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From the ionization chamber to the on-line microstrip devices:

the status of the art of detectors in proton therapy under the experience gained at the CATANA facility

Scuola Nazionale "Rivelatori ed Elettronica per Fisica delle Alte Energie, Astrofisica, Applicazioni Spaziali e Fisica Medica" INFN Laboratori Nazionali di Legnaro, 20-24 Aprile 2009

TALK OUTLINE

- 1. Absolute and relative dosimetry in proton therapy: starting point for the dosimetric commissioning
- 2. Ten years of detector characterisation
- 3. What future for clinical proton beam detectors?



DETECTOR DEVELOPMENT AND CHARACTERIZATION

STILL A NEED IN RADIATION THERAPY AND, IN PARTICULAR, IN A YET PIONEERING TECHNIQUE LIKE PROTON THERAPY

ABSOLUTE AND RELATIVE DOSIMETRY

(Dosimetric commissioning)

CONTINOUS R&D WORK

A complete description of each detector is impossible!

Dosimetric commissioning: absolute & relative dosimetry

Absolute Dosimetry: Energy Released in Water (Gray)

Relative Dosimetry: Three dimensional dose distribution measurements

Considering the high gradient dose, conformation and small fields often used the detectors have to be kindly characterized in terms of spatial resolution, energy or fluence dependence to be use in hadrontherapy.

Relative and Absolute Dosimetry are fundamental for:

Customizing of TPS Monitor Unit Calculation Quality Control

DETECTOR OVERVIEW

Activity started in 1996 with the first test on 30 MeV protons and continued with 62 MeV protons

Gas detector (ionisation chambers)

Film detectors

Solid state detectors

10 year of work 43 papers in review journals 61 conference proceeding Farmer Exradin

Pin Point

Markus

Advanced Markus

Radiographic films GAF Chromic films (MD52, EBT, ...

•Plastic scintillators

Diamond detector

•TLD

•MOSfet

•Silicon diode

•Silicon microstrip

62 MeV experimental proton beam

FULL ENERGY BEAM





62 MeV experimental proton beam



WHAT WE NEED FROM A DOSEMETER?

- Tissue equivalence (<Z> = 6, <Z/A> = 0.5)
- Linearity vs absorbed dose
- Independence from dose rate
- Independence from energy
- Radiation hardness
- Spatial resolution (< 1 mm)
- Small size: no perturbation of the proton fluency (Bragg-Gray theory)

MEASUREMENT CONFIGURATION



Detector placed at isocenter on a special table mounted on the treatment chair

MEASUREMENT CONFIGURATION



Depth in Eye Tissue (mm)

Detector placed at isocenter on a special table mounted on the treatment chair

ABSOLUTE DOSIMETRY

ICRU 59 AND TRS 398 IAEA RECOMMENDATION

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"FOR MEASUREMENTS OF DEPTH-DOSE DISTRIBUTION IN PROTON BEAMS

THE USE OF PLANE-PARALLEL CHAMBERS IS RECOMMENDED"

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Parallel plate MARKUS PTW is the golden standard for depth dose measurements

Dosimetric commissioning: absolute & relative dosimetry

ADVANCED MARKUS CHAMBER



- $V_P = 400 V \rightarrow V \times cm^{-1} = 4000 \rightarrow k_s = 1.00 (1 \div 100 Gy/min.).$
- **2** Pressure equilibrium ≤ 10 sec
- **3** Temperature equilibrium = 2-3 min./K
- **4** Response: 670 pC/Gy
- **5 Directional dependence:** smaller than 0.1% for tilting of the chamber by up to 10°
- **6** Electrode Acrylic (PMMA), graphite coated 5 mm Ø
- Deakage current ± 4 fA



Ionisation, free air monitor chamber calibrated against the Markus: Sensibility: 5 pC Reprucibility: 0.5 %

DETECTOR FOR RELATIVE DOSIMETRY













Film Kodak: XV and EDR2 2) TLD 3) Radiochromic Film
Scanditronix Diode 5) PTW Natural Diamond 6) Mosfet
In collaboration with ISS (S. Onori..) and DFC Florence (M. Bucciolini...)

RELATIVE DOSIMETRY: FILM DETECTORS

GAFCHROMIC® EBT Configuration

CLEAR POLYESTER - 97 microns

ACTIVE LAYER - 17 microns

SURFACE LAYER - 6 microns

ACTIVE LAYER - 17 microns

CLEAR POLYESTER - 97 microns

1) Transmission densitometer 2) Colour scanner

SPECTRA OF NEW AND OLD RADIOCHROMIC FILMS



ENERGY INDEPENDENCE HS GAF



DOSE CALIBRATION



Radiochromic Film Calibration Exposures



EBT GAF



EPSON flatbed color scanner RED CHANNEL

RELATIVE DOSIMETRY: plastic scintilators



Alumina (Al2O3)



Caesium Iodide (CsI)



Plastic scintillator BC 400

- Alumina: poor in light efficiency;
- Caesium Iodide: excellent light efficiency, but scarce homogeneity;
- BC 400: good brightness and homogeneity;

BC 400 has been chosen for our experimental purposes.

