTC) | p        | 2004          | 1431          | Dec-11        |                     |
| USA, MA.     | Boston (NPTC)               | p        | 2001          | 5562          | Oct-11        |                     |
| USA, TX.     | Houston (MD Anderson)       | p        | 2006          | 3400          | Feb-12        |                     |
| USA, FL      | Jacksonville (UFPTI)        | p        | 2006          | 3461          | Dec-11        |                     |
| USA, OK.     | Oklahoma City (ProCure PTC) | p        | 2009          | 623           | Dec-11        |                     |
| USA, PA.     | Philadelphia Upenn          | p        | 2010          | 433           | Dec-11        |                     |
| USA, IL.     | CDH Warrenville             | p        | 2010          | 367           | Dec-11        |                     |
| USA, VA.     | Hampton (HUPTI)             | p        | 2010          |               |               | no data available   |
|              |                             |          |               | 77191         | Total         |                     |



# Contents

## The Proton Computed Tomography (pCT)

- Where?
- Why?
  - Positioning of the patient
  - Measurement of the proton energy loss
- How?
- Conclusion

# Positioning of the patient

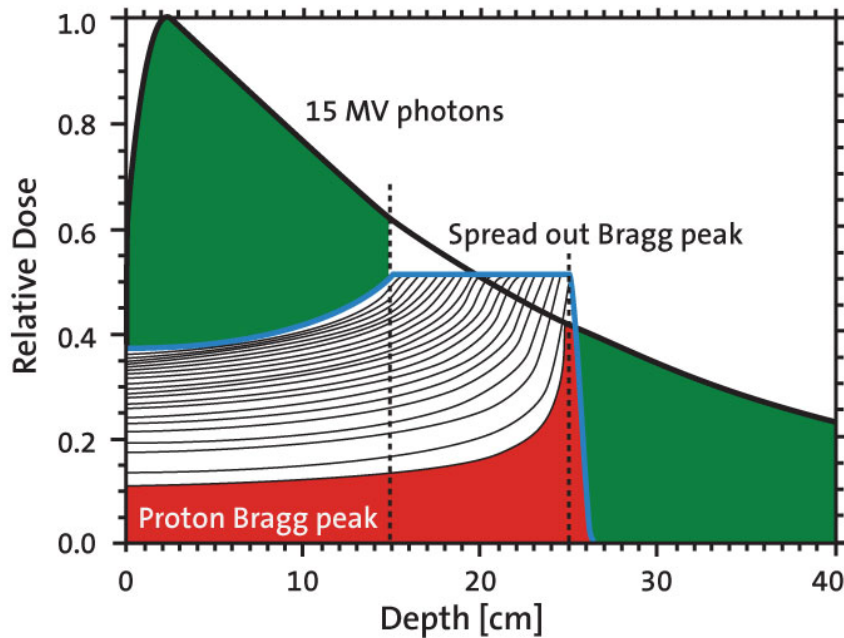
At present the positioning of the tumor is done in with X-radiographies compared with the XCT used for the treatment planning

pCT would permit to directly 3D locate the tumor with the same beam that it would be used for the treatment

Which is the present uncertainty?  
How much are we going to improve?

# Measurement of the proton energy loss

Proton Therapy is successful only if  
the p energy loss in the patient is precisely known



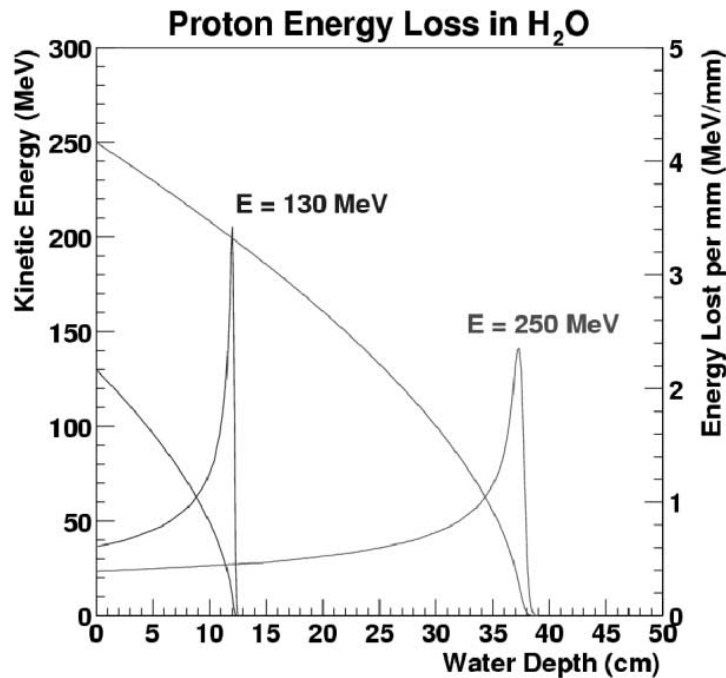
The beam energy is modulated  
in order to :

1. cover the tumor
2. save the closed critical tissue

# The proton energy loss (stopping power)

Protons lose most of their energy with the inelastic collisions with the outer atomic electrons (ionizations and excitations).

The **Bethe-Bloch theory** describes the proton stopping power:



$$-\frac{dE}{dx}(\bar{r}) = \eta_e(\bar{r}) S(I(\bar{r}), E(\bar{r}))$$

$$S(I, E) = K \frac{1}{\beta^2} \left[ \ln \left( \frac{2m_e c^2}{I} \frac{\beta^2}{1 - \beta^2} \right) - \beta^2 \right]$$

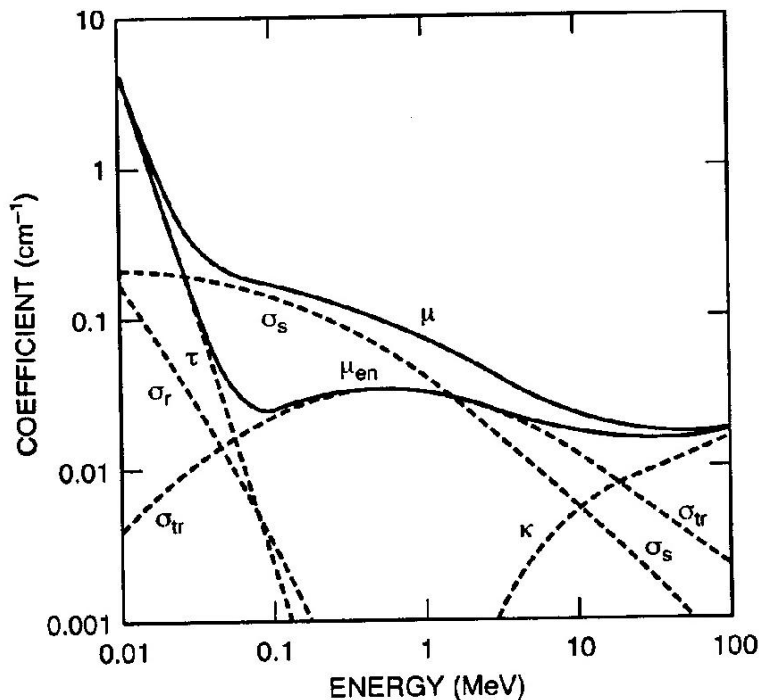
$$\eta_e(r) = \rho_e / \rho_{e, \text{water}} = \text{relative electron density}$$

$$I(\bar{r}) = \text{mean excitation potential} (= 75 \text{ eV for water})$$

$$K = 4\pi r_e c^2 \rho_{e, \text{water}} = 0.170 \text{ MeV/cm}$$

# At present, electron density derived from XCT

XCT measures the **attenuation coefficient of X-rays**  
which also depends from the electron density.



