

# Physics and Dosimetry of Hadron Therapy

*Lembit Sihver*

*Nuclear Physics Institute of the CAS, Prague, Czech Republic*

*Chalmers University of Technology, Gothenburg, Sweden*

*TU Wien, Wien, Austria*

# **Ionizing Radiation of Interest for External Radiation Therapy**

- **Photons ( $\gamma$ , X ray)**
  - **Charge: 0**
  - **Indirect Ionization**



# **Ionizing Radiation of Interest for External Radiation Therapy**

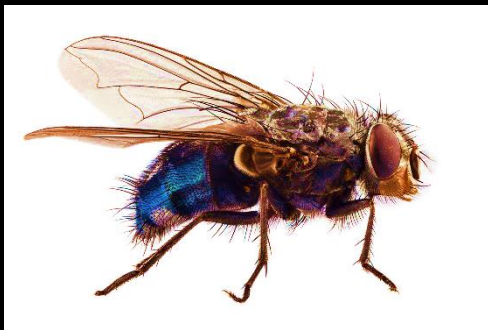
## ■ **Photons ( $\gamma$ , X ray)**

- **Charge: 0**
- **Indirect Ionization**



## ■ **Electrons**

- **Charge: -1**
- **Direct Ionization**
- **Mass: 0.512 MeV**



# **Ionizing Radiation of Interest for External Radiation Therapy**

## ▪ **Photons ( $\gamma$ , X ray)**

- Charge: 0
- Indirect Ionization



## ▪ **Electrons**

- Charge: -1
- Direct Ionization
- Mass: 0.512 MeV



## ▪ **Pions<sup>-</sup> ( $\pi^-$ )**

- Charge: -1
- Unstable
- The pion is absorbed by the nucleus, and the  $\pi^-$  rest mass of 140 MeV appears in the form of kinetic energy of nuclear fragments, except for about 40 MeV, which is used in overcoming the binding energy of the nucleus.
- Mass: 138 MeV ( $273 \times m_e$ )



# Particles of Interest for External Radiation Therapy

## ■ Protons

- Charge: +1
- Direct Ionization
- Mass  $\sim 938 \text{ MeV}$  ( $2,000 \times m_e$ )

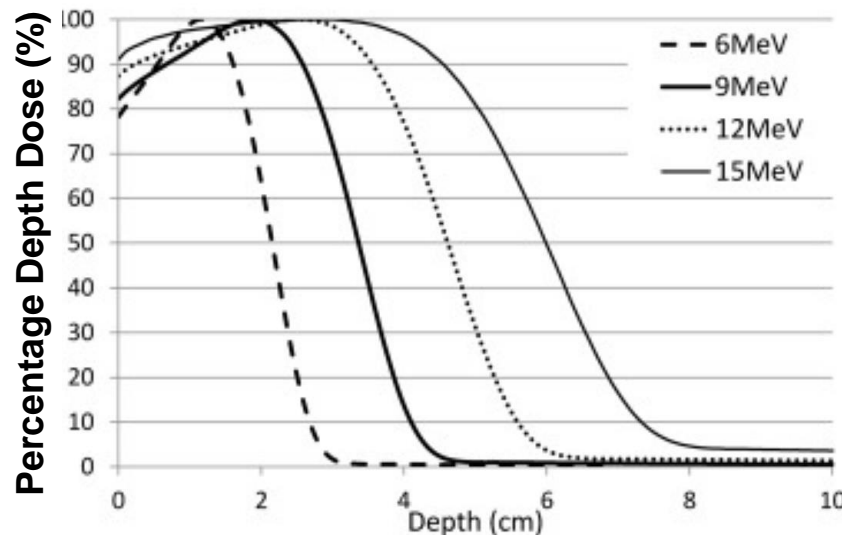
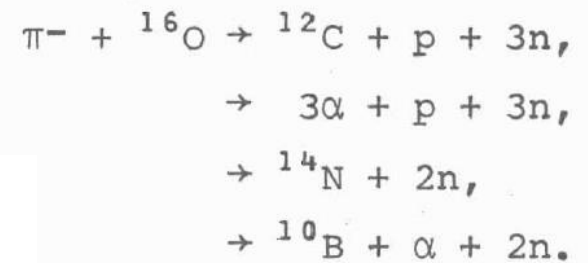
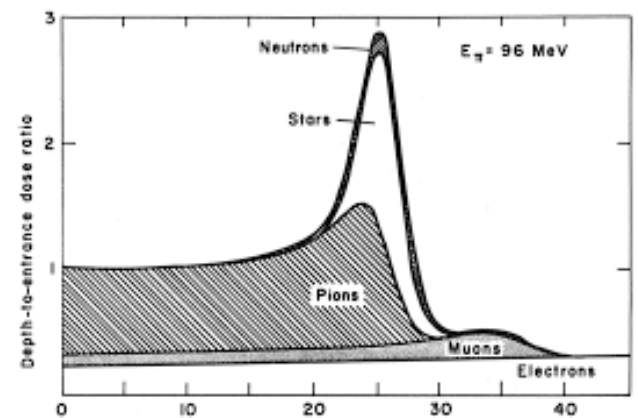
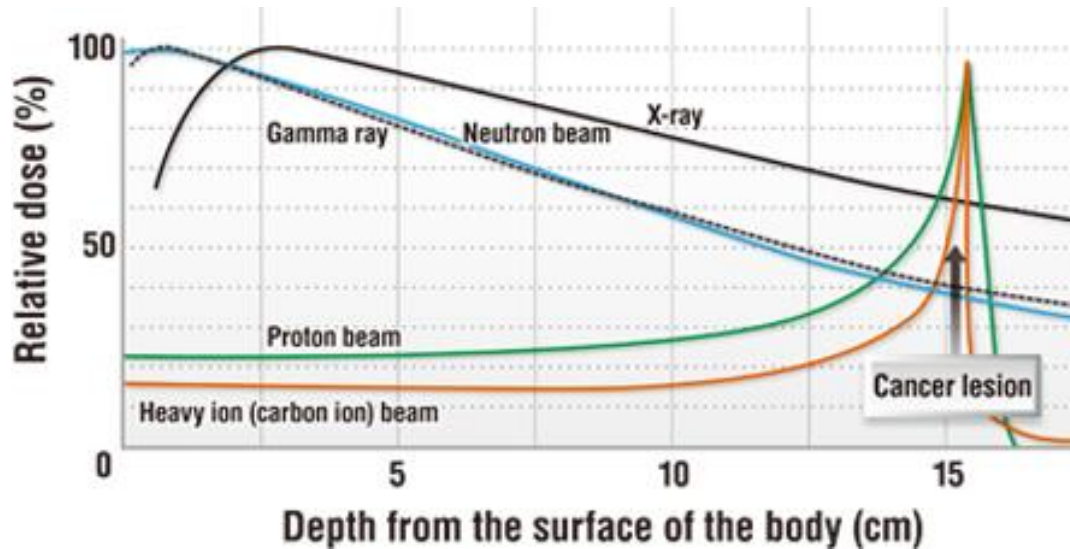


## ■ Carbon ions

- Charge: +6
- Direct Ionization
- Mass  $\sim 12 \times m_p$



# Different Deth-Dose Distributions



# Ionizing Radiation of Interest for External Radiation Therapy

## ▪ Neutrons

- Charge: 0
- Indirect Ionization
- Mass  $\sim 938 \text{ MeV}$  ( $2,000 \times m_e$ )



## ▪ Protons

- Charge: +1
- Direct Ionization
- Mass  $\sim 938 \text{ MeV}$  ( $2,000 \times m_e$ )



# HADRONS

Subatomic particles made of two or more quarks held together by the strong interaction

## ▪ Carbon ion

- Charge: +6
- Direct Ionization
- Mass  $\sim 12 \times m_p$





# Particles of Interest for External Radiation Therapy

## ■ Protons

- Charge: +1
- Direct Ionization
- Mass  $\sim 938 \text{ MeV}$  ( $2,000 \times m_e$ )



## ■ Carbon ions

- Charge: +6
- Direct Ionization
- Mass  $\sim 12 \times m_p$





# Dosimetry

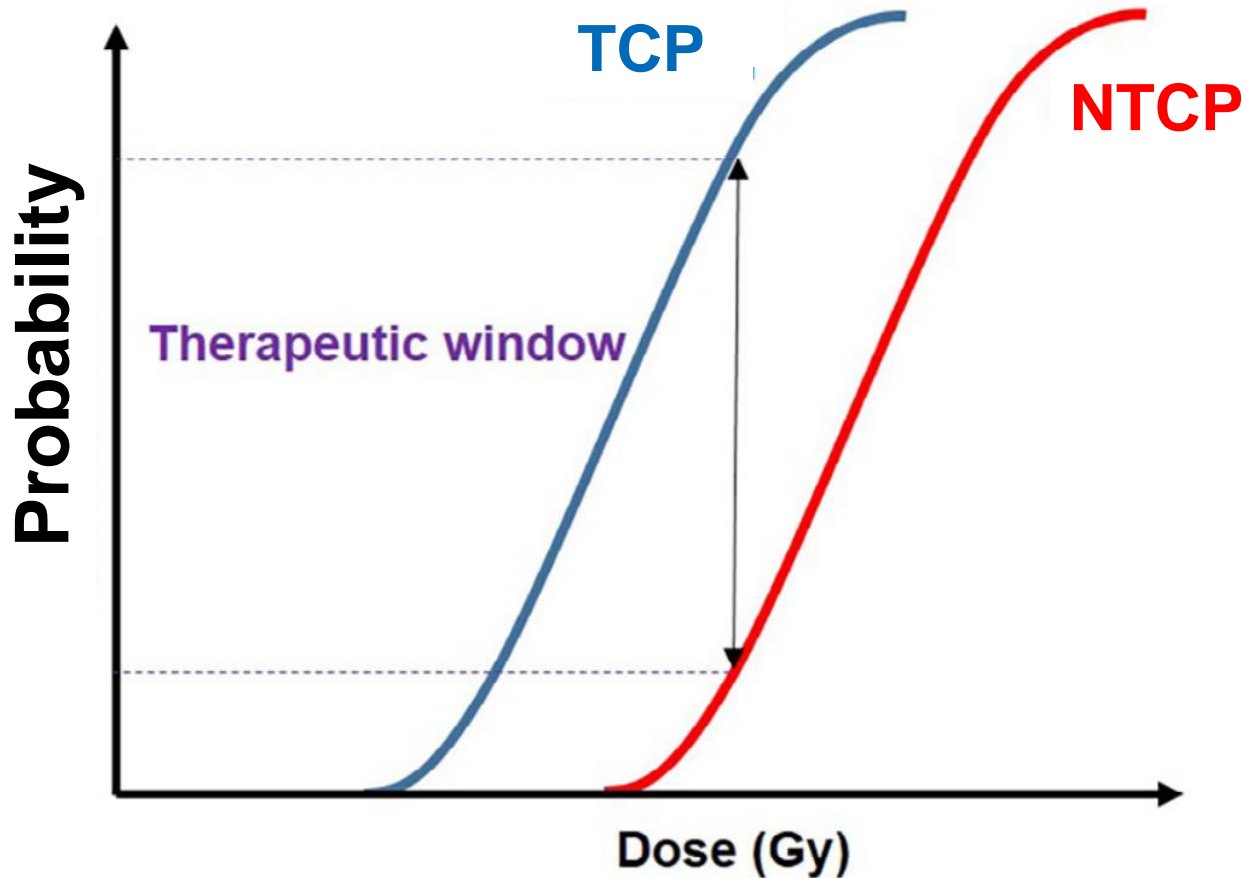
Dosimetry is the determination of absorbed dose in matter or tissue resulting from exposure to ionizing

$$D = \frac{dE}{dm} \left[ \frac{\text{J}}{\text{kg}} = \text{Gy} \right]$$

# **Dosimetry**

**Why do we care about the  
dosimetry?**

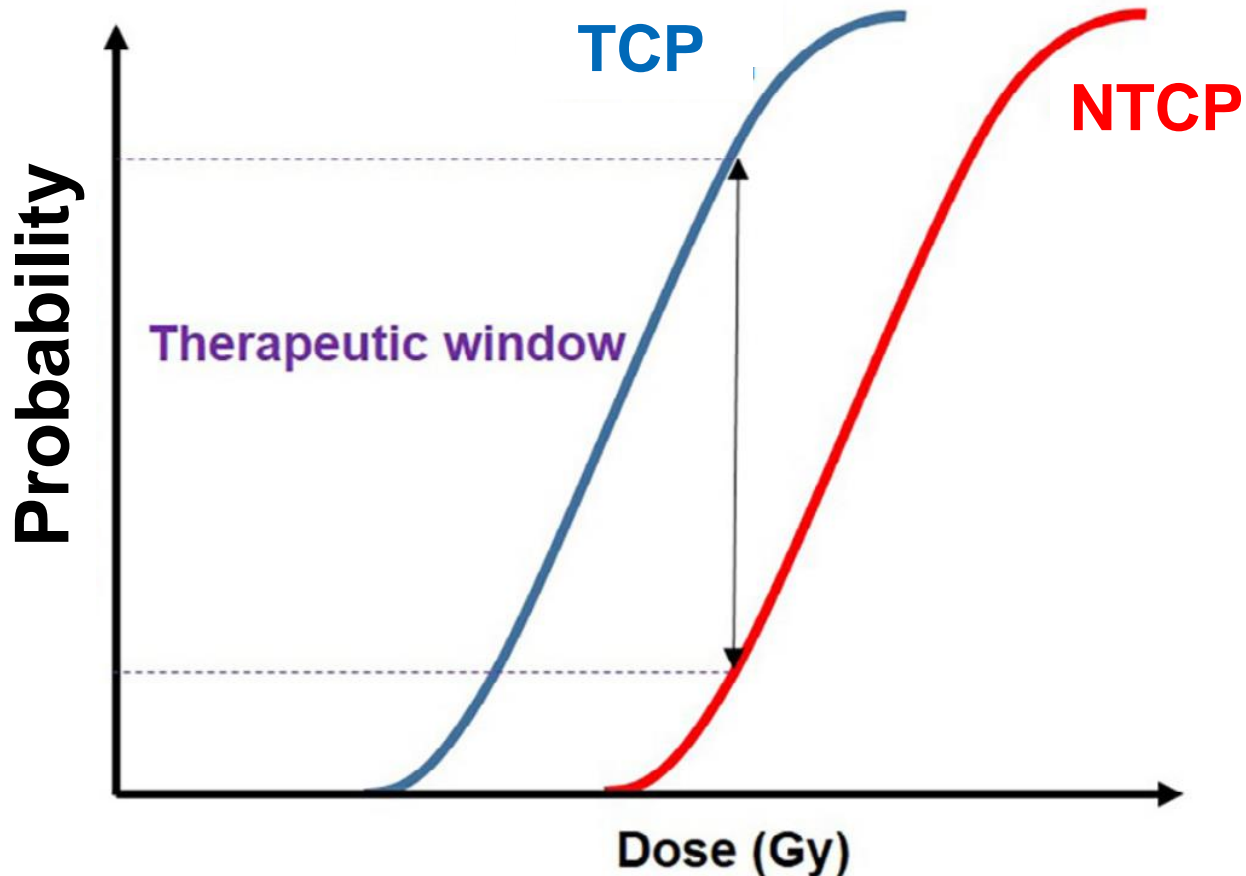
# The Need for Dosimetry



**TCP = Tumor Control Probability**

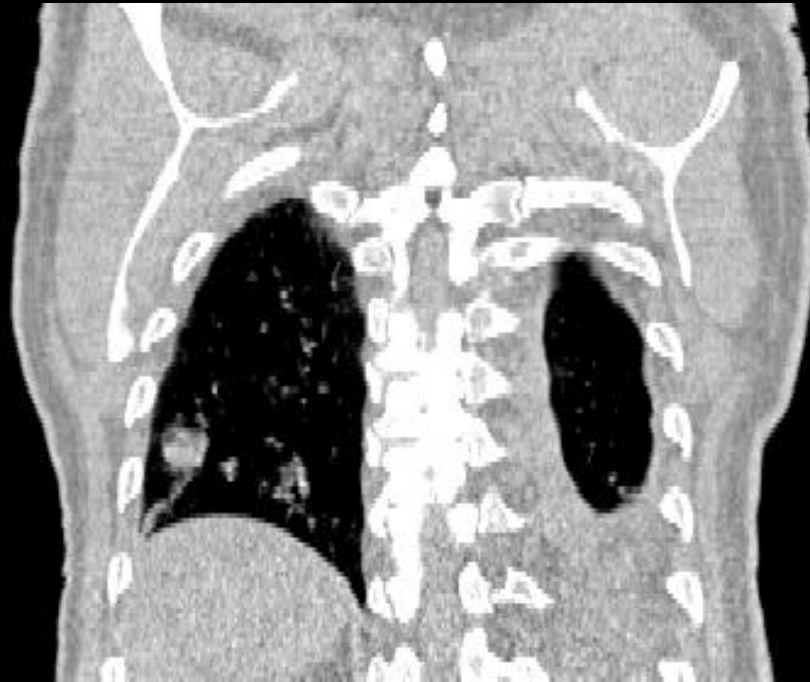
**NTCP = Normal Tissue Complication Probability**

# The Need for Dosimetry



Due to the steep slopes of TCP and NTCP, a 5% dose error can lead to a TCP change of 10-20% and even more for NTCP!!

**The movements and changes  
in the body makes it especially  
challenging!**



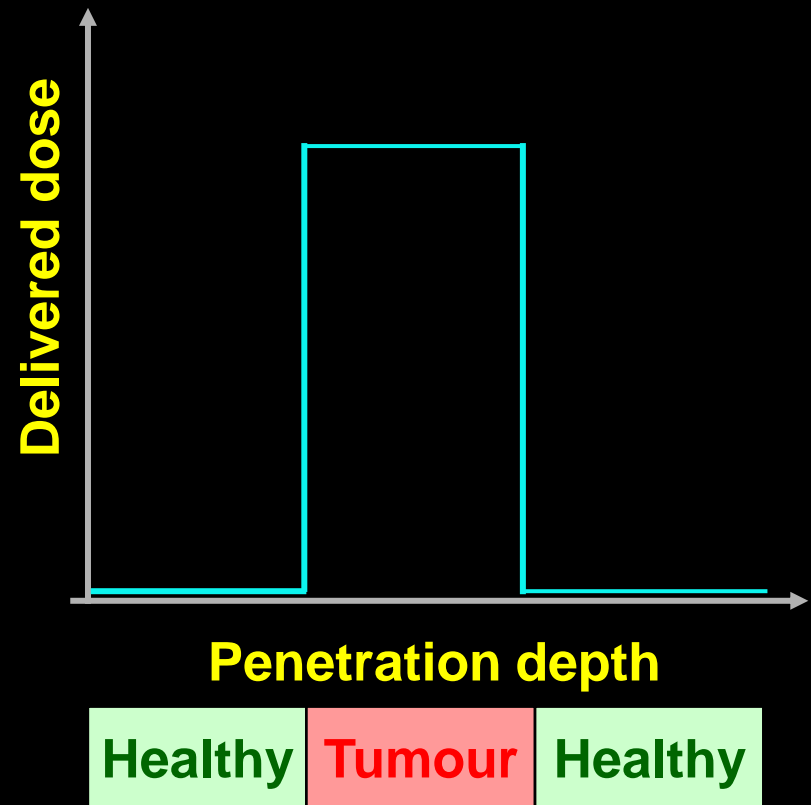
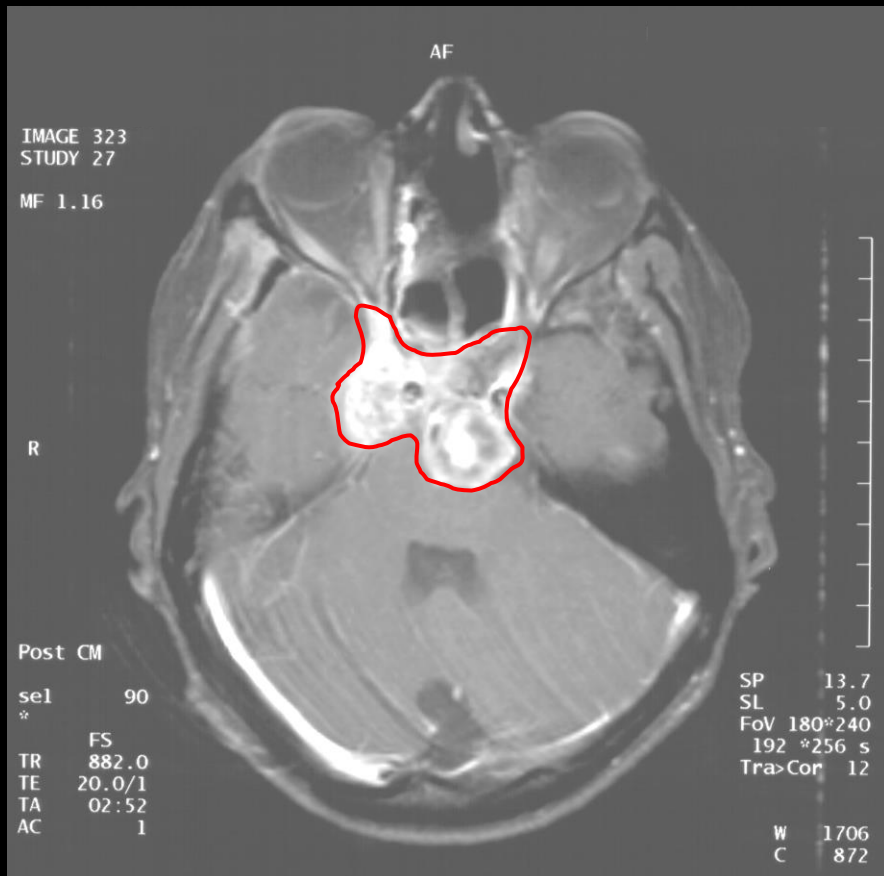
**Before we will go into the dosimetry and its challenges in hadron therapy, let's look at the basic physics of hadron therapy**

