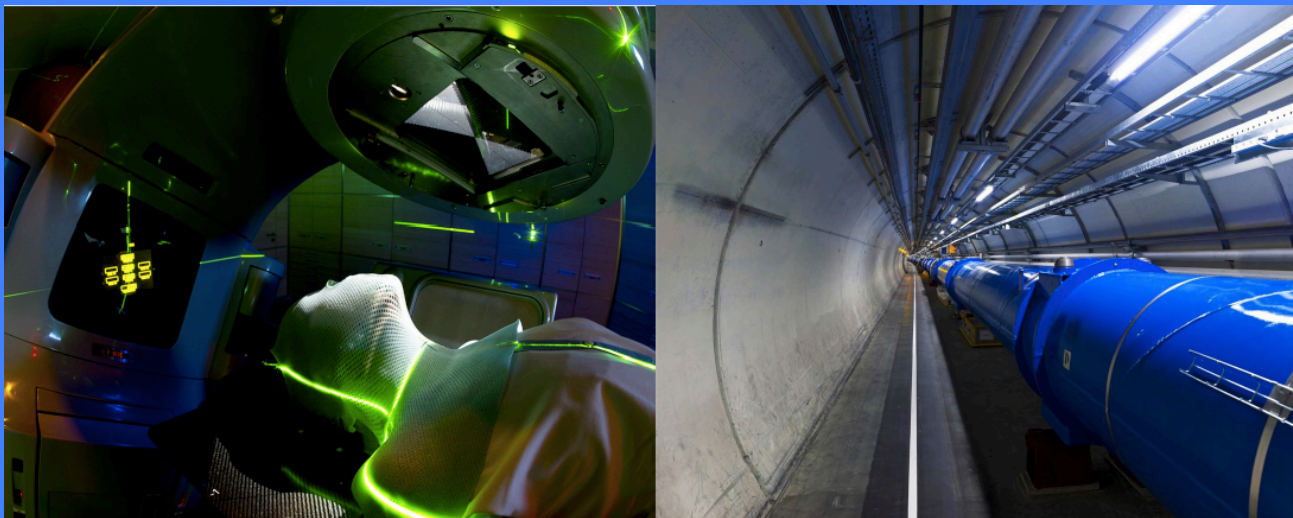




# CERN: From Particle Physics to Medical Applications



Evangelos N. Gazis

Professor of Particle Physics  
National Technical University of Athens

CERN High School Teachers, 28-31 Aug 2019



# Outline

- Medical Applications
- From X-rays to Medical Physics
- Natural Radioactivity
- Synergy of sciences
- CERN Collaboration for the diseases
- Cancer: Diagnosis – Treatment
- Medical Imaging
- PET Imaging
- Medical Imaging Collaborations: Crystal Clear & Medipix
- Digital Imaging
- Multimodality Imaging CT – PET
- MRI
- Accelerators for cancer : diagnosis - treatment
- Hadron Therapy - Timeline
- Carbon Ions – HIT
- PIMMS – CNAO
- ISOLDE – Isotopes Production



# Medical Applications

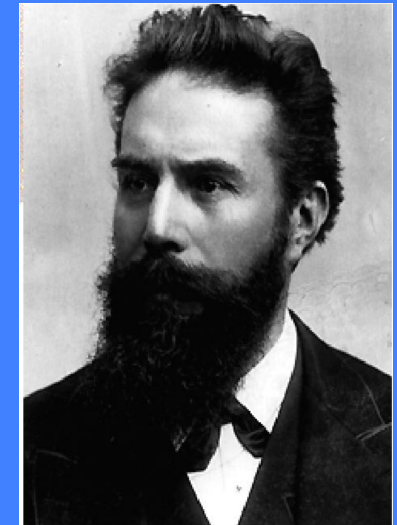
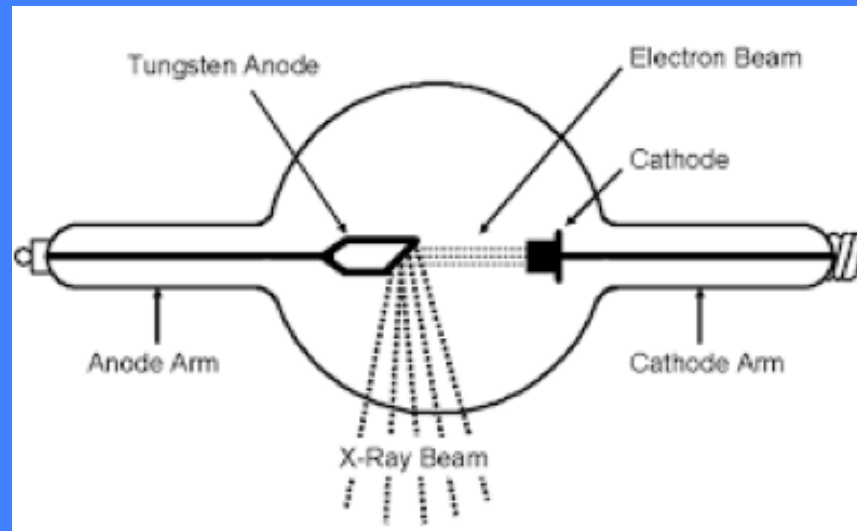
- Fundamentals of Radiation Physics
- Medical Diagnostic Techniques
- Imaging technics (basic)
- Radiation Therapy

## **Not covered in this talk:**

- Advanced Imaging
- Radiation Protection and Dosimetry
- Radiobiology
- Anatomy and Physiology
- Molecular and cellular oncology

# X-rays

## Beginning of Modern Medical Physics

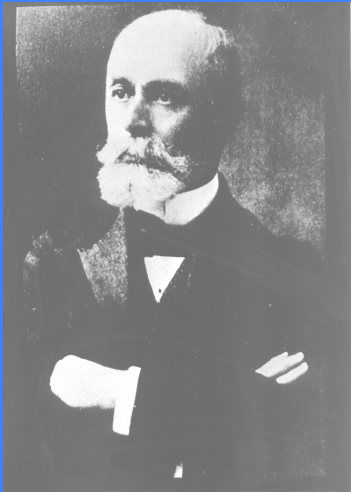


- ◆ November 8, 1895 **Wilhelm Röntgen** discovered X-Rays
- ◆ December 22, 1895 he takes the first image of his wife's hand
- ◆ Röntgen received the first Nobel prize in physics in 1901



# Natural Radioactivity

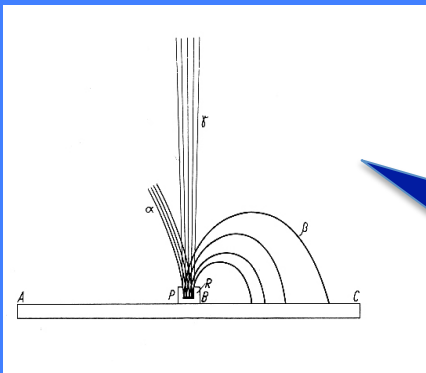
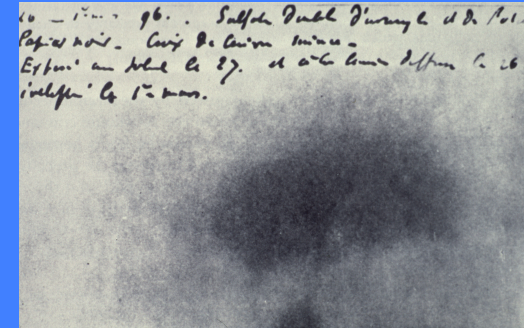
## Beginning of Modern Medical Physics



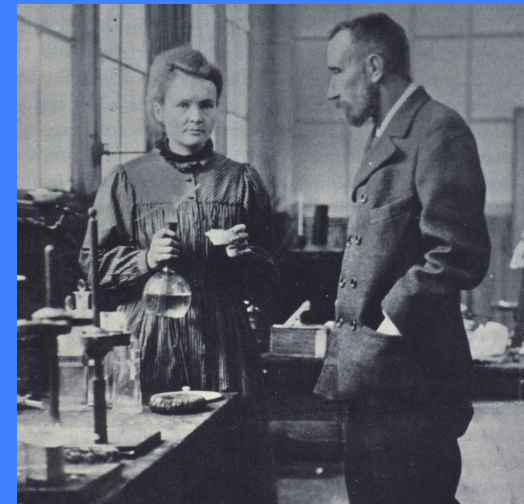
Henri Becquerel

1896: Discovery of natural radioactivity

1898: Discovery of radium used immediately for “Brachytherapy”



Thesis of Mme. Curie – 1904  
 $\alpha$ ,  $\beta$ ,  $\gamma$  in magnetic field



Marie Curie Pierre Curie  
(1867 – 1934) (1859 – 1906)



# What is the added value of the Physics of Medicine?

....or why should YOU study medical physics?

...or why should senior physicists care about medical research?

...and why care medical researchers/industry about physics???



# 1<sup>st</sup> Answer: Synergy

Accelerator physics

X-ray source physics

detector physics

image reconstruction

medical doctors

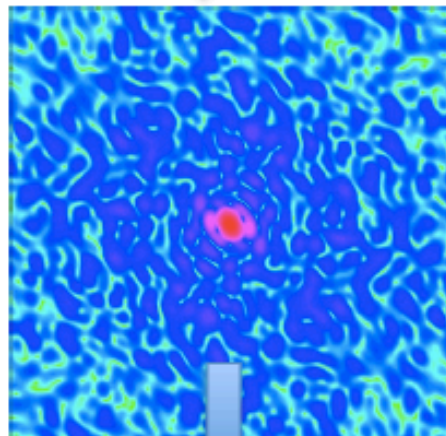
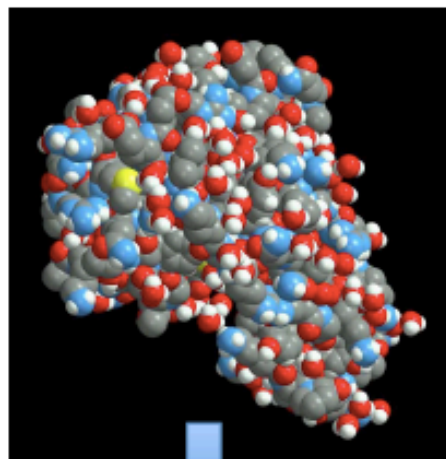




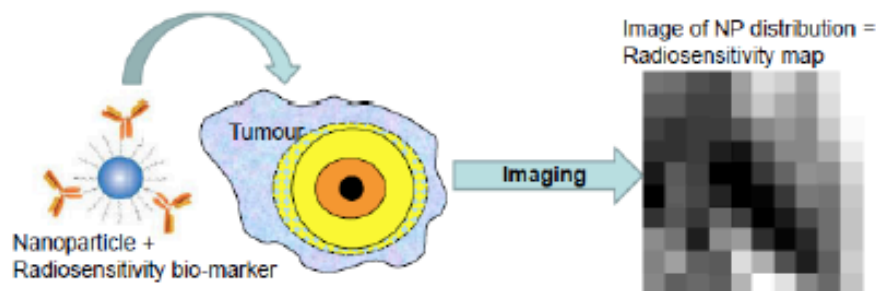


## 2<sup>nd</sup> Answer: Overcoming Limits!

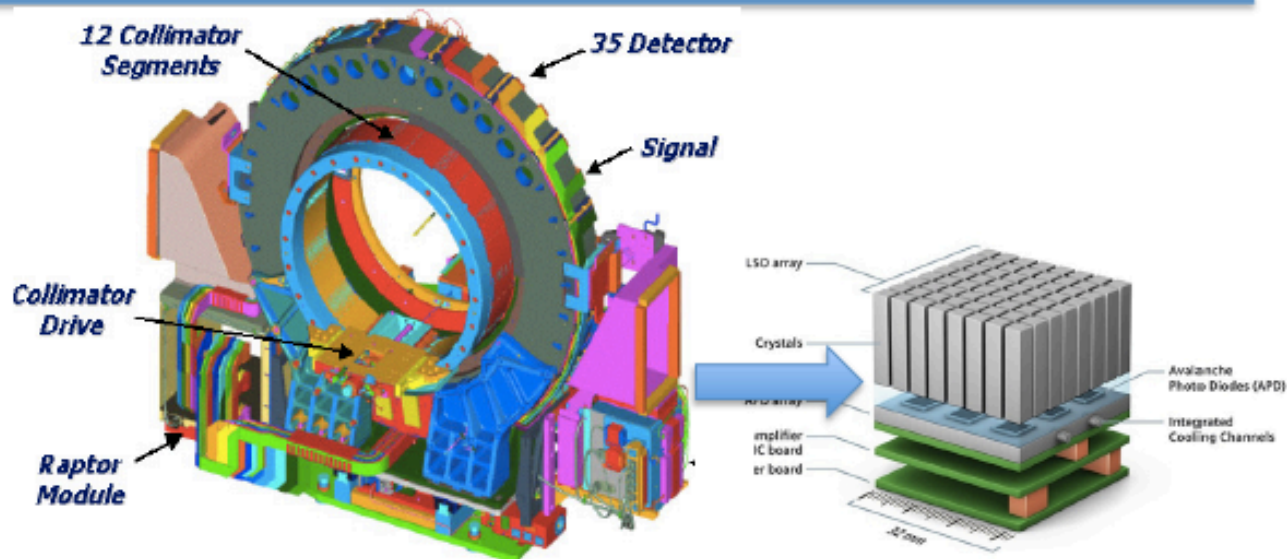
### single molecule imaging



3D protein structure



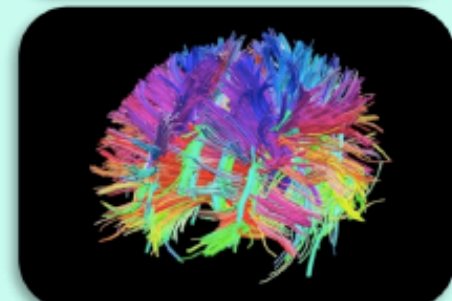
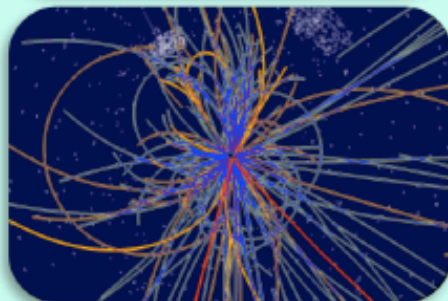
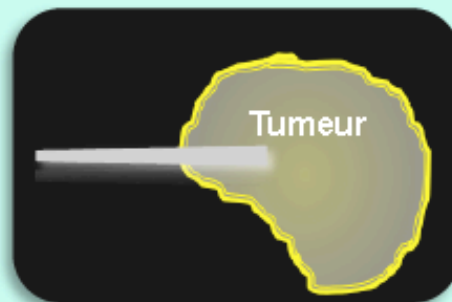
imaging of diagnostic agents not possible in-vivo