DETECTOR ARRANGEMENT

Mirror forming 45° with the beam direction.

Scintillating screen lodged on a support, perpendicularly to the beam axis.

Camera forming 90° with the beam axis, framing the image reflected by the mirror.

LabView Environment



CHARACTERIZATION

LINEARITY





Linearity in the treatment range $(10 \div 20 \text{ Gy/min})$



Castleman says: SNR (N)= N^0.5*SNR(1)

Experimental results are in good agreement with theoretical considerations

RELATIVE DOSIMETRY: diamond detectors (synthetic and natural)

PROPRIETA'	DIAMANTE	SILICIO
Gap [eV]	5.5	1.12
Campo di rottura [V/cm]	10 ⁷	3·10 ⁵
Hole mobility [cm ² /Vs]	1200	450
Velocità di saturazione [cm/s]	2.2·10 ⁷	0.8·10 ⁷
Mobilità elettronica [cm ² /Vs]	1800	1450
Vita media dei portatori minoritari [s]	10 ⁻⁹	2.5·10 ⁻³
Costante dielettrica ϵ_r	5.7	11.9
Numero atomico effettivo Z _{eff}	6	14
Energia per creare un coppia elettrone-lacuna [eV]	13	3.6
Energia di Wigner [eV]	43	13-20

RELATIVE DOSIMETRY: diamond detectors (synthetic and natural)

PROPRIETA'	DIAMANTE	SILICIO	Low dark
Gap [eV]	5.5	1.12	current
Campo di rottura [V/cm]	10 ⁷	3·10 ⁵	
Hole mobility [cm ² /Vs]	1200	450	
Velocità di saturazione [cm/s]	2.2·10 ⁷	0.8·10 ⁷	Fast
Mobilità elettronica [cm²/Vs]	1800	1450	responce
Vita media dei portatori minoritari [s]	10 ⁻⁹	2.5·10 ⁻³	time
Costante dielettrica ϵ_r	5.7	11.9	
Numero atomico effettivo Z _{eff}	6	14	l issue
Energia per creare un coppia elettrone-lacuna [eV]	13	3.6	equivalence
Energia di Wigner [eV]	43	13-20	Radiation hardness

ON-LINE CONFIGURATION (CVD diamond)

De Beer's diamond sample with two bonding solution





ON LINE CONFIGURATION: CVD vs NATURAL



RESPONCE vs ABSORBED DOSE (CVD diamond)



PTW NATURAL DIAMOND

TWO PTW DIFFERENT DETECTORS HAVE BEEN STUDIED



In photon and electron beams the relative differences have been studied and already published (see De Angelis C et al. Med. Phys. 2002; 29(2): 248-254.) In proton beams the measured repeatibility is 0.1%

PTW NATURAL DIAMOND

PTW Diamond Linearity (protons, photons, electrons)



PTW NATURAL DIAMOND

Bragg Peak Measurements



TLD characterization in proton beam



The linear region of the calibration curves covers the same dose range (0.2 ÷ 2 Gy) for all energies tested

Good agreement for lateral profile between TLD and radiographic film



RELATIVE DOSIMETRY: surface TLD detector

- 0.27mm mean thickness,
- 7 detectors in each set,
- 10 detector sets.



PHANTOM & DETECTORS



6 sets irradiated! in dose range 4.5 – 18Gy

READOUTS



SILICON DIODE

SCANDITRONIX Silicon Diode

Two different p-type Scanditronix detectors having the following features have been studied:

- •Thickness: 0.06 mm
- •Sensitive Area: 0.28 mm²
- •Sensitive Volume: 0.017 mm³
- •Materials tickness in front of the detectors: 0.42/0.54 mm
- •Center Distance from Detector Surface: 0.546/0.684

Both samples were preirradiated in proton beams, before shipping.

Measured repeatibility (10 meas. at 4 Gy): 0.2% Sensitivity change after 300 Gy irradiation 0.7%

SILICON DIODE



SILICON DIODE: full energy Bragg peak

The two detectors give different positions of the peak.

An overestimation of the peak has been noted (up to 6%).