# Physical Advantages of Ion Beams

The request of the radiooncologist

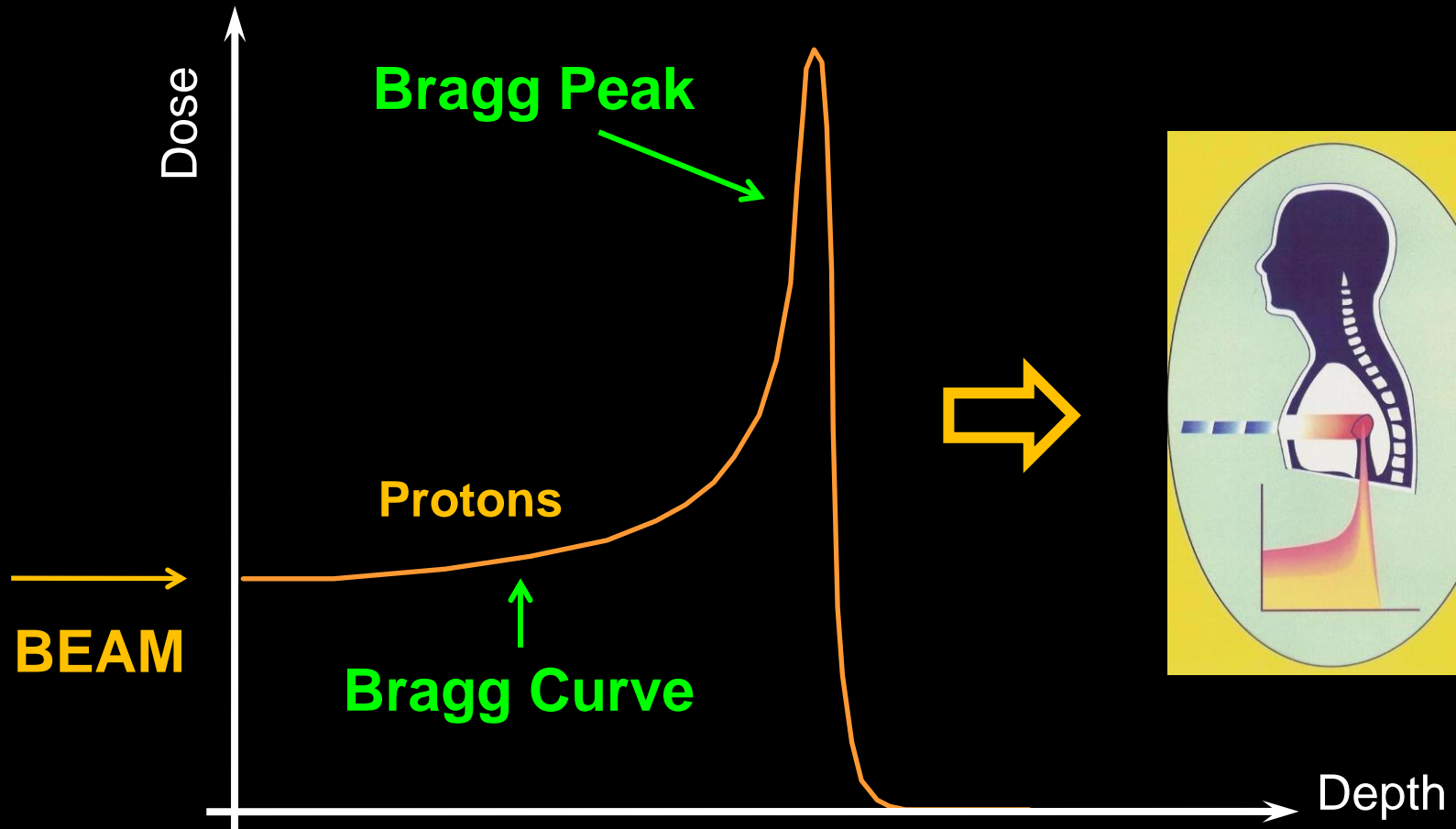
- destroy the tumour tissue

- spare the healthy tissue



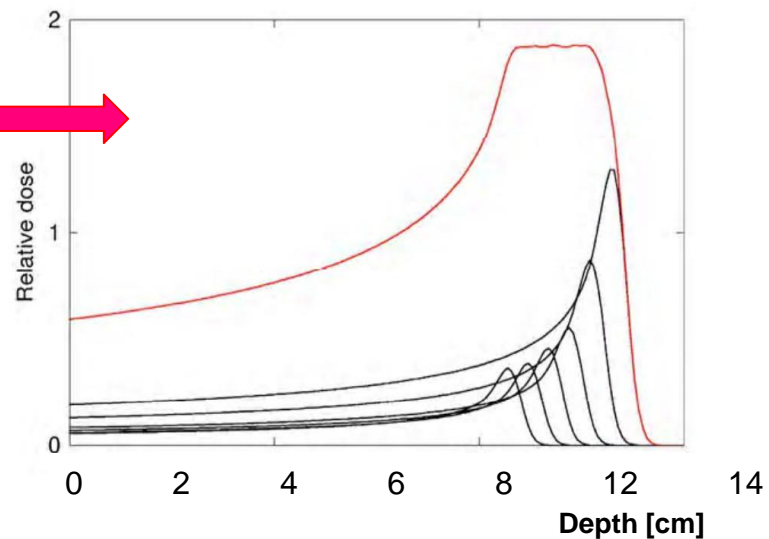
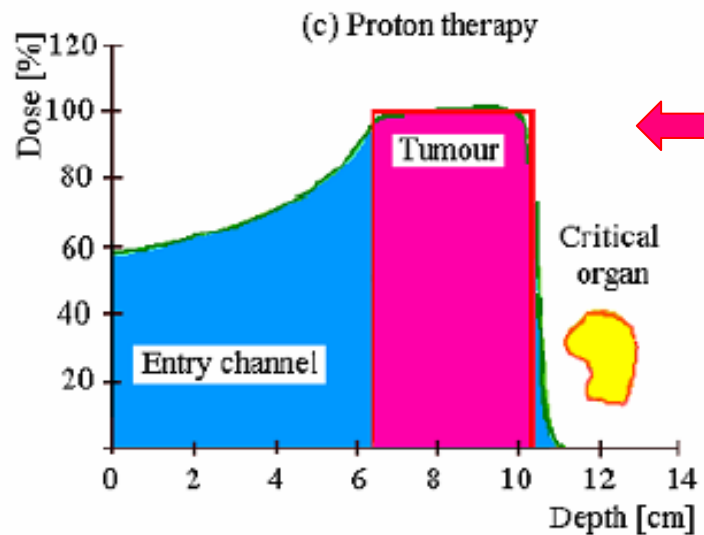
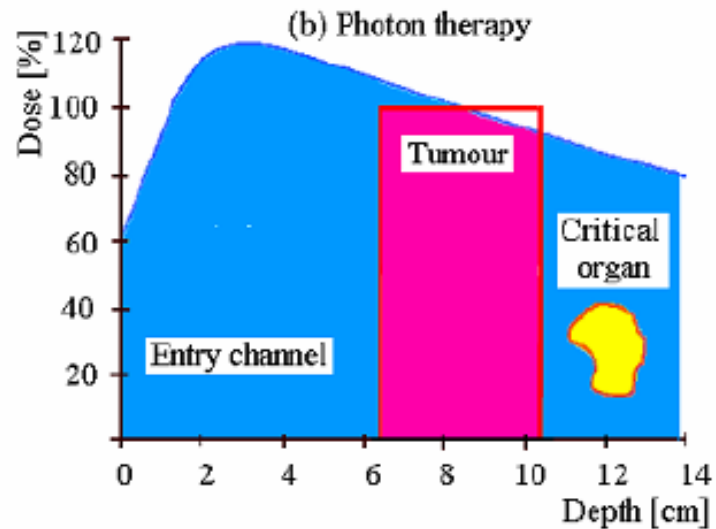
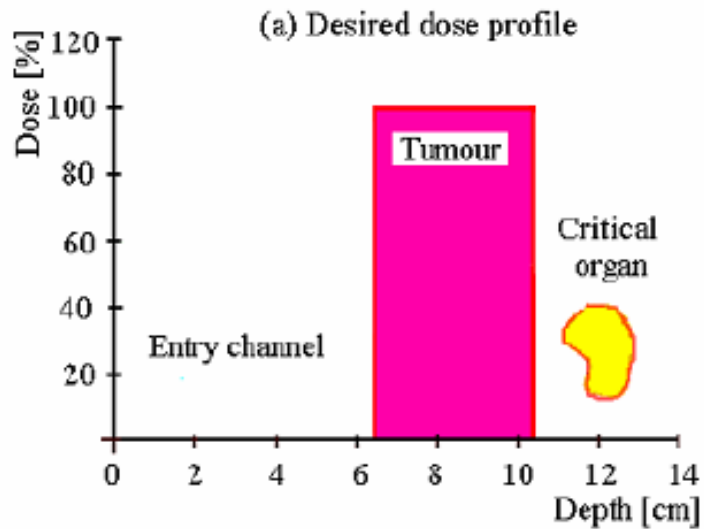


# Why use Proton Beams for Radiotherapy ?



Good dose localization  
(depth-dose distributions)

# Physical Advantages of Ion Beams



# Proton Interactions with Matter

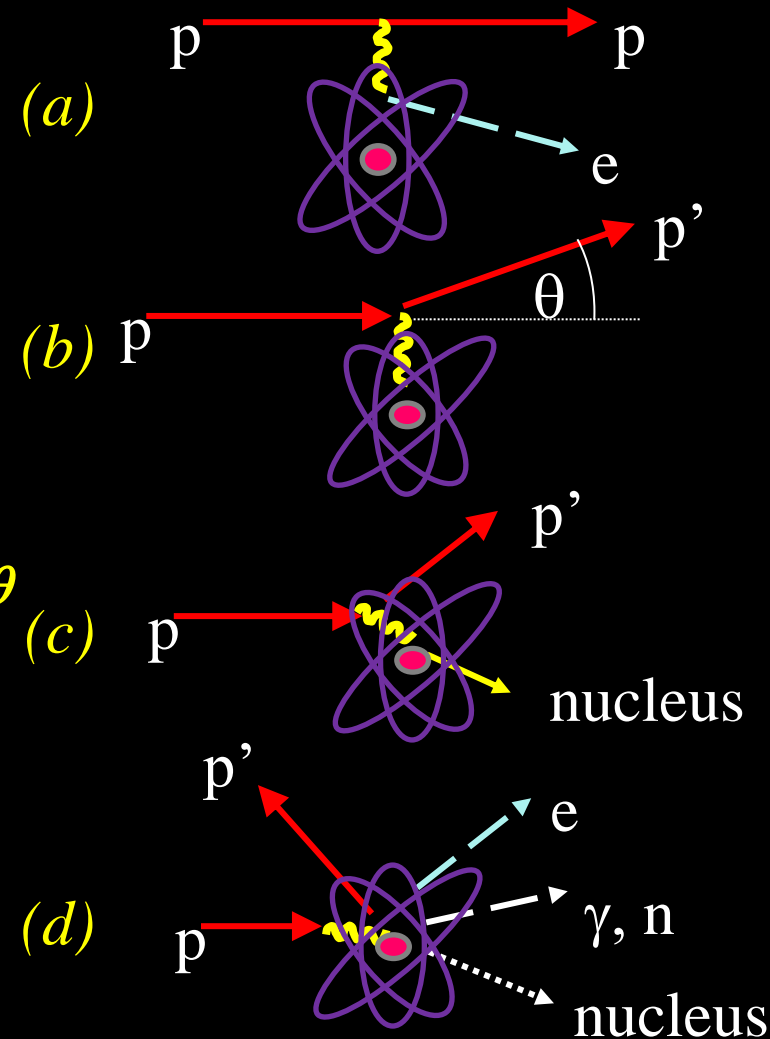
- With electrons mediated by Coulomb force (a)

- Excitation
- Ionization

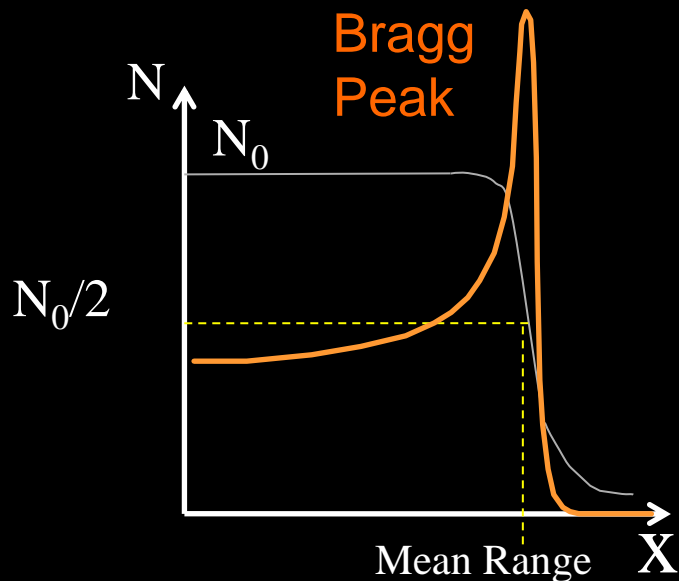
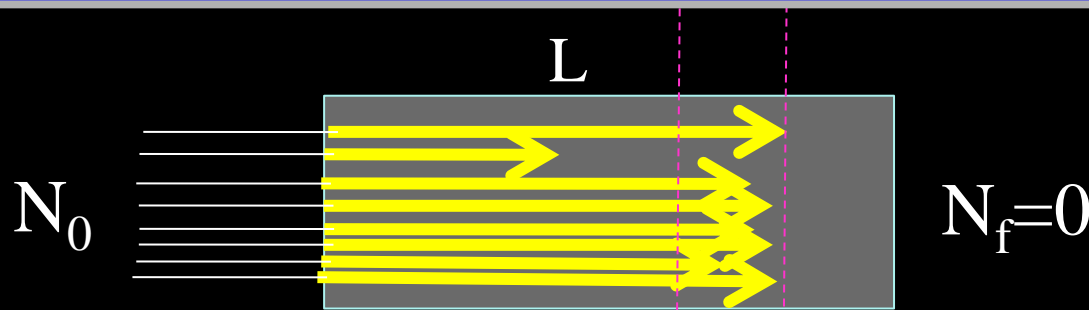
- With nucleus mediated by Coulomb & nuclear forces (b-d)

- Multiple Coulomb scattering (b), small  $\theta$
- Elastic nuclear collision (c), large  $\theta$
- Inelastic nuclear interaction (d)

Mean electron energy  $E_{\text{mean}}$  very low ( $m_p \gg m_e$ )  
 $E_{\text{mean}}$  independent of proton kinetic energy  
Interaction probability higher for slower protons

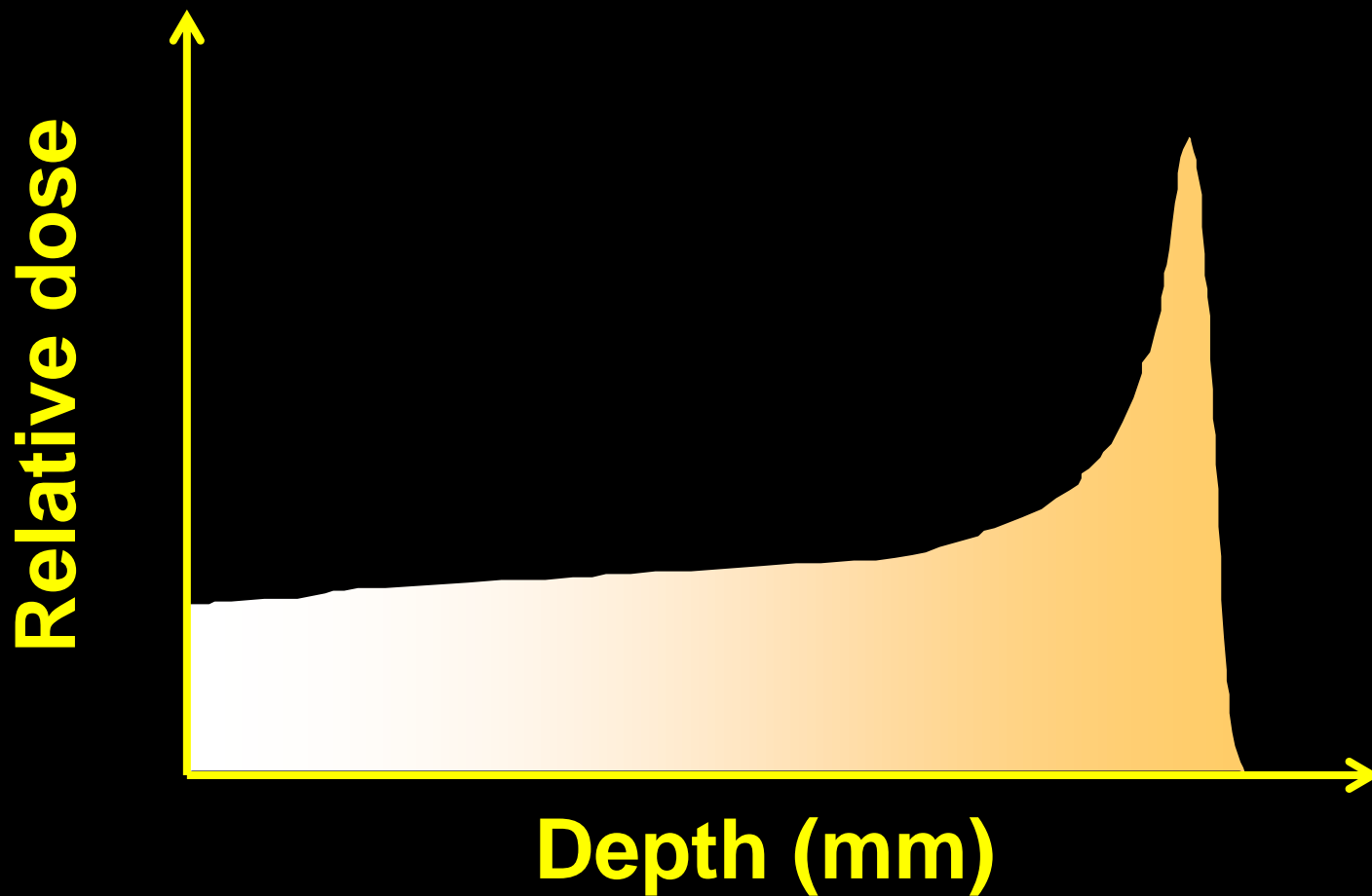


# Particle Beams – Nearly No Attenuation



- A heavy charged particle endures multiple interactions through matter, but “stays” in the beam, because it is deflected only slightly.
- It loses only a small fraction of its energy in each interaction (except in “rare” nuclear interactions) until it stops, i.e., continuous slowing down.
- It deposits most energy near the end!!

**How do we calc. the 1D depth-dose distribution (“Bragg Curve”)**

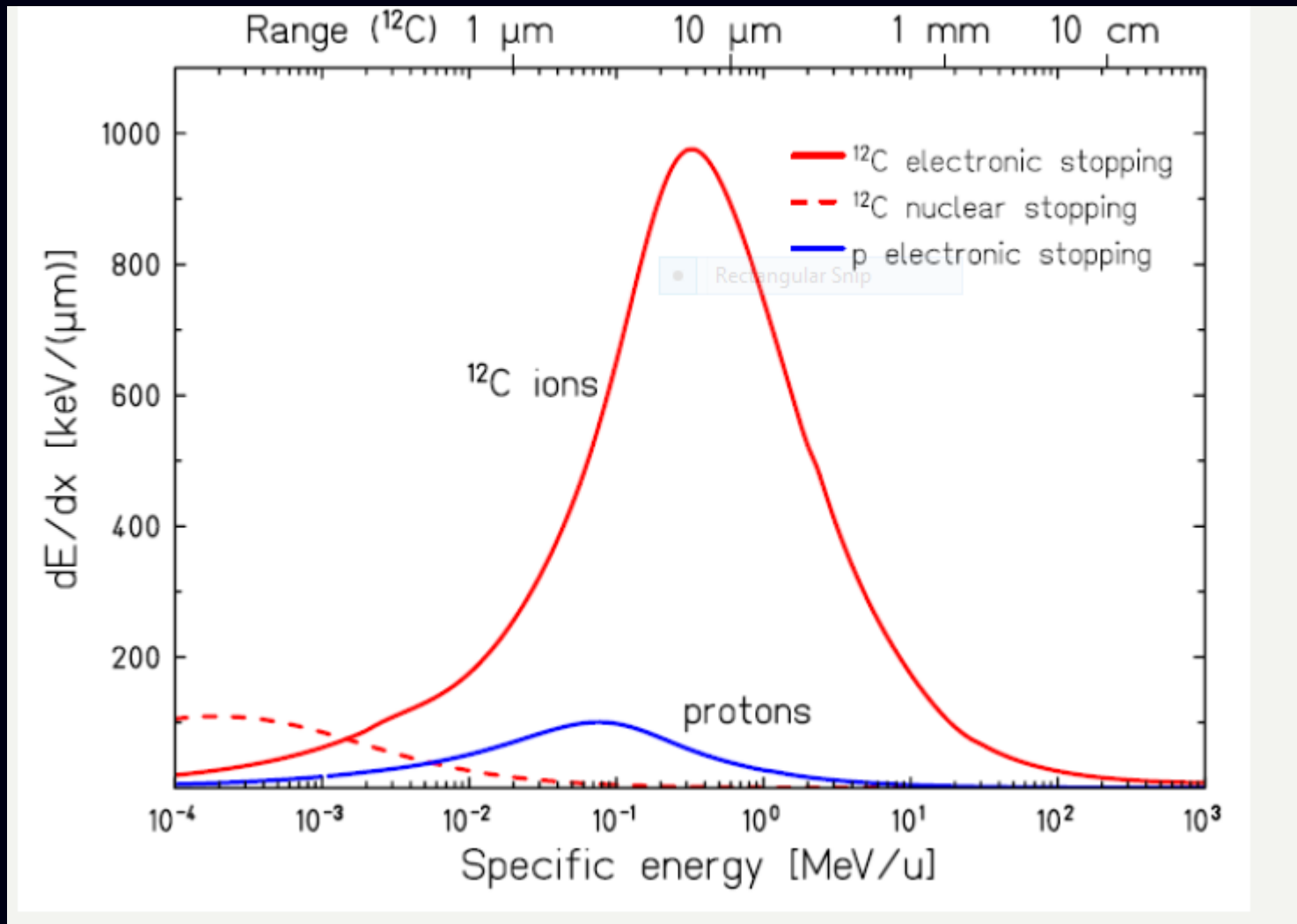


# Linear Stopping Power (S)

$$S = - [(dE/dx)_{el} + (dE/dx)_{nuc} + (dE/dx)_{rad}]$$

- **S** is the loss of energy ( $E$ ) of a charged particle per unit path length ( $x$ )
- **S** has the unit MeV/cm (or more common keV/ $\mu\text{m}$ )
  - $(dE/dx)_{el}$  : *electronic or collision stopping power*
  - $(dE/dx)_{nuc}$  : *nuclear stopping due to elastic Coulomb scattering*
  - $(dE/dx)_{rad}$  : *radiative stopping power due to the emission of bremsstrahlung in the electric fields of the particles in the material traversed*
    - *only important for ions at extremely high energies*

# Linear Stopping Power (S)





## Linear Stopping Power (S)

- Within the range of therapeutically relevant energies for protons and carbon ions, the process of energy loss is dominated by *electronic collisions* and can be described by the:

*Bethe-Bloch Formula* ( $E > \sim 1 \text{ MeV/nucleon}$ )

# Bethe-Bloch Formula for Stopping Power

$$-\left(\frac{dE}{dx}\right)_{coll} = 2\pi N_A r_e^2 m_e c^2 \rho \frac{Z}{A} \frac{z^2}{\beta^2} \left[ \ln\left(\frac{2m_e \gamma^2 v^2 W_{max}}{I^2}\right) - 2\beta^2 - \delta - 2\frac{C}{Z} \right]$$

$c$  : speed of light ( $3 \cdot 10^8$  m / s)

$r_e$  : classical electron radius ( $2.82 \cdot 10^{-15}$  m);  $m_e$  : electron mass (511 keV /  $c^2$ )

$N_A$  : Avogadro constant ( $6.02 \cdot 10^{23}$  molecules / mol)

$I$  : mean excitation potential of the medium;  $I \cong Z I_0$ , with  $I_0 = 10$  eV

$Z, A, \rho$  : atomic number, mass number and mass density of the absorbing medium

$z, v$  : charge (in units of  $e$ ) and speed of the incident particle

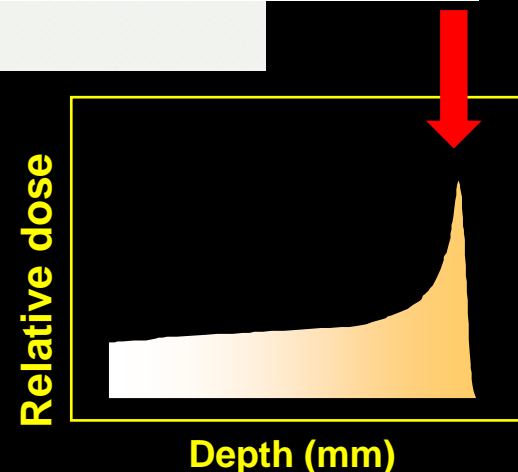
$\beta = v / c$ ;  $\gamma = 1 / (1 - \beta^2)^{1/2}$

$\delta$  : density correction (due to polarization effects in dense med)

$C$  : shell corrections (relevant for  $v \approx$  orbital velocity of the bound electrons; cannot be longer treated as stationary)

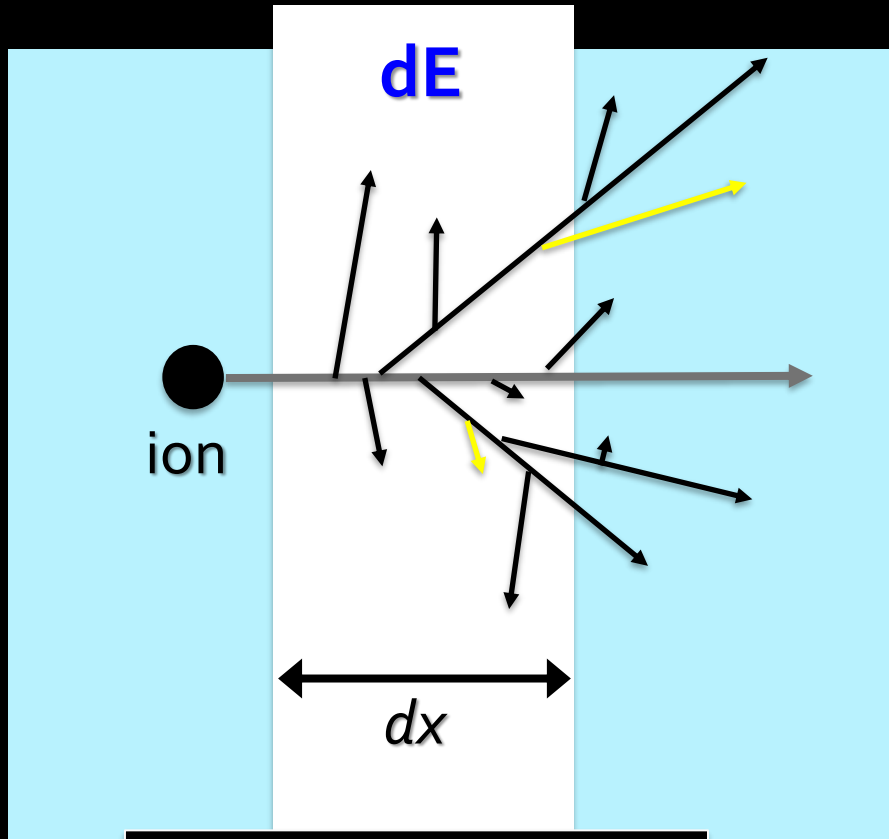
$W_{max}$  : maximum energy transfer in a single collision

Decreased velocity  $\rightarrow$   
increased value of  $(-dE/dx)$



# Linear Energy Transfer (LET)

Macroscopic:  $LET_{\infty} = (dE/dx)_{\infty}$



$LET_{\infty}$  is the unrestricted linear energy transfer

$LET_{\infty}$  is the amount of energy deposited per unit length of a material as a charged particle traverses the material

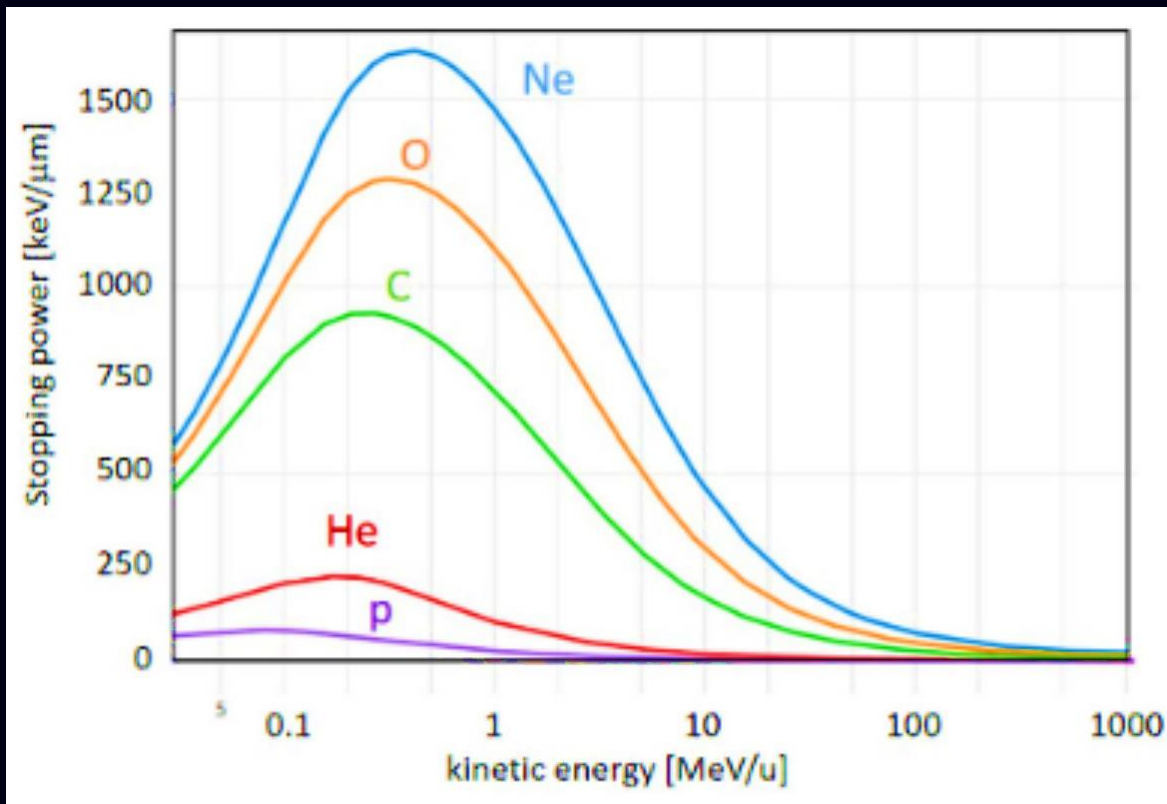
$LET_{\infty}$  has the unit MeV/cm (or more common keV/ $\mu\text{m}$ )