CERN-sized detector reduced to patient-size

# The 4<sup>th</sup> Pillar of Technology

## Collaboration

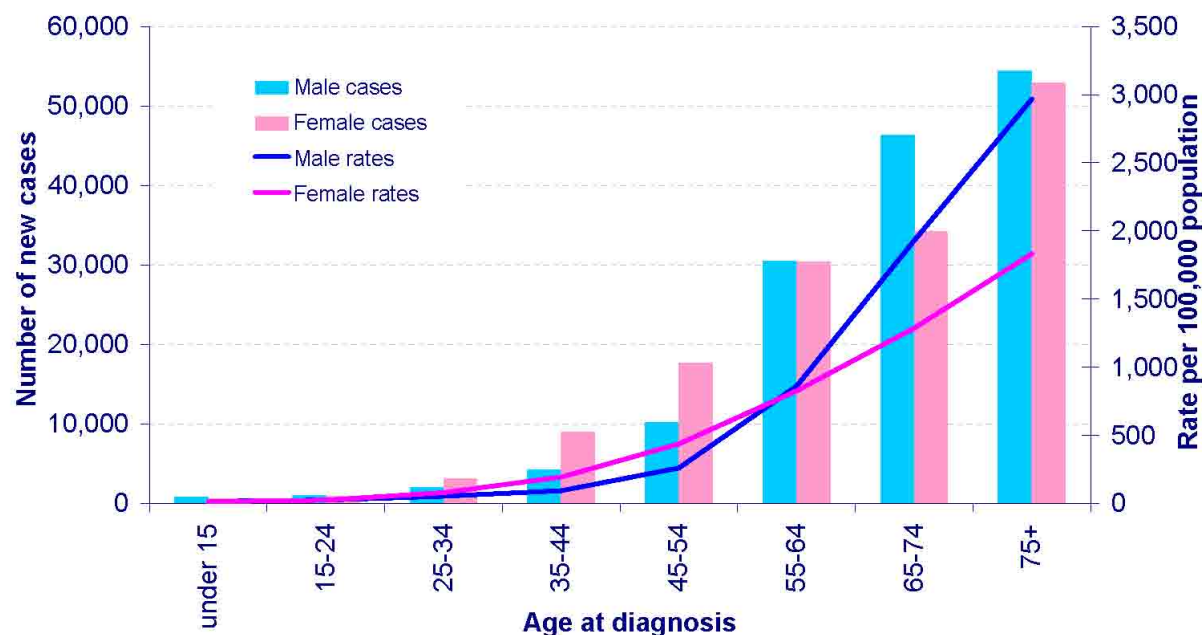


# CANCER a growing challenge

More than 3 million new cancer cases in Europe each year and 1.75 million associated deaths

Increase by 2030: 75% in developed countries and 90% in developing countries

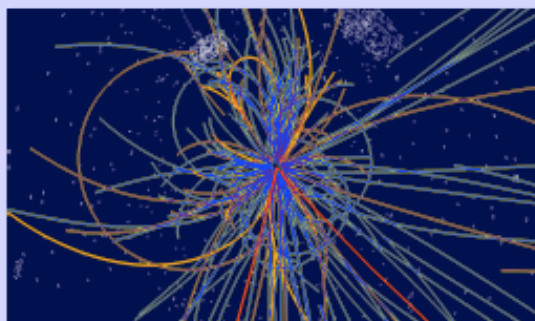
**Figure 2.1: Number of new cases and rates, by age and sex, all malignant neoplasms (exc NMSC), UK, 2007**





# CERN Technologies & Innovation

Accelerating  
particle beams



Detecting  
particles



**CANCER**

Large-scale  
computing (Grid)



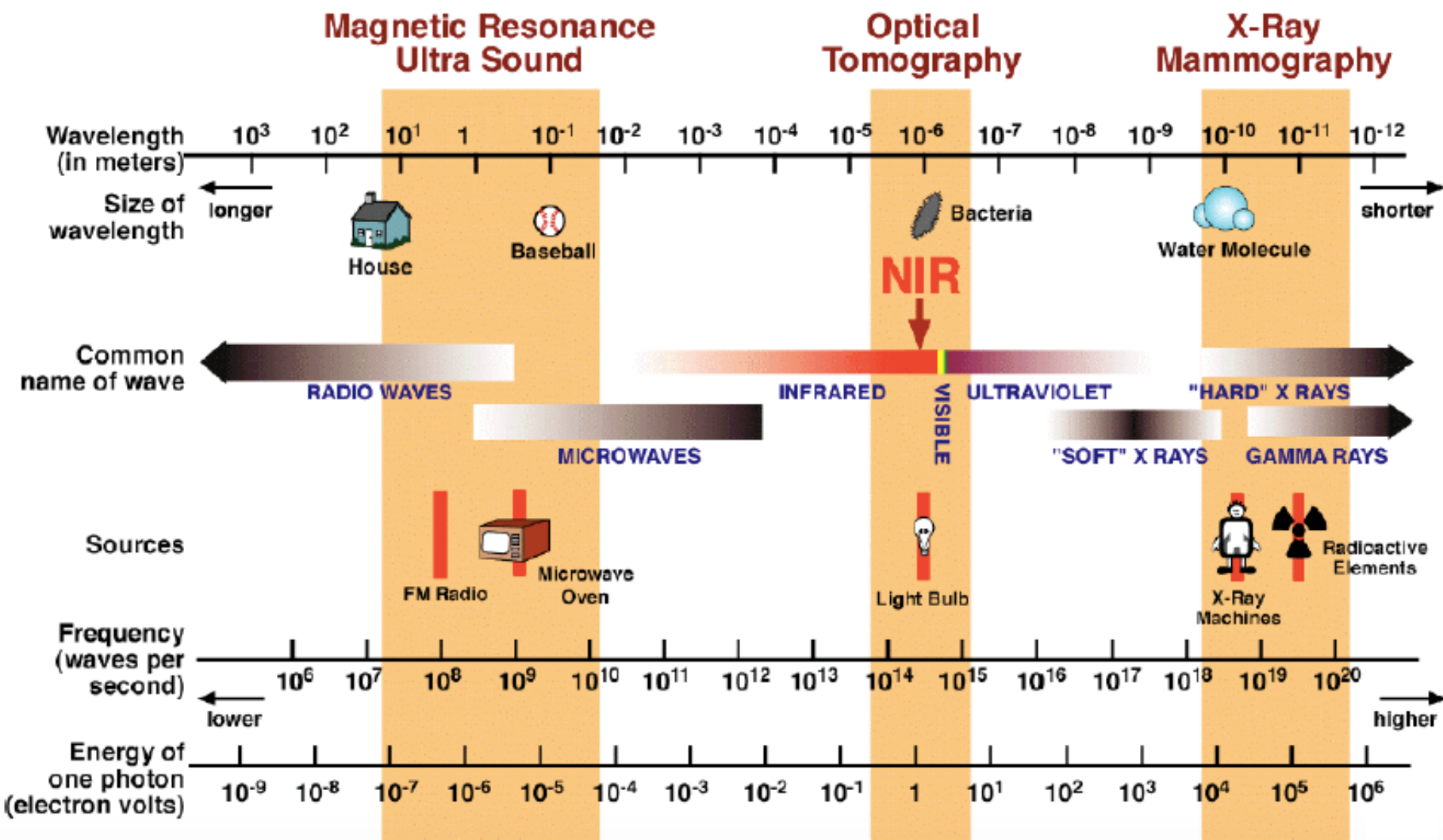


# Medical Imaging

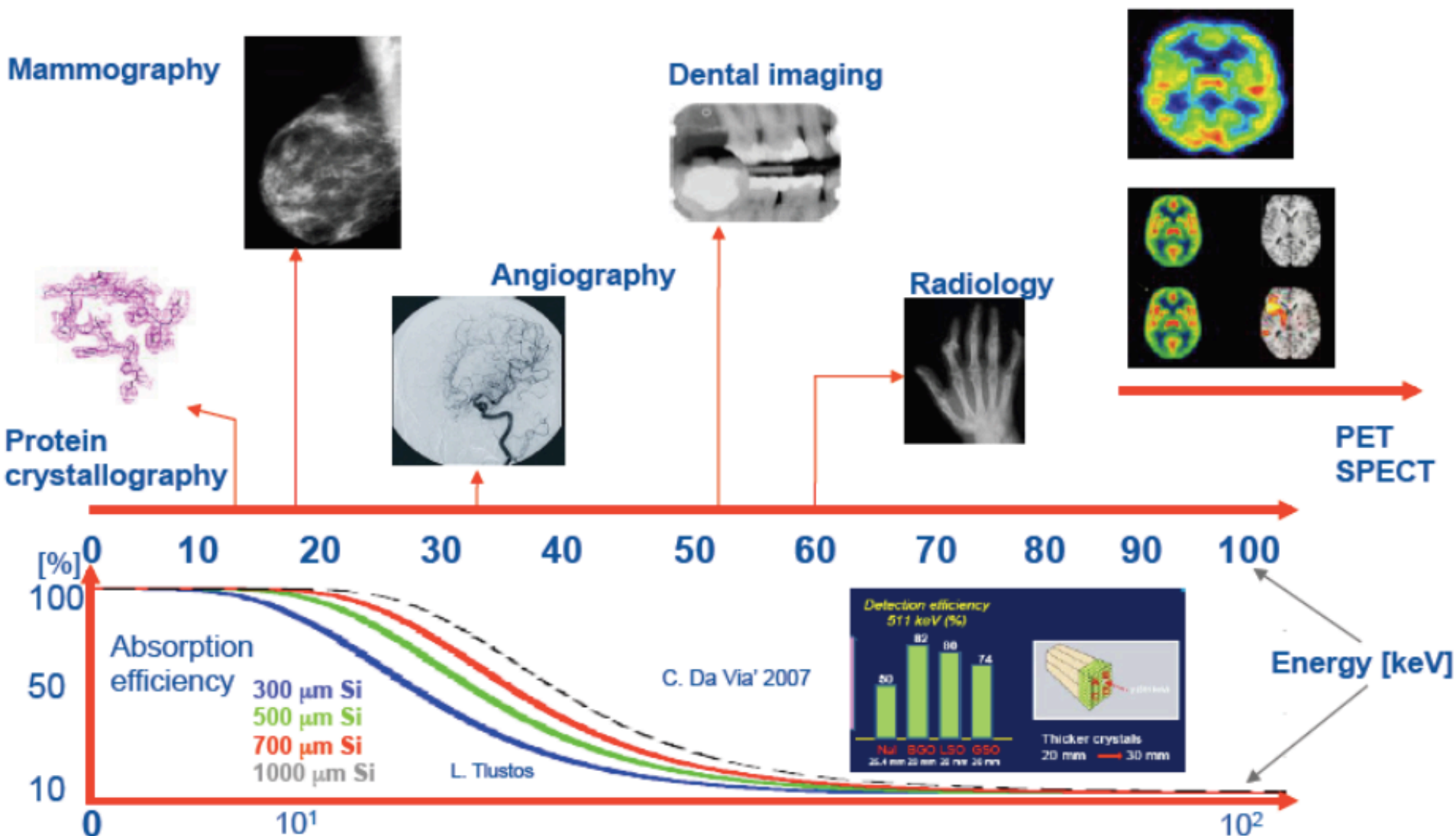


# Electromagnetic Waves Used in Medical imaging

larger than  $1 \text{ \AA}$  high attenuation from the body,  
shorter than  $10^{-2} \text{ \AA}$  = too high energy ( $>1\text{MeV}$ ) for direct detection



# Medical Imaging



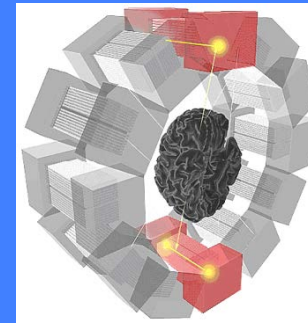


# Medical Imaging then Treatment

## Particle Detection



## Imaging

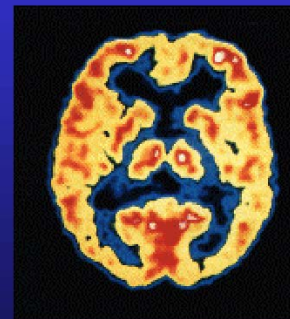


PET Scanner

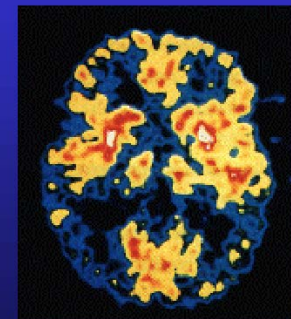
## Breast imaging (Clear PEM)



## Brain Metabolism in Alzheimer's Disease: PET Scan



Normal Brain

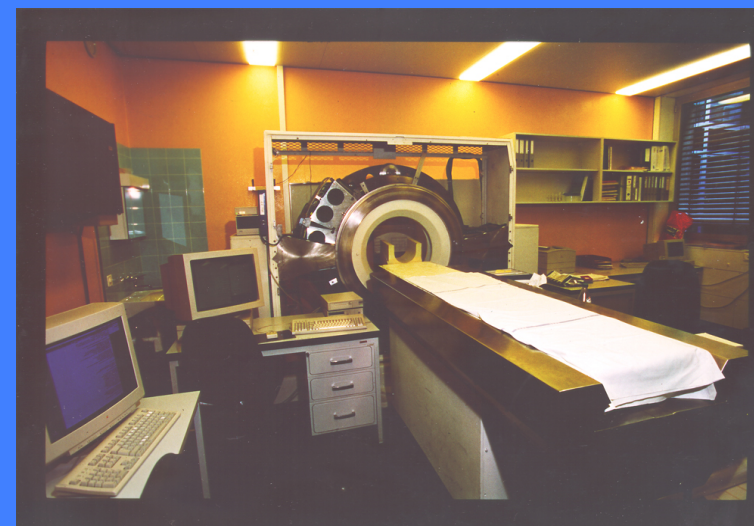
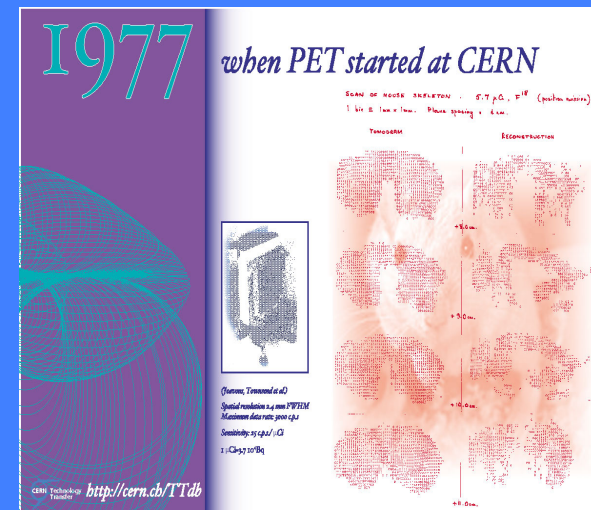