Scanditronix Diode characterization in proton beam



Detector diameter: 0.6 mm (t = 60 μm)
 Detector Material: Hi-pSi, high doped p-type silicon (*preirradiated* for use in proton beams)

SILICON DIODE: Spread Out Bragg Peak



RELATIVE DOSIMETRY: silicon diode and MOSFET





MOSFET (Metal Oxide Semiconductor Field Effect Transistor)



Correct Source-Drain I_{ds} V_{th} V_{th} Tensione di gate V_G [mV]

Why a MOSFET:

- > stable, reproducible
- > linear vs absorbed dose
- > dose-rate independence
- > on-line reading
- temperature indipendence
- > no fluence perturbation
- > small size

It is OK for *in-vivo* dosimetry

INSTRMENTATION

•Autosense™ reader

· MOSFET TN-502RD

"Dual Sensitivity Bias Supply" (up to 5 MOSFETs)



2 sensibility modality: *High e Standard*

A 15:28





AutoSense™ PCSoftware V. 1.1

RESPONCE vs ABSORBED DOSE



A STUDY OF THE OUTPUT FACTORS

Output Factor



Output Factor All dosimeters

THE MOPI ONLINE MONITOR

2 ionization chambers with anode segmented in strips (x,y)



Sensitive area
Total thickness
Number of strips/chamber
Strip width
Pitch
Readout rate



THE MOPI ONLINE MONITOR: TEST SET-UP



ionization chambers

THE MOPI ONLINE MONITOR

Three different currents of the beam steering magnet



What is the future?

- Proton computed tomography (PCT)
- MEDIPIX
- On line PET (DOPET)



WHY proton Computed Tomography (pCT)?

TOMOGRAPHIC IMAGE RECONSTRUCTION USING HIGH ENERGY PROTON BEAMS

MAIN IUSSUES IN PROTON THERAPY QUALITY

Patient positioning



Actually TPS are based on the xCT images as input and this bring a sensible amount of imprecision

THE ROLE OF MONTE CARLO

Study of the proton paths inside objects

•Study of the algorithms for image reconstructions

GEANT4 tomographic image



200 MeV, 179 projections at 1°, 5M Histories, 20 cm circular phantom

EXPERIMENTAL SOLUTION FOR SINGLE TRAKING

All the studies and prototypes developed in the last years are based on the principle of

follow each single proton traversing the medium to investigate



The "single tracking" approach should permit to improve the spatial risolution up to 1 mm or less

$$\int_{L} \eta_{e}(\vec{r}) d\vec{r} = K \int_{E_{out}}^{E_{in}} \frac{dE}{S(E)}$$

THE PRIMA (PRoton IMAging DETECTOR



Single tracking

Acquisition rate up to 1 MHz

Detector: 2 orthogonal microstrip; 200 um picth x 256 strips; about 5x5 cm of active area

Trigger from the calorimeter

THE ROLE OF MONTE CARLO

Experiment: 250 MeV proton beams (LLUMC)



Our Geant4 application and results on detector study



THE ROLE OF MONTE CARLO



The detector is based on a 300 µm thick silicon pixel detector bump-bonded to the Medipix2 readout chip (MPX2MXR version) to form an assembly of 256 x 256 square pixels at a pitch of 55 µm. The active area of the detector is 2 cm². Each cell of the read-out chip comprises a low noise preamplifier, two pulse height discriminators and a 14 bit counter. The maximum counting rate per pixel is 1 MHz. The chip operates in single event counting mode, that is only the events releasing an energy above a threshold are counted. The threshold is adjustable at the pixel level with 3-bit resolution.



The tests have been performed with the 62 MeV proton beam. In the first test the count rate measured by our detector has been compared to the one acquired by a reference detector based on a YAP scintillator. Good agreement between the two counters (the measured count rate of both detectors was 550 events/mm^2 pA),

G. Bisogni et al. INFN Pisa

SET-UP

- Fascio Catana protoni da 62 MeV senza modulazione
- Rivelatore MPX2 (Si 300 um spessore, 55 um pixel, single event counter)
- Collimatori in ottone
 - Diametro 5 mm, uno e due fori



MISURE EFFETTUATE

- Stima del rate di protoni sul rivelatore
 - Misura effettuata confrontando il rate misurato da uno scintillatore YAP di Nunzio con il rate misurato da MPX2
- Misure di conteggi in funzione del tempo di acquisizione
- Misure di conteggi in funzione di dose e dose rate
- Misure di allargamento dei profili dopo vari spessori di lucite