Independent of radial dose distribution

# Linear Stopping Power (S)

$$S = -LET_{el,\infty} = -(dE/dx)_{el,\infty}$$

From now on labelled  $LET_{\infty} = (dE/dx)_{\infty}$



# Mass Stopping Power

$S/\rho$  = mass stopping power [MeV,cm<sup>2</sup>/g]

$\rho$  = density

$$S/\rho = \left( \frac{dE}{\rho dx} \right) \text{ [MeV,cm}^2\text{/g]}$$

The dose from charged particles in some medium (e.g. water):

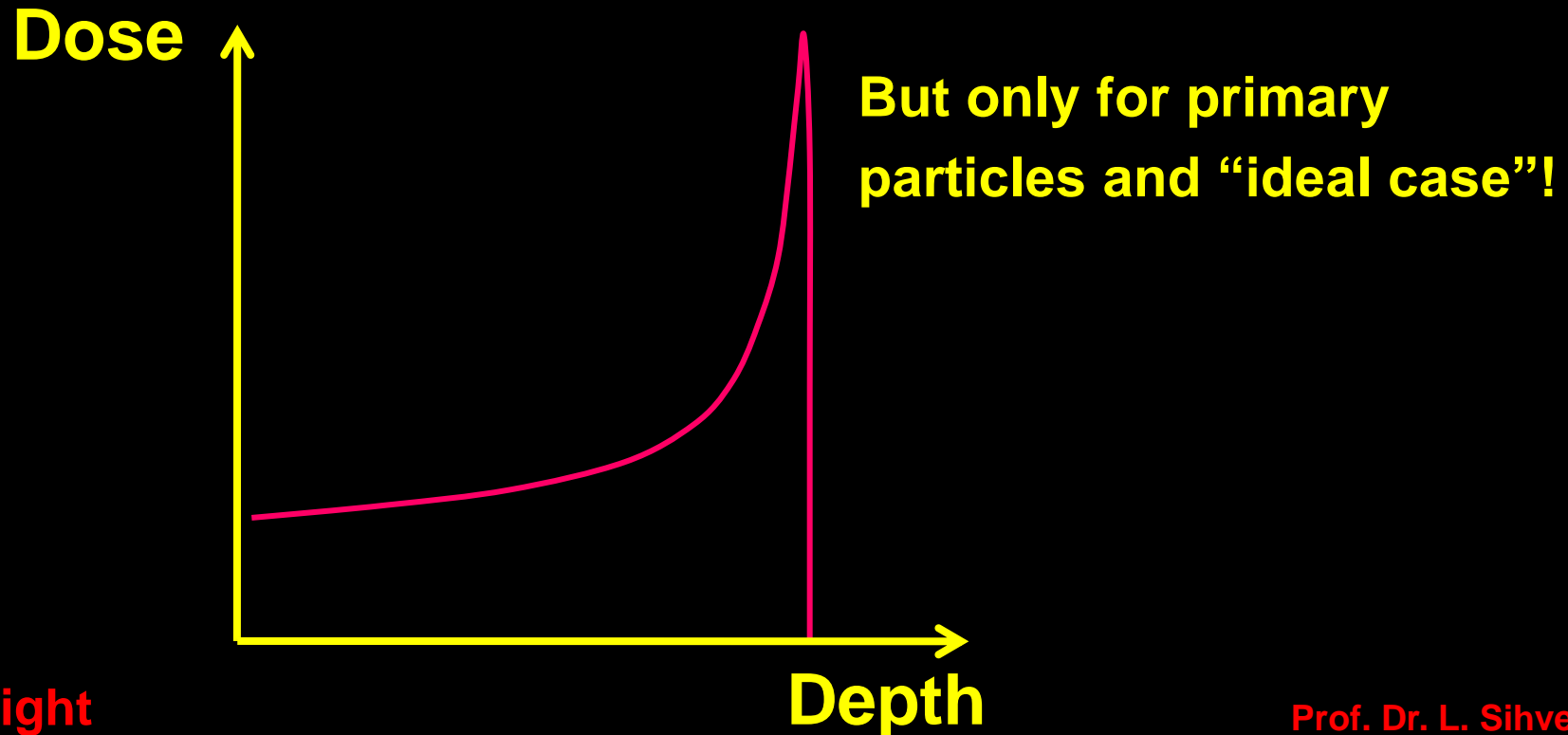
$$D_w = \int_0^{E_0} \phi_E \left( \frac{dE}{\rho dx} \right)_w dE \text{ [Gy]}$$

$\phi_E$  = the particle fluence

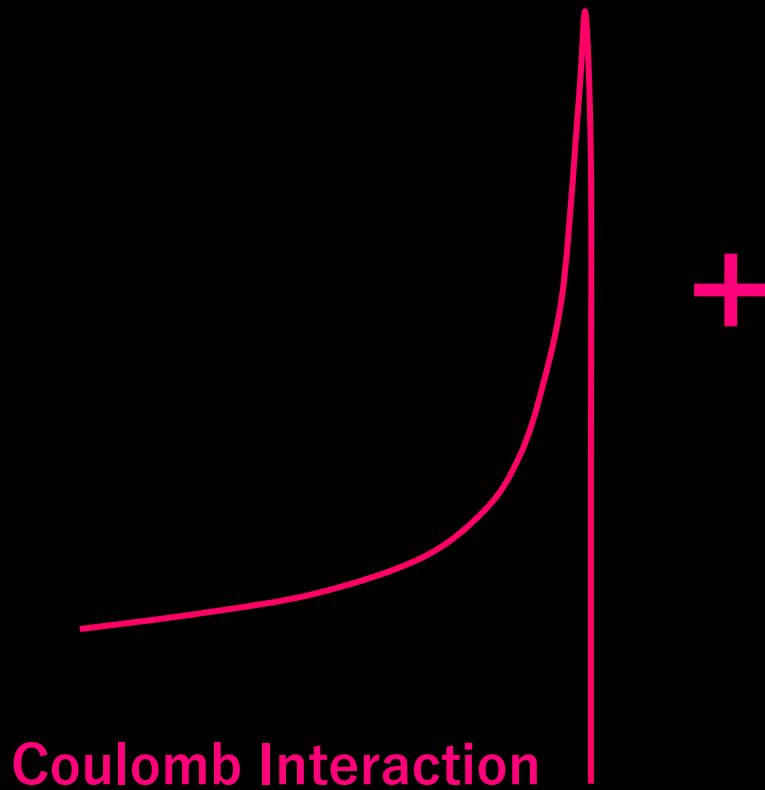
# 1D Depth Dose Distribution - Bragg Curve

Now we can calc. depth-dose distributions:

$$D_w = \int_0^{E_0} \phi_E \left( \frac{dE}{\rho dx} \right)_w dE \quad [\text{Gy}]$$



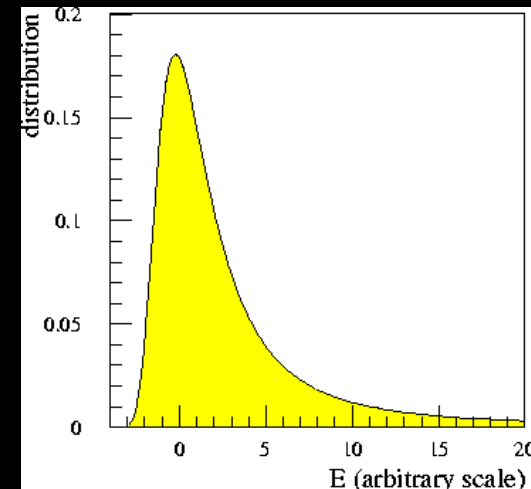
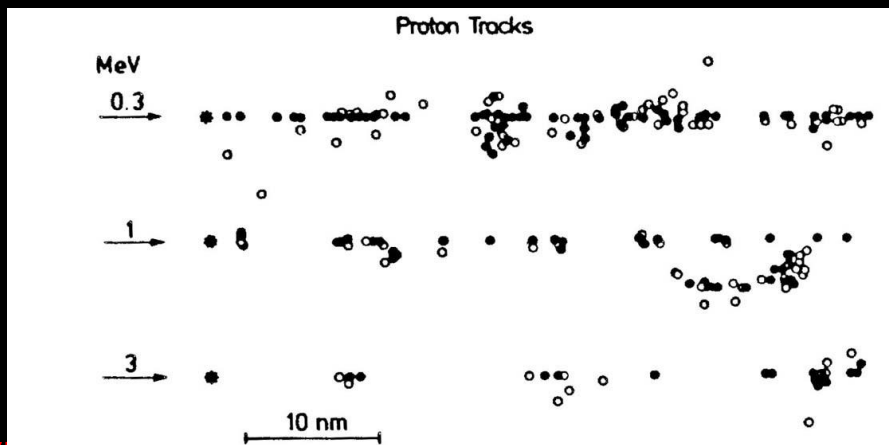
# 1D Depth Dose Distribution - Bragg Curve





# Energy Straggling

- So far we used the **Continuously Slowing Down Approximation (CSDA)**
- The Bethe-Bloch formula gives *the mean energy* lost per unit path length
- In reality, ions lose their energy in individual collisions with electrons
  - Actual energy loss will scatter around the mean value
  - Energy loss distribution is not Gaussian around mean
- The stochastic behavior of energy deposition in matter (*energy straggling*) is described by stochastic distributions such as the Landau-Vavilov distribution



# Range Straggling

Energy straggling → range straggling

Theoretical  
w/o Straggling

Range Straggling  
Distribution

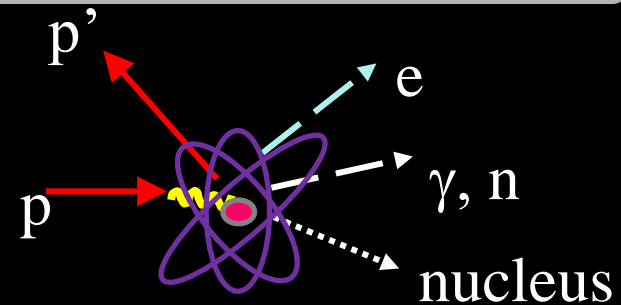
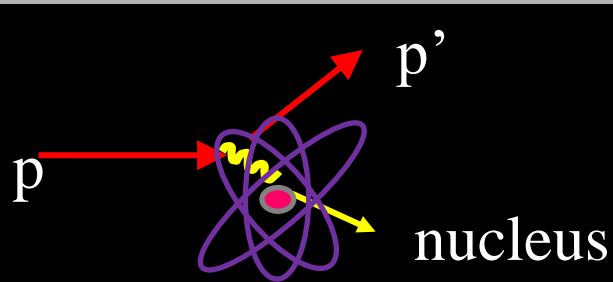
+

+ ?

Coulomb Interaction

Range Straggling

# Nuclear Interactions of Protons



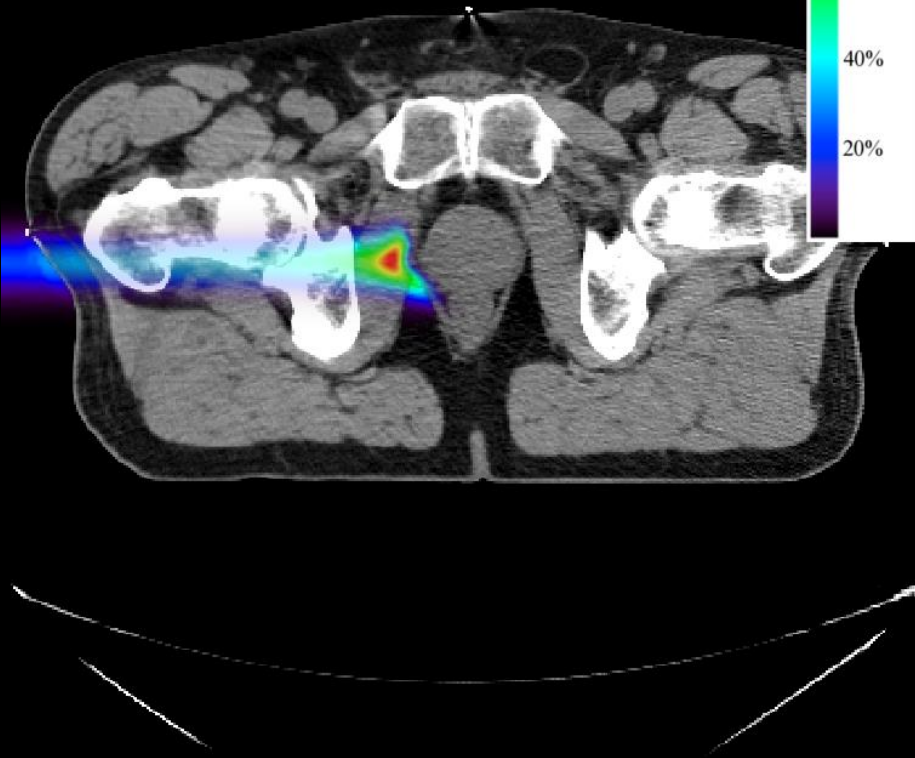
**Elastic collision (large  $q$ )**

**Nuclear interaction**

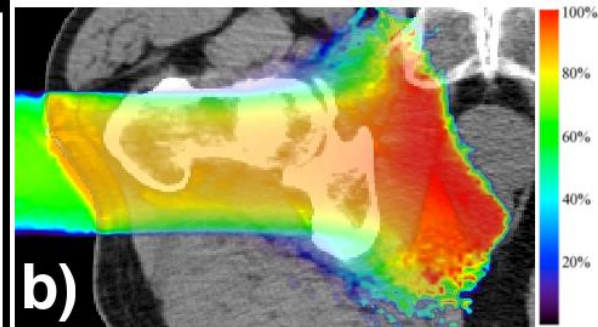
- A certain fraction of protons have nuclear interactions in tissue (about 1% of all protons per cm of penetration)
- Mostly with oxygen and carbon nucleus
- Nuclear interactions cause a decrease in primary proton fluence
- Nuclear interactions lead to secondary particles and thus to local and non-local dose depositions (neutrons!)
- The dose from nuclear interactions is negligible in the Bragg peak
- Target fragments have high LET and therefore high relative biological effectiveness (RBE) and can cause normal tissue complications