Alzheimer's Disease

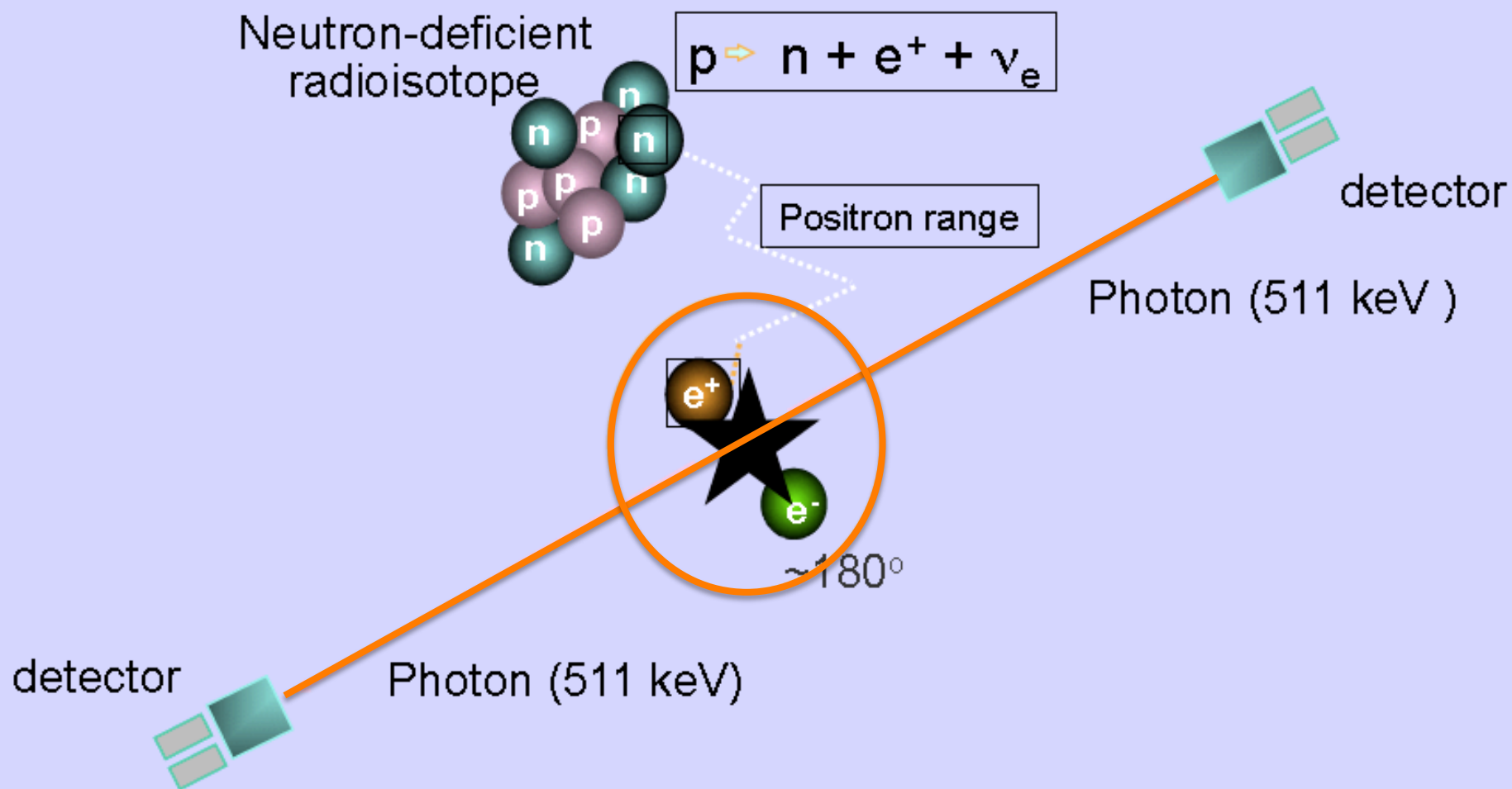


# PET Imaging

Idea of PET

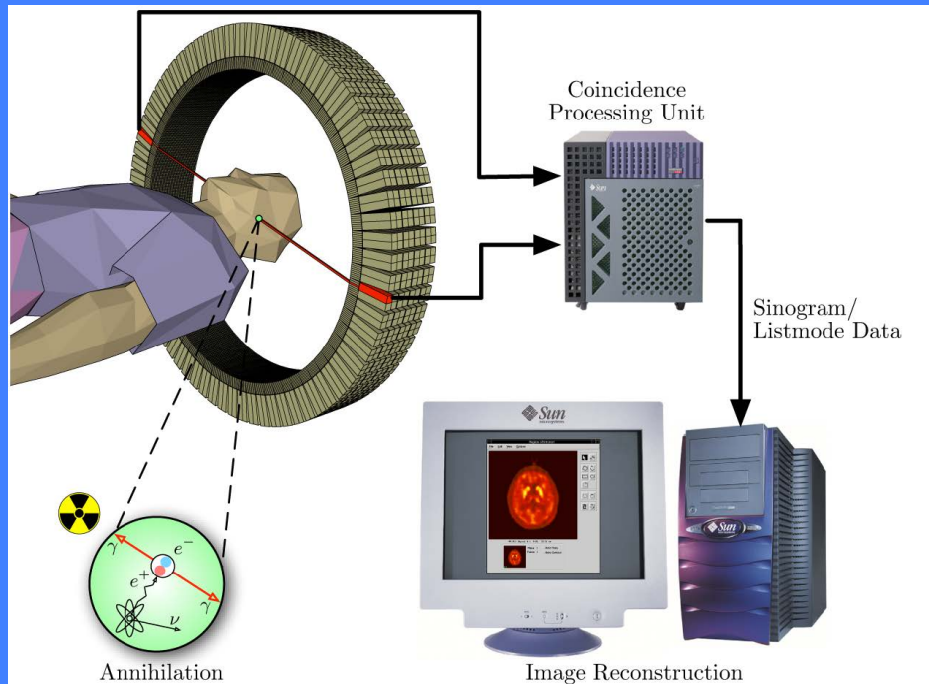


# PET : Operation

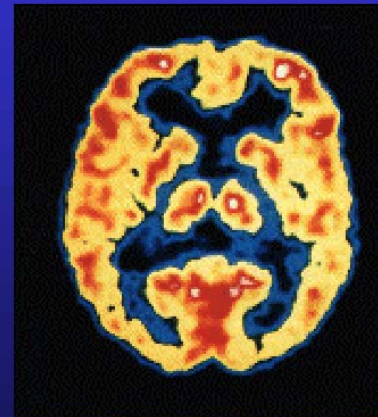




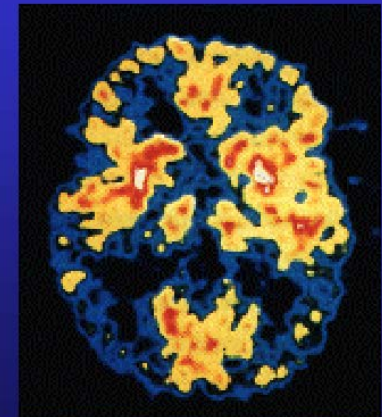
# PET : Scan



## Brain Metabolism in Alzheimer's Disease: PET Scan



Normal Brain



Alzheimer's Disease





# Crystal Clear Collaboration

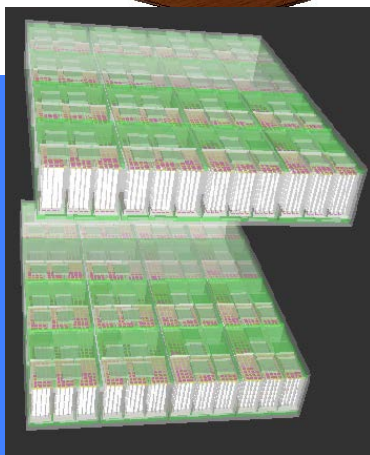
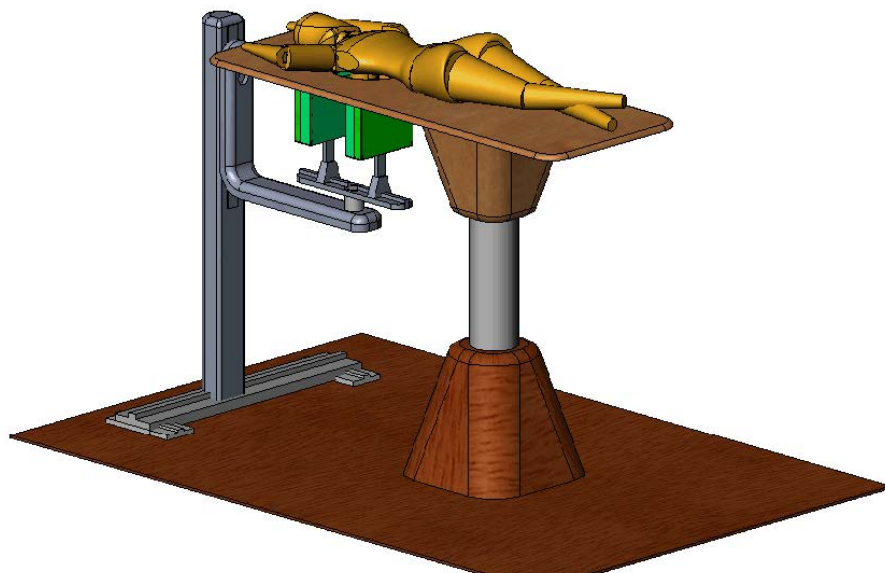


- New **scintillating** materials
  - LuAP, phoswich LuAP-LSO (CERN patent)
  - Other crystals
- New **photodetectors** (Avalanche Photo-Diodes)
- New **low noise** electronics

- ❖ New **intelligent DAQ** systems with pipeline and parallel architectures
- ❖ better **simulation** GEANT 4
- ❖ better **reconstruction** algorithms



# Crystal Clear Collaboration (PEM)



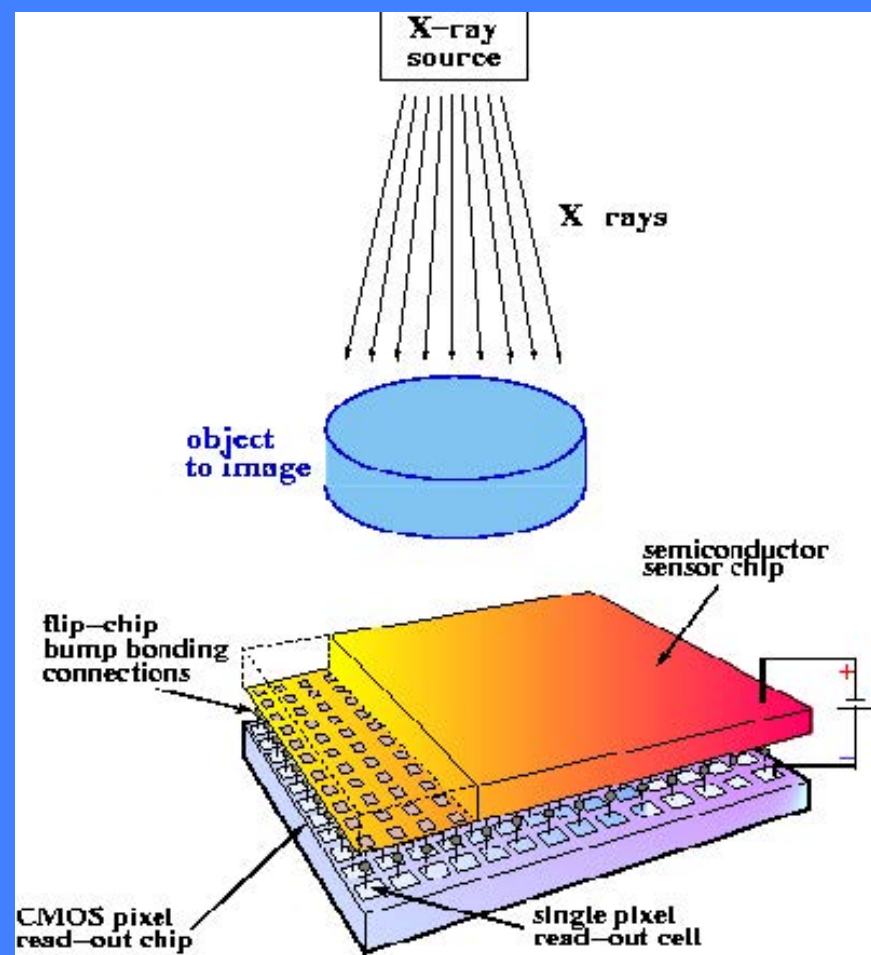
- PET dedicated to breast cancer screening
  - Extremely sensitive to small tumor masses
- Spatial resolution (1-2mm)
- High counting sensitivity
- Short PET exam
- Coupled to ultrasound

- High Energy Physics original development:
  - Particle track detectors
- Allows counting of **single photons** in contrast to traditional charge integrating devices like **film** or **CCD**
- Main properties:
  - Fully **digital** device
  - Very **high space** resolution
  - Very **fast photon** counting
  - Good **conversion efficiency** of low energy X-rays



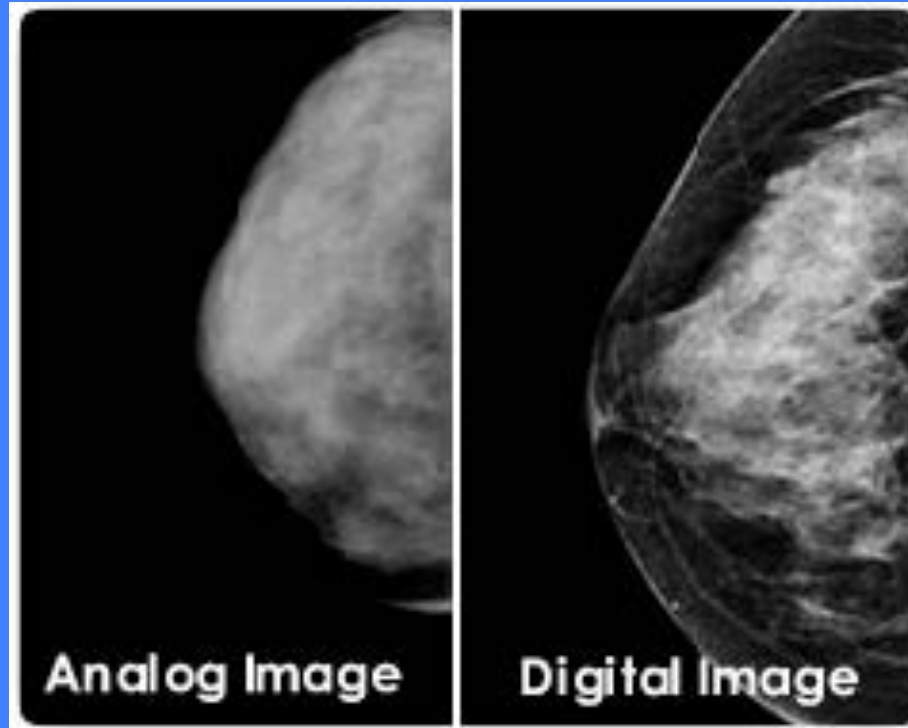
# WHAT IS MEDIPIX ?