$$N(l) = N_0 e^{-\mu l}$$

$$\mu = \rho_e^\gamma \left( \sigma^{ph} + \sigma^{coh} + \sigma^{incoh} \right)$$

XCT gives HU  
we need the calibration :

$$HU \Rightarrow \eta_e^\gamma$$

$$HU \Rightarrow \eta_e$$

# Calibration for radiotherapy from XCT

Phantoms with known materials are used to estimate

$$\eta_e^\gamma$$

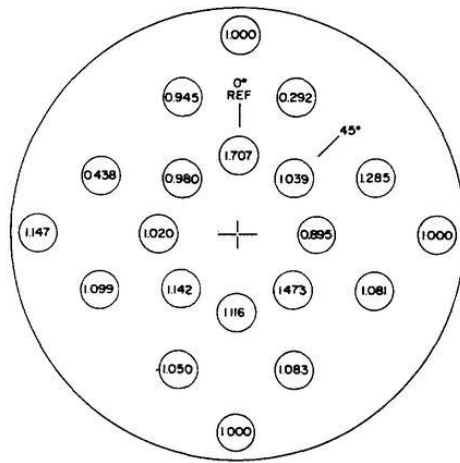
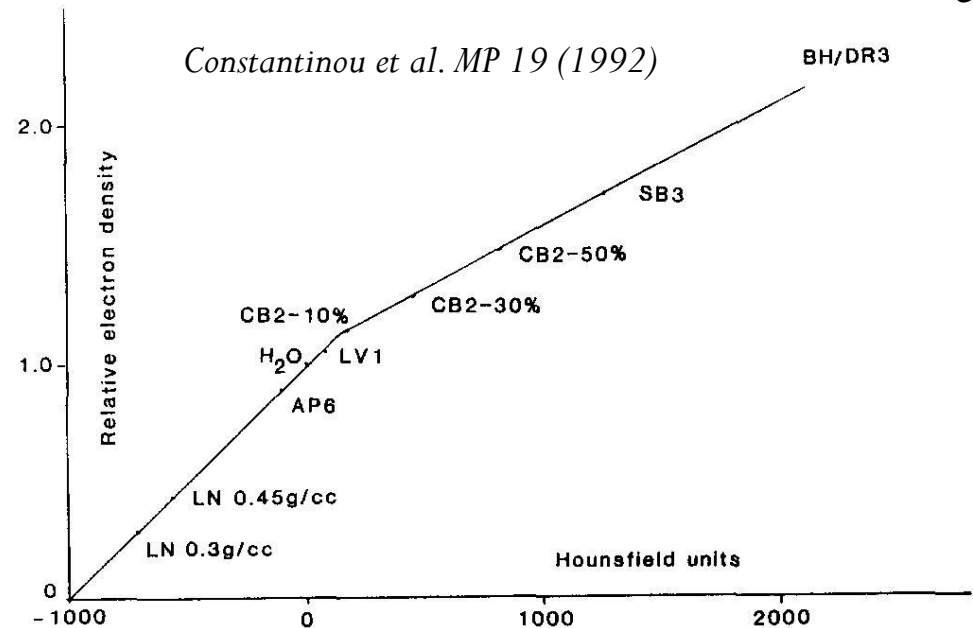


FIG. 1. The 33-cm-diam electron density calibration phantom. The numbers represent electron density relative to water for each rod. (Manufactured by RMI).

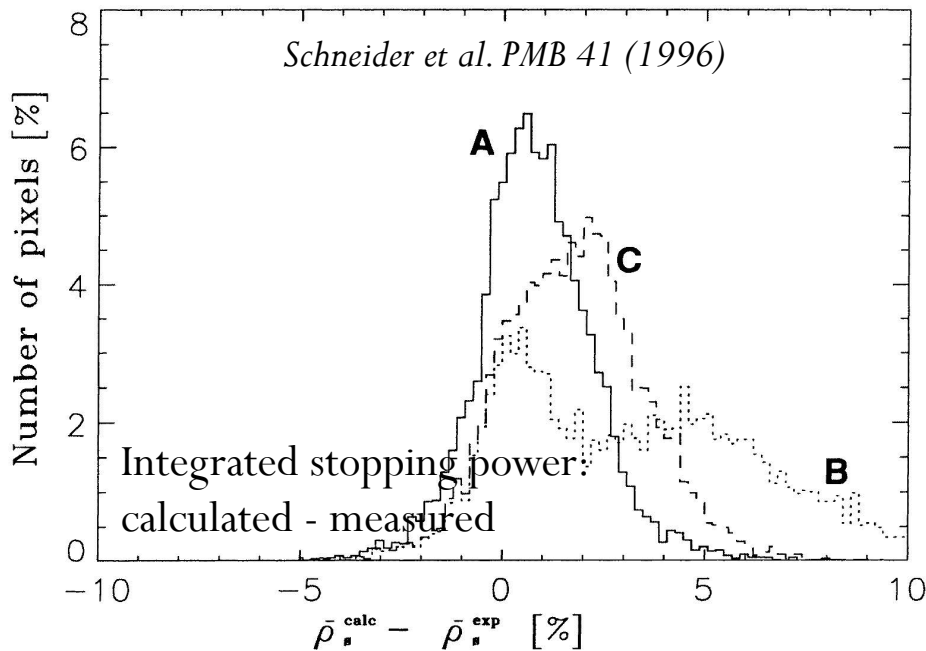


**XCT has several sources of uncertainty :** (Schneider et al. PMB 41 1996)

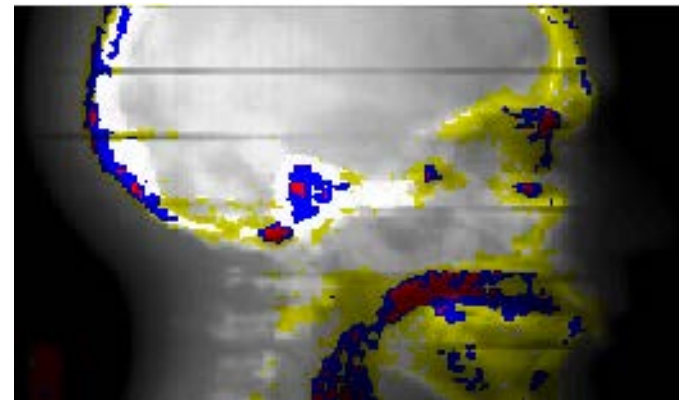
- HU variations of 1-2% in homogenous materials
- HU variations till 3% as function of the position
- HU variation of 10% as function of the scanner
- Errors due to the approximation real tissues/substitute tissues