# Spatial Distribution

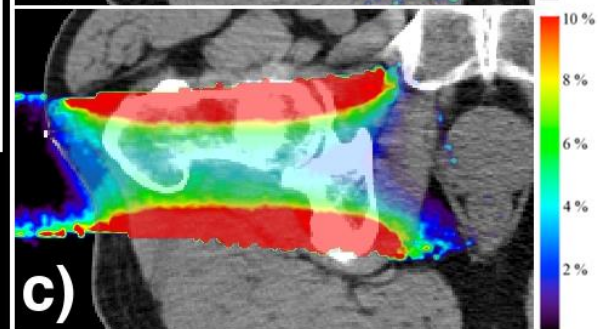
a) total energy deposited



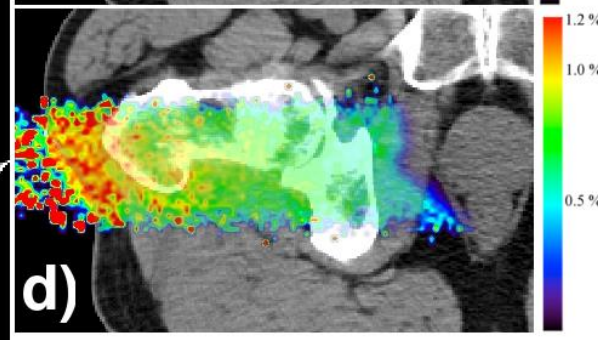
Contribution in %



primary protons



secondary protons



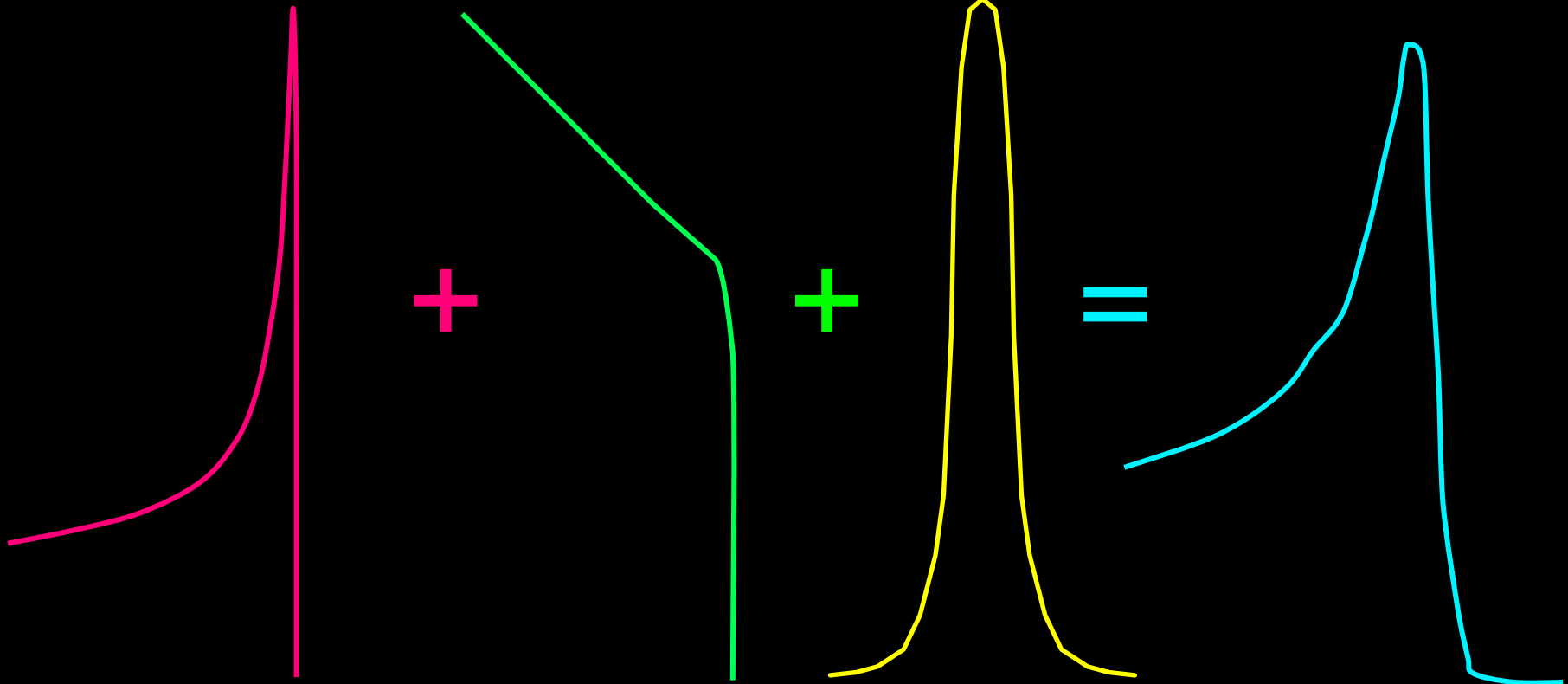
alphas & recoils

# This makes the 1D Bragg Curve!



Nuclear Reaction

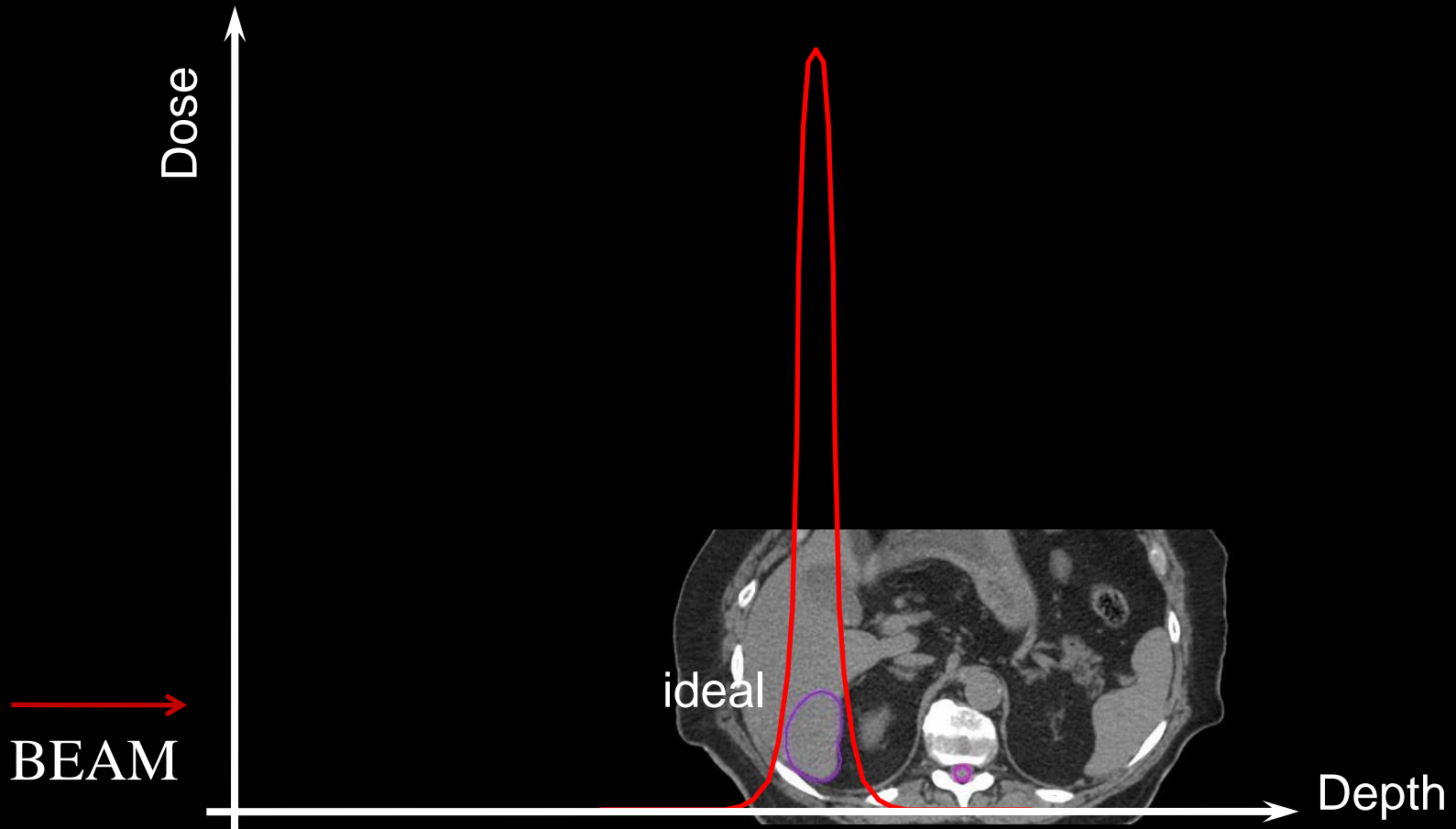
Bragg Curve



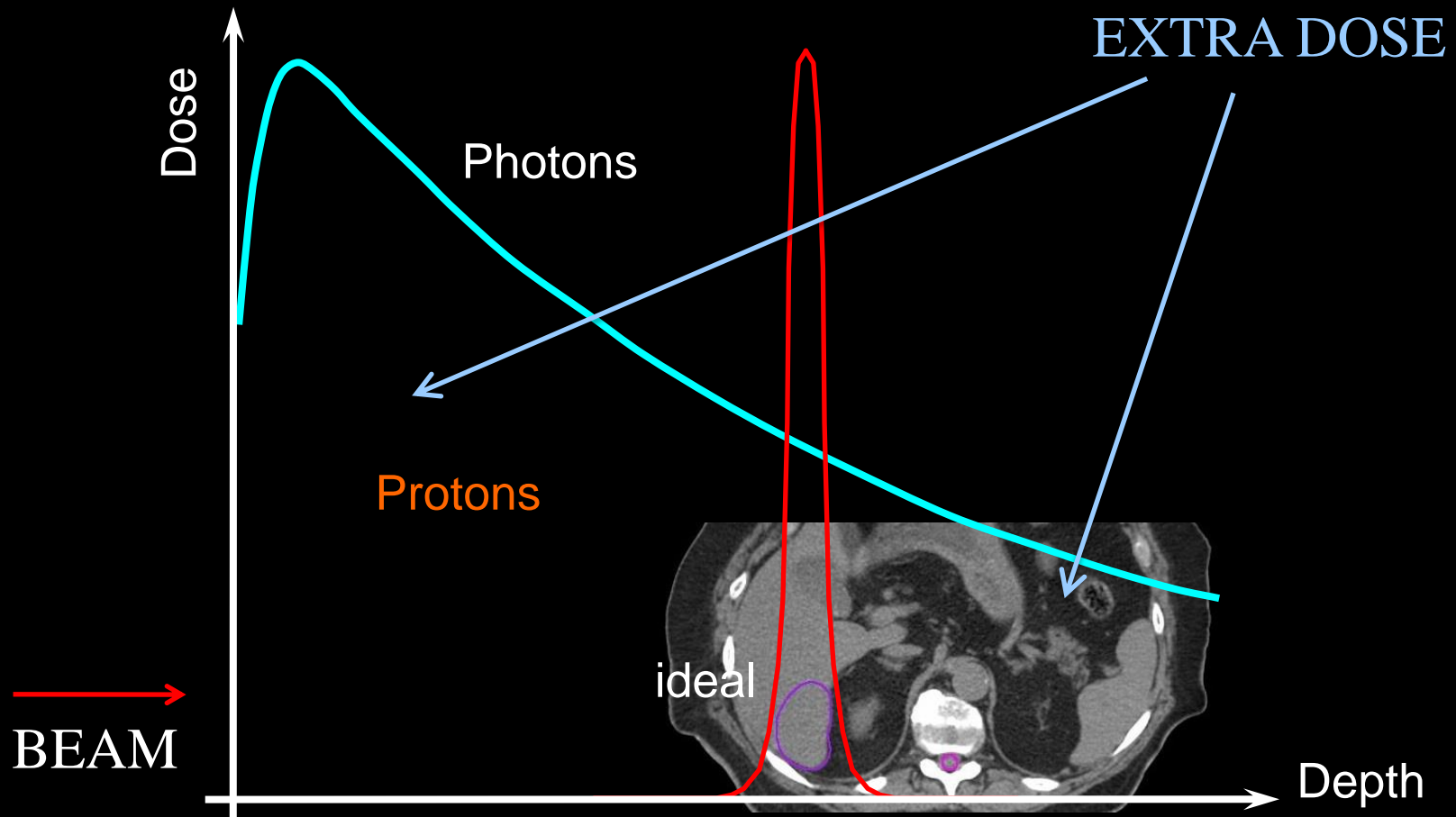
Coulomb Interaction

Range Straggling

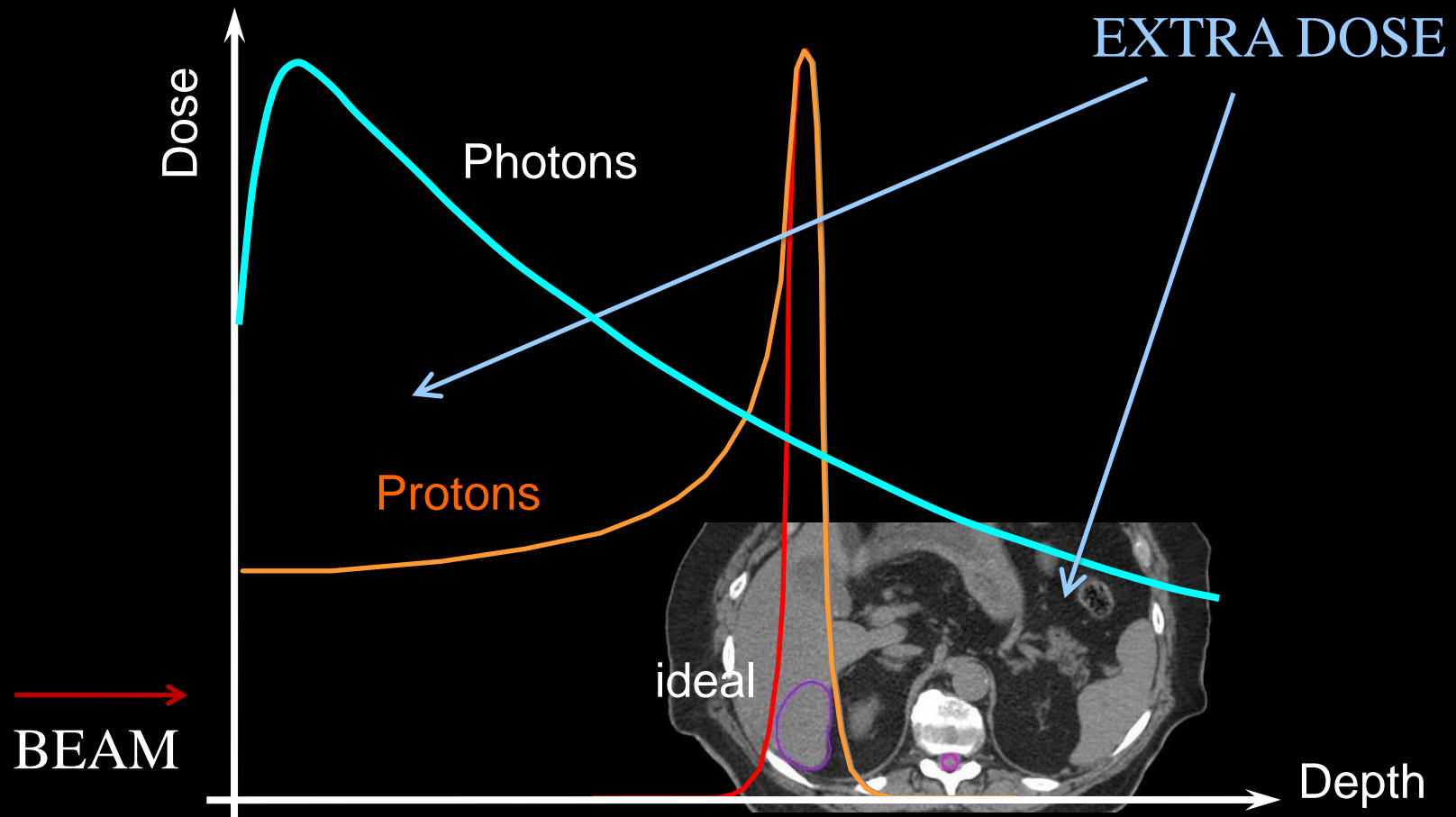
# External Beam Therapy



# External Beam Therapy

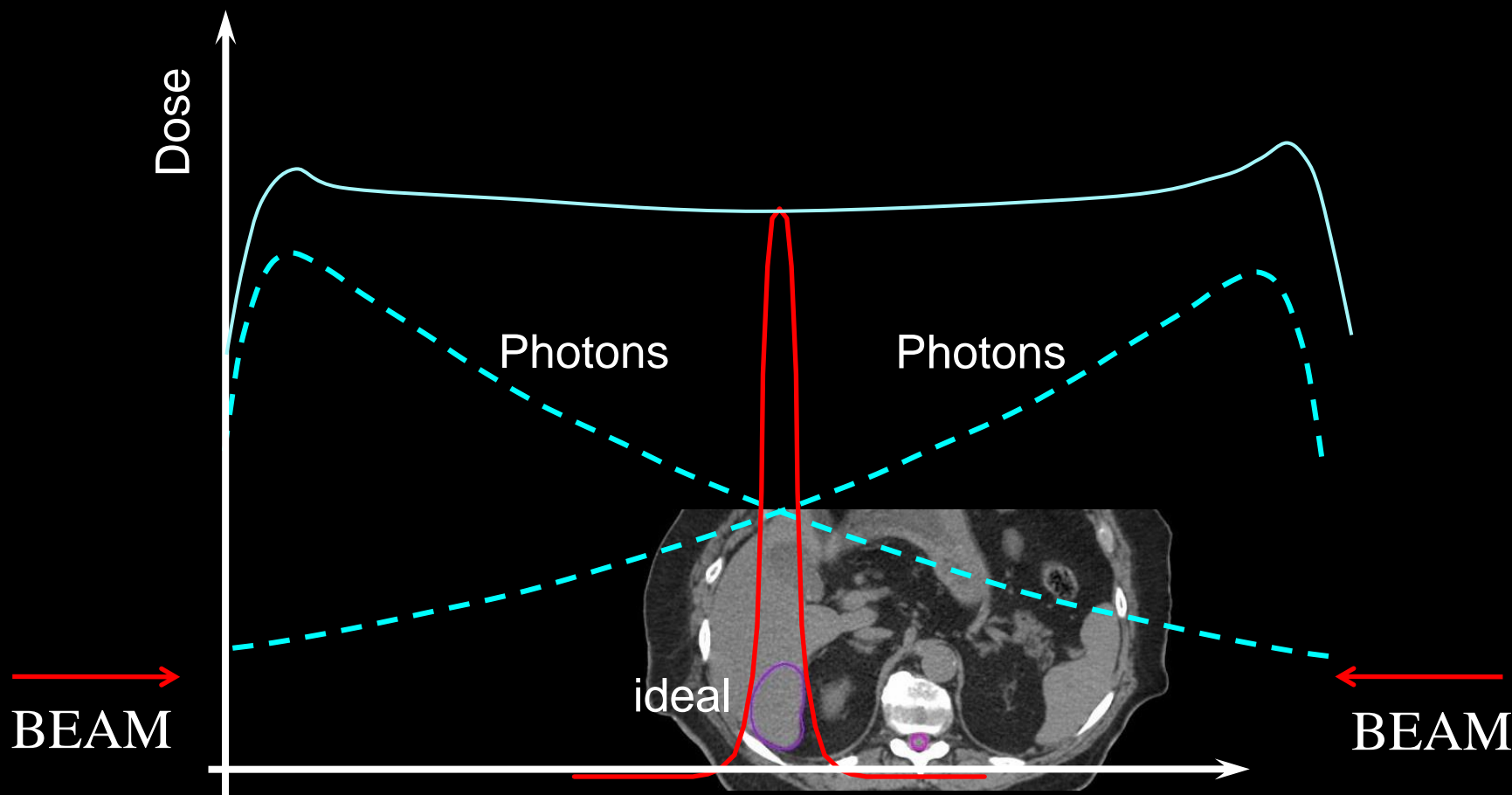


# External Beam Therapy



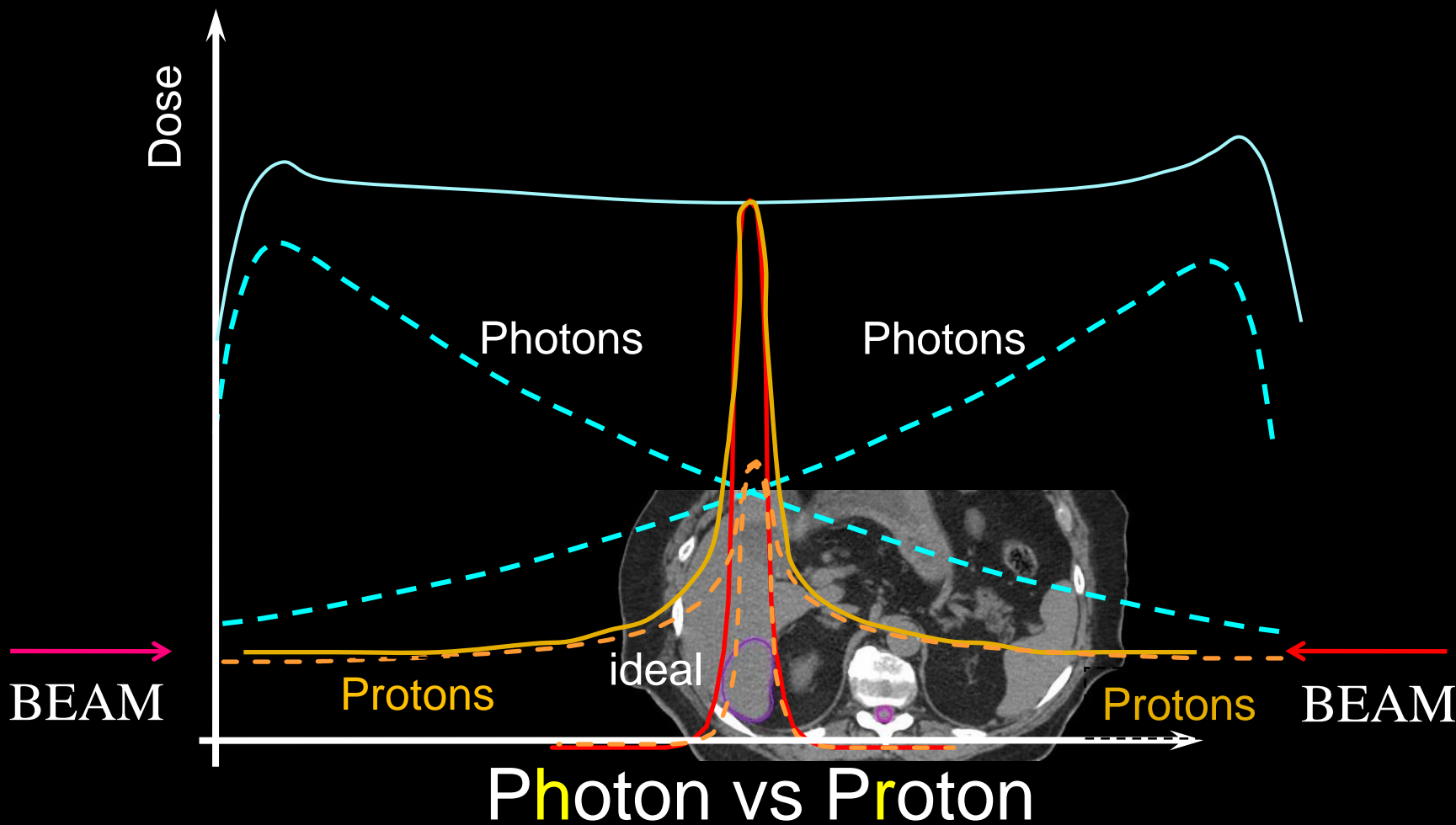


# Proton Beam Advantage



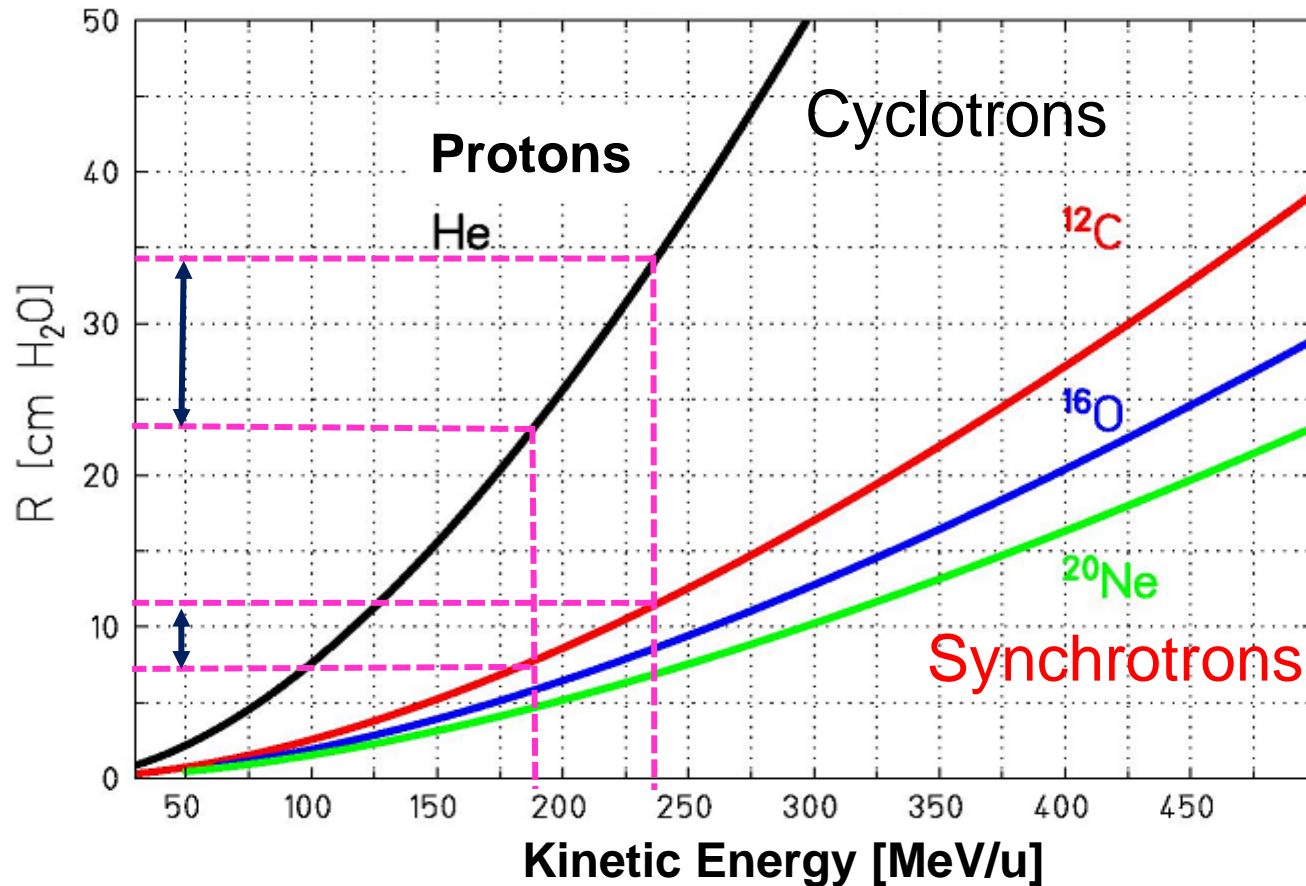
**Beam for beam you can always do a better job with particles (except at the surface).** *Flanz '09*

# Proton Beam Advantage



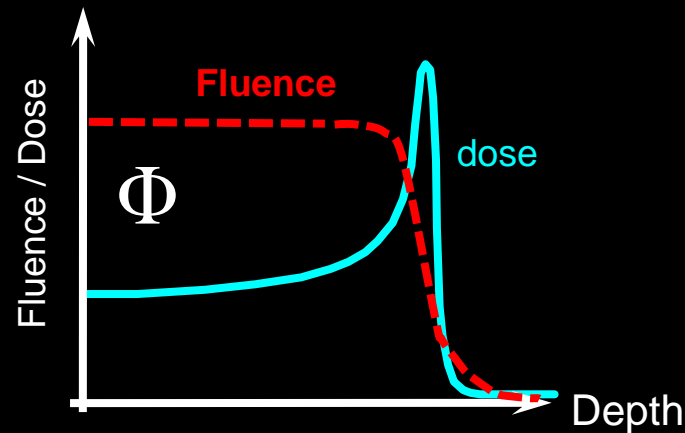
Beam for beam you can always do a better job with particles (except at the surface).

# Particle Range vs Energy



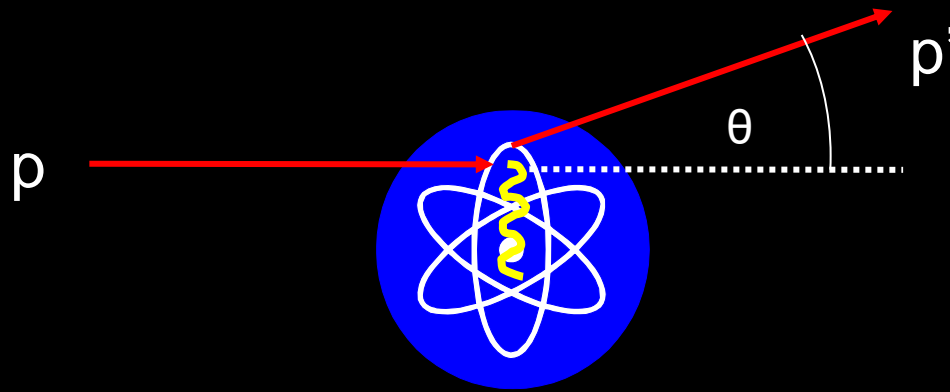
$$R(E) = \int_0^E \left( -\frac{dE}{dx} \right)^{-1} \cdot dE$$

# Dose Dependence on Depth

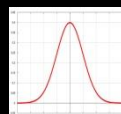
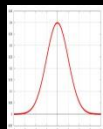


- Nearly no attenuation, fluence stays constant except near the end of the ranges of the ions
- Ions lose energy gradually
- Energy loss per ion pair stays same
- Ion pairs per unit length increases
- Increase in LET, and in the ratio between biological dose and physical dose, i.e., increase in RBE, at the end
- Electron energy low  $\rightarrow$  no build up

# Multiple Coulomb Scattering (MSC)



- Ions are deflected in the electric field of the nuclei
- In general, multiple deflections will occur for each ion
- Play key role in determining lateral dose distribution



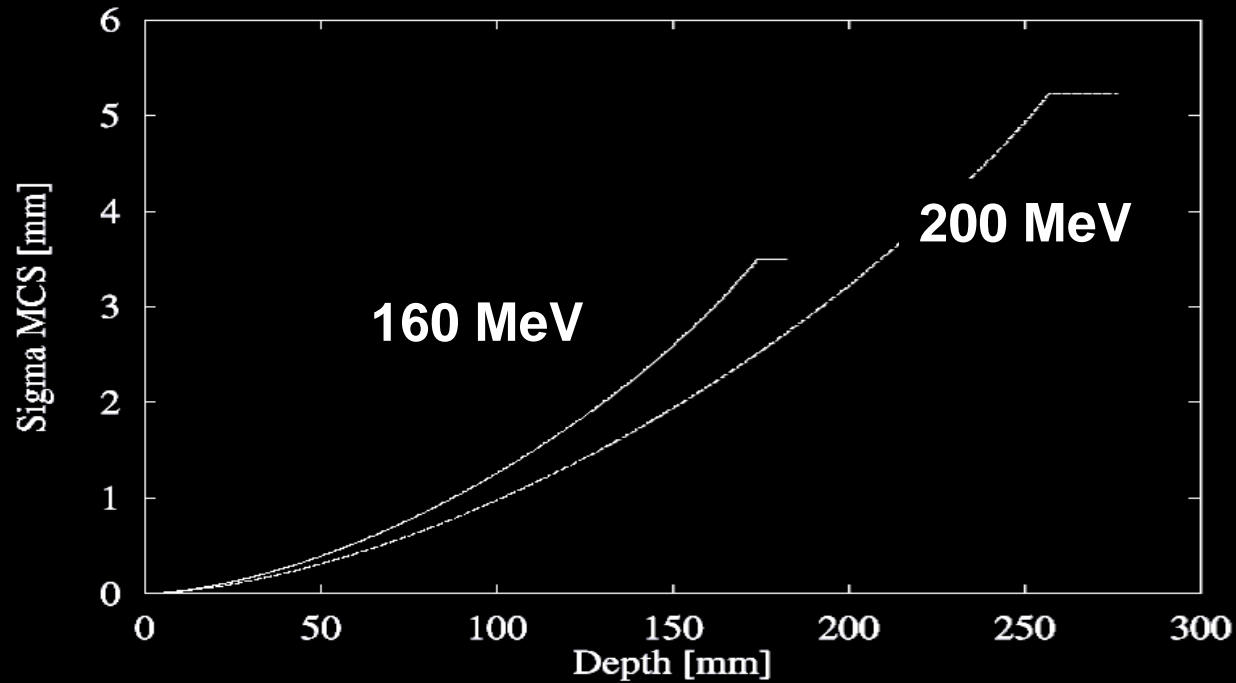
# Multiple Coulomb Scattering (MSC)

## Moliere Theory

- The definitive theory of multiple Coulomb scattering was published by Molière in 1947
  - It has *no* empirical parameters and covers *arbitrarily thick* scatterers
  - The angular distribution at large angles falls off roughly as  $1/\theta^4$  but is nearly Gaussian for small angles

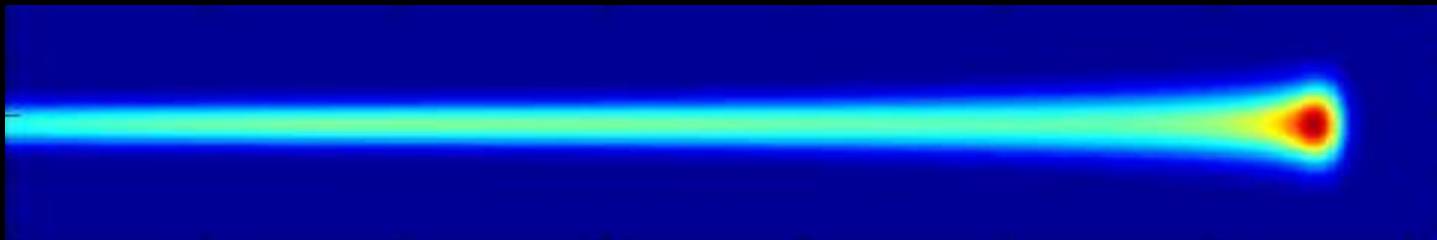
# MCS Dependence on Beam Energy

## Protons



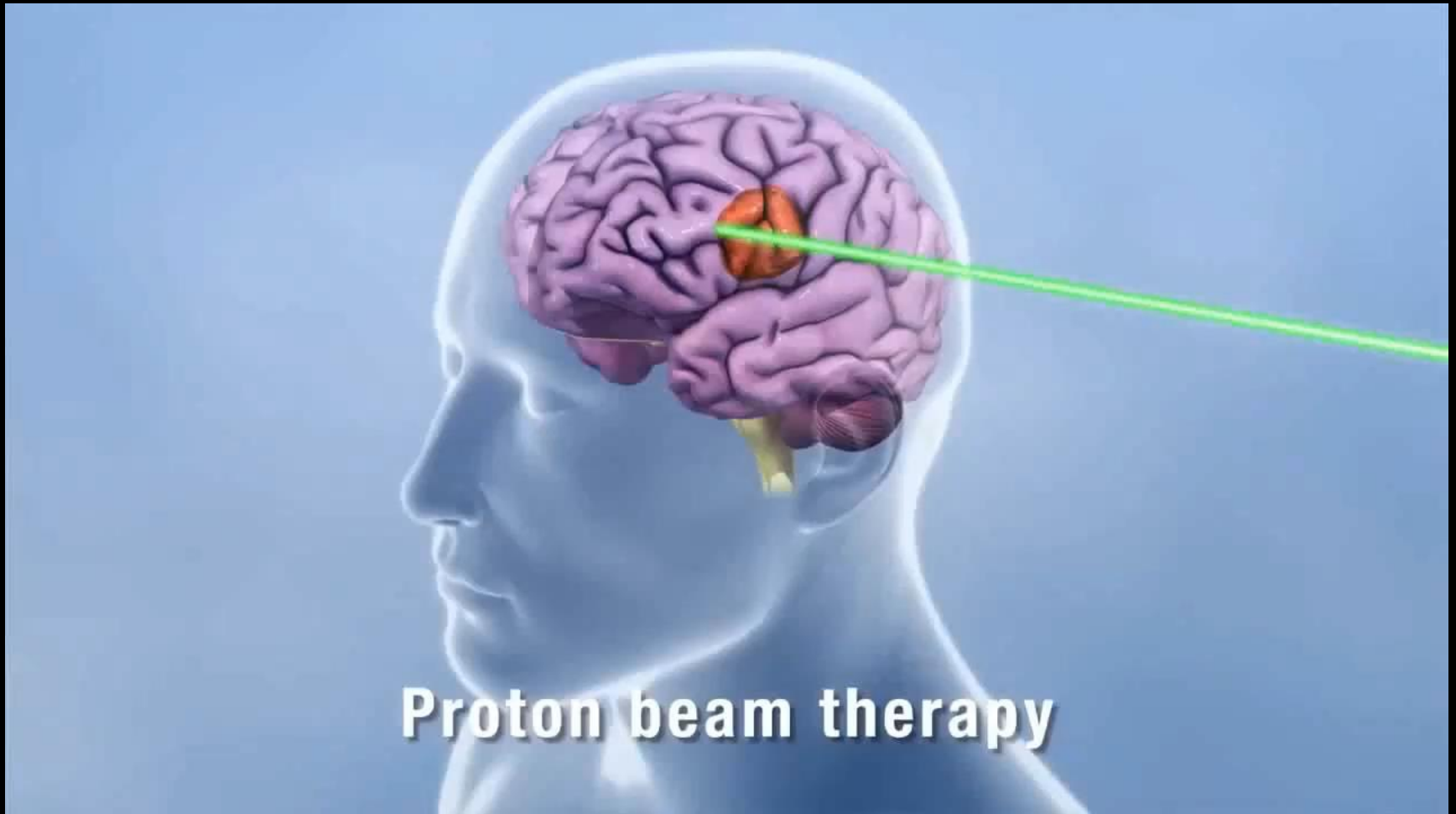
# Lateral Penumbra Changes in Depth

- Ions experiences MCS
- Each time deflected by a small angle, but the particle stays in the beam
- Effect of deflection accumulates
- Ions spread out laterally Gaussian flattens out
- Beam penumbra increases
- At the same time, ion energy decreases and deflection angle increases for each interaction
- Beam penumbra increases faster near the end of beam range