- an electronic chip similar to the electronic imaging chip in a digital camera
  - sensitive to x-rays instead of visible light
- it can create the first true color images with x-rays.
  - it permits us to move from black and white x-ray images to full color x-ray images
- can be read out very rapidly.
  - allows the use of the chip for color x-ray digital movies or for fast color x-ray CT scans



## Current

- Limited contrast
- High dose
- Restricted screening
- Limited access to preventive health care



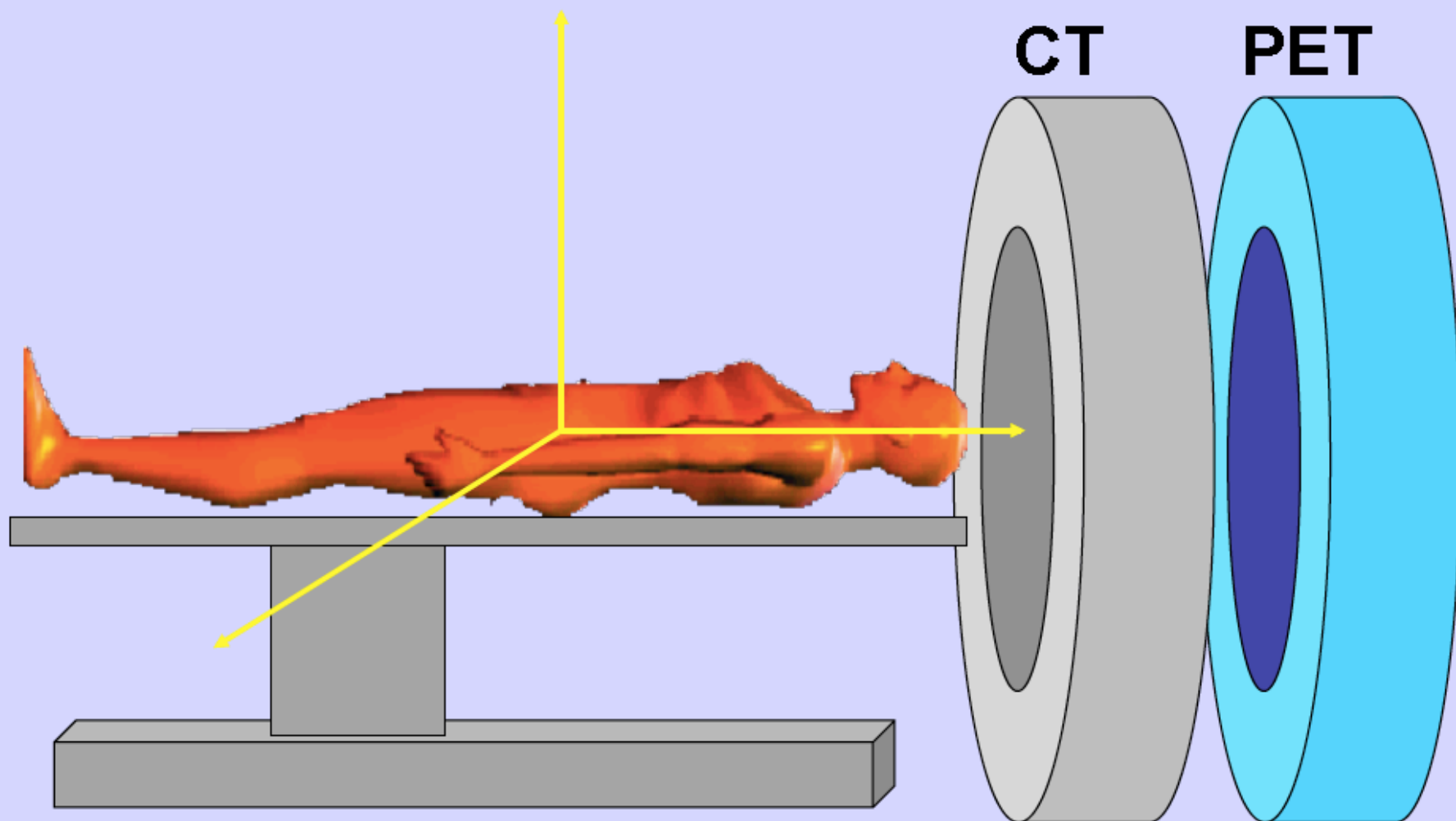
## Digital

- High contrast
- Lower dose
- Opportunity for screening
- Access to preventive health care



# Multimodality Imaging

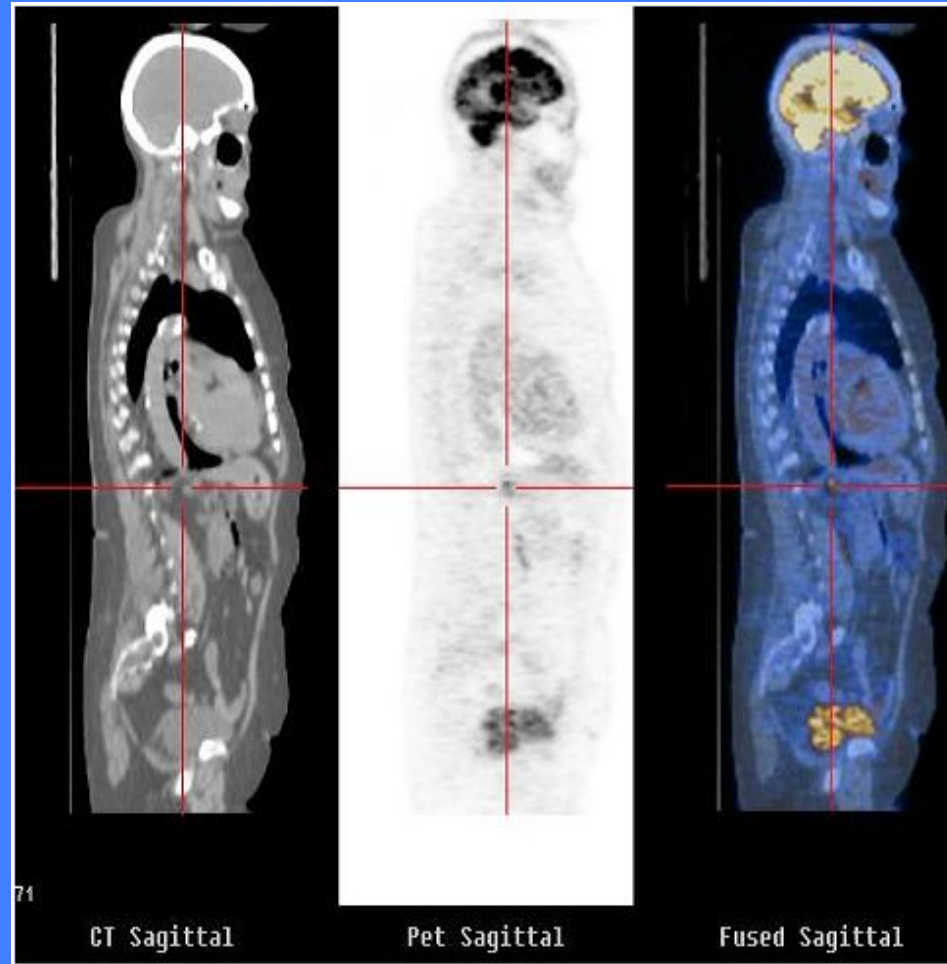
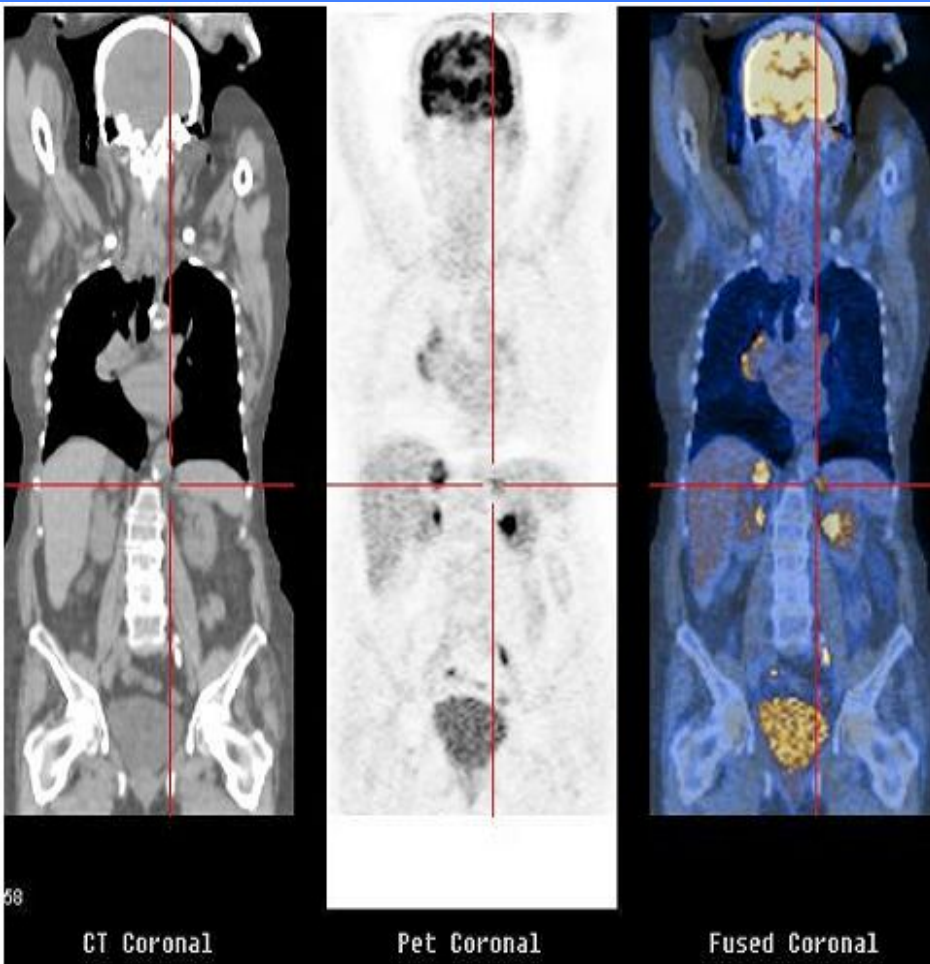
## PET – CT David Townsend





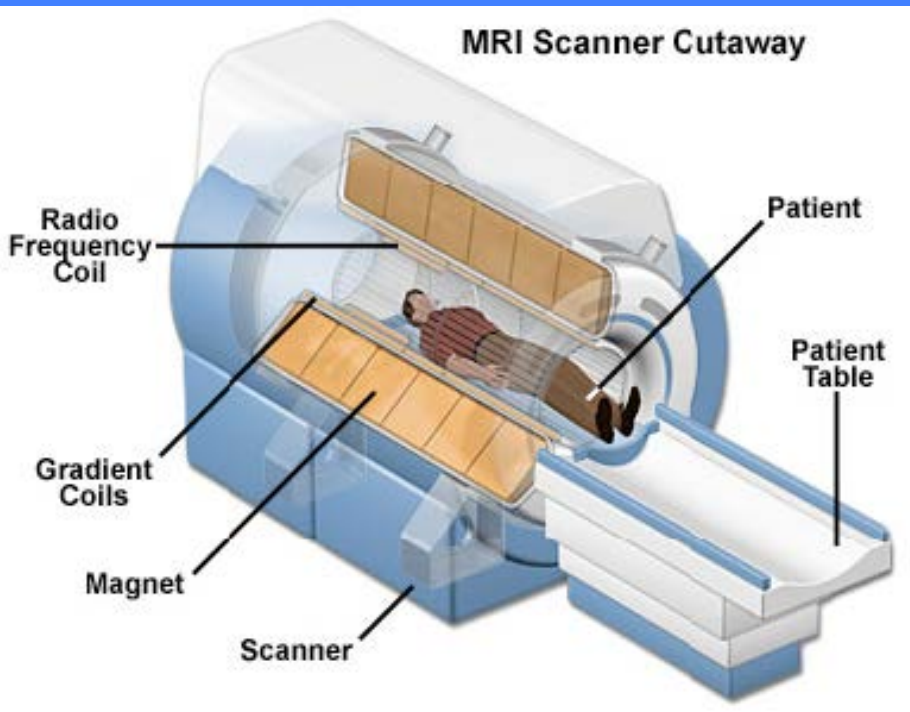
Morphology

Metabolism

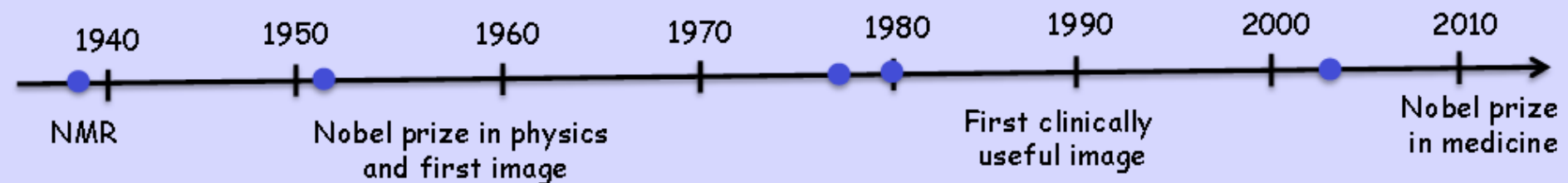
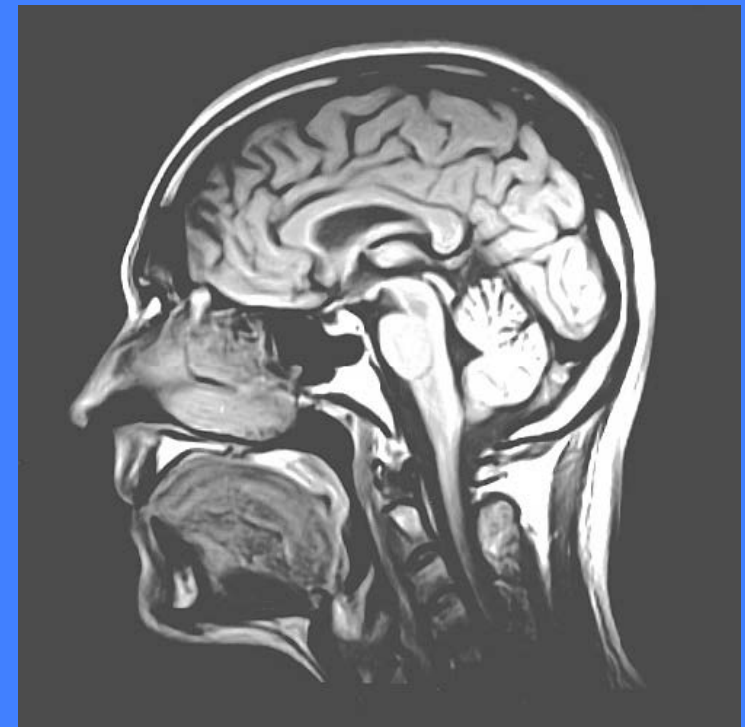