# Calibration of stopping power for pCT from HU

Different methods = different results



Range uncertainty in inhomogeneous densities due to Coulomb scattering



*Schneider et al. MP 22 (1995)*

The **Bragg peak position** is predicted to only 3-4% of the proton range in tissue or less in complicated tissue-air tissues-bone interfaces

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  - The method
  - Protons interactions, more precisely
  - Examples of pCT designs
- Conclusion



# The method

Main goal of pCT : to determine the volume electron density by measuring the energy loss of protons after traversing the object

Proton stopping power

$$-\frac{dE}{dx} = \eta_e \cdot S(I, E)$$

Volume electron density

$$\eta_e = -\frac{1}{S(I, E)} \frac{dE}{dx}$$

Integral over the proton path

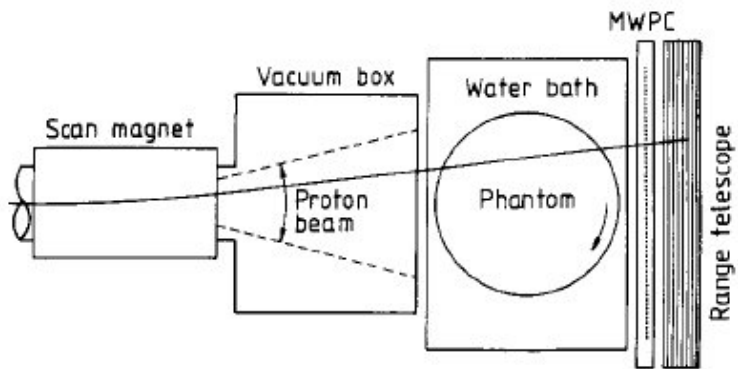


Radon transform of  $\eta$

$$\int_L \eta_e dx = \int_L -\frac{1}{S(I, E)} \frac{dE}{dx} dx = -\int_{E_{in}}^{E_{out}} \frac{1}{S(I, E)} dE$$

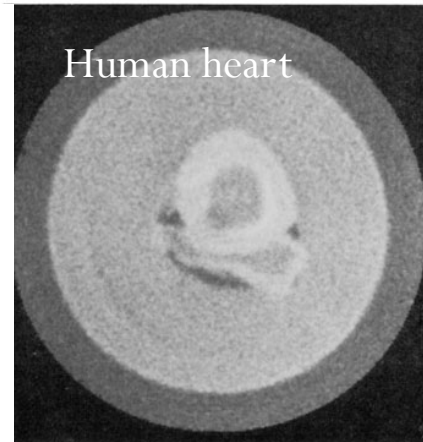
# First trails in the '80

Assume a **straight path L** and **only  $E_{out}$ ,  $x_{in}$  and  $x_{out}$**  measured

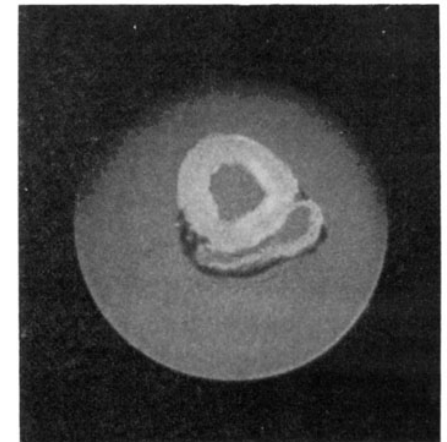


**Figure 1.** Schematic layout of apparatus.

*Hanson et al. MPB 27 (1982)*



pCT



XCT

**Results are deceiving** with respect to XCT, **loss of interest for pCT**

But more and more proton therapy centers,  
in the 90' people try to do better ...

# pCT, more precisely

In pCT, **proton energy** sufficient **to traverse the body**

- 200 MeV ( $R=25.8$  cm) for adult skull (20 cm)
- 250 MeV ( $R=37.7$  cm) for adult trunk (34 cm)

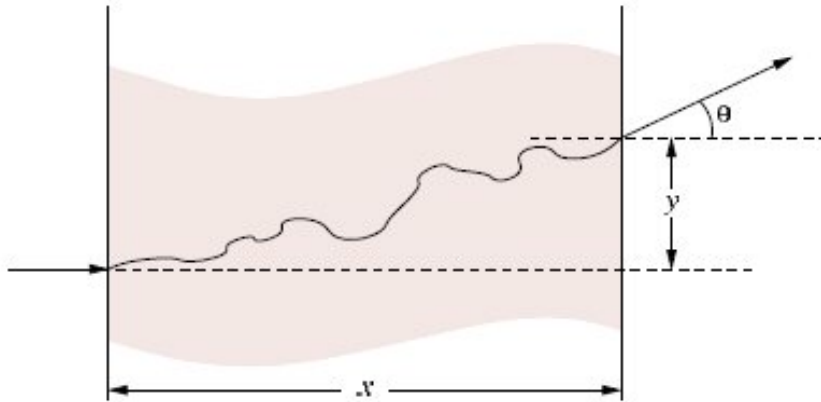
**Reconstruction** is made **track by track** (list mode)

Three phenomena define **the intrinsic limitation of pCT**:

1. Coulomb scattering → limiting spatial resolution
2. Energy loss straggling → limiting electron density resolution
3. Nuclear interactions → noise and additive dose

# Multiple-Coulomb scattering

Protons undergo many individual **elastic interactions** that **change their final direction and position**



$$\sigma_{\Theta} \propto \frac{\sqrt{x / X_0}}{v^2} \propto \frac{Z}{v^2} \sqrt{x(\text{cm})}$$

$$\sigma_y = \frac{1}{\sqrt{3}} x \sigma_{\theta}$$

**Protons don't follow straight lines!**

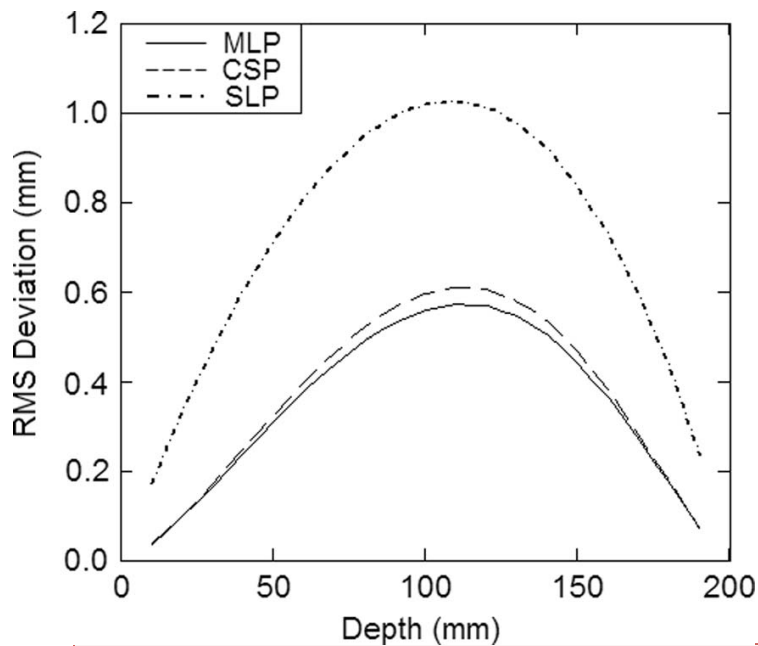


pCT reconstruction must include  
a mathematical formalism to take into account MCS