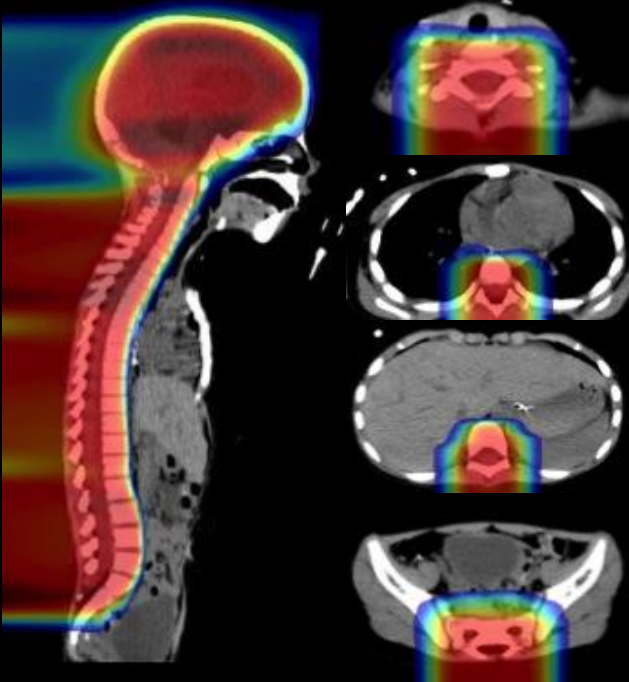


# Proton Pencil Beam Scanning

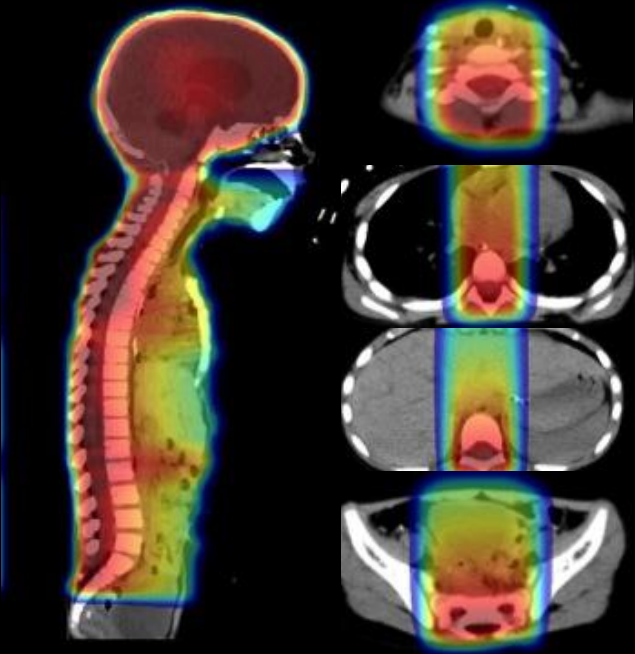


# Pediatric cancer (e.g. medulloblastoma) - representative for the superior dose distribution using particles

**Proton beam**



**X-ray beam**



**Absolute risk of secondary cancer**

Method	Risk	X-ray/Proton
Proton	0.05	1.0
X-ray (Standard)	0.75	15.0
X-ray (IMRT)	0.43	8.6

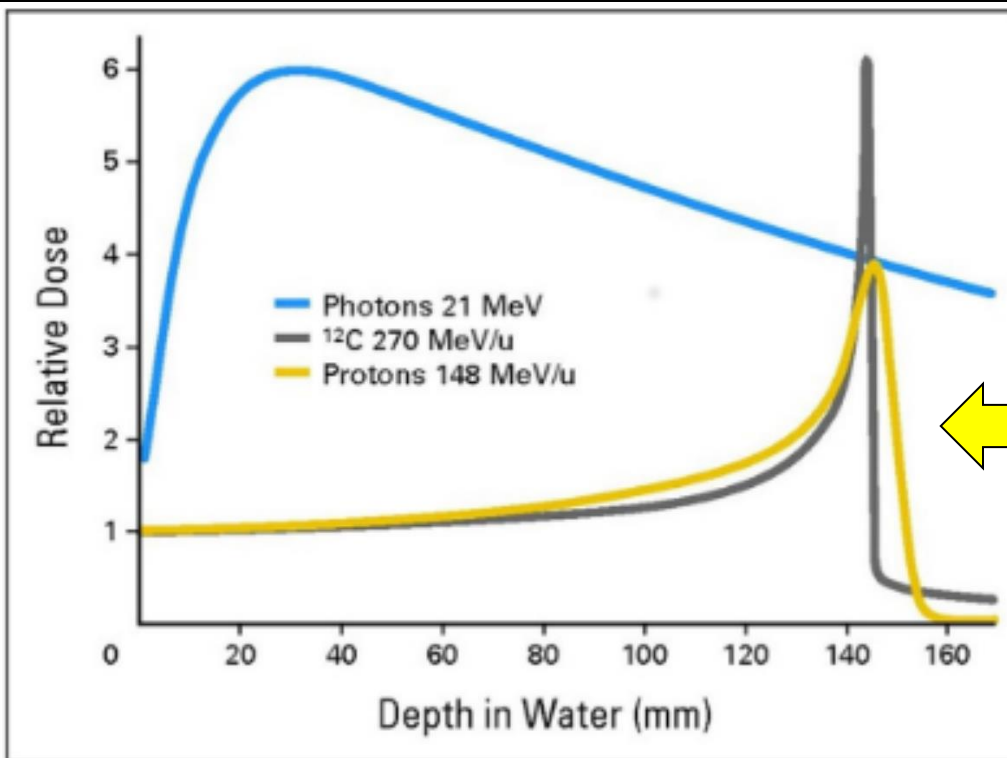
*Intention to treat spinal cord and brain only!*

Miralbell R et al., Int J Radiat Oncol Biol Phys 54: 824-829, (2002) Prof. Dr. L. Sihver

# Why Carbon Therapy?



# Physical Advantages of Carbon Beams



The variance of the range Straggling,  $\sigma_R^2$ , is related to the energy losses,  $\sigma_E^2$ .

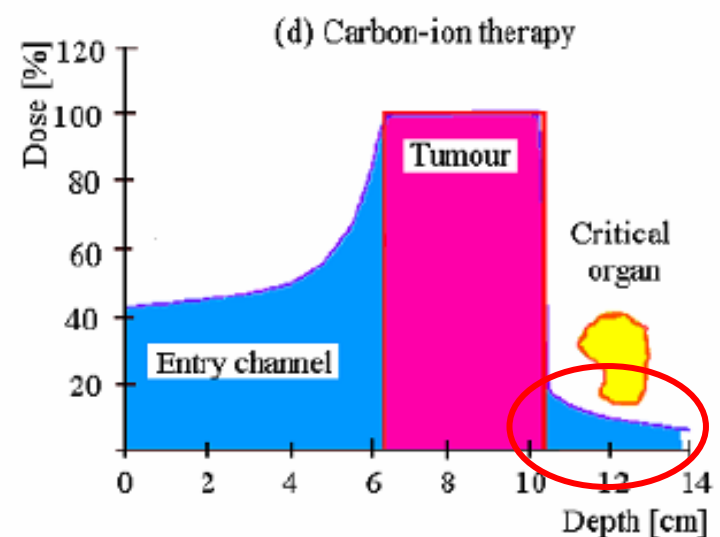
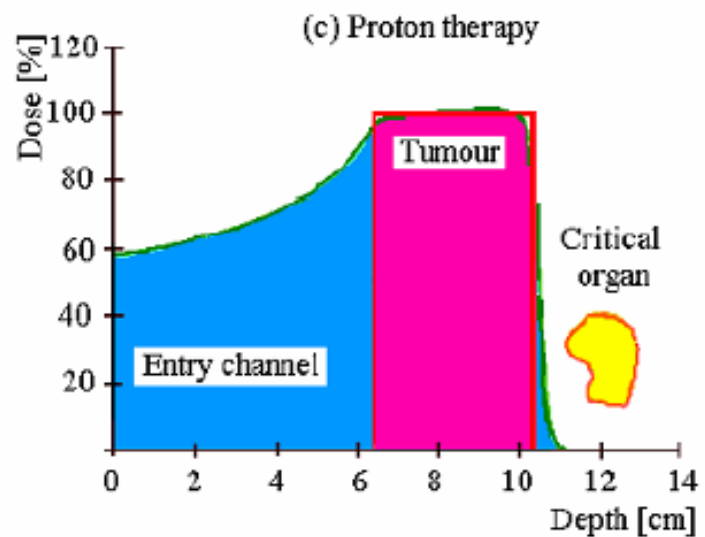
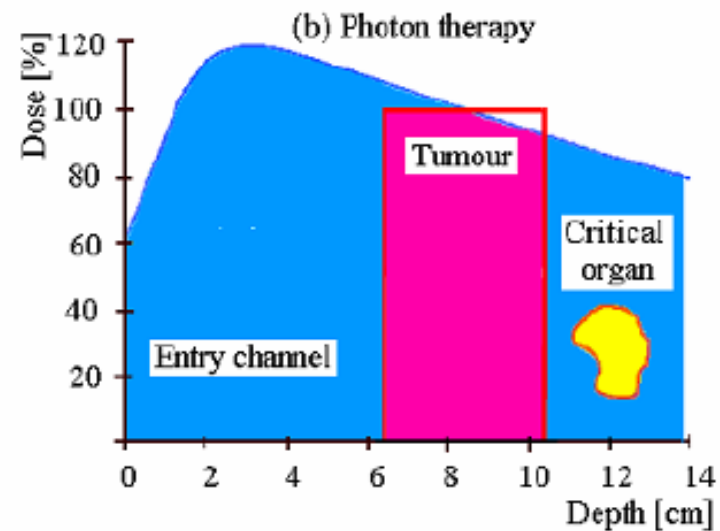
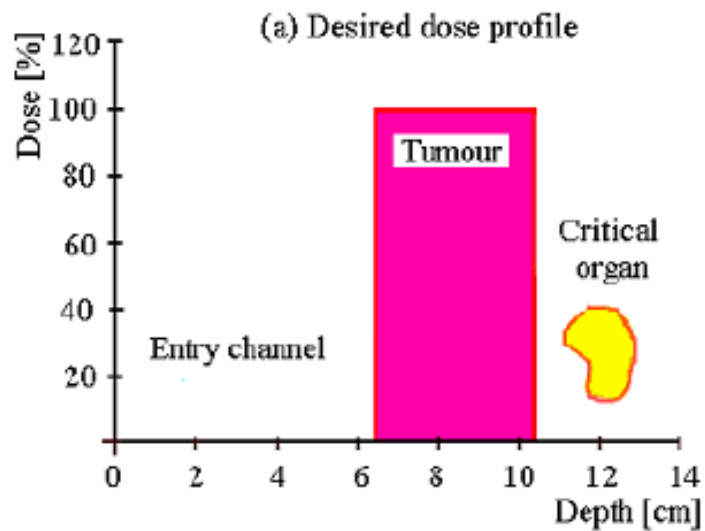
The width of the range straggling can be expressed by:

$$\sigma_R = \frac{R}{\sqrt{m}} f\left(\frac{E}{mc^2}\right),$$

where  $m$  and  $E$  are the projectile's mass and Energy. The  $1/\sqrt{m}$  dependence causes protons to have a higher straggling than light ions, by a factor of 3.5 with respect to carbon ions.

Jäkel O. Physical advantages of particles: protons and light ions. Br J Radiol 2020; 93: 20190428.

# Physical Advantages of Ion Beams



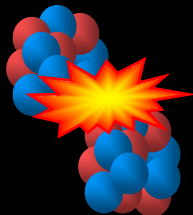
# Nuclear Reactions

Carbon ion therapy: 120 - 400 MeV/u

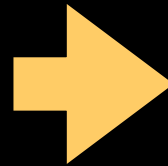
For the therapy we have to know all interaction events,  
i.e. *particles (all generations) fluences vs. energies, etc.*

# Nuclear Reactions

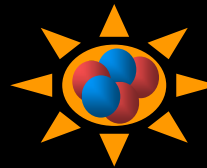
projectile



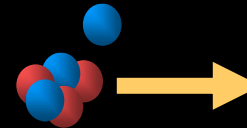
target



target fragments



projectile fragments



**New mixed radiation field !**

Interaction of the radiation with the tissue and organs in the body...

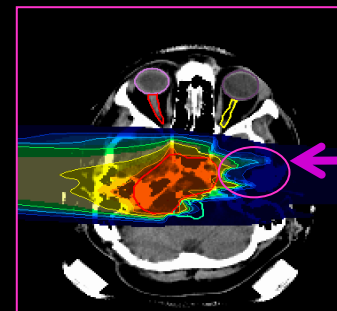
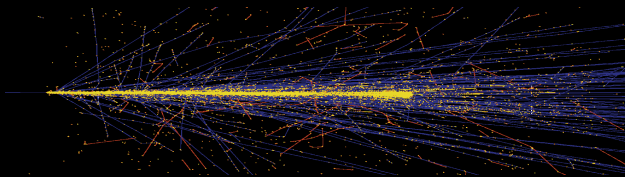
*Target Fragments*

*Projectile fragments*

... lower charges than target  
... high LETs  
... short ranges

... lower charges than primaries  
... mixed LETs  
... long ranges

High-energy carbon beam stopping in water



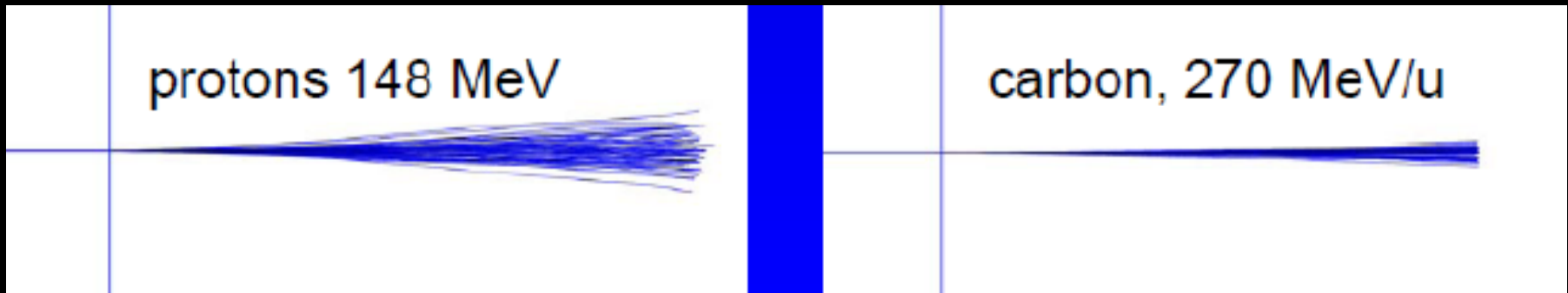
Caused by projectile fragments

**Mixed radiation field with  
charged particles, neutrons and  
gamma rays is a challenge for  
the dosimetry!**

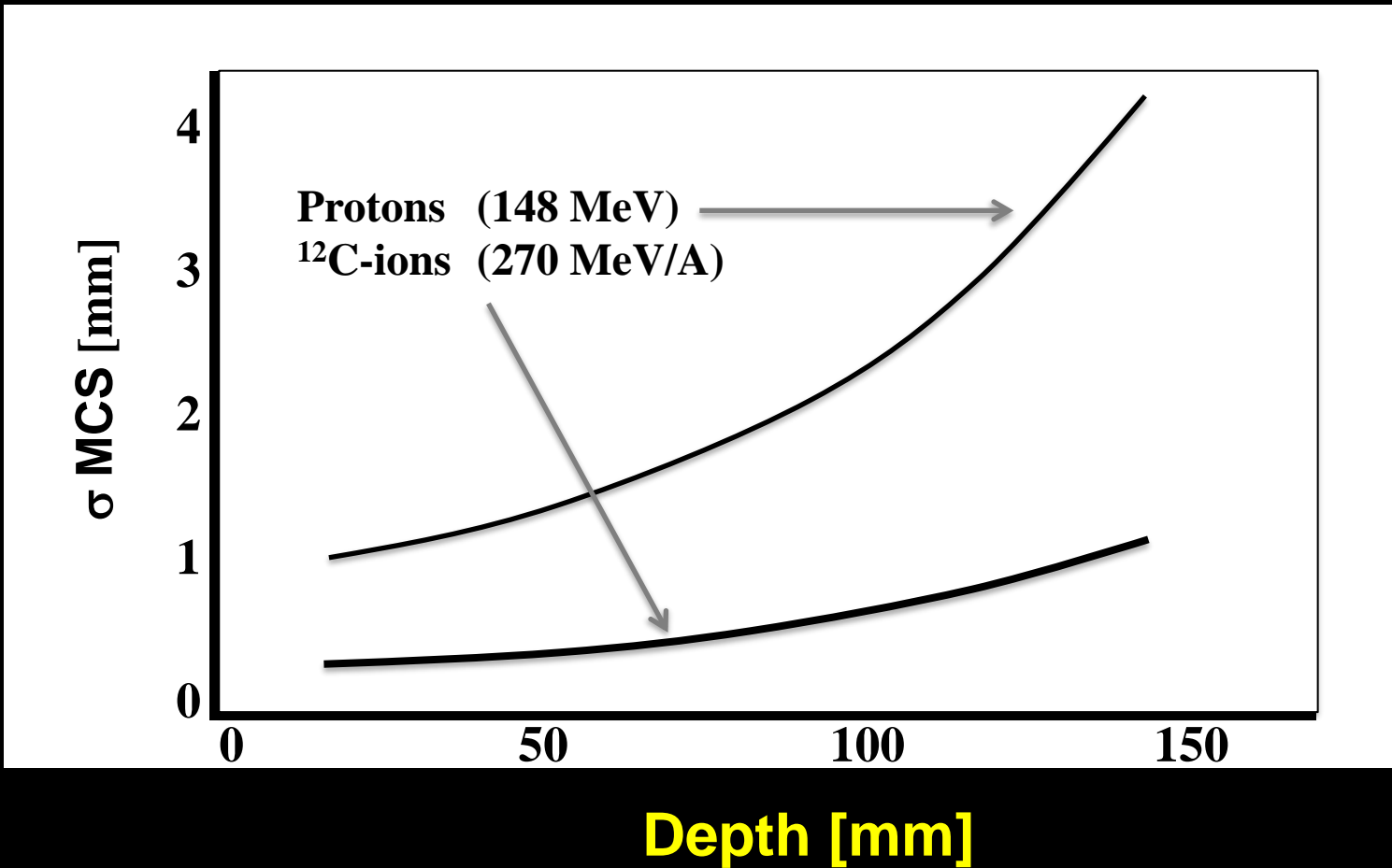


# Multiple Coulomb Scattering (MSC)

Heavy ions exhibit more precise physical dose distributions than protons because angular and range scattering are inversely proportional to the square of the atomic number.



# MCS Dependence on Particle Charge



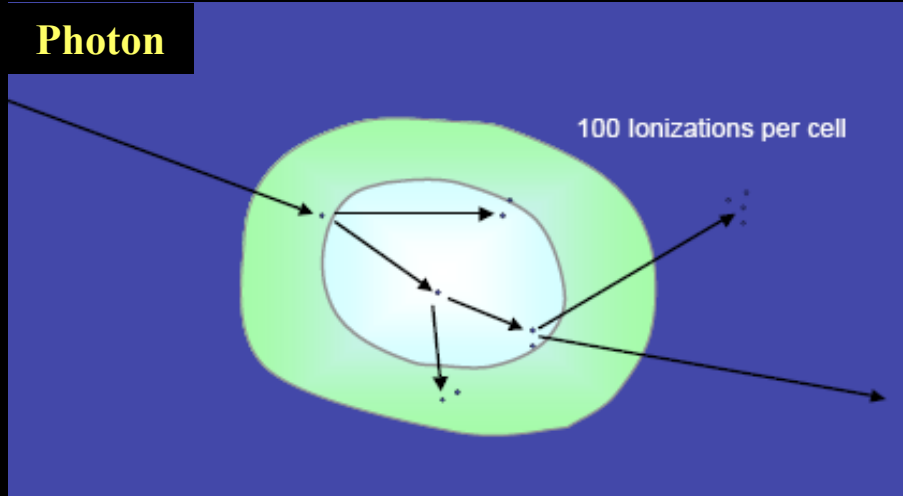
**Biological effects are controlled  
by the differences in physics  
and chemistry of different  
ionizing radiation!**

# Distribution of Ionizing Events

## Low LET radiation

Photon

100 Ionizations per cell

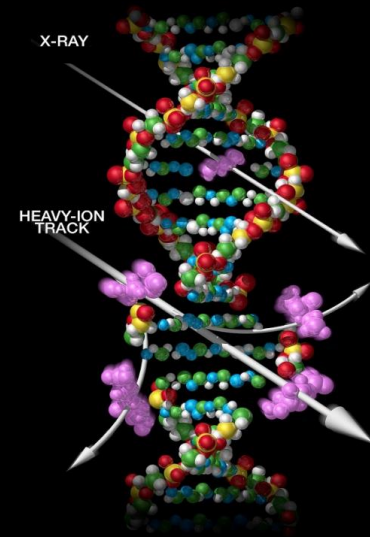
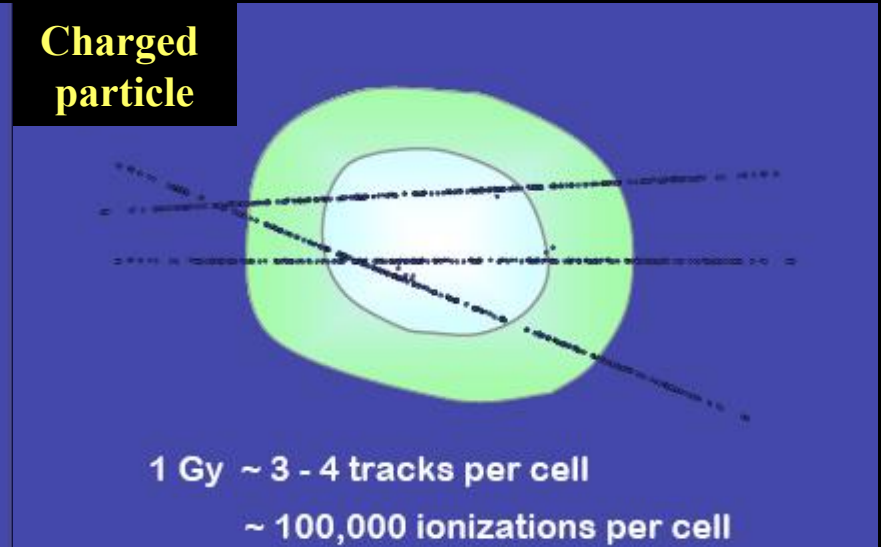