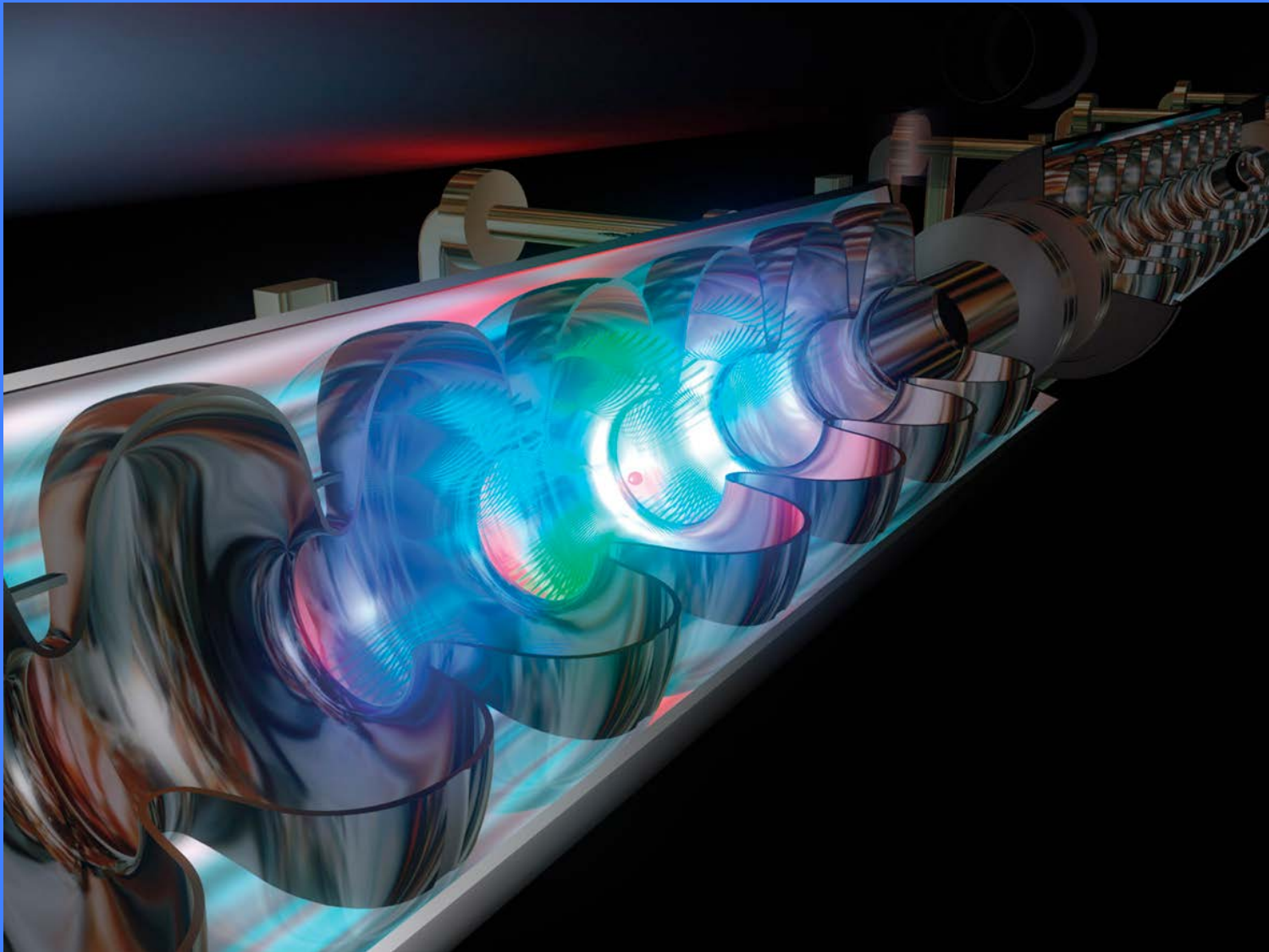
# Magnetic Resonance Imaging - MRI



First human body scan

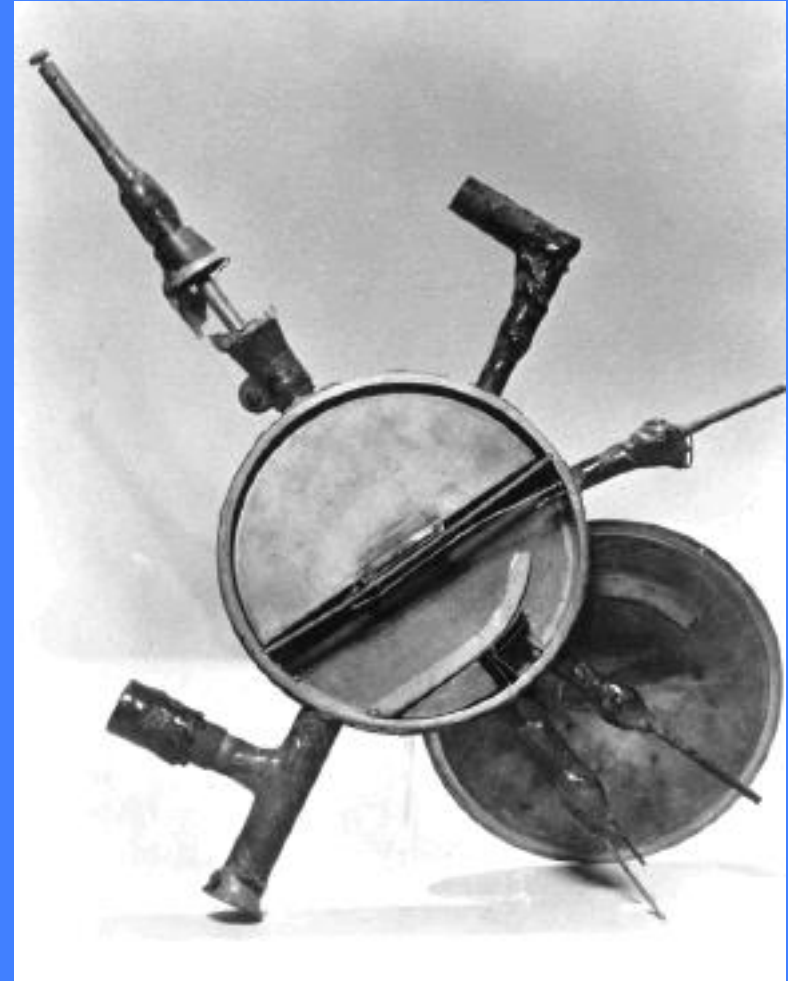
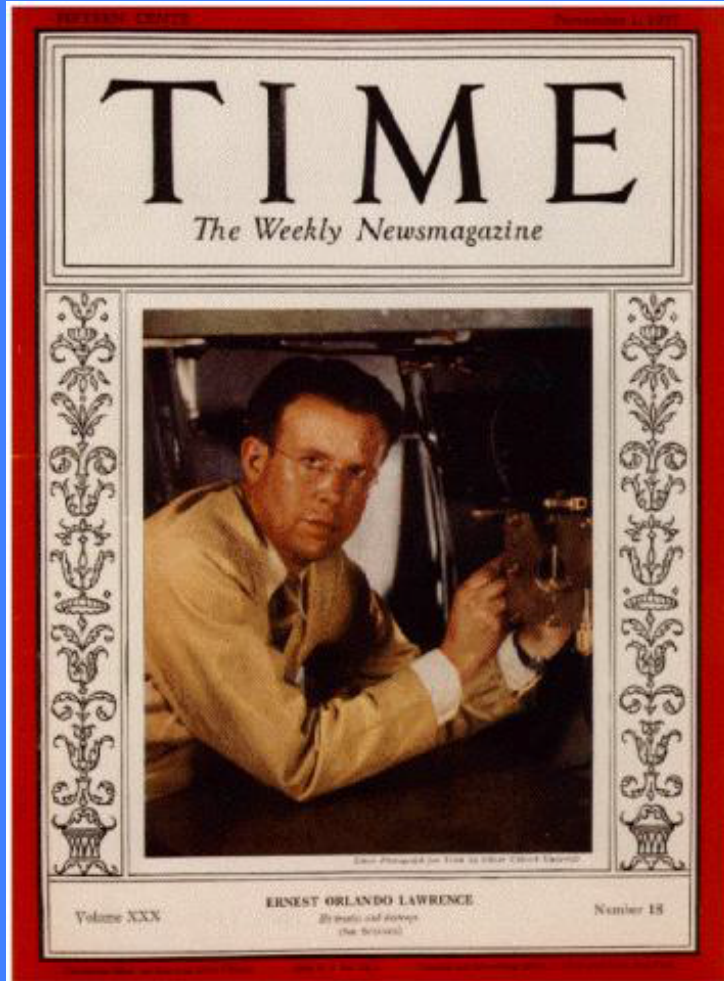


# Accelerators for Cancer Treatment



# The 1<sup>st</sup> Cyclotron

E. O. Lawrence is awarded Nobel Prize in 1939  
For inventing the cyclotron 1937





# Use of Accelerators Today

## General industrial use:

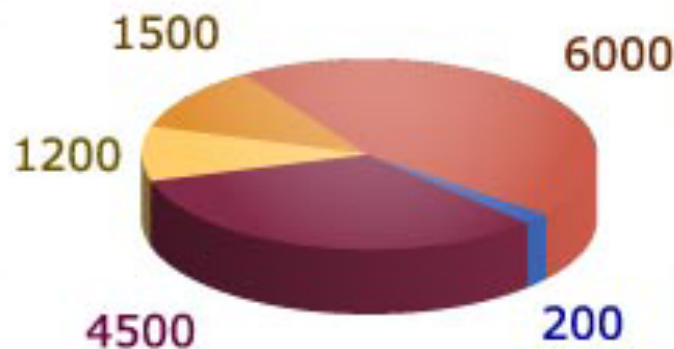
Sterilisation, imaging

## Research accelerators:

Particles, synchrotron light used in biomedical, physics, chemistry, biology, material research

## Radiotherapy:

Cancer treatment with X-rays, protons and other particles



## Ion implantation, surface modifications:

Controlled semiconductor doping; Changing properties of surfaces

## Radioisotope production:

Cancer treatment; imaging organs for medical use

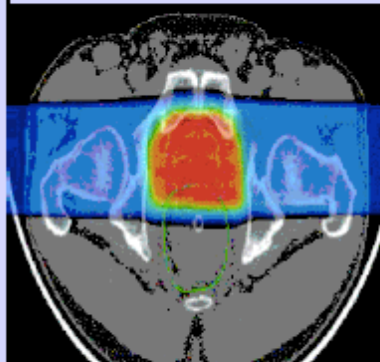
# Cancer a growing societal challenge !

Over 3 million cancer cases in Europe each year

Surgery



Radiotherapy



*X-ray, IMRT, Brachytherapy,  
Hadrontherapy*

Chemotherapy & others



*Hormones; Immunotherapy;  
Cell therapy; Genetic treatments;  
Novel specific targets (genetics..)*

Local control

Local control

Limited Local control

Survival  
Quality of life

After diagnosis some diseases like hyperthyroidism, cancer, blood disorders, etc... can be treated using **radiotherapy**

Three main methods:

- **Unsealed source** radiotherapy
- **Brachytherapy** (sealed source therapy)
- **External beam**: x-rays, electrons, p, n, heavy ions

Stages in the radiotherapy process:

QA, Imaging, Planning, Simulation,  
Treatment, Verification, Modelling Outcome

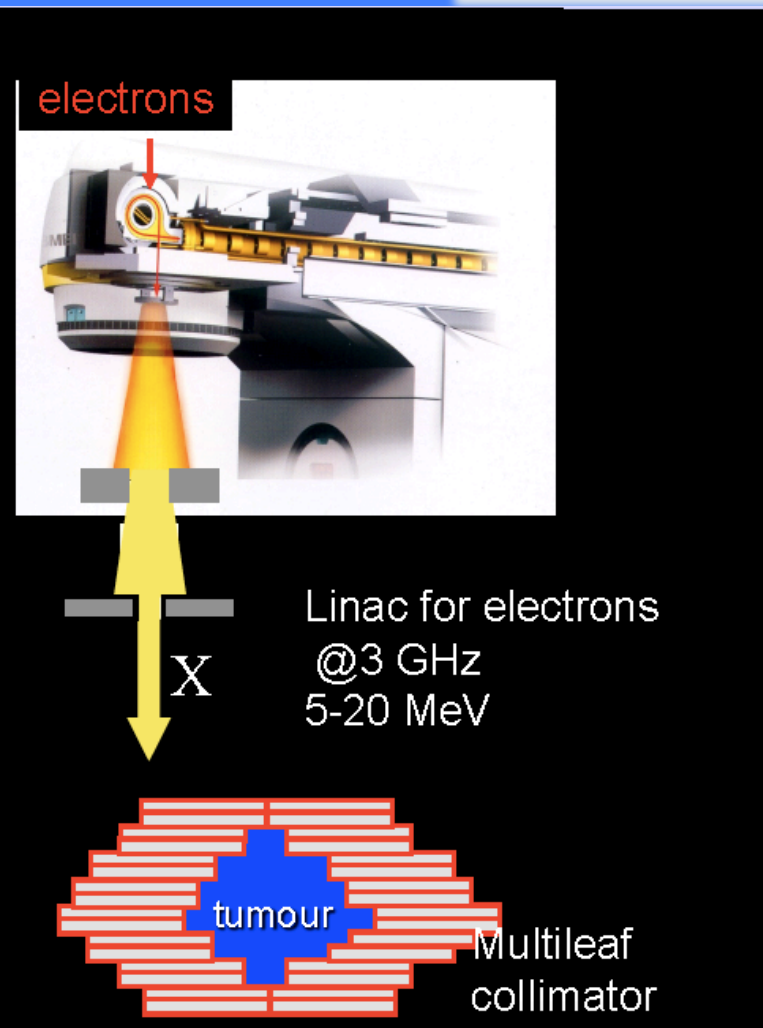
Physics, engineering, imaging, technology based



"seeds"- small radioactive rods implanted directly into the tumor



# Conventional Radiotherapy



More than 7000 linacs in the world for radiotherapy



20 000 patients per year every  
10 tumor million inhabitants  
1 linac every <250,000 inhabitants

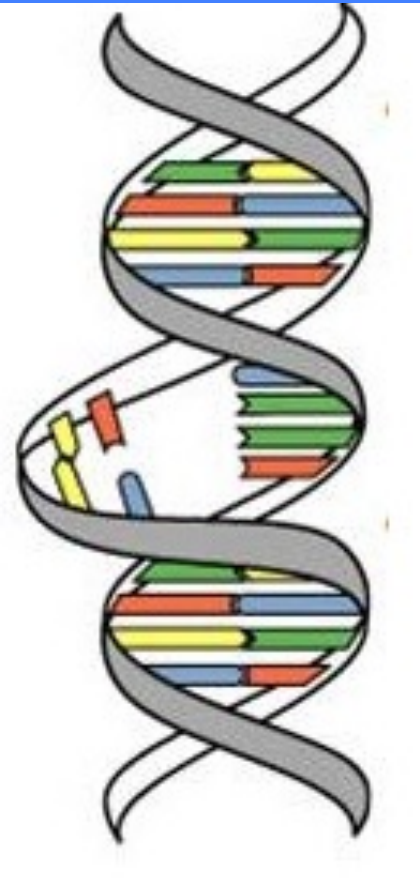


## 3 "Cs" of Radiation

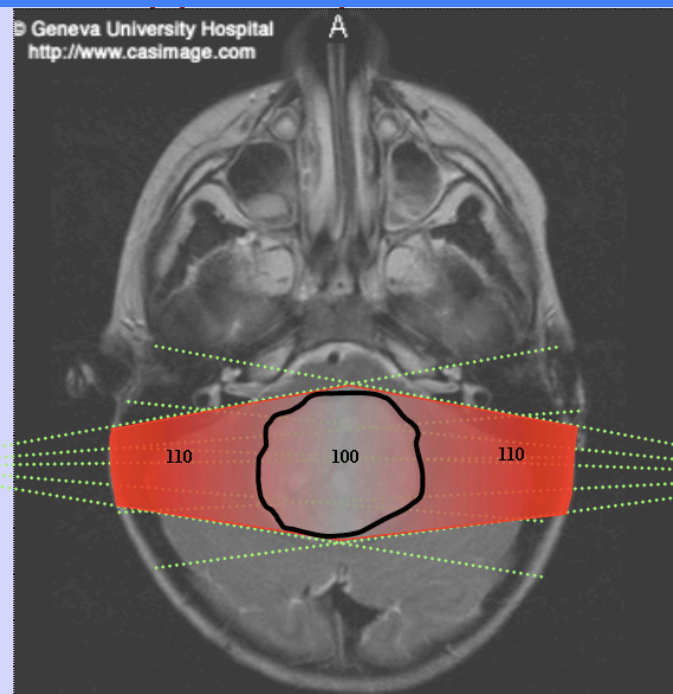
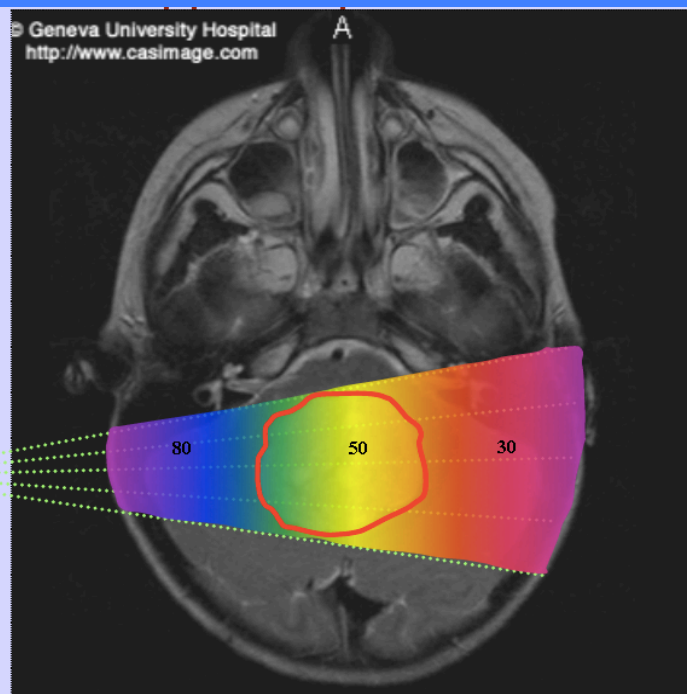
- **C**ure (~ 45% cancer cases are cured)
  - **C**onservative (non-invasive, few side effects)
  - **C**heap (~ 5% of total cost of cancer on radiation)
- There is no substitute for RT in the near future
  - The rate of patients treated with RT is increasing