# Most Likely Path (MLP)

Algorithm based on :

- proton **position**, **energy** and **direction**
- modeling of MCS

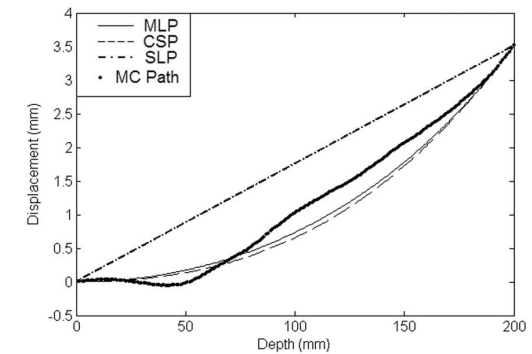
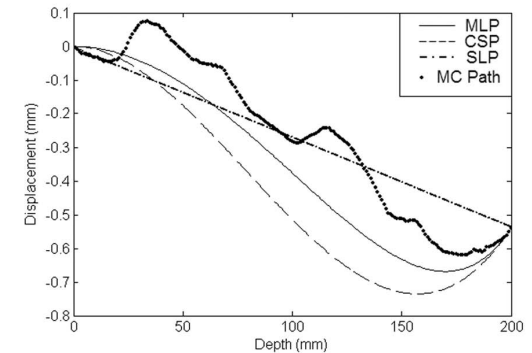
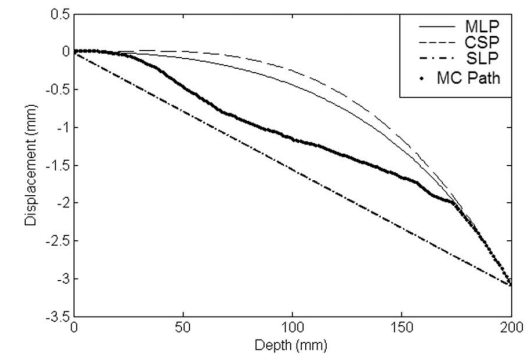


*Shulte et al. MP 21 (1995)*

*Williams PMB 49 (2004)*

*Li et al. MP 33 (2006)*

MLP = Most Likely Path  
CSP = Cubic Spline Path  
SLP = Straight Line Path

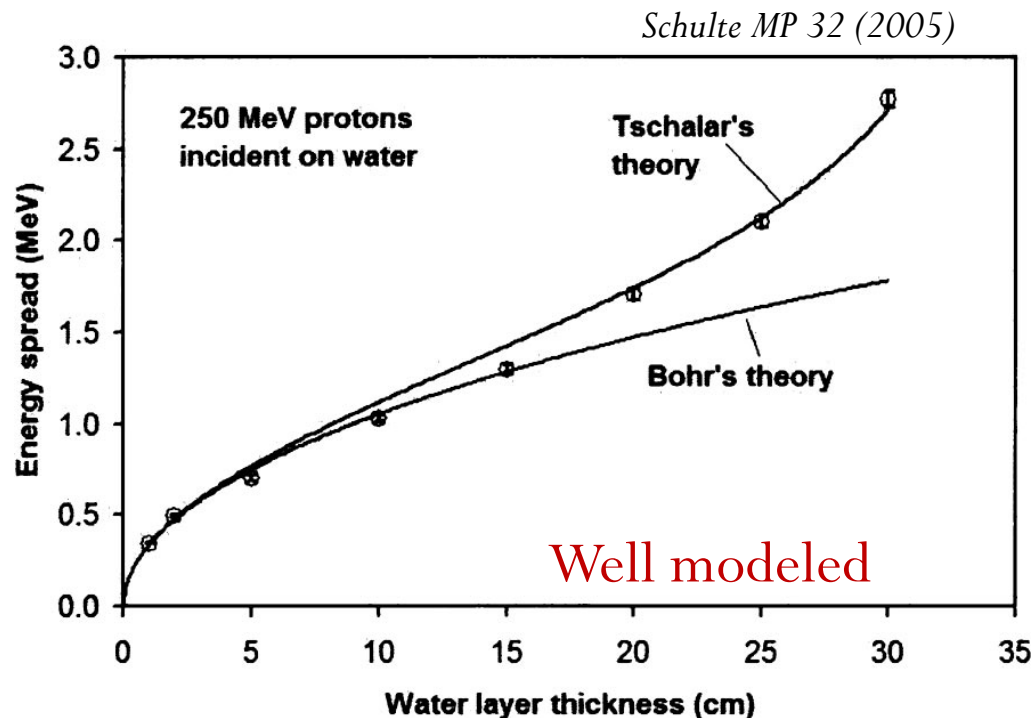


**MCS limits intrinsically the pCT spatial resolution to  $\sim 1$  mm**

# Energy loss straggling

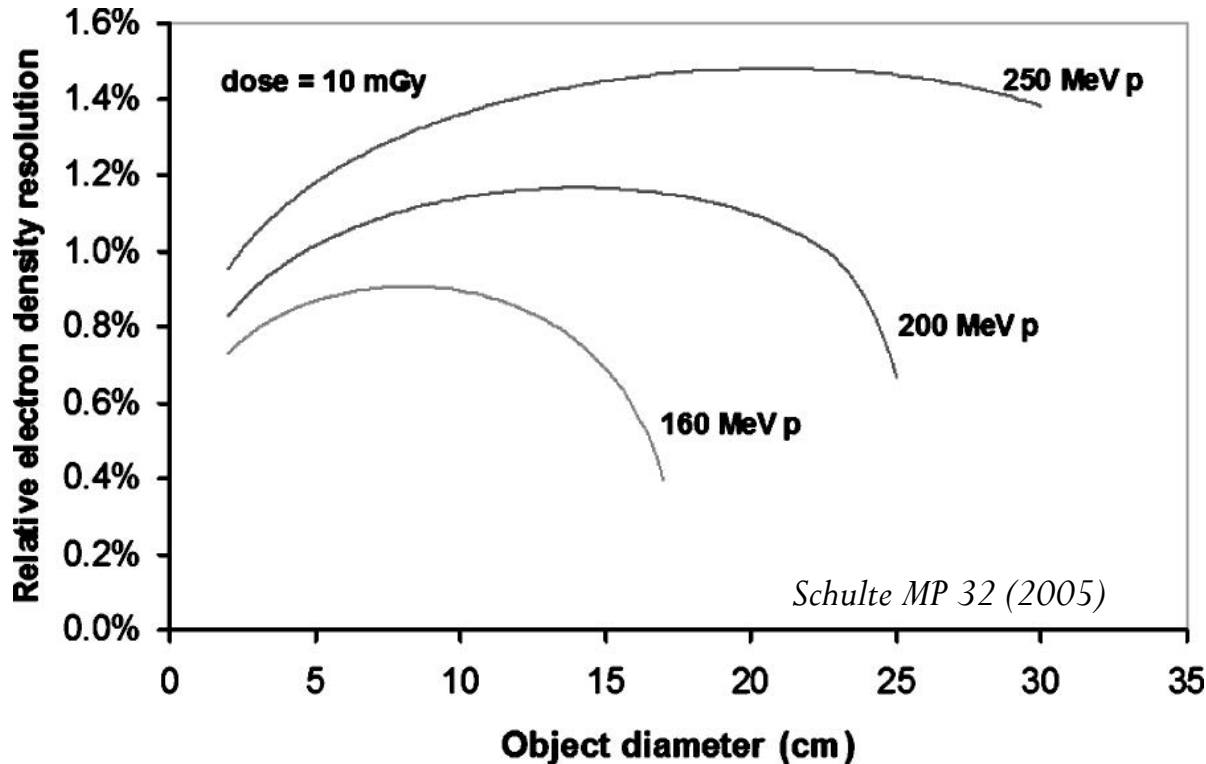
Is due to :