## High LET radiation

Charged particle

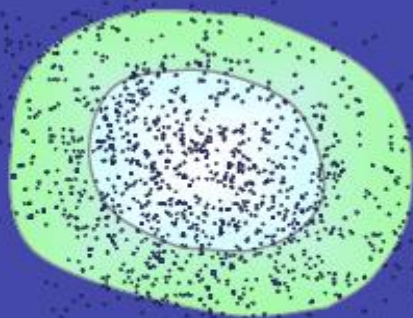
1 Gy ~ 3 - 4 tracks per cell

~ 100,000 ionizations per cell

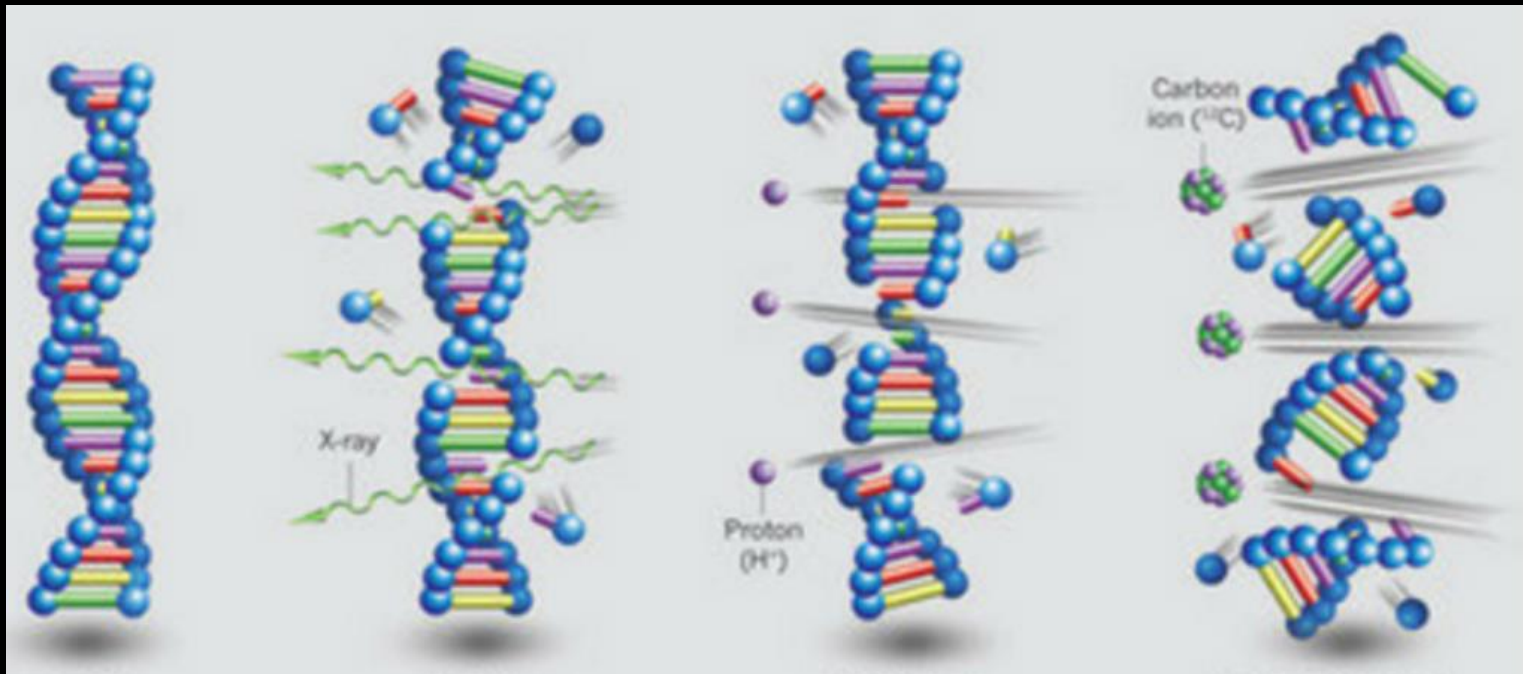


1 Gy ~ 1000 tracks per cell

~ 100,000 ionizations per cell



# Radiation Types and their Damage to DNA



**DNA**

**X-ray**

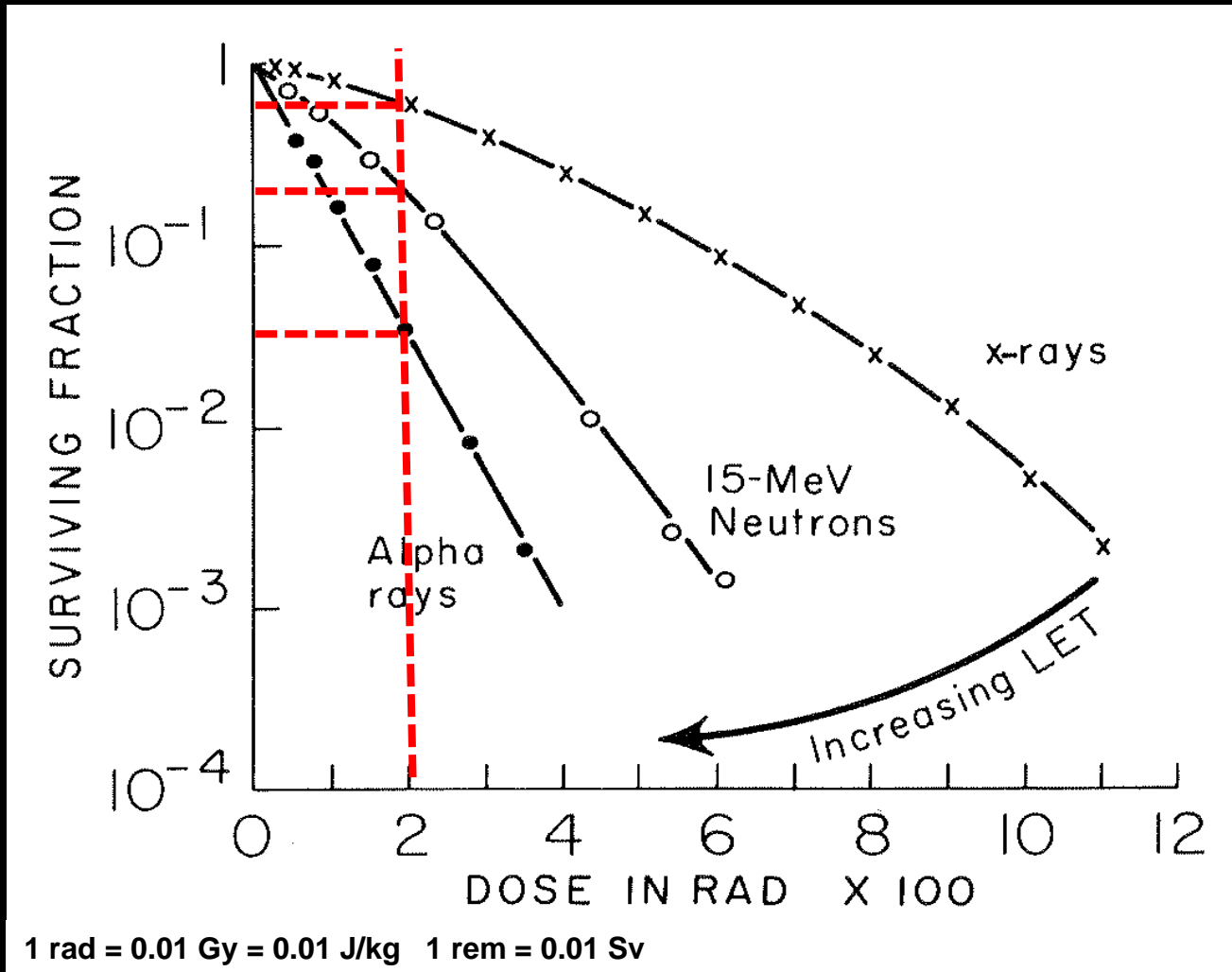
**Proton  
Beam**

**Carbon Ion  
Beam**

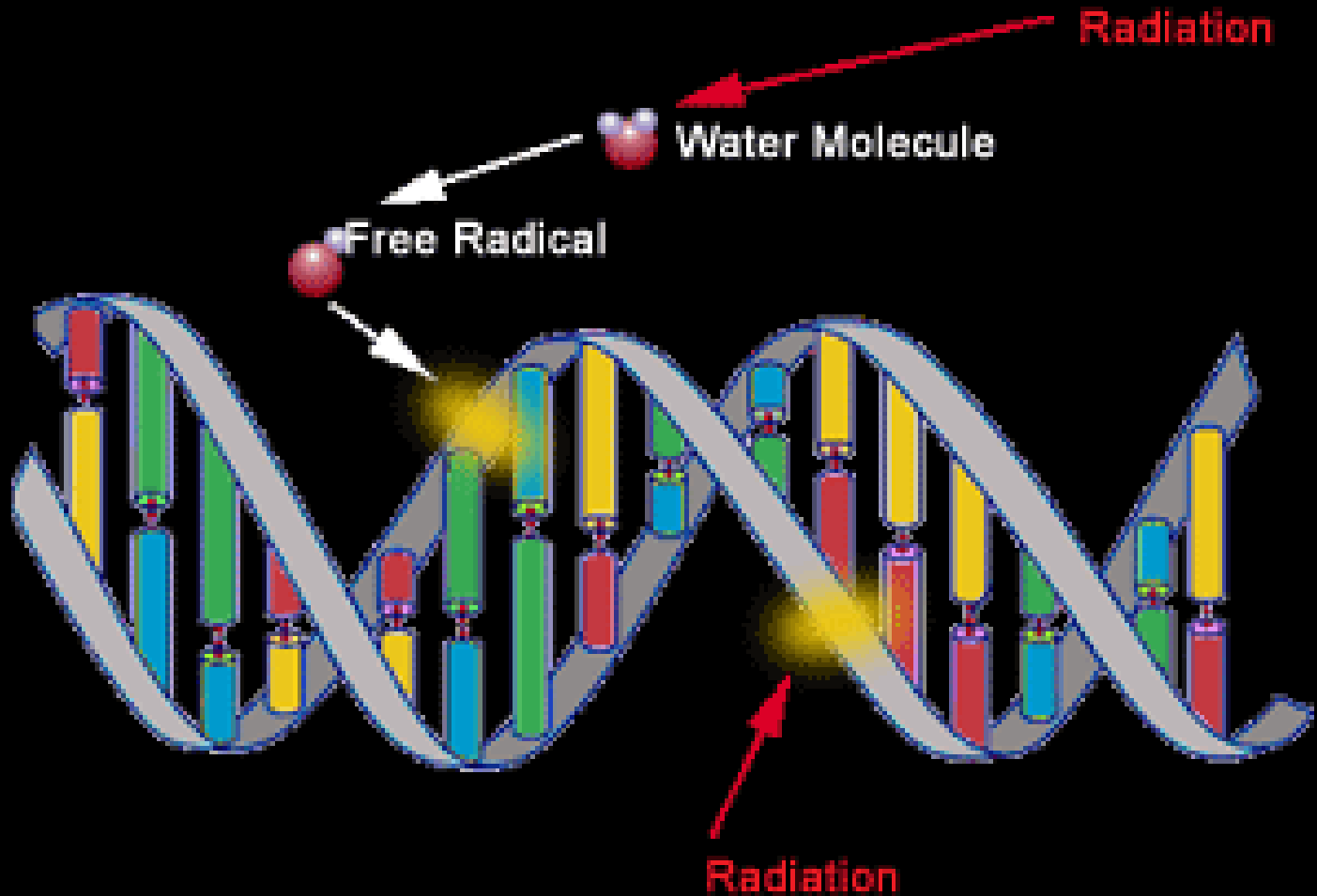
**Low-LET radiation:**  
*Repairable* single/double  
strand breaks

**High-LET radiation:**  
*Complex* DNA lesions  
Multiple DNA pathways  
More difficult to repair  
Enhances cell death

# Cell Survival and LET

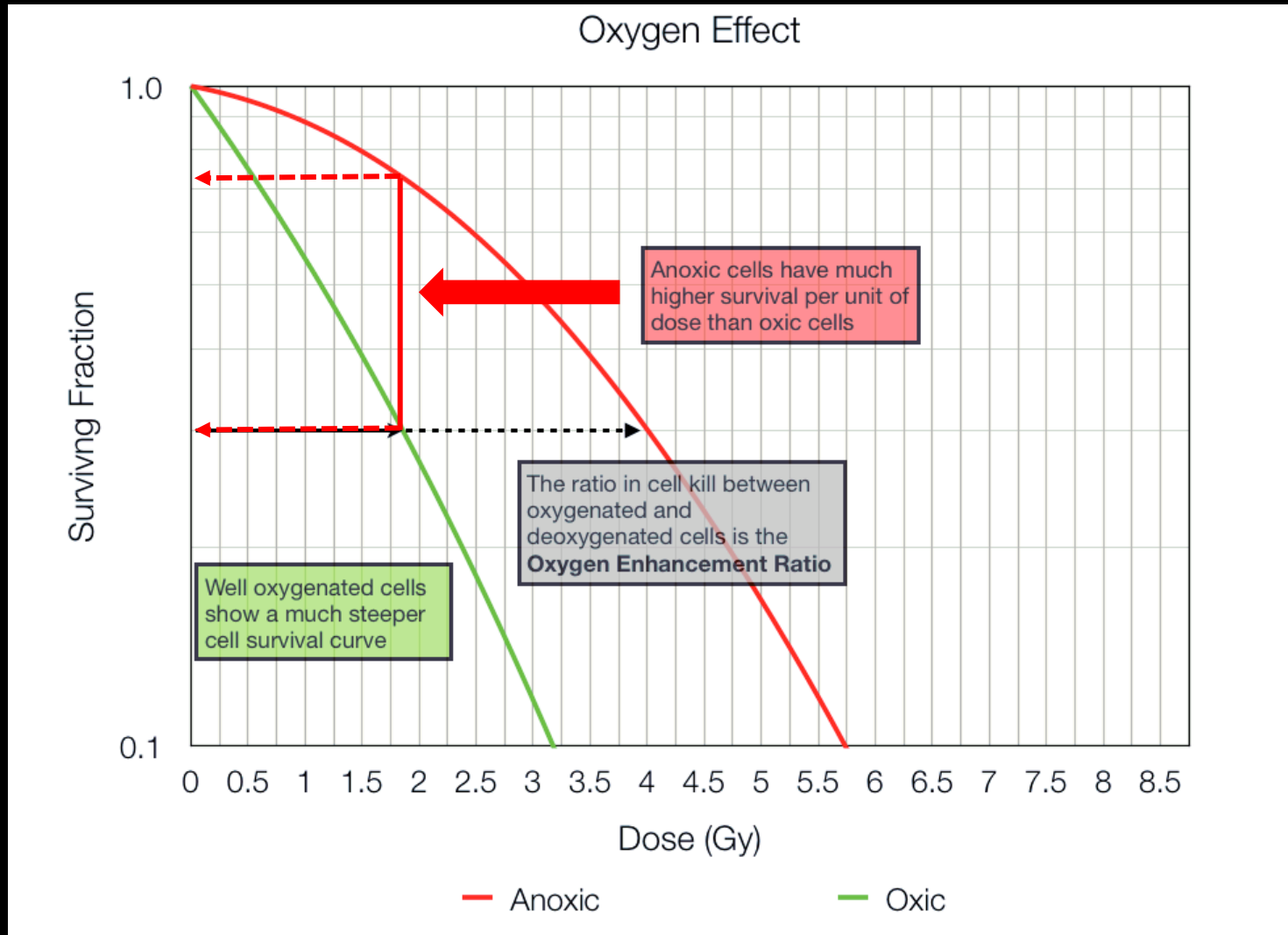


# Indirect effects



# Direct effects

# Oxygen Enhancement Ratio (OER) for low LET radiation ( $\gamma$ and $e^-$ )

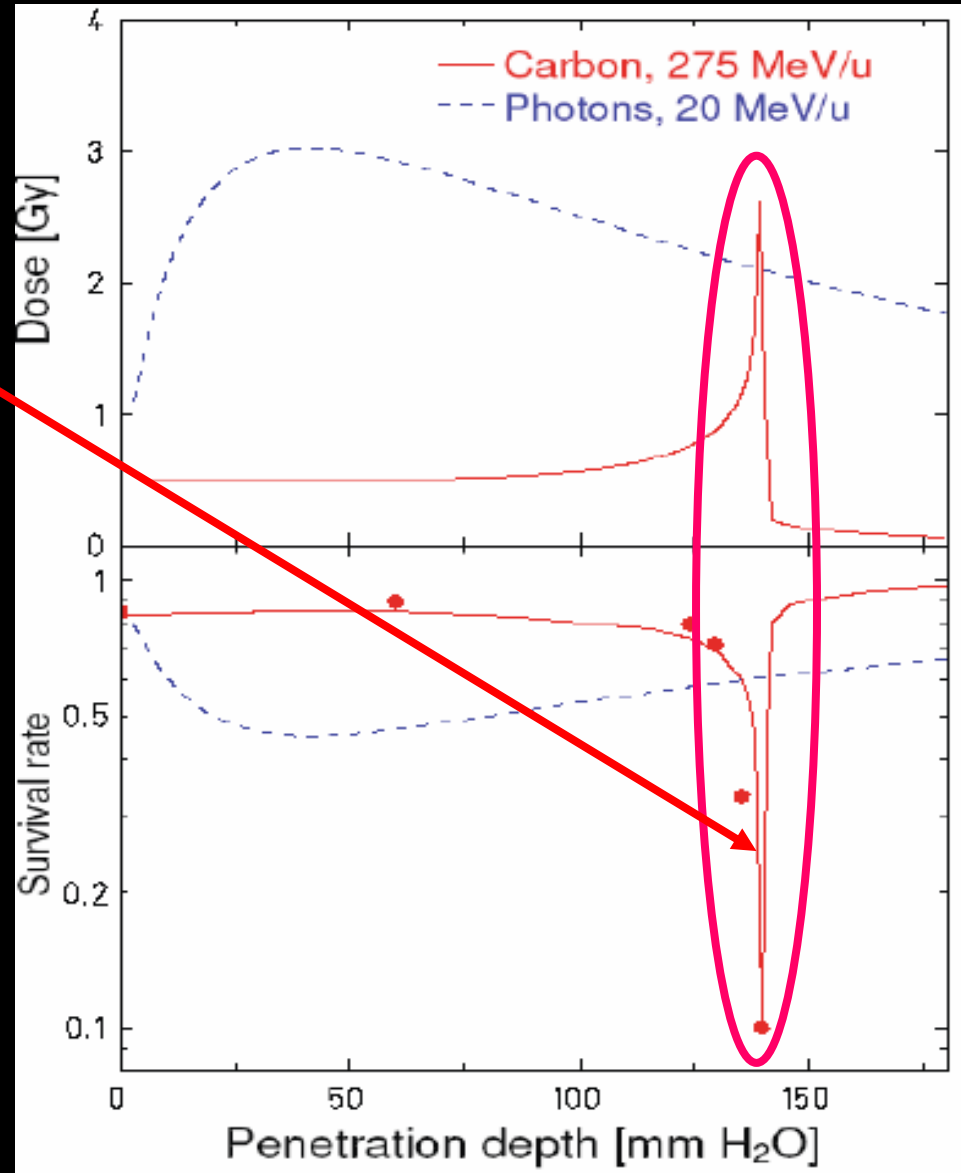


$$\text{OER} = \frac{\text{Dose to produce a certain effect under hypoxic condition}}{\text{Dose to produce the same effect under oxic condition}}$$

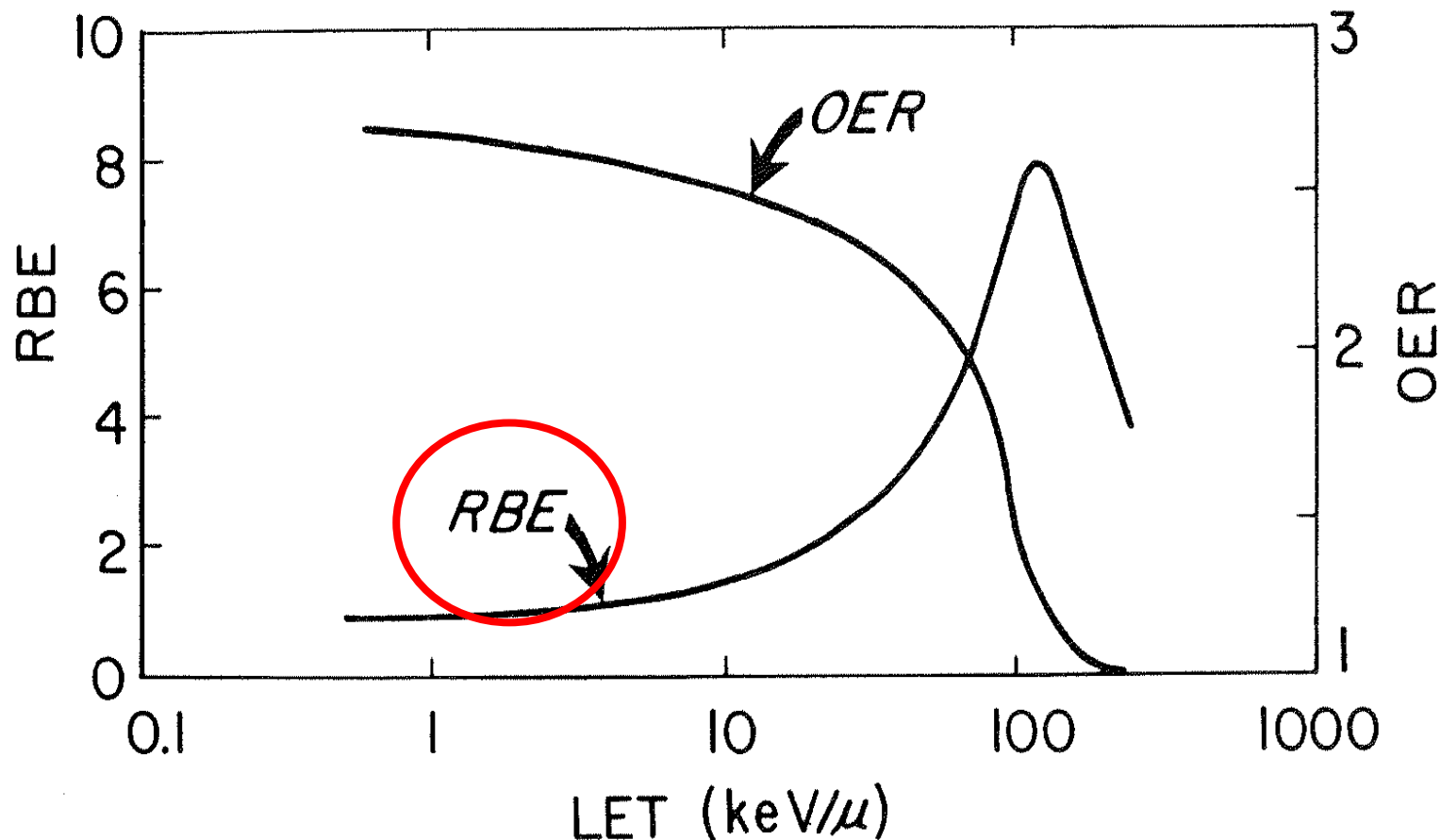


# Cell Survival

Low cell survival rate  
at Bragg Peak

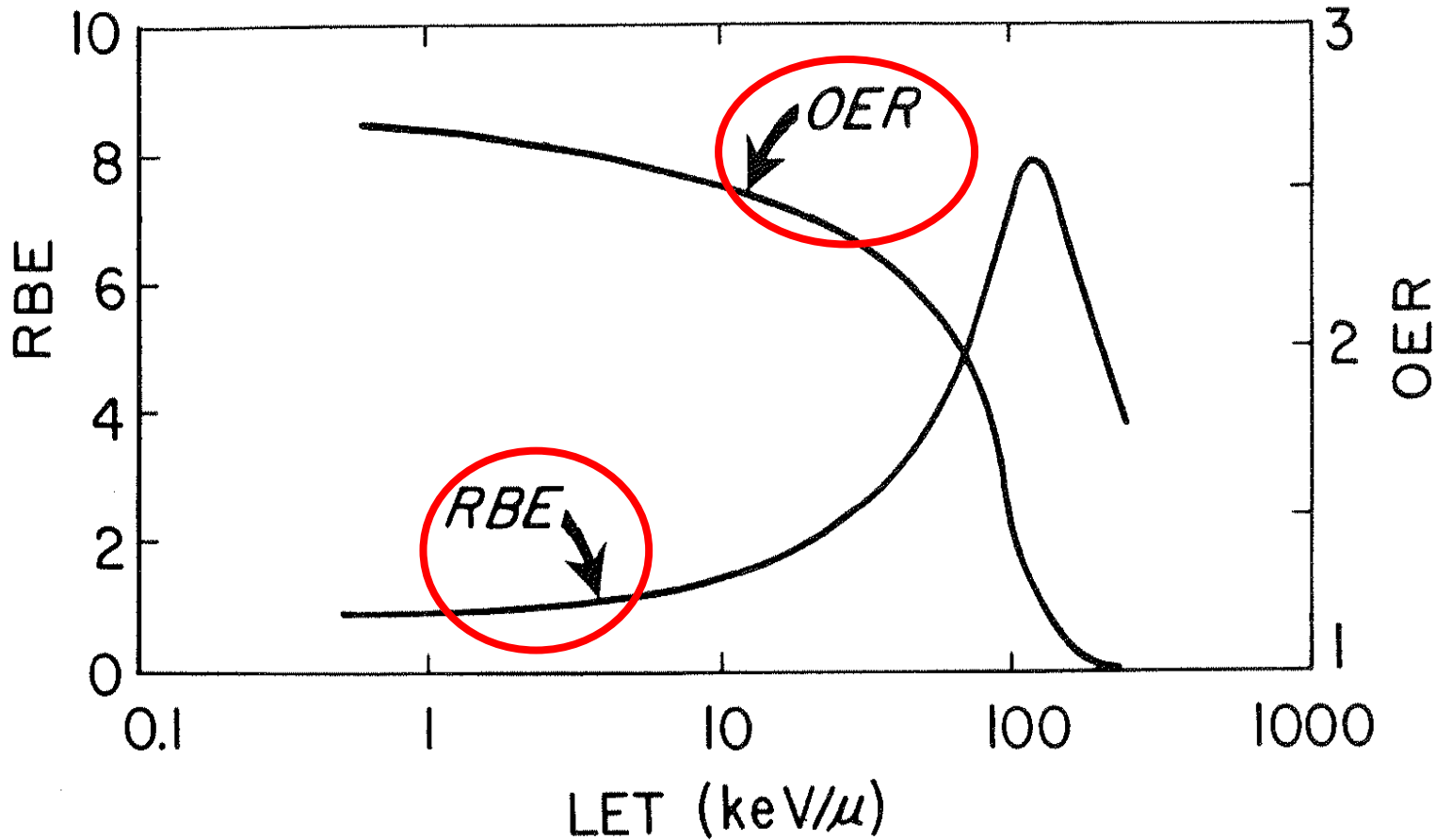


## Relative Biological Effectiveness (RBE) vs LET



$$RBE = \left[ \frac{D(X - \text{rays} / 250kV)}{D_T(\text{test} - \text{radiation})} \right]_{\text{iso effect}}$$

