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Résumé

Introduction

- radiations used for radiotherapy
- what to estimate to carry out heavy ion therapy

Macroscopic effect

nuclear reaction
 dose

Microscopic effect

- beam quality
 - biological effect
- spatial distribution
 - advanced irradiation
- microdosimetry
 - nature of heavy ion radiotherapy
- neutrons



Radiations used for tumor therapy

direct ionizing radiation

- electron
 proton *"Heavy ion"*helium, carbon, neon, silicon, ...
 indirect ionizing radiation
 photon
 neutron



HIMAC (Heavy Ion Medical Accelerator in Chiba)

Established in 1994.Aimed at finding optimal heavy ion therapy scheme.





ion species maximum energy beam intensity repetition cycle H,He,C,Ne,Si,Ar,Fe,Kr,Xe 800 MeV/n (=Z/A=1/2) 10⁰ ~ 10⁹ particles/pulse 0.3 / 0.5 Hz

Clinical trials at HIMAC

• Targets of carbon therapy





nasal passage cancer



Nov. 1, 2003: approved as a Highly Advanced Medical Technology ¥3.14M / treatment

Basis of clinical dose prescription

+

loss of capacity for

multiplication (inactivation)

cell survival \bullet



radiation



$$RBE = \frac{D_{reference_radiation}}{D_{test_radiation}} |_{same_effect}$$

survival

carbon: 2~3

Characteristic of heavy-ion therapy

Fragmentation of incident particles



Projectile fragments have almost

-...same velocity -...same direction Therapeutic beam is contaminated by fragments!

with those of primaries.

Why?

Effect by fragmentation from clinical point of view

Production of fragment particles...



✓ ...causes unwanted dose beyond the range
 ✓ ...makes estimation of biological effect complex.
 ✓ ...makes it possible to monitor beam range.



T. Nishio (NCC-east), private comm.



M. Scholz et al., Radiat. Environ. Biophys., 36, 59 (1997).

Macroscopic effect

Depth dose distribution

Depth-dose distribution

Dose......"quantity of radiation"

Dose = J / kg[Gy]

the most fundamental factor to be controlled on radiotherapy

physical factors

reaction cross section

stopping power multiple scattering and straggling

Disintegration of primaries
 loss of dominant dose carrier

Production of fragments
 Center dose but form 'fragment tail' beyond the range

Depth-dose distribution

Depth-dose distribution in water measured at HIMAC



Dose can be controlled in clinically enough precision.

basis of ongoing carbon therapy

Macroscopic to Microscopic What? – particle identification

Importance of P. I.

■ LET and particle species dependency of RBE (CHO cell)



M. Scholz and G. Kraft, Rad. Prot. Dos., 52, 29 (1994)

How should we take into account this complexity? Radiation quality (fluence and energy)



How many? - fluence

■ ¹²C-290MeV/n

Calculation: hibrac (old)



Comparison with PHITS





Heavy-ion therapy sites



Where? – spatial distribution

Importance of the spatial distribution

Scanning irradiation with pencil beam

stopping power multiple scattering production cross section reaction cross section momentum transfer



(picture from **SIEMENS**)

lateral nonuniformity & Solution of the operation of the oper

More microscopic! - microdosimetric approach

Microdosimetric problem

 Random energy deposition in cell nucleus



FIG. 1. Energy density as a function of the mass for which energy density is determined. The horizontal line overn the region in which the absorbed dose can be established in a single measurement. The shaded portion represents the range where statistical fluctuations are important.

•amorphous (averaged) track



•actual (sparse) track



Fig. 5. Comparison of the microscopic structure of carbon tracks at different energies with a simplified depiction of a DNA molecule. Ionisation and consequently the damage to the DNA is low at high particle energies but increases significantly at lower particle energies yielding clustered damage that is difficult to repair.

What? – neutrons

Neutrons in therapy room

¹²C-290MeV/n 10000 particles (simulation with PHITS)



Geometry: from Nose (IHI)

Neutrons in therapy room

¹²C-290MeV/n 10000 particles (simulation with PHITS)

Neutron dose distribution





FIGURE 16.6. Quality factors for neutrons, that is, the maximum dose equivalent divided by the absorbed dose at the same depth in the body. The curve represents the recommendation of the ICRP, (ICRP, 1971. References in the figure are given in that report. Reproduced with perministion freen Pergamen Press, Loh.)

•Risk estimation for the induction of the secondary cancer

Dependency to irradiation scheme

Optimal treatment method

Summary

- Macroscopic effect
- Charge-changing cross section
 Depth-dose distribution is given in good precision

Dose is delivered to tumor accurately

Need further investigation for heavier elements

Summary Microscopic effects ✓ Fluence and LET distribution (broad beam) Feedback to biology (ex. survival simulation) ✓ Spatial distribution (pencil beam) Input data for RTP of scanning irradiation Angular distribution, double differential production cross section, momentum transfer, ... Development of simulation code including spatial

information, advanced RTP (inhomogeneous structure)

✓ Microdosimetric approach

Understanding of the nature of heavy ion therapy

Thank you for your attention.



Mt. Fuji from Chiba city

A part of this work was carried out as the Research Project with Heavy Ions at HIMAC.