- the varying number of collisions
- the energy transfer fluctuations



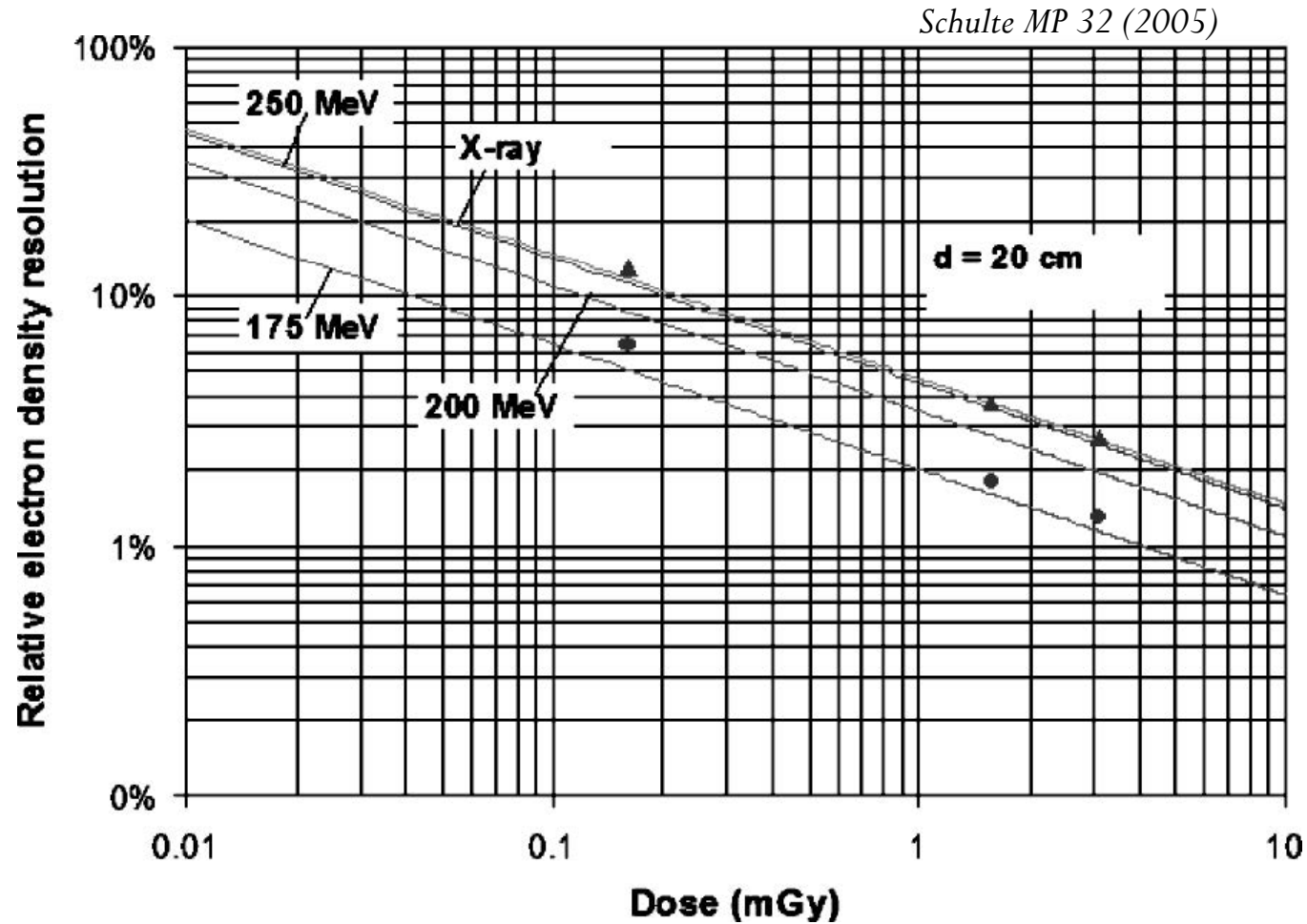
# Energy loss straggling

$$\sigma_{\eta_e} = \frac{\pi \sigma_{E_{out}}}{\sqrt{D \cdot F(S(E_{out}), d, d_{voxel})}}$$



Energy straggling limits intrinsically the measure of  $\eta_e$  to  $\sim 1\%$

# Energy loss straggling



pCT seems to be potentially better than XCT at  $E < 250$  MeV

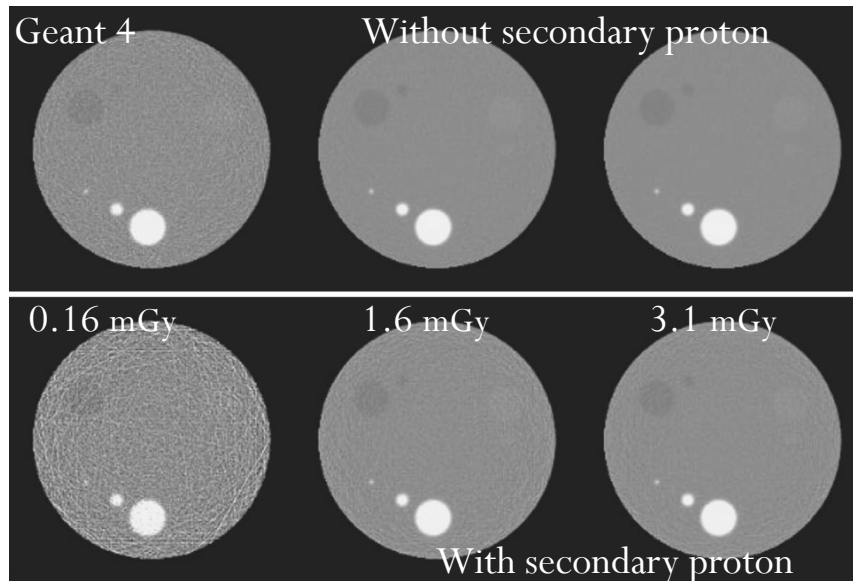


# Non elastic-nuclear interactions

Loss of the primary proton and reduction of the p fluence

$$\Phi(x) = \Phi_0 \exp(-kx) \quad \begin{array}{l} k \sim 0.01 \text{ cm}^{-1} \\ 100 \text{ MeV} < E < 300 \text{ MeV} \end{array}$$

~10% (~20%) reduction after 10 (20) cm water



These protons induce noise in pCT,  
They are eliminated with  $3\sigma$  cut

Which error in the treatment plan?