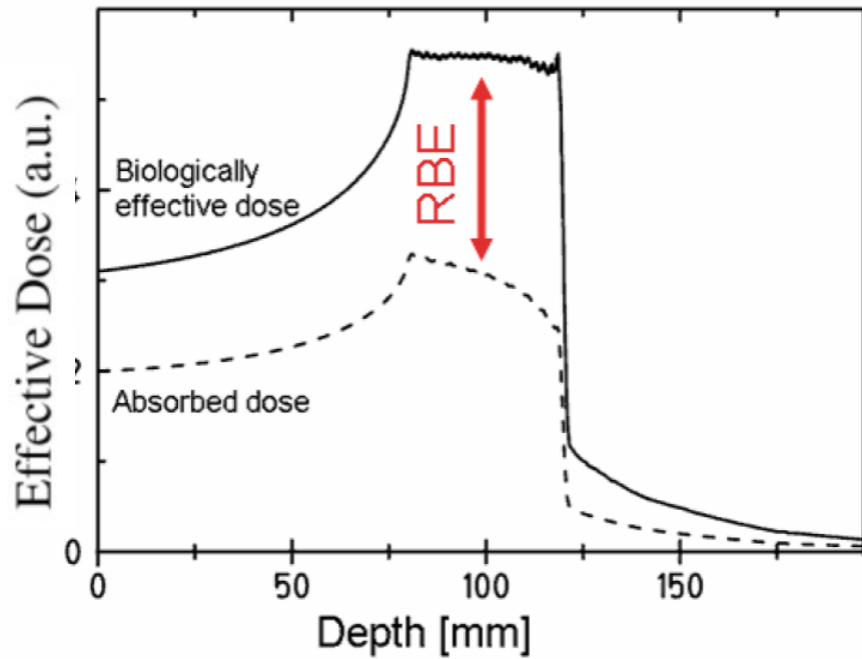
## Oxygen Enhancement Ratio (OER) vs LET



$$RBE = \left[ \frac{D(X - \text{rays} / 250kV)}{D_T(\text{test} - \text{radiation})} \right]_{\text{iso effect}}$$

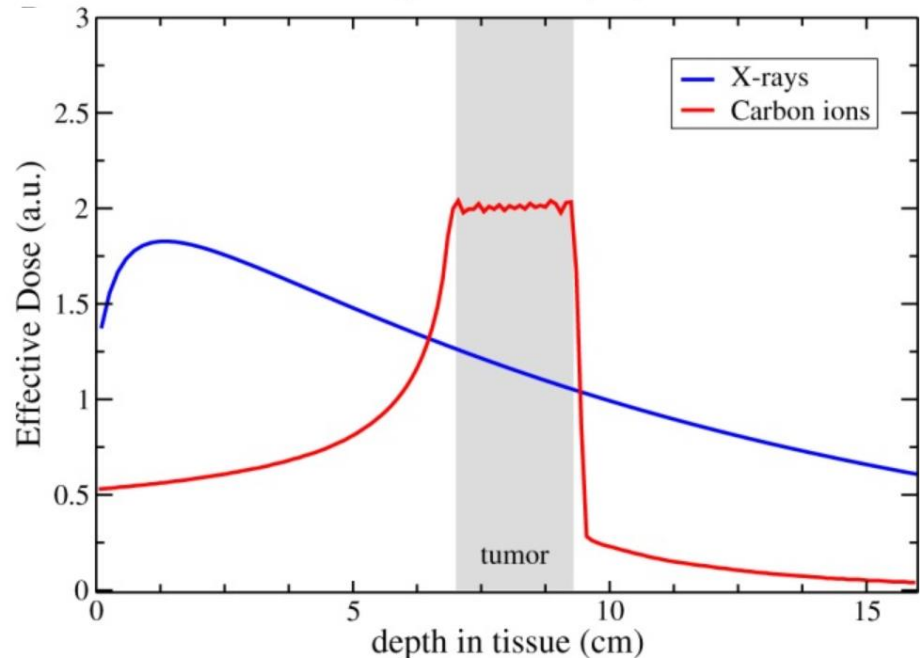
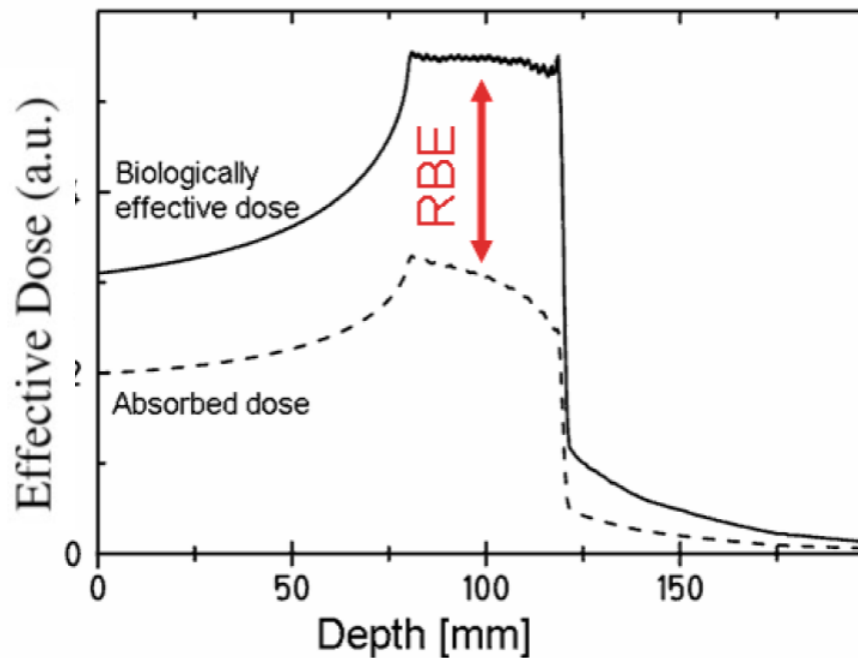
# Biologically Effective Dose

$$\text{Biologically Effective Dose} = \text{Physical Dose} \times \text{RBE}$$



# Biologically Effective Dose

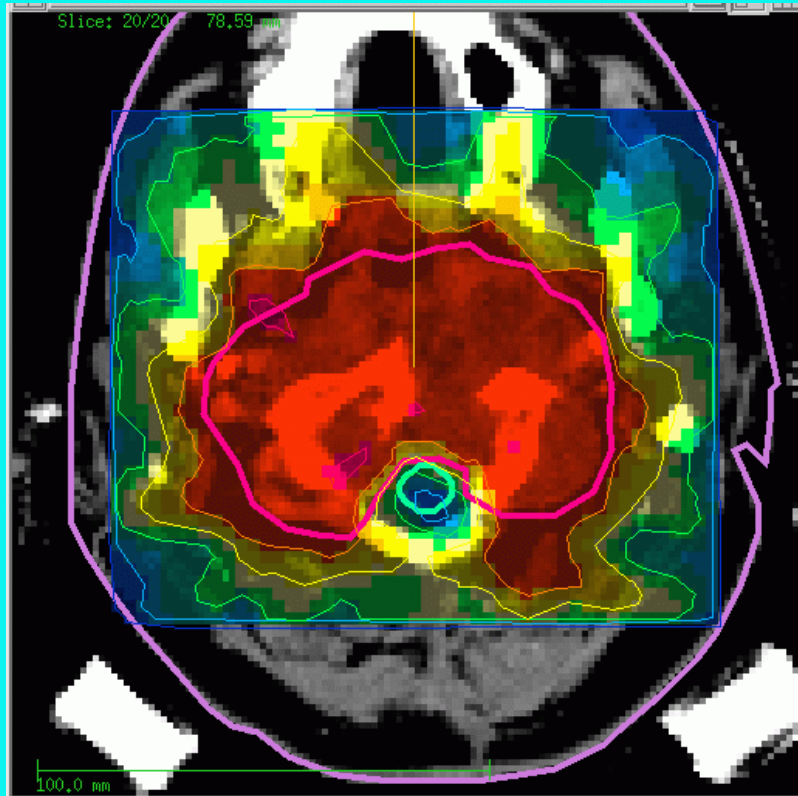
$$\text{Biologically Effective Dose} = \text{Physical Dose} \times \text{RBE}$$



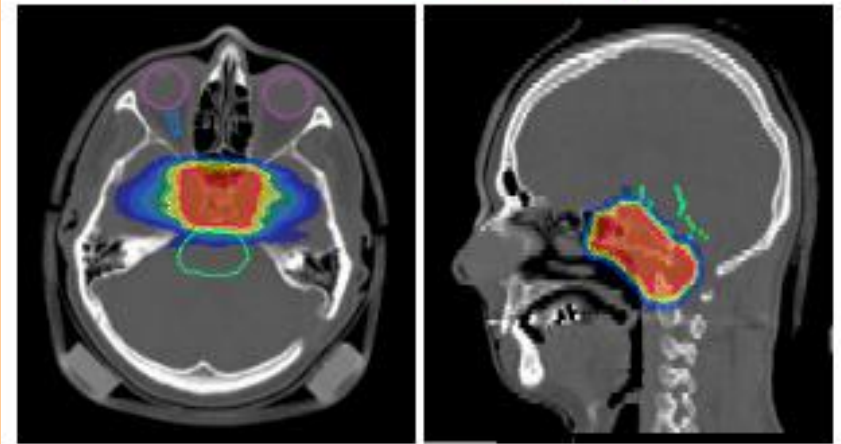
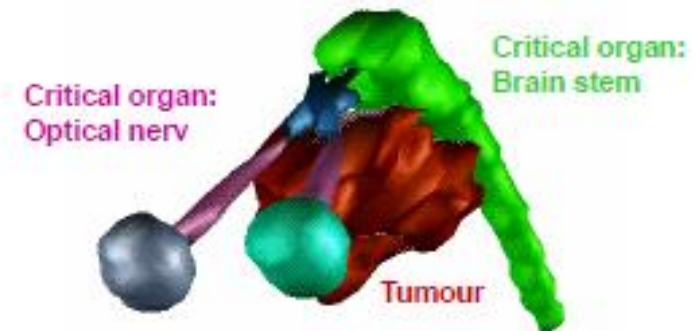
# The superior dose-conformality of ion beams

## Biological Dose

### Photon IMRT



### Carbon ions: 2 Fields



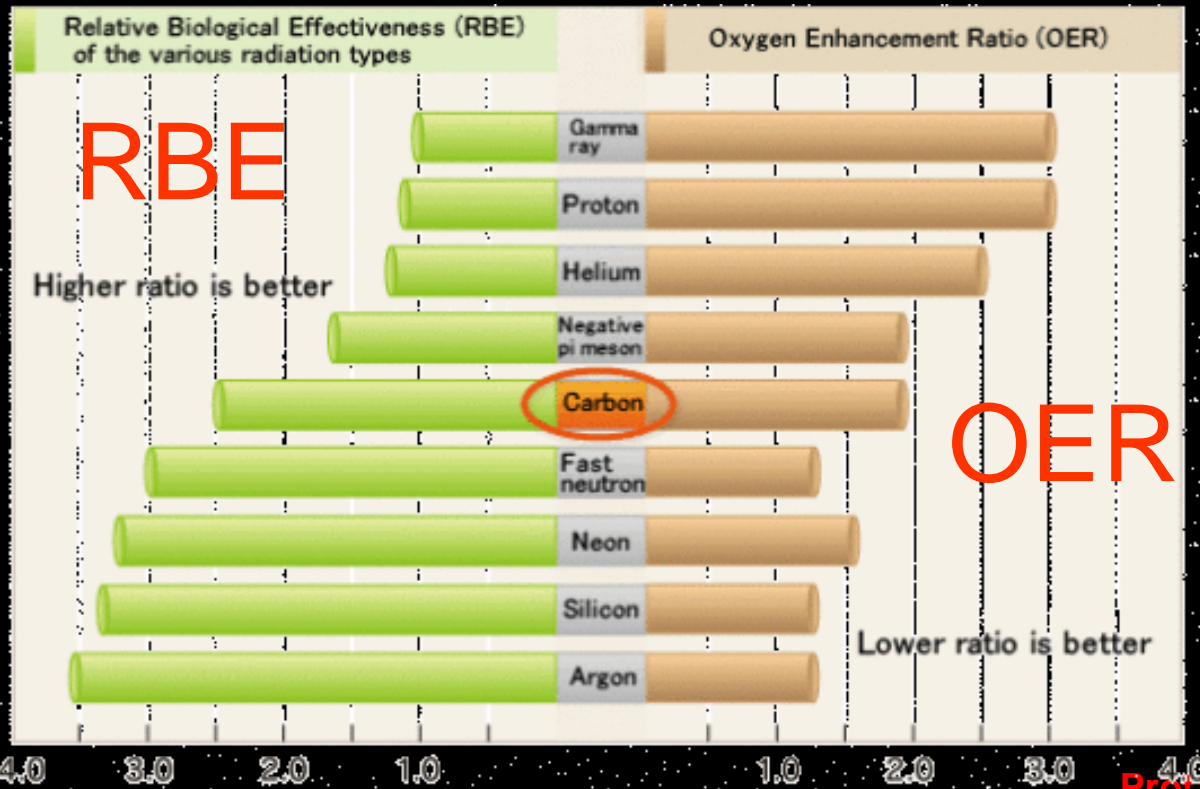
T. Bortfeld, S. Nill, U. Oelfke, O. Jäkel, DKFZ Heidelberg

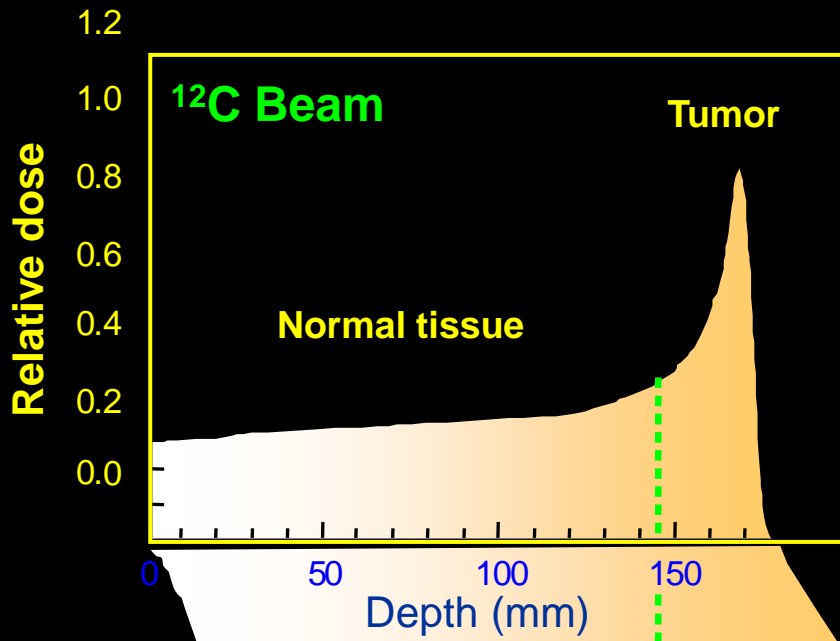
Copyright

Prof. Dr. L. Sihver

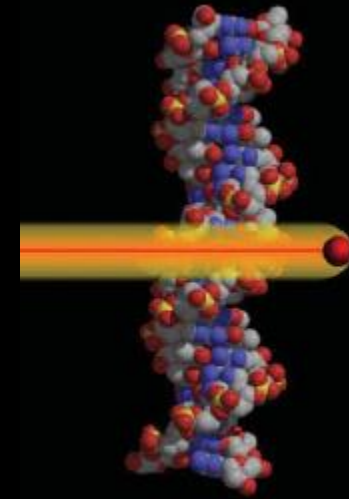
# Why use carbon beams for radiotherapy ?

- High Relative Biological Effectiveness (RBE)
- Low Oxygen Enhancement Ratio (OER)
  - Effective against hypoxic (“lack of oxygen”) tumor cells
- Carbon ions have max in RBE close to the Bragg Peak
- Limited amount of projectile and target fragmentation, etc...





Durante & Loeffler,  
*Nat Rev Clin Oncol* 2010



**Potential advantages**

- High tumor dose, normal tissue sparing
- Effective for radioresistant tumors
- Effective against hypoxic tumor cells
- Fractionation spares normal tissue more than tumor
- Increased lethality in the target because cells in radioresistant (S) phase are sensitized

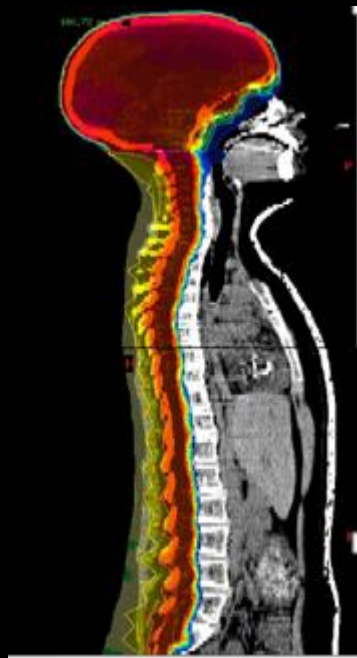
- Energy**
- LET**
- Dose**
- RBE**
- OER**
- Fractionation dependence**
- Cell-cycle dependence**
- Angiogenesis**  
(development of new blood vessels.)
- Cell migration**

Parameter	Normal tissue (0-150 mm)	Tumor (150-170 mm)
Energy	high	low
LET	low	high
Dose	low	high
RBE	≈ 1	> 1
OER	≈ 3	< 3
Fractionation dependence	high	low
Cell-cycle dependence	high	low
Angiogenesis	↑	↓
Cell migration	↑	↓

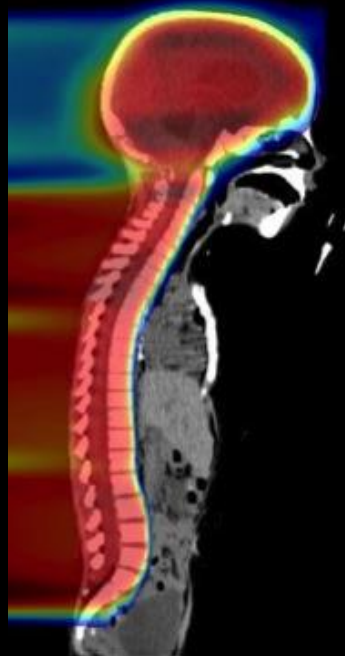


**Pediatric cancer (e.g. medulloblastoma) - representative  
for the superior dose distribution using particles**

Carbon beam



Proton beam



X-ray beam



***Intention to treat spinal cord  
and brain only!***

Miralbell R et al., Int J Radiat Oncol  
Biol Phys 54: 824-829, 2002)

**But we need to be sure that we can  
measure the absorbed dose correctly!**

# Properties of an Ideal Dosimeter

- Linear over whole dose range
- Dose rate independence (non-linear effects at higher dose rates, e.g. recombination effects, no fading for passive detectors)
- No energy/particle dependent response
- No directional dependence
- High spatial resolution (image, small effective volume)
- Online active readout
- Easy to use, easy to set up, reliable, ...

# Commonly used Dosimeters

- **Calorimeter (real time, absolute dosimetry)**
- **Fricke dosimeter (passive detector)**
- **Ionization chambers (active detector)**
- **Radiochromic films (passive detector for relative profile and homogeneity control)**
- **TLDs / OSL (passive detectors)**
- **Si diodes / Diamonds ...(active detectors)**
- **Scintillators (active detectors)**
- **...**

# Dosimetry

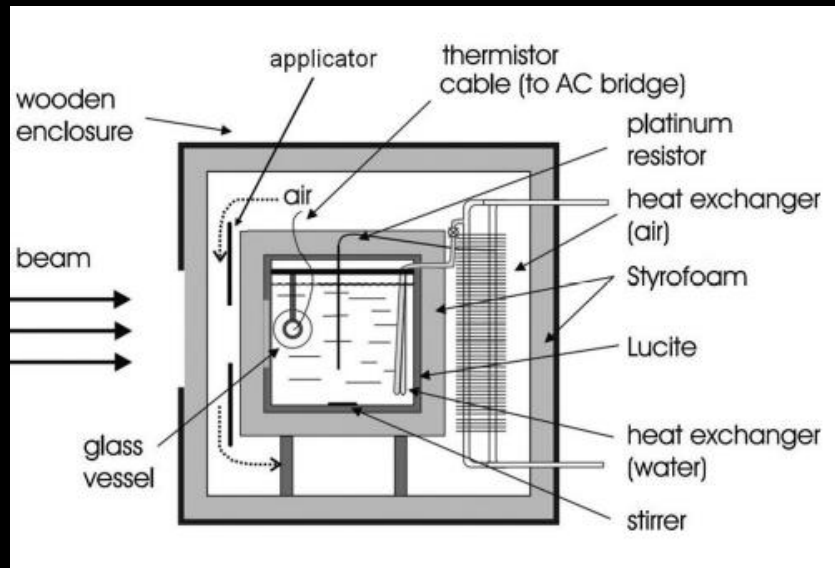
## Standards to measure absorbed dose to water

- The only solid state dosimeters used for absolute dosimetry in radiotherapy
  - Calorimeter
  - Fricke chemical detector
    - Fricke solution – (1mM  $\text{FeSO}_4$  or  $\text{Fe}(\text{NH}_4)_2(\text{SO}_4)_2$  + 0.8N  $\text{H}_2\text{SO}_4$  air saturated + 1mM NaCl)
    - $\text{Fe}^{2+}$  oxidizes to  $\text{Fe}^{3+}$ , when irradiated, which absorbs at 304 nm
- Ionization chamber

# Calorimeter

## Water calorimeter

- In a stagnant water calorimeter, absorbed dose at a point can be directly measured because of the relatively low thermal diffusivity of water:  $D_w = c_w \Delta T_w$   
 $\Delta T_w$  = temperature increase in undisturbed water solely due to radiation,  
 $c_w$  = the specific heat capacity of water (in J/kg,K)

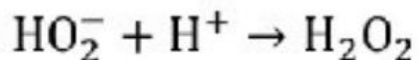
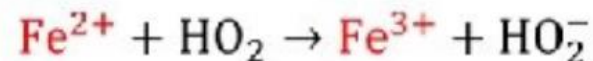
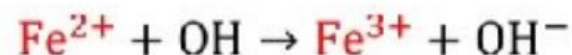
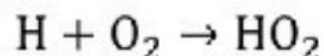
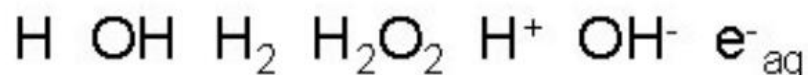


- Not practical for clinical use

# Fricke Dosimeter

Fricke dosimeter: 1 mM FeSO<sub>4</sub>, 1 mM NaCl, 0.8 N H<sub>2</sub>SO<sub>4</sub>

Ionizing radiation interacting with water produces a range of ions, radicals and molecules:



ferrous

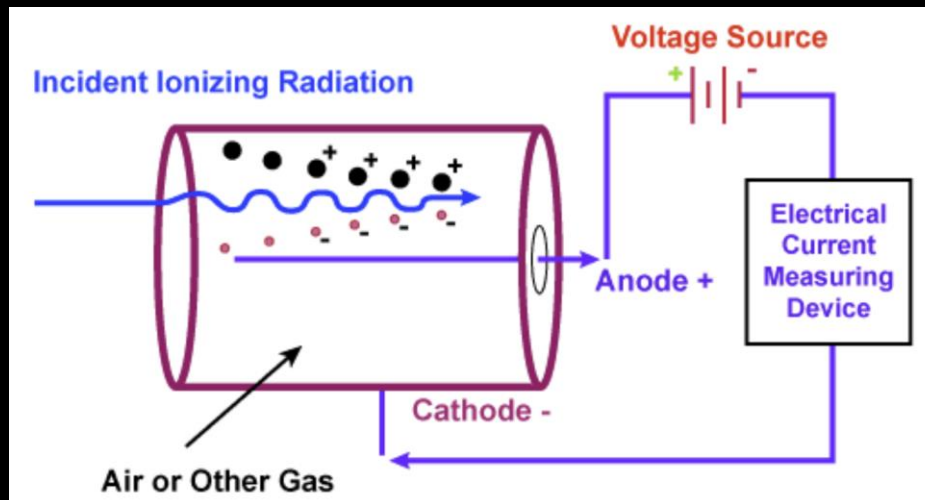


ferric

Oxidation of ferrous ions (Fe<sup>2+</sup>) to ferric ions (Fe<sup>3+</sup>) by ionizing radiation  
Fe<sup>3+</sup> absorbs at 304 nm

# Ionization Chamber (IC)

- The standard is an air filled ionization chamber.



- Ionization of the molecules of the gas occur.
- Within a high voltage field positive ions will be attracted to the negative side of the detector (the cathode) and the free electrons will travel to the positive side (the anode). These charges are measured.





## Dose in Water

The dose in water from charged particles:

$$D_w = \int_0^{E_0} \phi_E \left( \frac{dE}{\rho dx} \right)_w dE \quad [\text{Gy}]$$

# Ionization Chamber (IC)

The dose from charged particles:

$$D_w = \int_0^{E_0} \phi_E \left( \frac{dE}{\rho dx} \right)_w dE$$

- Not applicable in clinics, since fluence spectra are unknown
- We want dose to the tissue equivalent material (water), but IC is normally filled with air!

$$D_w = \frac{QW_{air}}{\rho V}$$

# Ionization Chamber (IC)

The dose from charged particles:

$$D_w = \int_0^{E_0} \phi_E e \left( \frac{dE}{\rho dx} \right)_w dE$$

- Not applicable in clinics, since fluence spectra are unknown
- We want dose to the tissue equivalent material (water), but IC is normally filled with air!

$$D_w = \frac{QW_{air}}{\rho V}$$

The volume of the gas filled cavity is not known with sufficient accuracy!

# Calibration of Ionization Chamber (IC)

- When an IC is used in a beam of quality  $Q$  different from that used for its calibration,  $Q_0$ , the absorbed dose to water is given by:

$$D_{w,Q} = M_Q N_{D,w,Q_0} k_{Q,Q_0}$$

Absorbed dose in water

Measured signal

Calibration factor at reference condition

Beam quality correction factor of the chamber to differentiate between the reference beam quality  $Q_0$  and the actual treatment beam quality  $Q$

# Calibration of Ionization Chamber (IC)

- When an IC is used in a beam of quality  $Q$  different from that used for its calibration,  $Q_0$ , the absorbed dose to water is given by:

$$D_{w,Q} = M_Q N_{D,w,Q_0} k_{Q,Q_0}$$



$$k_{Q,Q_0} = \frac{N_{D,w,Q}}{N_{D,w,Q_0}} = \frac{D_{w,Q} / M_Q}{D_{w,Q_0} / M_{Q_0}}$$

# Calibration of Ionization Chamber (IC)

A general expression of  $k_{Q,Q_0}$  is:

$$k_{Q,Q_0} = \frac{(s_{w,air})_Q}{(s_{w,air})_{Q_0}} \frac{(W_{air})_Q}{(W_{air})_{Q_0}} \frac{p_Q}{p_{Q_0}}$$

Stopping-power ratios  
signal water-to-air

Mean energies to produce  
an electron-ion-pair in air

Overall perturbation factors,  
including all departures from the  
ideal Bragg-Gray detector conditions,  
i.e.,  $p_{wall}$ ,  $p_{cav}$ ,  $p_{cel}$ ,  $p_{fluence}$ ,  $p_{dis}$ , ...