Present Limitation of RT:

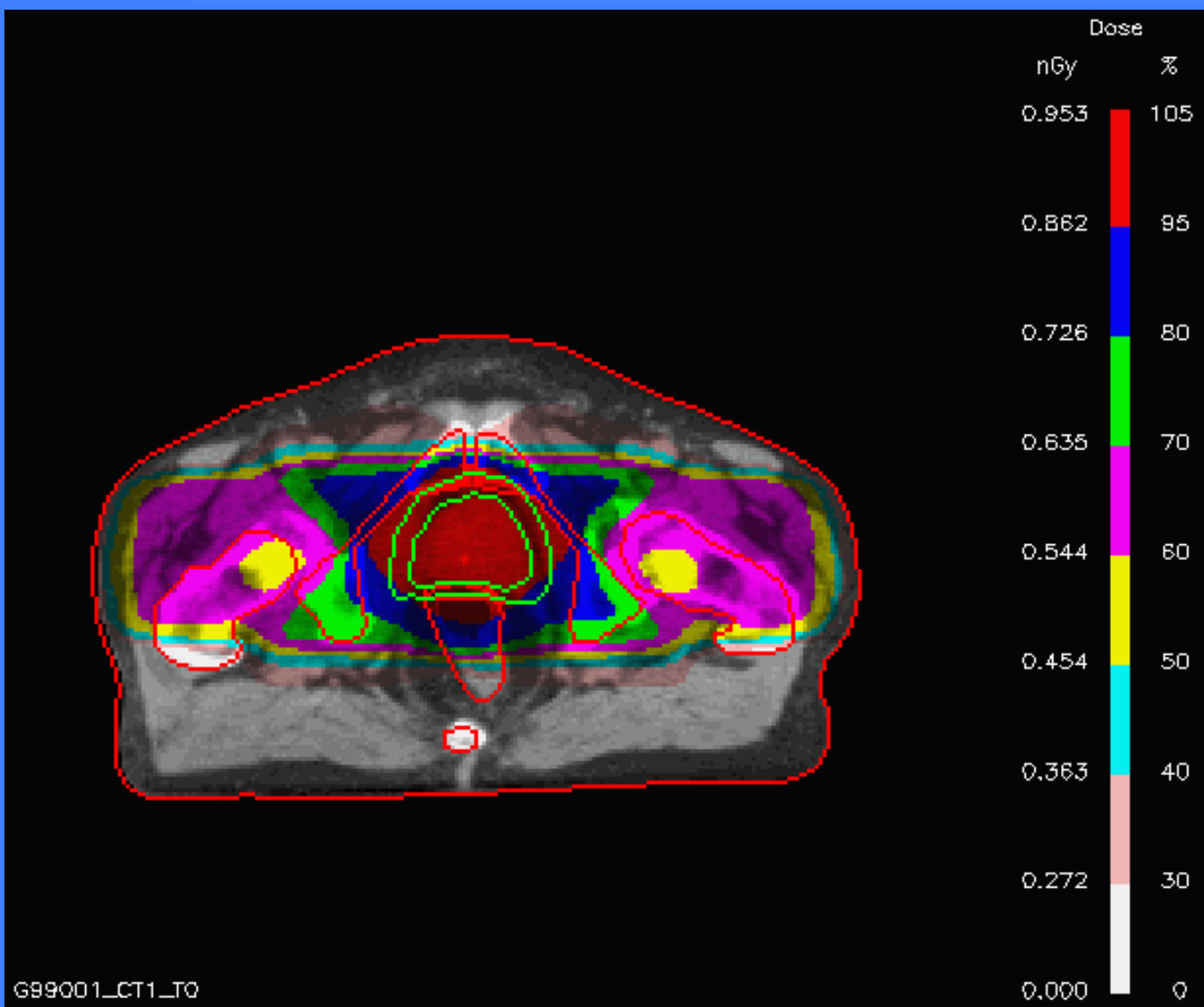
~30% of patients treatment fails locally



# Two Opposite Photon Beams



# Two Opposite Photon Beams





# How to decrease the failure rate?

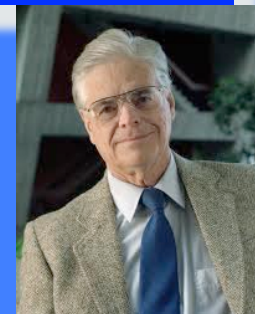
- Accelerator technologies to improve treatment : higher dose
- Detectors/imaging: accuracy, multimodality, real-time, organ motion
- Biology: fractionation, radio-resistance, radio-sensitization
- Data: storage, analysis and sharing
- Collaboration in this multidisciplinary field is key

Raymond Miralbell, HUG

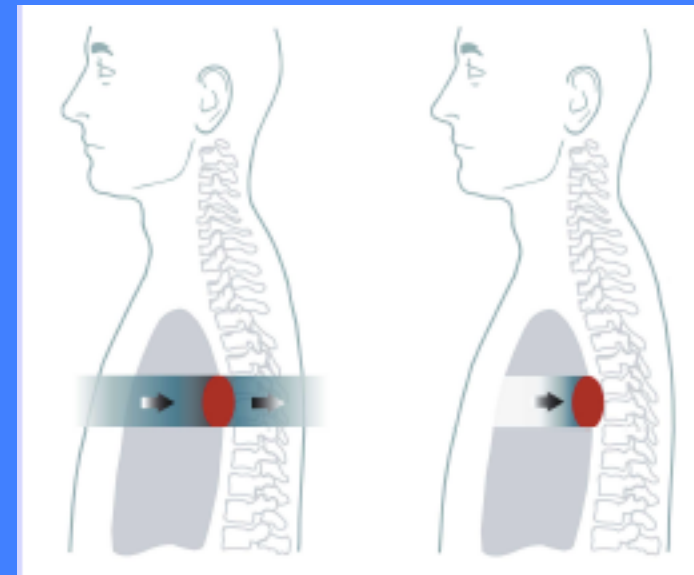
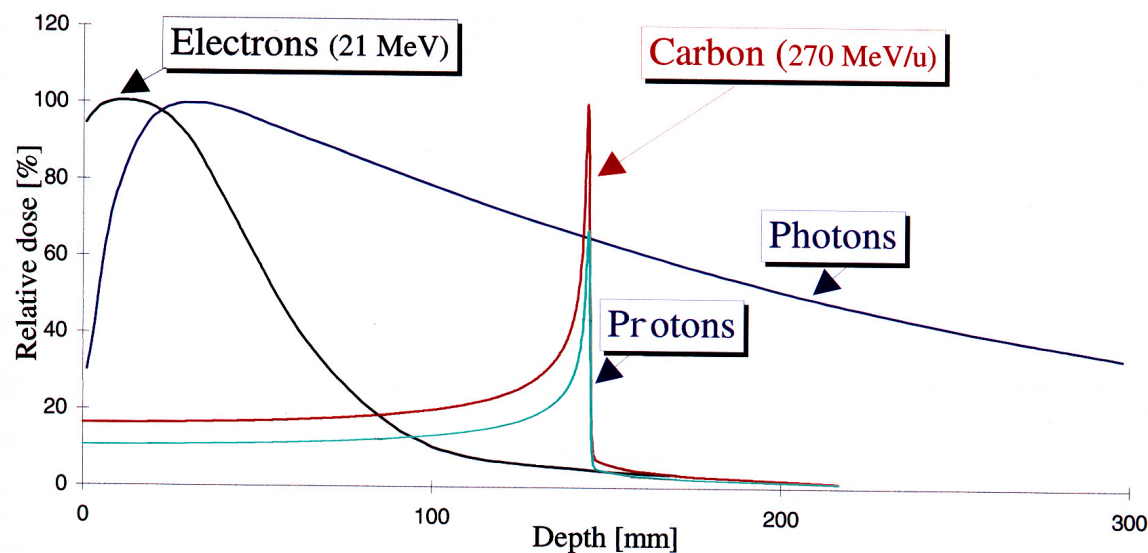
# Hadron Therapy : All started in 1946

In 1946 Robert Wilson:

- Protons can be used clinically
- Accelerators are available
- Maximum radiation dose can be placed into the tumor
- Particle therapy provides sparing of normal tissues



- Tumors near critical organs
- Tumor in children
- Radio-resistant tumors

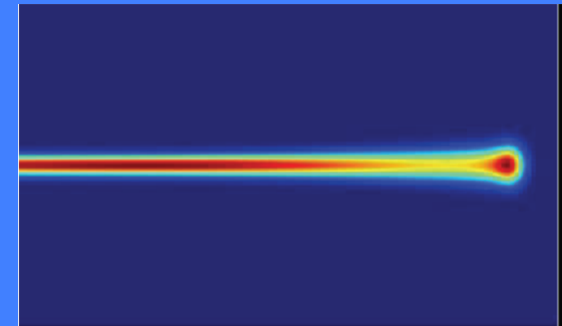
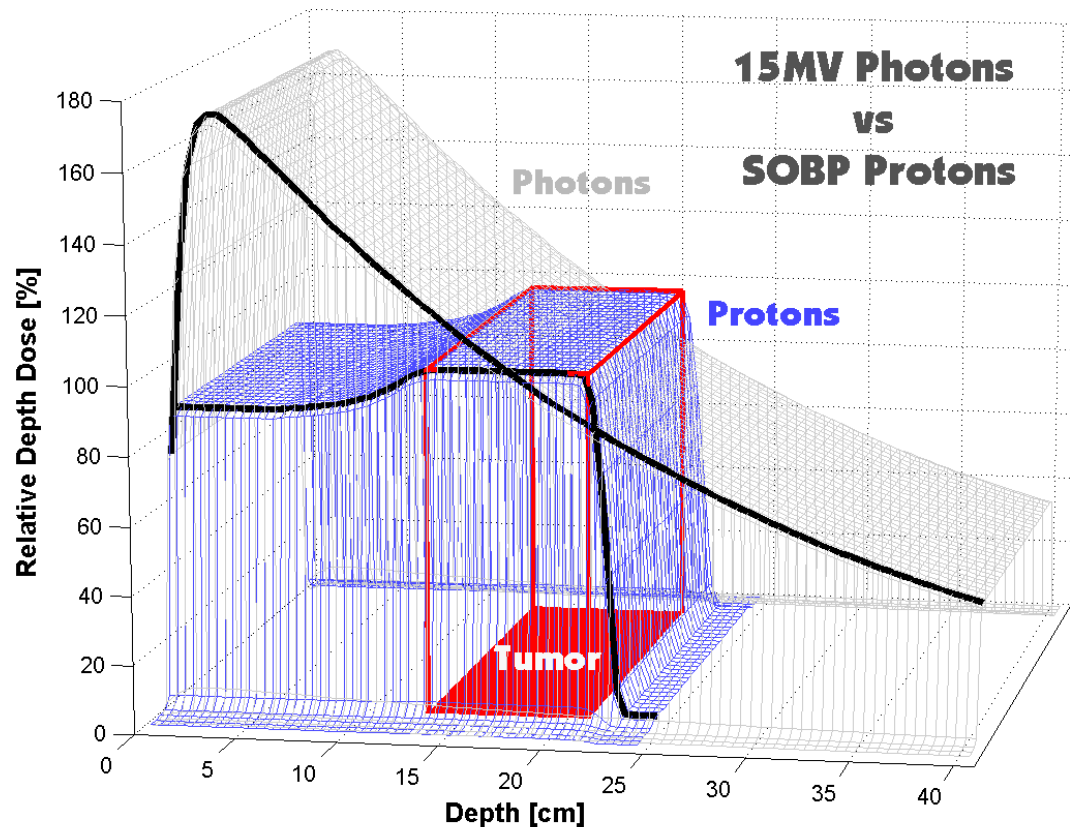