# pCT design: summary

| Category        | Parameter                           | Value  |
|-----------------|-------------------------------------|--|
| Proton source   | Energy                              | ≈200 MeV (head)<br>≈250 MeV (trunk)                          |
|                 | Energy spread                       | ≈ 0.1%   |
|                 | Beam intensity                      | $10^3 - 10^7$ protons/sec                                    |
| Accuracy        | Spatial resolution                  | < 1 mm   |
|                 | Electron density resolution         | < 1%   |
| Time Efficiency | Installation time                   | < 10 min   |
|                 | Data acquisition time               | < 5 min  |
|                 | Reconstruction time                 | < 15 min (treatment planning)<br>< 5 min (dose verification) |
| Reliability     | Detector radiation hardness         | > 1000 Gy<br>< 1%  |
|                 | Measurement stability               |  |
| Safety          | Maximum dose per scan               | < 5 cGy  |
|                 | Minimum distance to patient surface | 10 cm  |



Measure of  $x$ ,  $p$ ,  $E$  with  
 $\sigma_x < 1\text{mm}$   $\sigma_E < 1\%$



**MHz DAQ :**  
 A head with 100 p, 1 mm voxel  
 $7 \cdot 10^8$  p: 10 kHz = 20 hrs  
 2 MHz = 6 min  
**GPU reconstruction**

*Schulte TNS 51 (2004)*

# Present designs

| Group                              | Tracker                      | Energy detector              |
|------------------------------------|------------------------------|------------------------------|
| Firenze/LNS (Italy)                | Silicon strip detector       | YAG:Ce crystals              |
| LLU-UCSC-NIU (USA)                 | Silicon strip detector       | CsI crystals                 |
| NIU/FNAL (USA)                     | Scintillating Fibers+ SiPM   | Range + WLSF+ SiPM           |
| TERA/CERN (Italy)                  | Gas electrons multipliers    | Range + WLSF+ SiPM           |
| GSI/HIT (Germany): Ion radiography | Stack of Ionisation chambers | Stack of ionisation chambers |

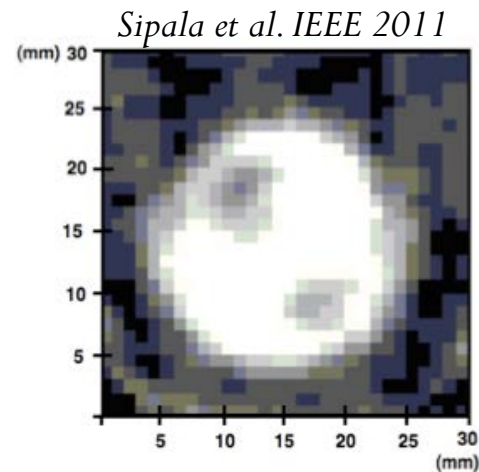
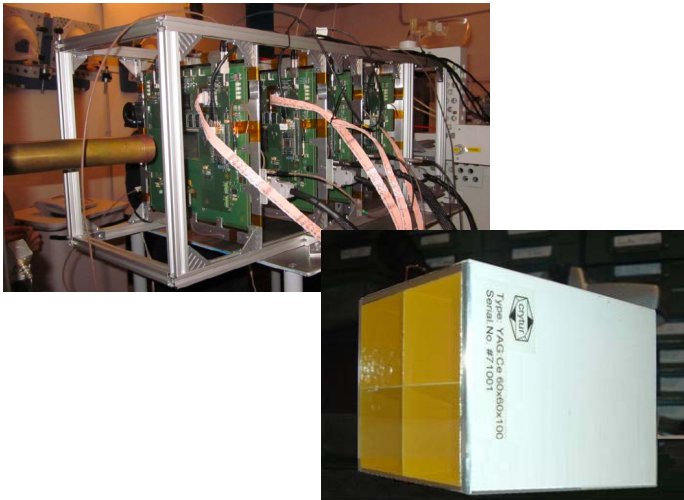
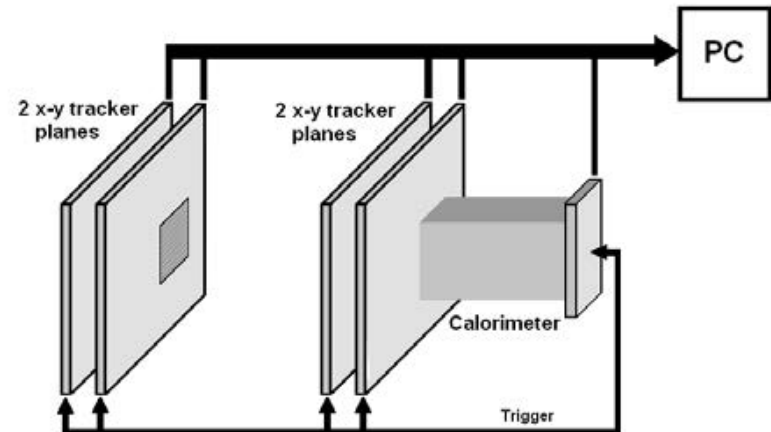
# PROton IMAGING (PRIMA) Firenze/LNS

## Tracker:

- Silicon strips, 200  $\mu\text{m}$  pitch
- Active area 51 x 51  $\text{mm}^2$
- RAM for  $10^6$  events

## Calorimeter:

- 4 scint. crystals 30 x 30  $\text{mm}^2$
- 4 PM Hamamatsu 1.8 x 1.8  $\text{mm}^2$
- 1 MHz



First tomographic image  
with 60 MeV p, 1  $\text{mm}^2$  voxel, (MLP  
+FDK)