# Uncertainty in the Measured Dose

- It has been estimated\* that the overall uncertainty is
  - ≈ 2% of the dose for proton beams
  - ≈ 3% for carbon ion beams

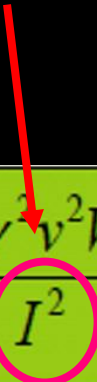
which are high compared to the 1% of photon beams

- In particular, the largest contribution comes from the uncertainty in the ratio of stopping powers between water and air ( $s_{w,air}$ )

	$k_{Q,Q_0} = \frac{(s_{w,air})_Q}{(s_{w,air})_{Q_0}} \frac{(W_{air})_Q}{(W_{air})_{Q_0}} \frac{p_Q}{p_{Q_0}}$			Total Uncertainty
	↓	↓	↓	
<sup>60</sup> Co	0.5 %	0.2 %	0.6 %	~ 0.82 %
Protons	1.0 %	0.4 %	~ 0.5 %	~ 2.0 %
Ions	2.0 %	1.5 %	1 %	~ 3 %

# Uncertainty in the Measured Dose

- Unknown electronic stopping powers remains as the primary source of uncertainty for IC dosimetry
- Uncertainties in the mean excitation ( $I$ ) values for water and compounds are important

$$-\left(\frac{dE}{dx}\right)_{coll} = 2\pi N_A r_e^2 m_e c^2 \rho \frac{Z}{A} \frac{z^2}{\beta^2} \left[ \ln\left(\frac{2m_e \gamma^2 v^2 W_{max}}{I^2}\right) - 2\beta^2 - \delta - 2\frac{C}{Z} \right]$$




# Thermoluminescence Detectors (TLD)



Prof. Dr. L. Sihver

# Thermoluminescence Detectors (TLD)

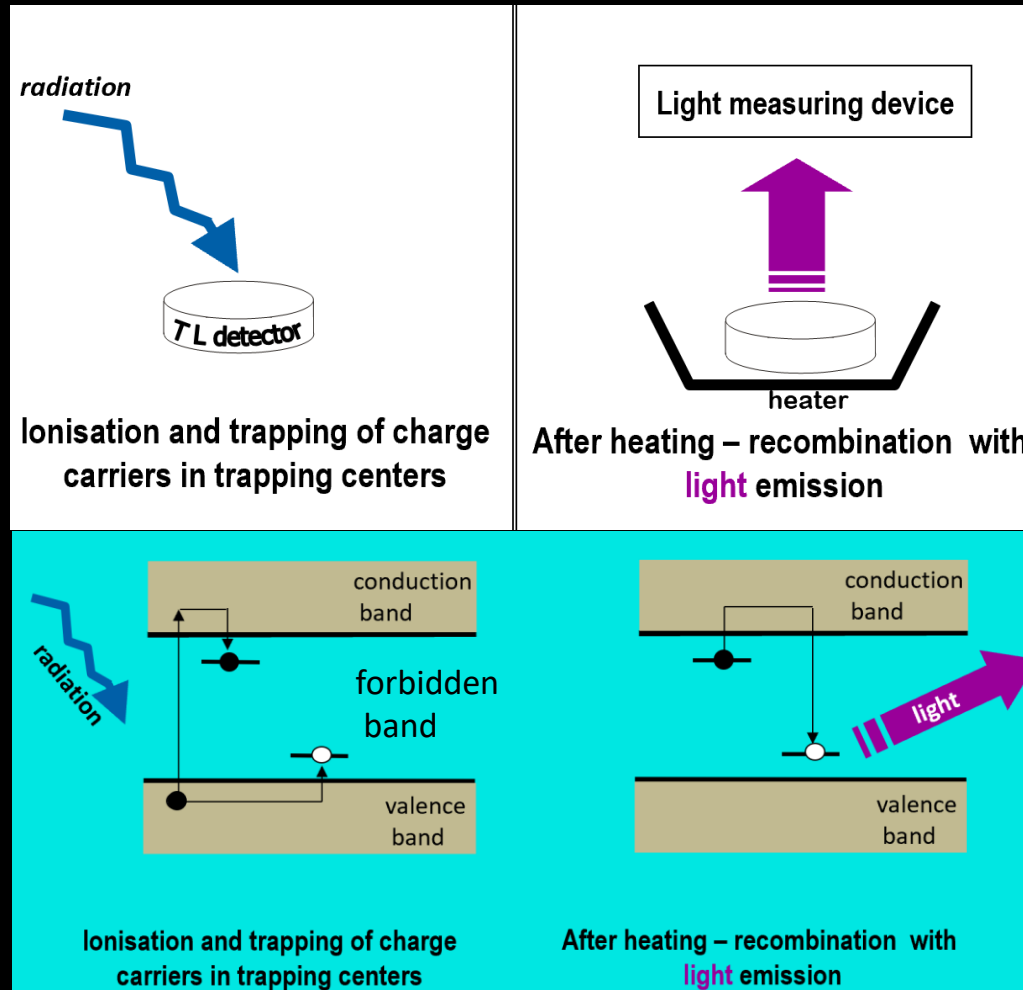


Typical TL materials

Lithium fluoride  
Calcium sulphate  
Calcium fluoride  
Aluminium oxide

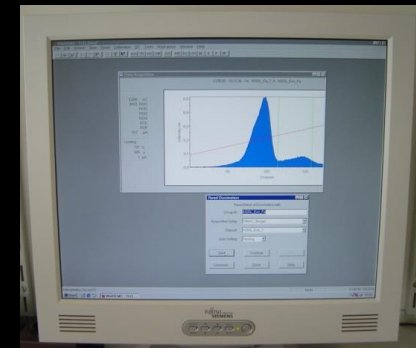
# Thermoluminescence Detectors (TLD)

Thermoluminescence (TL) is a 2-stage process

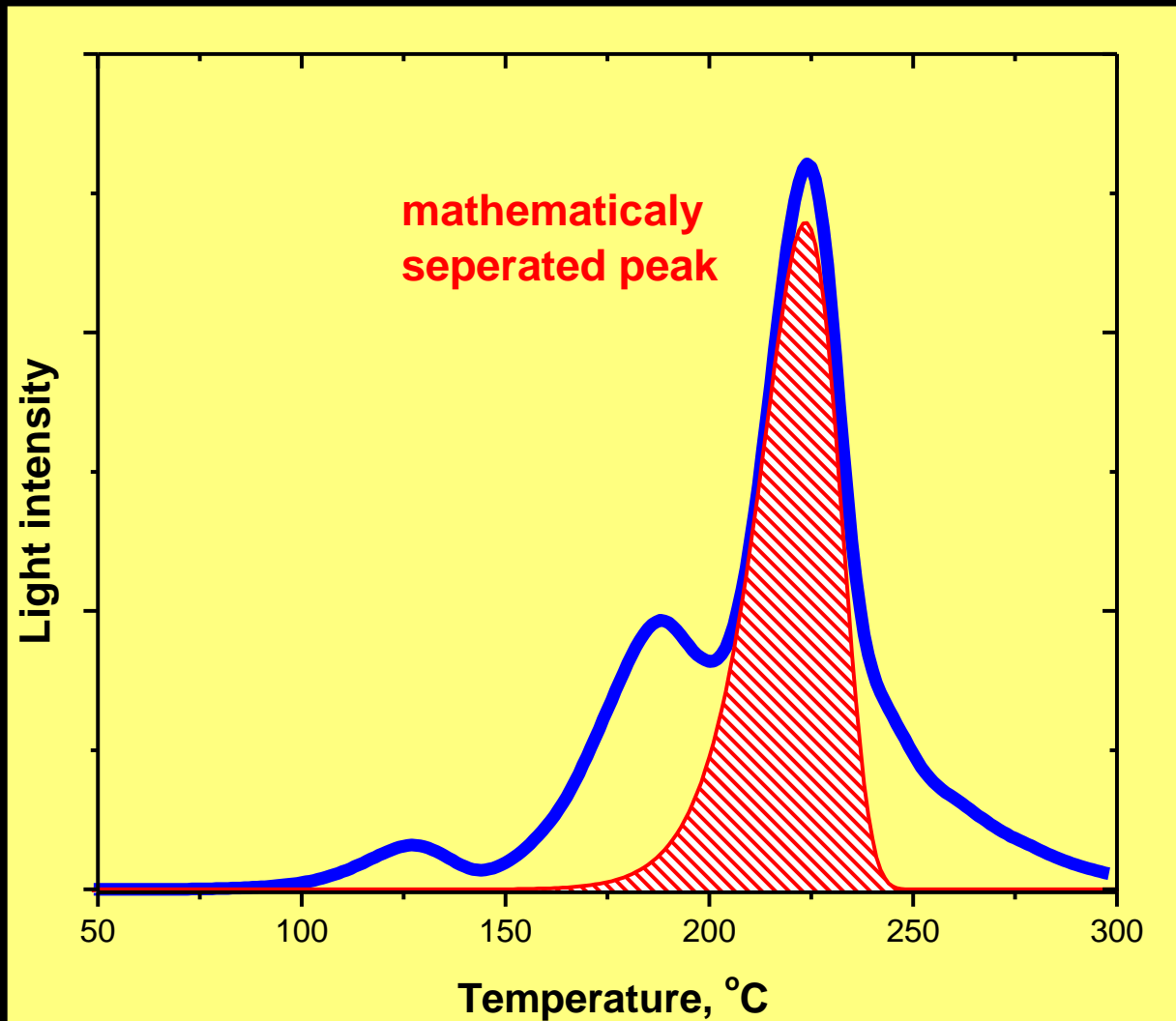


# Thermoluminescence Detectors (TLD)

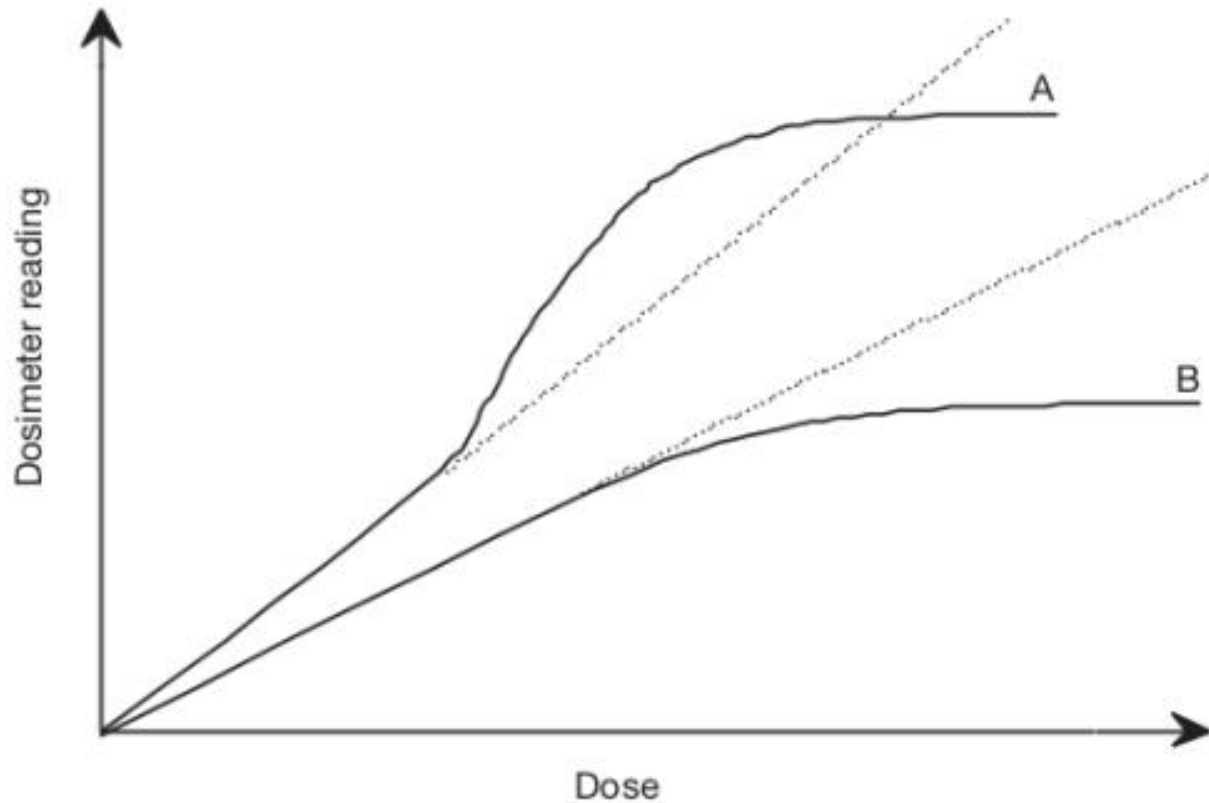
- TL – detector is place on a heating plate
- TL – detector is heated
- The light emission is measured depending on the temperature with a photomultiplier
- This leads to the so called “Glow Curve“



# TL Glow Curve



# Linearity of a Dosimeter



- A: Supralinearity and saturation
- B: Sublinearity / saturation

# Relative Effectiveness

- Different types of radiation produce different effects in a dosimeter for the same endpoint
- “*Relative effectiveness*” (*RE*) characterizes this effect and is usually defined as

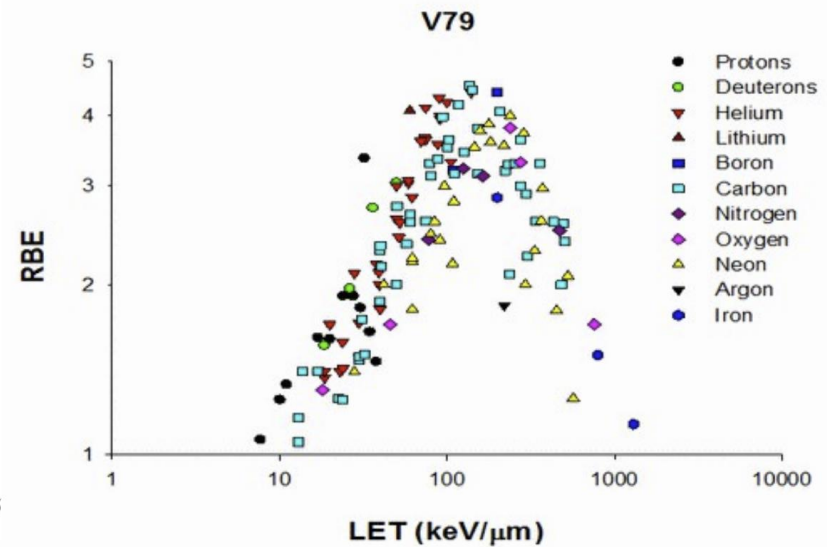
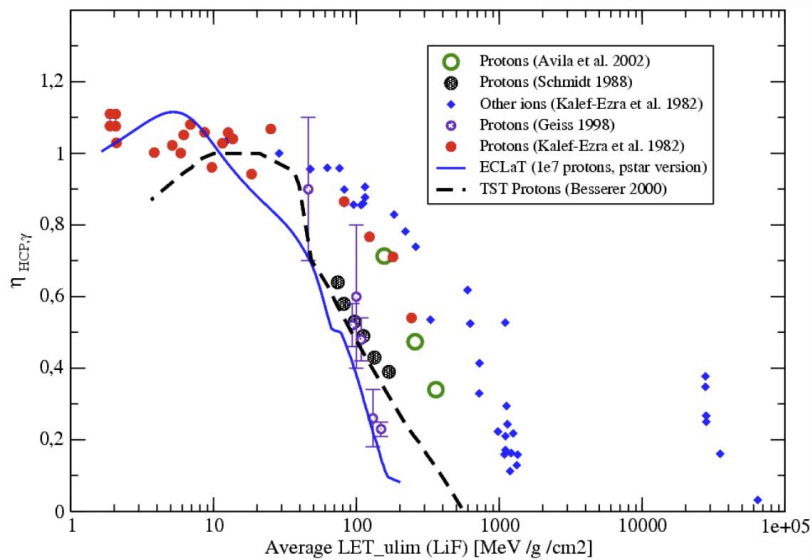
$$RE = D_{\gamma, \text{isoeffect}} / D_{\text{ion}, \text{isoeffect}}$$

# Dosimetry

Detectors

## Compare!

Radiobiology



$$\eta_{Q_0, Q} = \left. \frac{D_{Q_0}}{D_Q} \right|_{\text{iso-effect}}$$

$$\text{RBE} = \left. \frac{D_{Q_0}}{D_Q} \right|_{\text{iso-effect}}$$

Niels Bassler – Aarhus Particle Therapy Group

Copyright

Prof. Dr. L. Sihver



# Modelling of Detector Response Function

$$D_w = \int_0^{E_0} \phi_E^e \left( \frac{dE}{\rho dx} \right)_w dE$$

Measurements

Monte Carlo Simulations

Dose

Fluence

LET

...

Dose

Fluence

LET

...

Detector Response Function for  
the Actual Given Radiation Field

Detector Response

## MC Modelling $k_Q$ factor of an IC using Response Factor

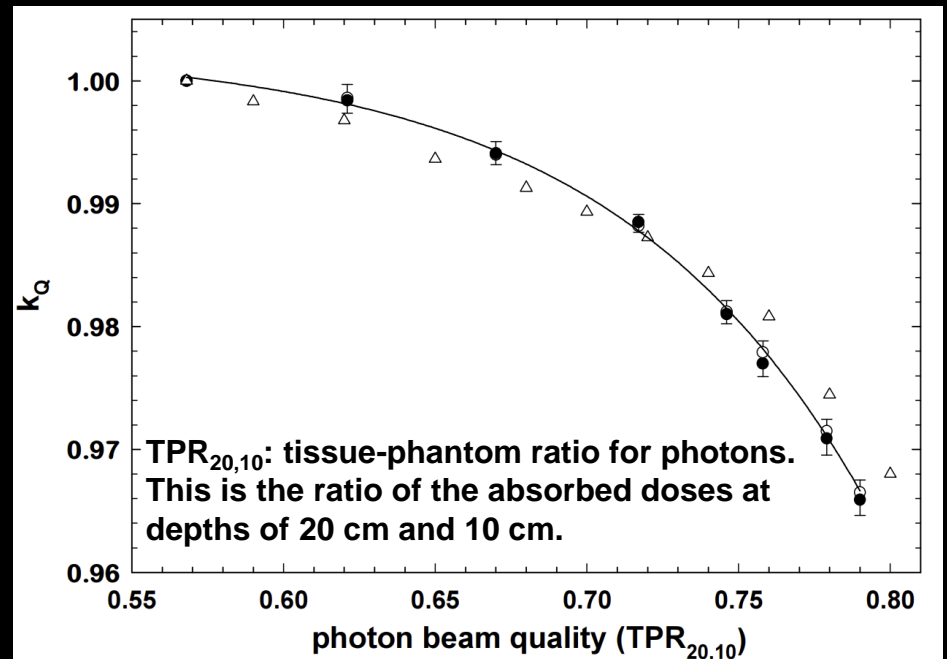
Calculation of the response factor ( $f_Q$ ) of an IC:

$$f_Q = \frac{D_w}{D_{det}} = (S_{w,air}) \rho_Q \quad \rho_Q: \text{perturbation factor}$$

Calculation of beam quality correction factor  $k_Q$ :

$$k_Q = \frac{f_Q}{f_{Q0}} = \frac{(W_{air})_Q}{(W_{air})_{Q0}}$$

- For clinical photon beams, there is a good agreement between measures and Monte Carlo calculated values for  $k_Q$ , but there is a lack of data for protons and heavier ions

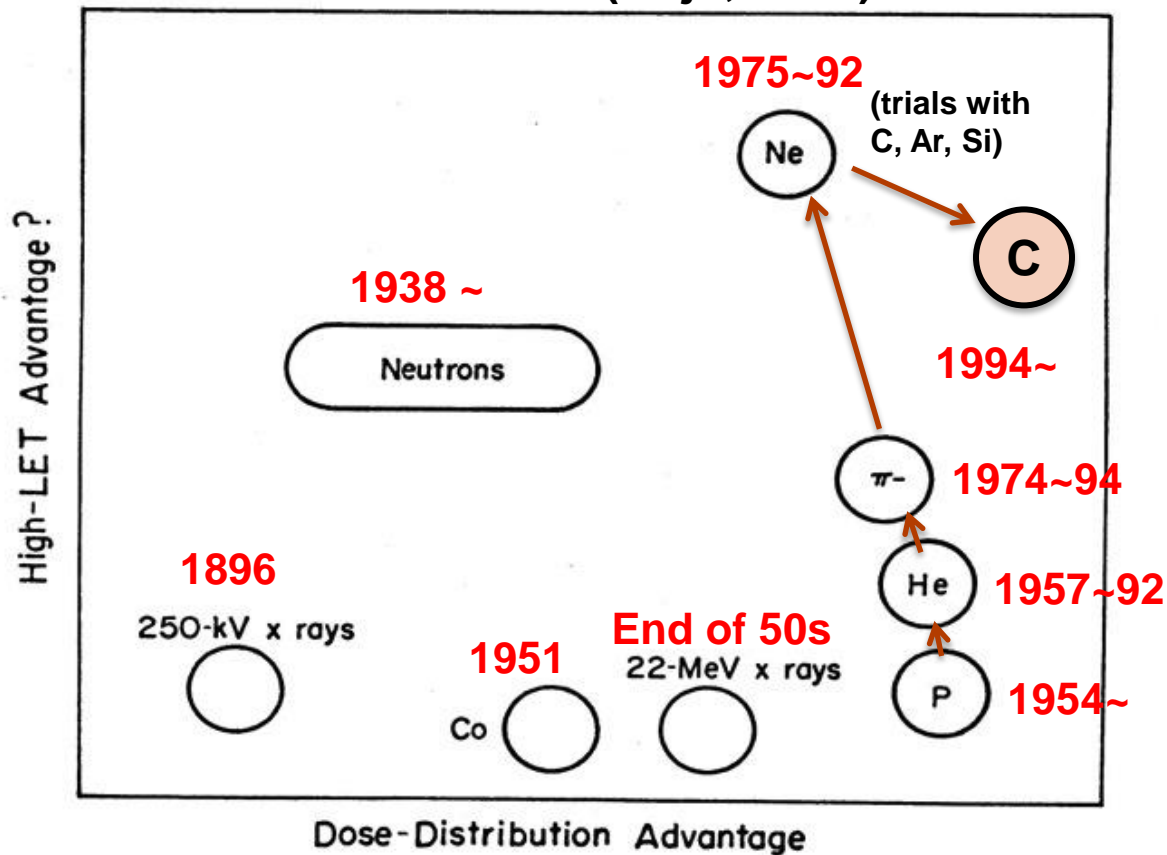


# Needed Dosimetry Improvements

- Improved accuracy of the ionization potential/range in water and other tissue/bone equivalent materials
- Micro- and nanodosimetric ionization density (ID)/track structure measurements
- On-line detection of dose and LET/ID
- For non-gaseous detectors, the sensitivity depends on the ID so when ID increase at the Bragg peak, the sensitivity decreases (quenching of the response)
- Dosimetry in strong magnetic fields during combined MRI and RT
- Dosimetry for FLASH radiotherapy (RT)
  - ultra-high dose rate (UHDR),  $\geq 40$  Gy/s

....

## Schematic Comparison of different Types of Radiations (Raju, 1974)



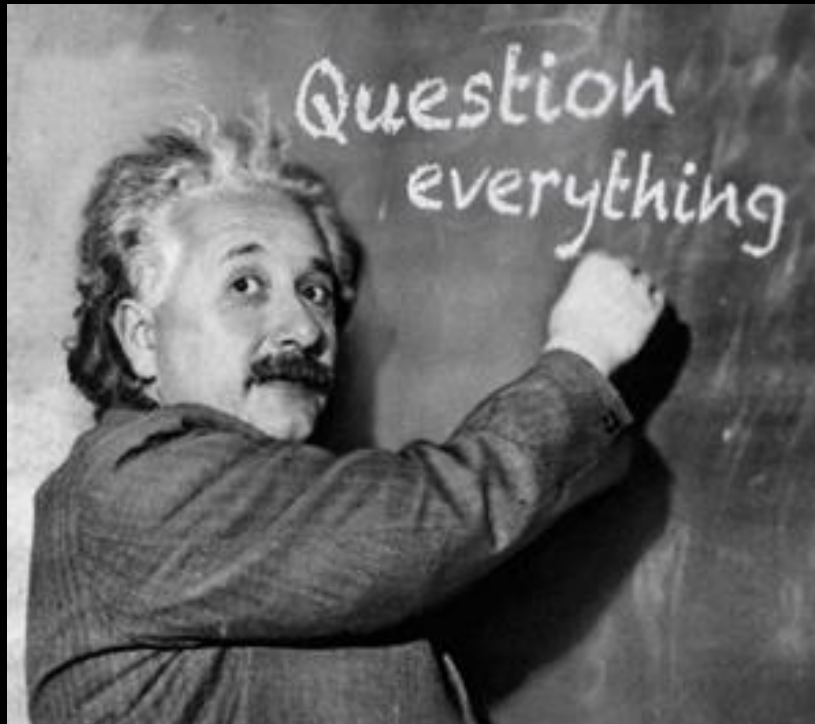
- **1946** Ion beam therapy proposal by R.Wilson
- **1968** CT invented
- **??** FLASH RT

Fig. 17. Schematic comparison of different types of radiations of interest in radiotherapy. Although based on experimental results, this comparison is an oversimplification. It is recognized that the relative positions of the particles in this schematic diagram can be argued and should be used for clarity purposes only. RBE is not taken into consideration in the dose-distribution advantage (abscissa). This idea of representing the relative merits of particles was suggested by Mr. A. M. Koehler (Raju *et al.*, 1974b).

***Thank you very much for  
your kind attention!!***



# Questions?



# Comments?



**"This material was prepared and presented within the HITRIplus Specialised Course on Heavy Ion Therapy Research, and it is intended for personal educational purposes to help students; people interested in using any of the material for any other purposes (such as other lectures, courses etc.) are requested to please contact the author:**

**Prof. Dr. Lembit Sihver**

**Email: [lembit.sihver@tuwien.ac.at](mailto:lembit.sihver@tuwien.ac.at)  
[sihver@chalmers.se](mailto:sihver@chalmers.se)**

**Copyright**