Conventional: X-Rays    Ion Radiation



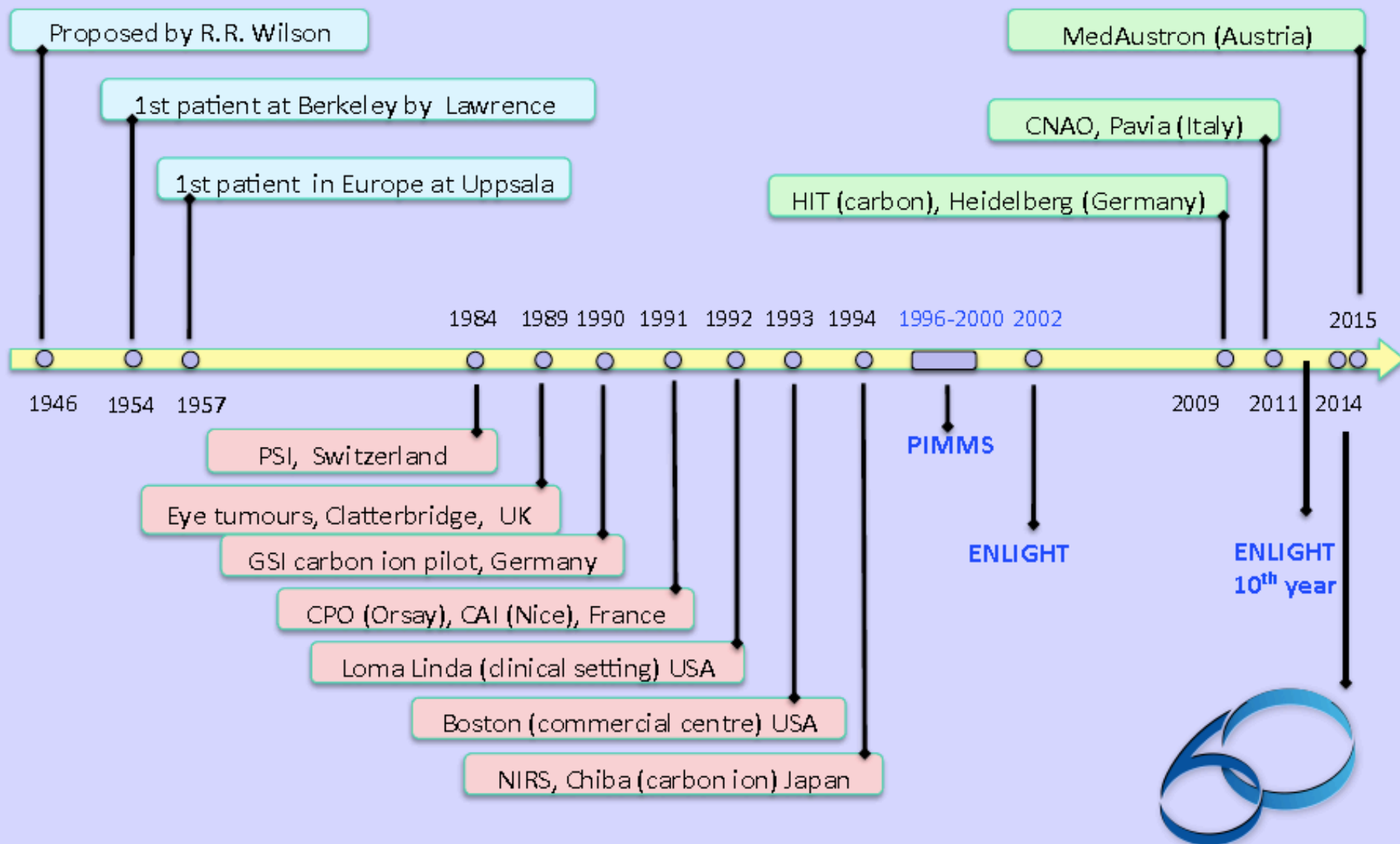
# Hadron Therapy

## Optimization of the natural distribution of dose

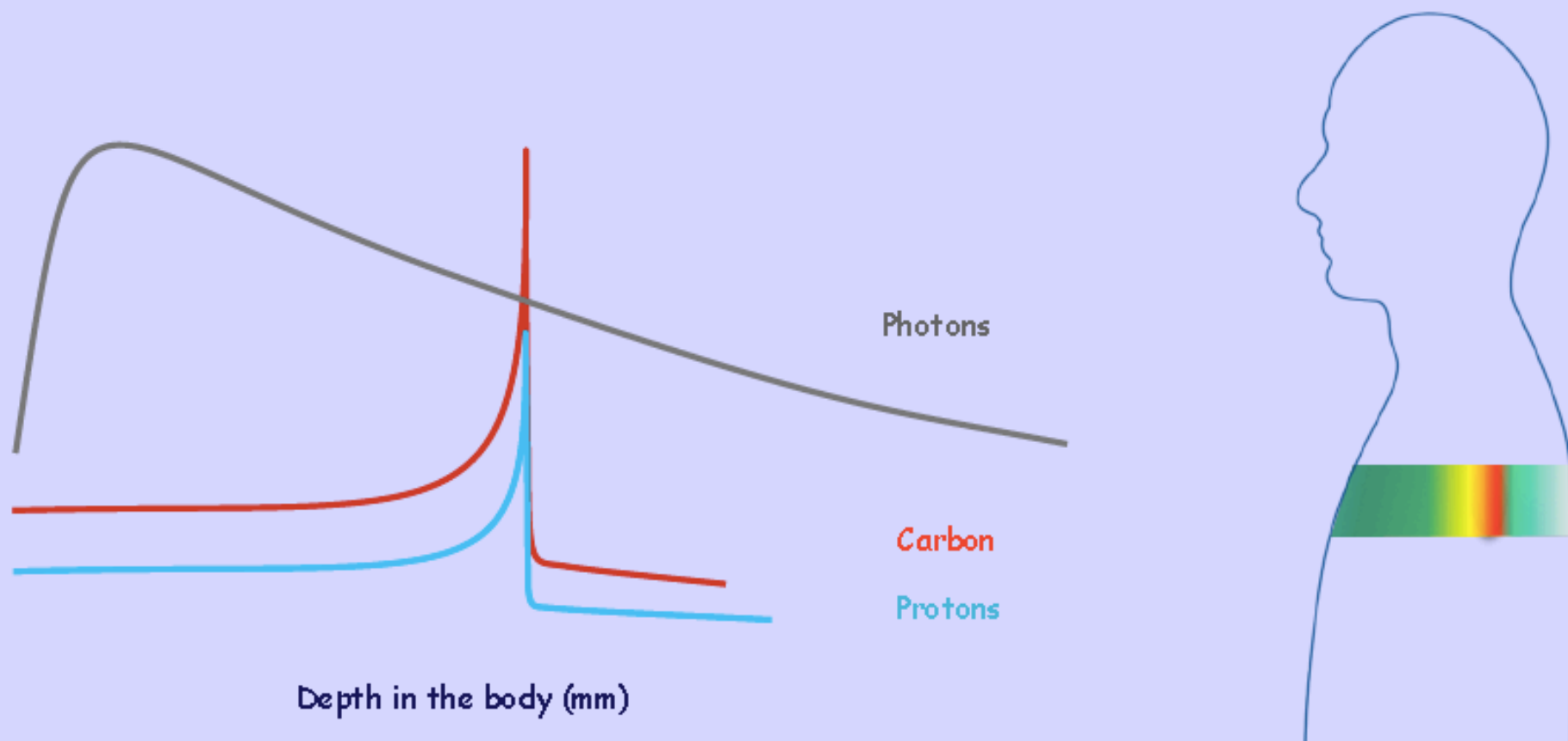




# Hadron Therapy Timeline

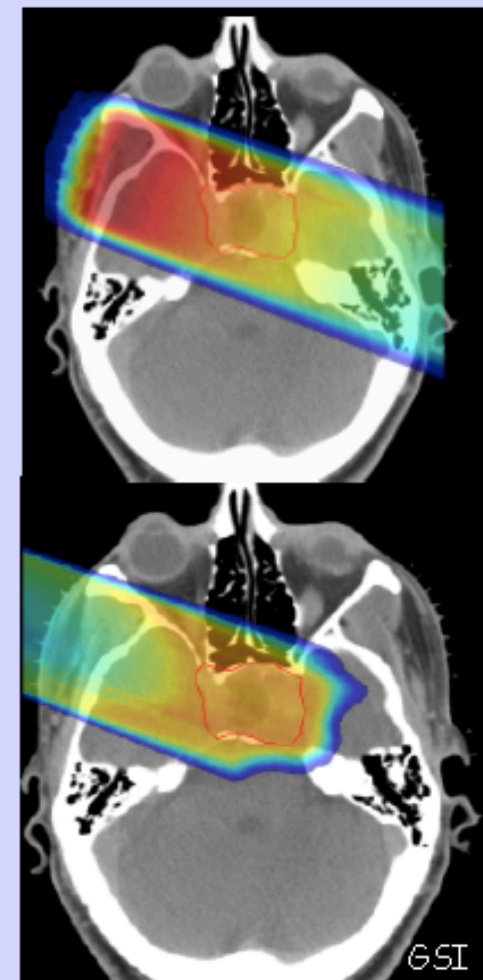
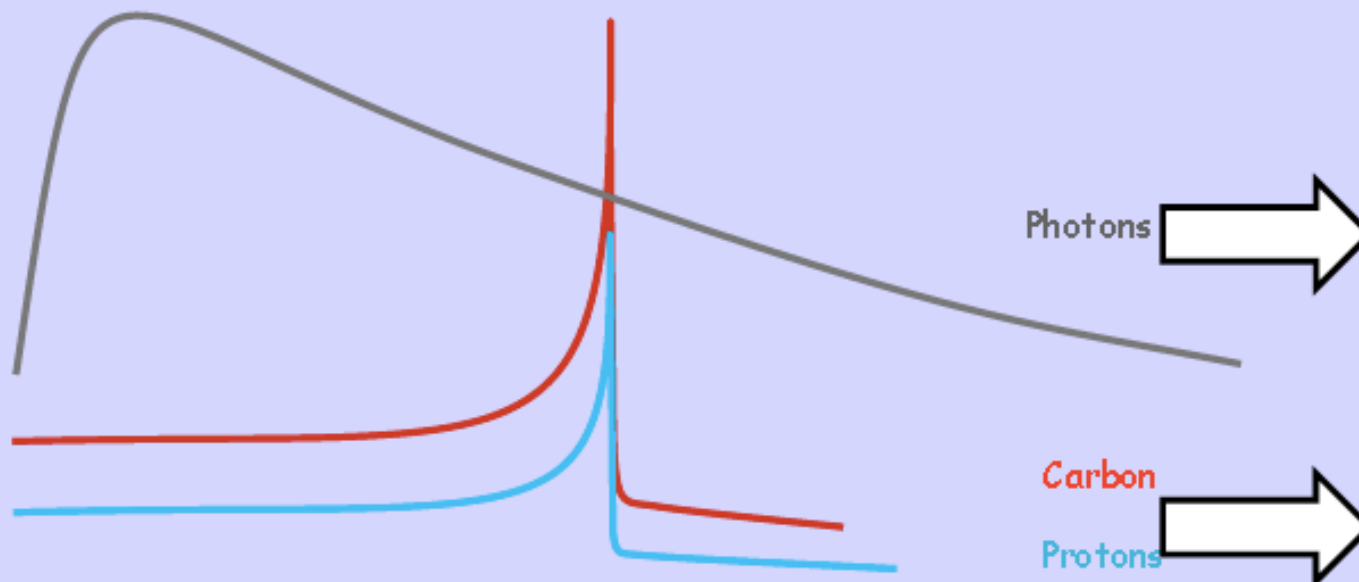


# Why Hadron Therapy



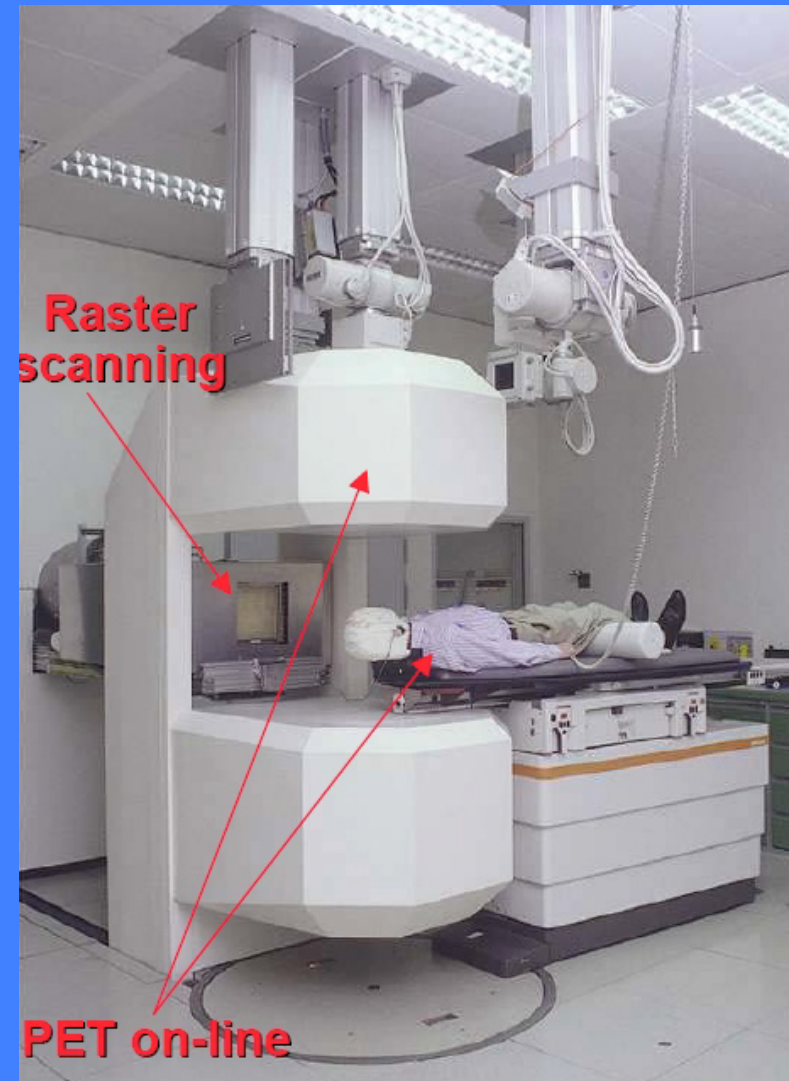
Tumors near critical organs  
Tumors in children  
Radio-resistant tumors

# Photons vs Protons

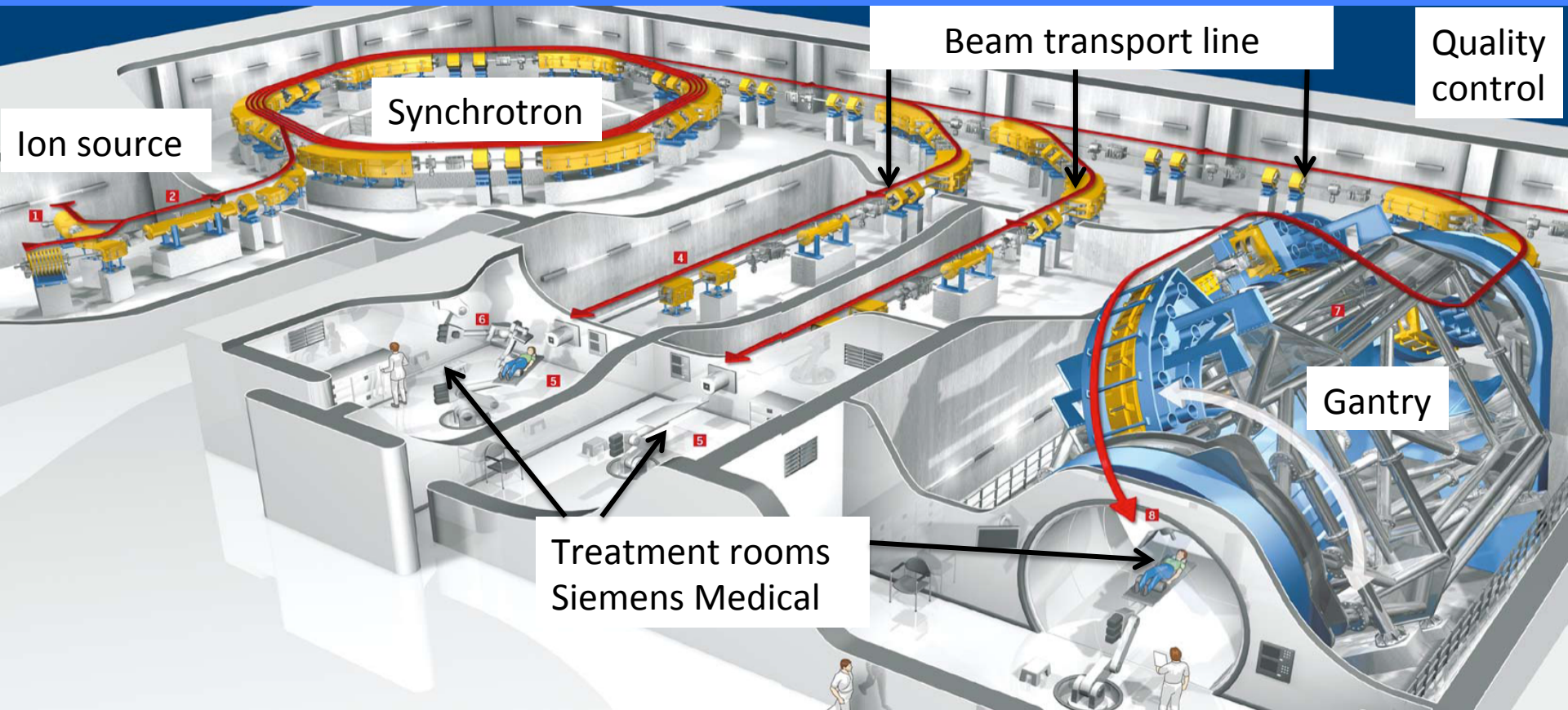


# Carbon Ions Project : pilot project in Europe

- GSI – Darmstadt (1997 – 2008)
- G. Kraft (GSI) & J. Debus (Heidelberg)  
– 450 patients treated with carbon ions







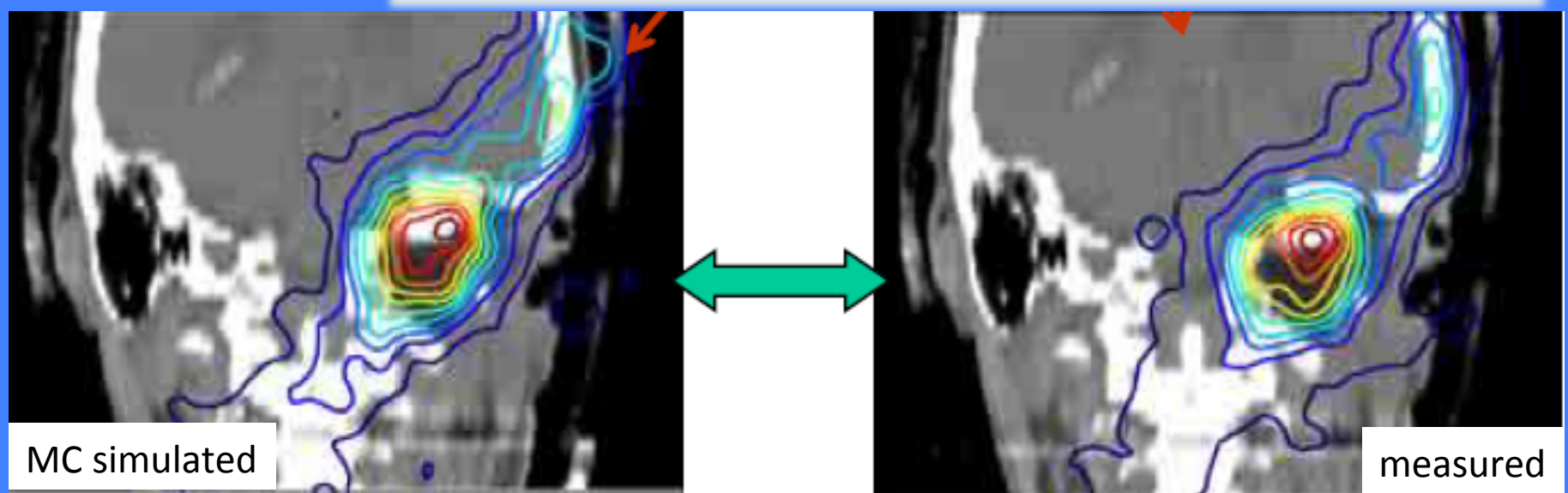


# HIT - Heidelberg





# Real time Monitoring



On-line determination of the dose delivered  
First time in 110 years !