# LLU-UCSC-NIU collaboration

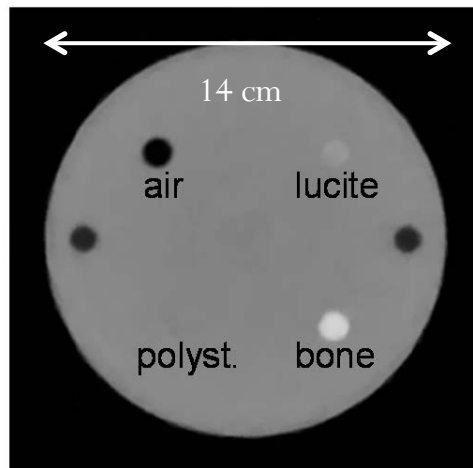
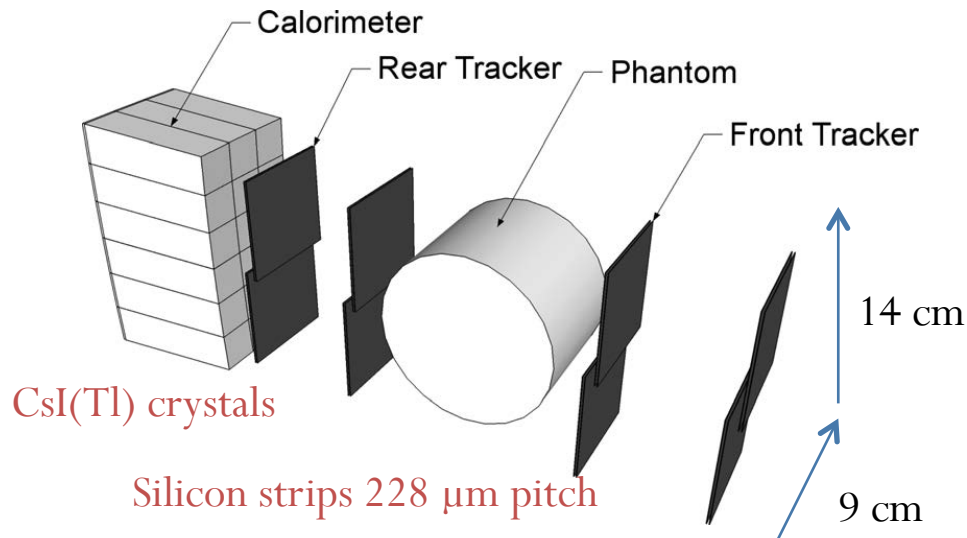


TABLE II. PREDICTED / RECONSTRUCTED RELATIVE STOPPING POWER RSP

| Material    | Predicted RSP | Reconstructed RSP |
|-------------|---------------|-------------------|
| Polystyrene | 1.037         | 1.035             |
| Bone        | 1.70          | 1.68              |
| Lucite      | 1.20          | 1.19              |
| Air         | 0.004         | 0.05              |

Tomographic image 0.65 mm<sup>2</sup> voxel,  
4 hrs at 20 kHz , reconstr. MLP+FDK+ART

*Sadrozinsky et al. IEEE (2011)*

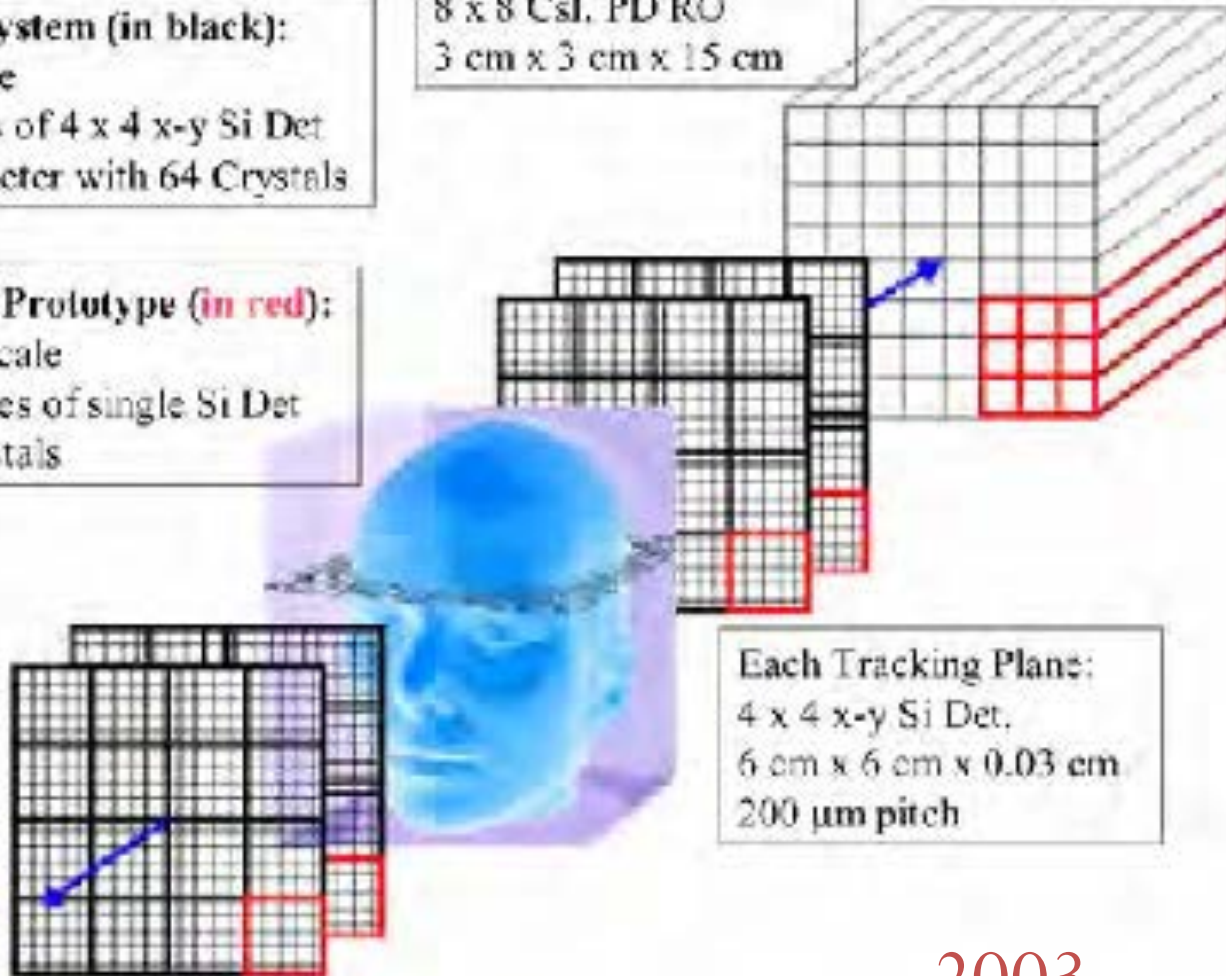
# LLU-UCSC-NIU collaboration

**Final System (in black):**  
full scale  
4 planes of 4 x 4 x-y Si Det  
Calorimeter with 64 Crystals

**Build Prototype (in red):**  
1/16 scale  
4 planes of single Si Det  
9 Crystals

Calorimeter:  
8 x 8 CsI, PD RO  
3 cm x 3 cm x 15 cm

Each Tracking Plane:  
4 x 4 x-y Si Det,  
6 cm x 6 cm x 0.03 cm  
200  $\mu$ m pitch

