Neutrons Secondary to Proton Therapy; **Problems & Solutions**

Eric J. Hall, D.Phil., D.Sc., FACR, FRCR, FASTRO Columbia University, New York, N.Y.

Radiation: The Two-Edged Sword

• Radiation therapy can cure cancer

• Radiation can induce cancer



PROTONS

 Because they have a limited range and deposit most energy in the Bragg peak, Protons are the logical next step in improving dose distributions, i.e. to maximize the dose to the tumor and minimize dose to normal tissue.

Proton IMPT, 3 Scanned Pencil Beams



Epithelioid Sarcoma Close to Vertebrae

The Herman Suit Doctrine

 Normal Tissues that are not Irradiated cannot develop Late Effects of Any Sort
 Fibrosis
 Cancer

PROTONS

Radiation-induced cancers outside the treatment volume (the problem with IMRT) should be essentially eliminated because of the reduction in the volume of normal tissues exposed.

- However.....





Protons

- Radiation induced cancers outside the treatment volume should be essentially eliminated because of the reduction in the volume of normal tissues exposed
- However...

Beaming Guides

Energetic Proton When a high-energy proton bombards a heavy atomic nucleus, causing it to become excited, neutrons are expelled

How to scan pencil proton beams over a large target...

Passive scanning



Active scanning





Neutron RBE

Photons have curved dose-response relations, while those for neutrons are straighter



It follows that the neutron RBE is dose dependent, with a constant maximal value (RBE_M) at low doses



Estimated low-doses RBE values for fission-neutron induced carcinogenesis in mice Data from Ullrich et al.

Mouse strain / Endpoint	Measured RBE _M
RFM / Thymic lymphoma	27 ± 26
RFM / Pituitary	59 ± 52
RFM / Harderian gland	36 ± 10
RFM / Lung tumor	6 ± 3
BALB/c Lung adenocarcinoma	19 ± 6
BALB/c Mammary carcinoma	33 ± 12
Overall	30 ± 17

Neutron RBEs – Dicentrics in Human Lymphocytes



Nolte et al., Radiation and Environmental Biophysics (2005)

Neutron RBE Must assume a value of 20 to 30

- Fission Neutrons in A-bomb Survivors; RBE=100 (Lower Limit 25)
- Fission Neutrons in Mice; variety of Cancers; Mean RBE=30

Chromosome Aberrations in Lymphocytes for a Neutron Spectrum Similar to MD Anderson; RBE=100 ??







Neutron Dose Equivalent vs. Distance from Field Edge



Distance from Field Edge (cm)

19

Northeast Proton Therapy Center

- Patient treated for lung cancer (72 Gy)
- Monte Carlo calculations to arrive at neutron organ doses (Paganetti et al. Phys Med Biol 2005)
- Cancer Risks calculated for External neutrons.
 Neutron RBE assumed to be 25 (Brenner & Hall 2007)

Calculated neutron organ doses for a three-field proton therapy plan at the passively-modulated NPTC facility, treating a lung tumor with a planned 72 Gy GTV dose

	Neutron	dose (mGy)
Organ	Internal	External
Red bone marrow	28	16
Colon	0	4
Lung (out of field)	39	34
Stomach	0	20
Bladder	0	3
Breast	1	24
Liver	1	32
Esophagus	2	29
Thyroid	1	32
Brain	0	12
Kidney	0	19
Pancreas	0	22

 Doses derived from detailed Monte-Carlo simulations of the NPTC beamline by Paganetti and colleagues



- For a 72 Gy proton therapy lung-tumor plan at the passively-modulated NPTC facility, assuming the patient is cured of his / her primary tumor. **
- **Doses derived from detailed Monte-Carlo simulations of the NPTC beamline** ** by Paganetti and colleagues
- **RBE assumed: 25** **

Paganetti, Suit, and colleagues are quite right....

"The major secondary dose contributors are neutrons from the proton treatment nozzle. These external neutrons account for a much higher secondary dose (by about two orders of magnitude) than the internal neutrons."

Jiang, Wang, Xu, Suit and Paganetti (2005)

Treatment nozzle for a passive scattering proton therapy beamline



© M. Goitein: Application of Physics in Radiation Oncology



Evaluated neutron production cross sections for 150-MeV protons



25

A polyethylene collimator (ρ =0.94 g/cc) will result in a significant reduction in the secondary neutron dose But it would have to be ~14" thick.



But a polyethylene ($\rho=0.94$ g/cc) collimator would have to be ~14" thick, which would be problematic



Using a 7¹/₂" thick SWX-207HD polyethylene collimator (0.8% boron: density =1.65 g/cc) will result in a significant reduction in the secondary neutron dose



A 7늘" thick collimator....

... may still be problematic, if it degrades the lateral penumbra

Solution 1: Taper the inside walls of the collimator (Polyethylene is very easy to machine)

Problems and Solutions. The Problem

Having spent about \$125 million on a proton facility to reduce dose to normal tissues, does it make sense to spray the patient with a total body dose of neutrons, the RBE of which is poorly known, and end up with a second cancer risk similar to IMRT ?

Problems and Solutions. Solutions;

 Use a Scanning Beam and avoid the problems of Passive Modulation.....
 However this is technically difficult, and introduces problems of its own

 Replace the Brass collimator with a collimator made of SWX-207HD Polyethylene (containing Boron), or a Hybrid collimator, made of part brass and part polyethylene.(Brenner, Hall & Paganetti, in press)

NASA Space Radiation Laboratory

At the 1GeV/nucleon Iron Ion beam at Brookhaven National Laboratory, the neutron contamination is minimized by the use of a massive collimator consisting of layers of LUCITE, **ALUMINUM and POLYETHYLENE.** This sort of technology could replace Brass in the clinical facilities.



CENTER FOR RADIOLOGICAL RESEARCH 2006

In most clinical configurations, almost all the external neutron dose comes from the final brass (or cerrobend) patient-specific collimator

Brass: Copper 61.5% Zinc 35.2% Lead 3.3%

Cerrobend: Bismuth 50% Lead 26.7% Tin 13.3% Cadmium 10%