## Modeling of beta+ emitters:

- Cross section
- Fragmentation cross section
- Prompt photon imaging
- Advance Monte Carlo codes





# PIMMS project at CERN (1996-2000)

- Proton Ion Medical Machine Study

400 MeV/u synchrotron

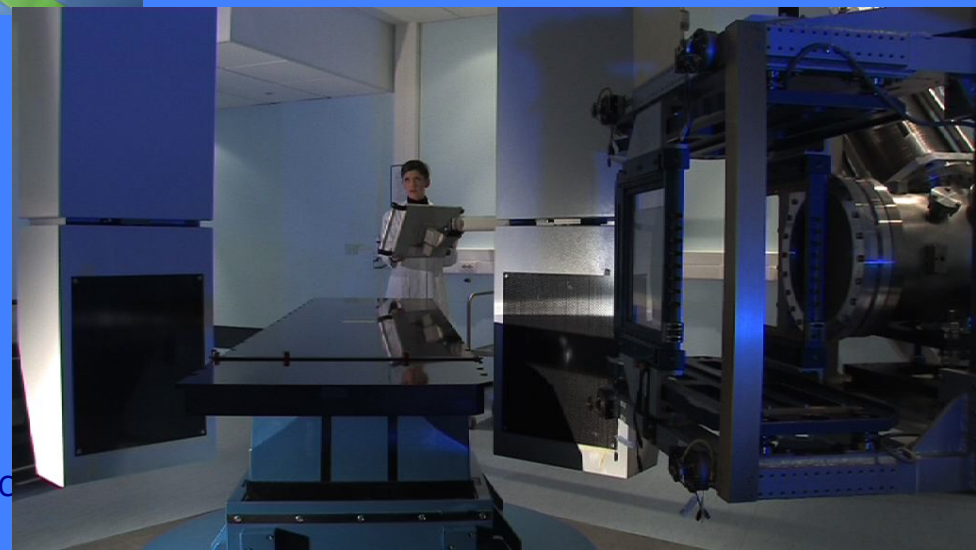
Circumference  $C = 76.84 \text{ m}$   
Tune horizontal  $Q_x = 1.67$   
Tune vertical  $Q_z = 1.72$







# CNAO – Pavia Italy



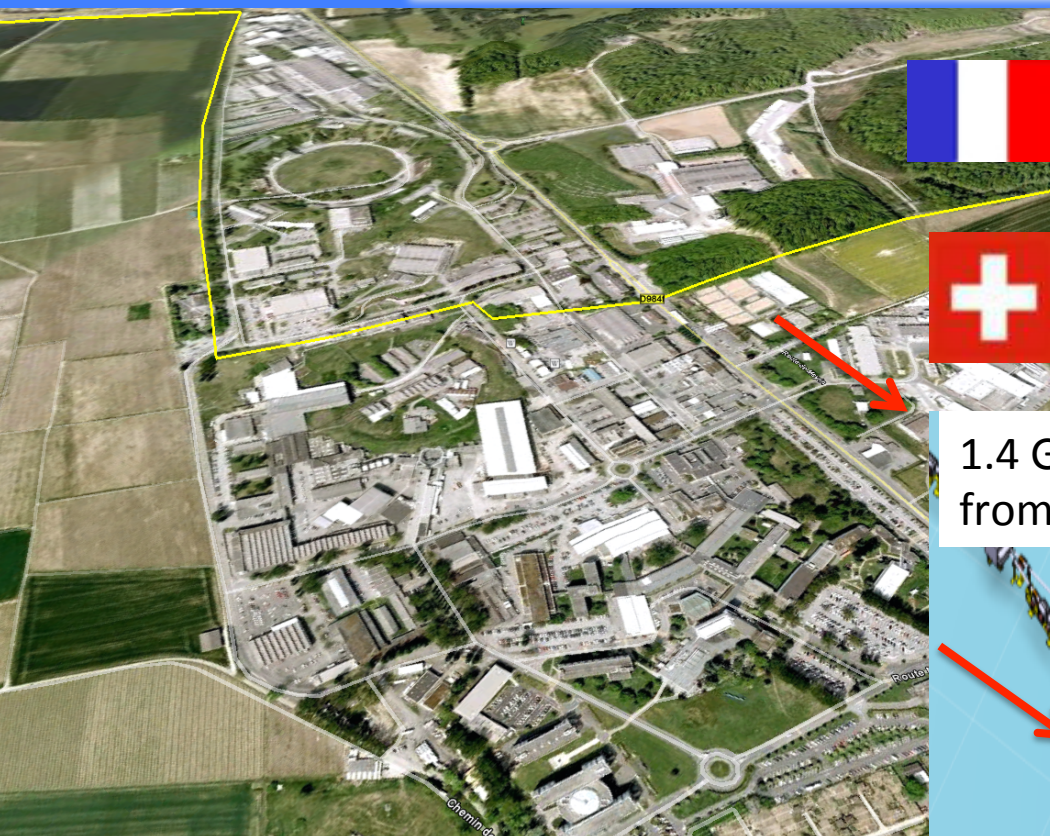
31/08/2019

E Gazis/C

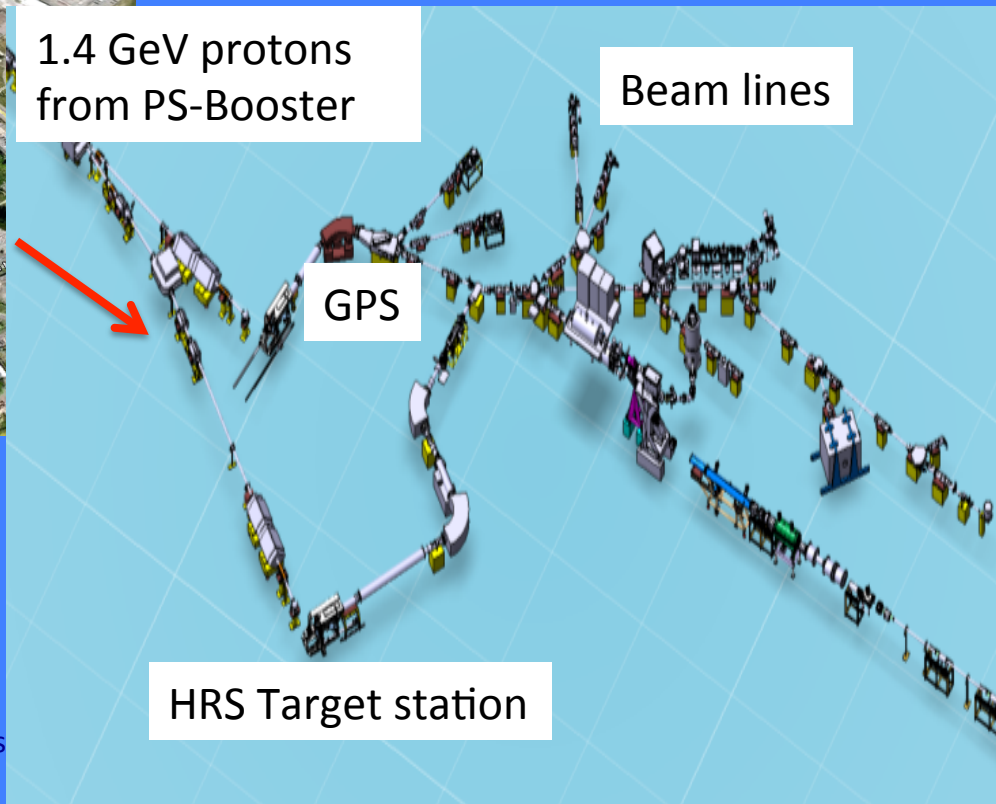


# ISOLDE

## Isotopes for detection and treatment



In collaboration with University Hospital Geneva





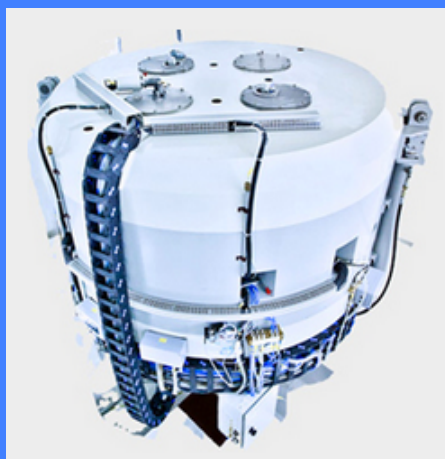
# Radio-Isotopes production

- Nuclear Reactions



- Accelerators

- Circular

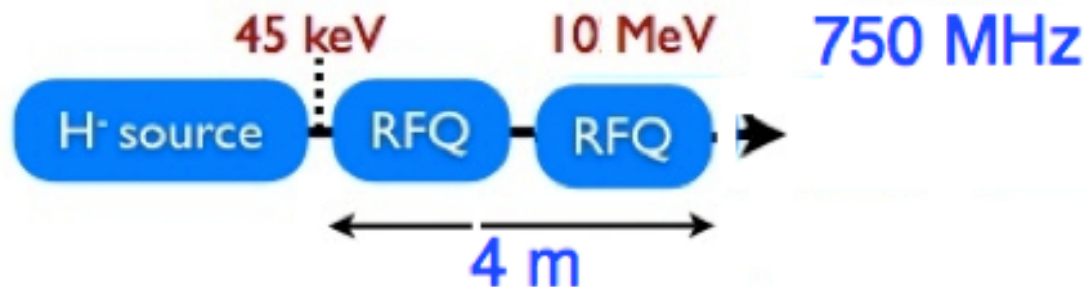


- Linacs



# LINAC for Radio-Isotopes production

The Linac basic architecture is shown here:



**RFQ:**  
CERN BE-ABP-  
HSL

The first linac designed in USA with energy 7 MeV

Medical isotopes

<sup>11</sup>C, <sup>15</sup>O, <sup>18</sup>F, <sup>111</sup>In, <sup>123</sup>I





# LINAC for Radio-Isotopes production

## Compact Linear Accelerator for Radio-Isotope production in Hospitals



### Advantages vs Cyclotrons:

- NO beam loss,
- NO radiation, min. shileding
- Minimum maintenance
- Light weight, Power (25 kW @ 1.5% duty)

A

2 RFQs  
Source  $W = 40$  KeV  
 $L = 3.2$  m  
Output  $W = 8$  MeV  
Freq. = 750  
Aver. current = 50 mA  
Duty cycle = 5 %  
Peak current = 1 mA

2 RFQs  
Source  $W > 70$  keV  
 $L = 5$  m  
Output  $W = 8$  MeV  
Freq. = 750  
Aver. current = 150 mA  
Duty cycle = 1.5 %  
Peak current = 10 mA

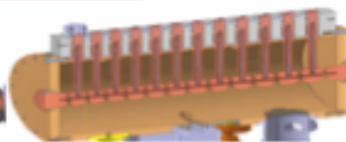
Being designed

B

### $^{99m}\text{Tc}$ for BRACHYTHERAPY



- 1 Klystron as RF power source
- Minimum maintenance, low loss
- Low cost (3 MCHF?)



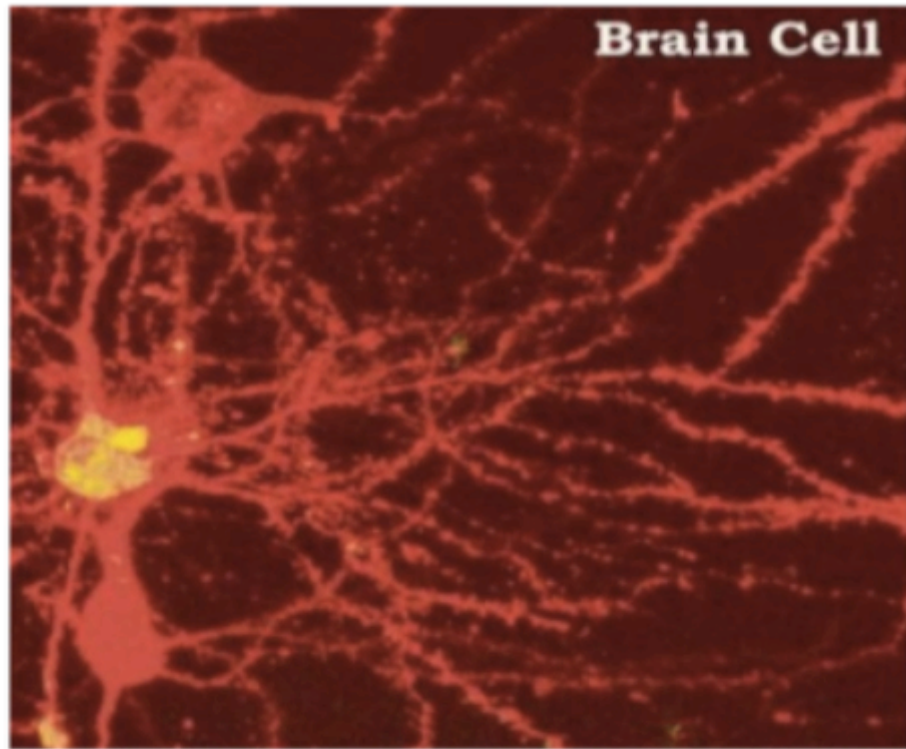
Option:  
 $d^+$ , 18 MeV, 15 m  
For  $^{137}\text{La}$  production

2 RFQs + 1 DTL  
Source  $W = 90$  KeV  
 $L = 10$  m  
Output  $W = 18$  MeV  
Freq. = 704 MHz  
Average current = 2 mA  
Peak current = 20 mA  
Duty cycle = 10 %

# Radio-Isotopes for PET

| Isotope      | Half -time      | Maximum Energy (MeV) | Range in H <sub>2</sub> O (mm) |
|--------------|-----------------|----------------------|--------------------------------|
| <b>F-18</b>  | <b>109.7min</b> | <b>0.635</b>         | <b>2.39</b>                    |
| <b>C-11</b>  | <b>20.4min</b>  | <b>0.96</b>          | <b>4.11</b>                    |
| <b>N-13</b>  | <b>9.96min</b>  | <b>1.19</b>          | <b>5.39</b>                    |
| <b>O-15</b>  | <b>2.07min</b>  | <b>1.72</b>          | <b>8.2</b>                     |
| <b>Rb-82</b> | <b>1,27min</b>  | <b>3,150</b>         | <b>15,50</b>                   |

# Thank you for your attention



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