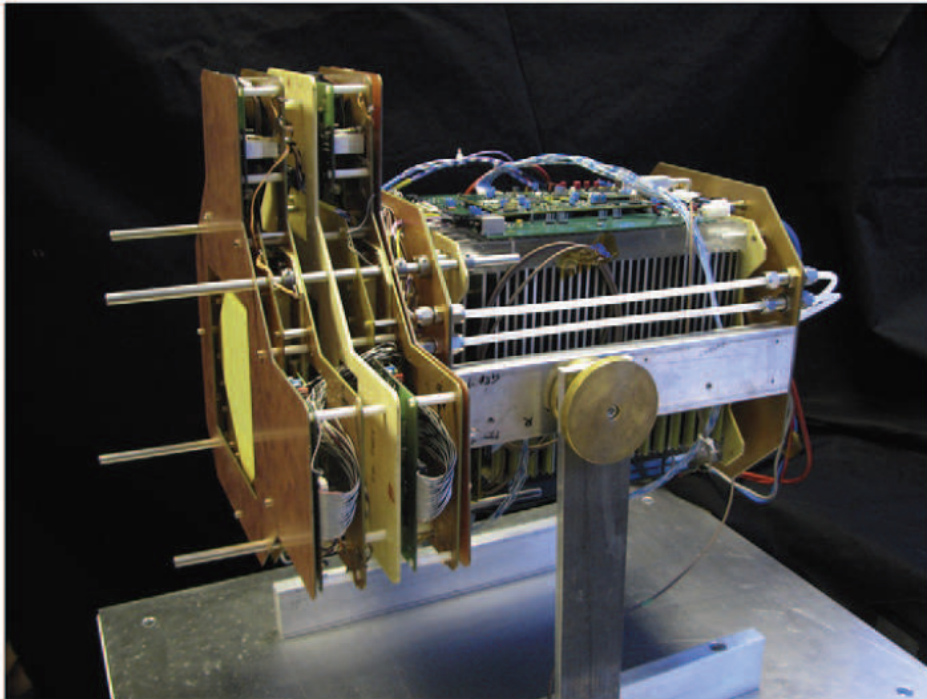


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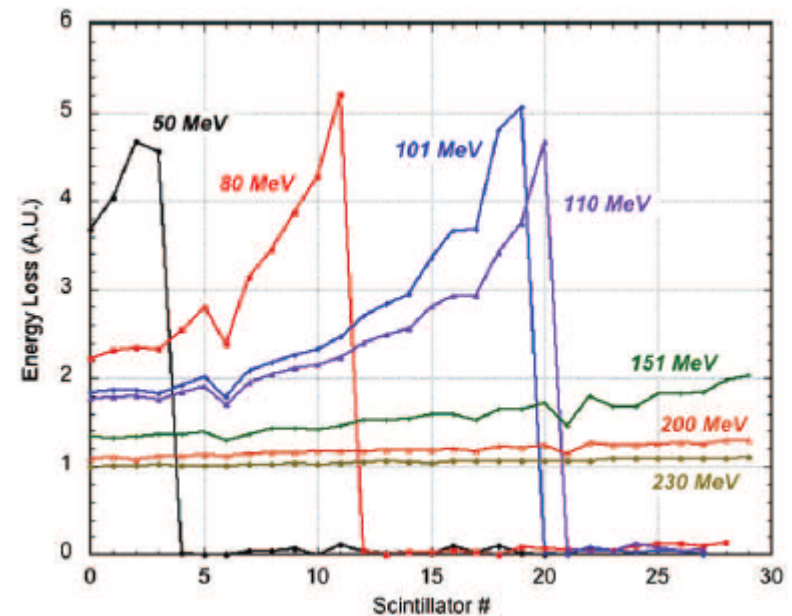
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# Proton Range Radiography (TERA)

F. Sauli et al. (NIMA629 (2011) 337)



- Tracking: 2 GEM detectors
- Range telescope: stack of 30 plastic scint., 3mm thick, read by SiPMs

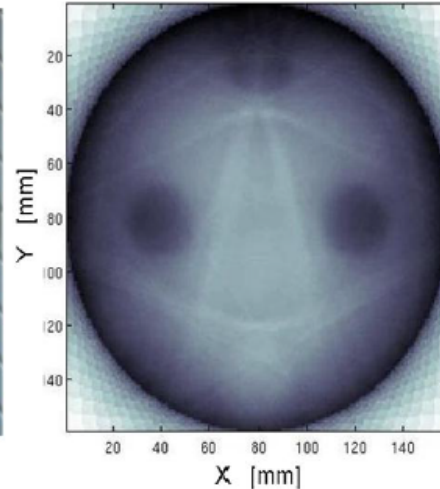
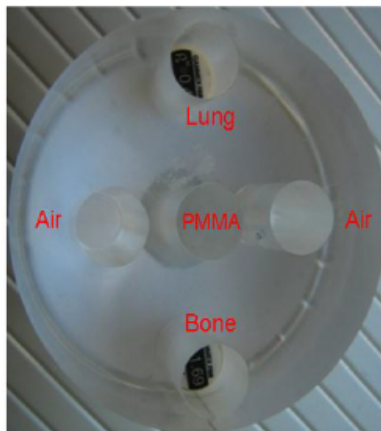
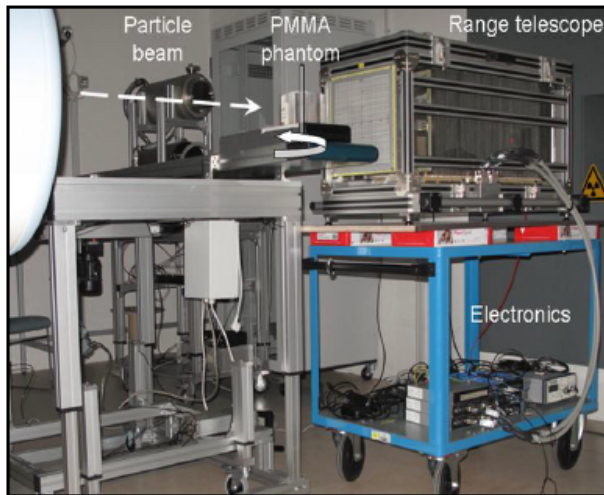


Range resolution : 1.7mm RMS

Expected count rate with suitable acquisition system :  $10^6$  Hz

30x30 cm<sup>2</sup> surface easily achievable

# Towards ion radiography / tomography at HIT



## Stack of ionization chambers (Voss et al, GSI) with new electronics

- Scanning  $0^\circ$ - $180^\circ$  in steps of  $5^\circ$   
 $^{12}\text{C}$  pencil-beam 400 MeV/u  
3.5 mm Gaussian FWHM  
 $5 \times 10^6$  pps
- PMMA phantom  $D=160$  mm  
tissue equivalent rods  $d=28$ mm
- Multi-channel electrometer  
electronics highly integrated
- Simple 2D back-projected  
reconstruction

**Proof-of-principle  $^{12}\text{C}$   
Heavy Ion Tomography**

Rinaldi Ph.D. research at HIT/DKFZ (in collaboration with B. Voss, GSI); Voss et al GSI Report 2010, in press

K. Parodi, 2011



# Conclusions

pCT must do better than 3-4% in proton range

but

can not be better than  $\sim 1\%$  due to intrinsic limitations

## Main challenges:

- Detector spatial resolution  $< 1$  mm
- Energy resolution  $< 1\%$
- Fast DAQ  $> 1$  MHz
- Iterative reconstruction in GPU



Thanks!