"RADIOGRAFIA" DEL FASCIO





 Collimatori fori da 5 mm
 Soglia discriminatore 190 keV
 Dose rate 0.15 Gy/sec

STIMA DEL RATE DI PROTONI SUL RIVELATORE

- La misura di rate con lo YAP e' stata fatta fissando la corrente del fascio a 3 pA
- II dose rate stimato e' di 3 *10⁻⁴ Gy/sec
- Tale valore e' stato scelto per minimizzare il contributo degli eventi di pile-up sul rivelatore
- Risultati:
 - Collimatore un foro da 5 mm:
 - Event rate medio = 10 kHz/pA
 - Collimatore due fori da 5 mm;
 - Event rate medio = 20 kHz/pA
 - Event rate atteso per pixel= 1.6 sec⁻¹ * pA⁻¹

EVENT RATE MPX2

- Corrente fissata a 3 pA
- II dose rate stimato e' di 3 *10⁻⁴ Gy/sec
- Soglia discriminazione 190 keV
 - Event rate misurato:
 - 1.7 sec⁻¹ * pA⁻¹

- La corrente portata a 1 nA
- Dose rate stimato 0.1 Gy/sec
- variata con continuità la soglia del chip da 10 keV fino a 250 KeV (lim max, saturazione preamplificatore)



EVENT RATE VS DURATA GATE

- Abbiamo fissato la soglia a 190 keV e la corrente del tubo a 1 nA
- Dose rate stimato = 0.1 Gy/sec
- Variata la durata del gate di acquisizione da 0.1 a 5 sec



EVENT RATE VS DOSE

- Fissata la soglia a 190 keV e la durata del gate di acquisizione a 10 sec
- Misurato event rate sul rivelatore in funzione della dose a diversi dose rate



IMMAGINI CON SPESSORI LUCITE

Immagini ottenute ponendo davanti al rivelatore spessori crescenti di lucite (da 5 a 20 mm)



Soglia 190 keV Dose in ingresso 30 cGy Dose rate 0.15 Gy/sec

PROFILI ORIZZONTALI

Profili orizzontali lungo la riga passante epr il centro delle immagini

Experimental NO PMMA









Vertical profile of the counts for the central column

Image of the beam cross section obtained by scanning our detector across the beam area. The image size is **30 x 30 mm ^2** and it has been obtained in 9 steps. The dose for each step was 30 cGy.



"PENOMBRA" 20-80

Confronto dati sperimentali con simulazione GEANT4



Stringent necessity of monitoring

The higher physical selectivity of ion therapy demands higher precision in the monitoring of the applied treatment due to:

- fractioned therapy
- shifts of the patient
- local tissue reduction

For these reasons in vivo information on the range of ions are desirable, but the complete stopping of the ions in patient prevents the application of electronic portal imaging methods as used in conventional radiotherapy.

Possibilities:

 radiographic imaging of high energy transmitted protons prior to therapeutic irradiation (hadron CT)

• In-beam PET (DOPET- INFN and University of Pisa)

A. Del Guerra, F. Attanasi, N. Belcari, S. Moehrs, V. Rosso, S. Vecchio, G.A.P. Cirrone, G. Cuttone, P. Lojacono, F. Romano, N. Lanconelli.

β⁺- emitter target and projectile fragments:

- Protons:
 - ¹⁶O (p,n) ¹⁵O
 ¹²C (p,n) ¹¹C
 T^{1/2}_{15-O}=121.8 s
 T^{1/2}_{11-C}=1222.8 s
 - 15-20 MeV threshold for p-induced nuclear reactions that cause poor spatial correlation between β^+ -activity and dose depth profile
- Carbon:
 - X (¹²C, ¹¹C+n) X X (¹²C, ¹⁰C+2n) X ¹⁶O (¹²C, X) ¹⁵O+n T^{1/2} _{10-C} =19.3 s
- ¹²C (¹²C, X) ¹¹C +n

- Superior biological effectivenes
- The induced β+activity can be measured with a PET tomograph.



The in-beam PET-tomograph prototype



Two planar heads, each with an active area of 45 mm x 45 mm
Distance between the heads: 7+7 cm

Nominal energy: 62 MeV Final collimator: 25 mm Ø Full energy Dose: 30Gy peak T-irr= 20sec. T-acquis = 20sec

LYSO crystal matrix, 21 x 21 pixels 2.152 mm x 2.152 mm each (Hilger Crystals)
Crystal thickness: 18 mm

The in-beam PET-tomograph prototype



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LYSO crystal matrix, 21 x 21 pixels 2.152 mm x 2.152 mm each (Hilger Crystals)
Crystal thickness: 18 mm
64-anode PMT (Hamamatsu)
"multiplexed" read-out electronics: 64-inputs/4 outputs

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Nominal energy: 62 MeV Final collimator: 25 mm Ø Full energy Dose: 30Gy peak T-irr= 20sec. T-acquis = 20sec The experiments on homogeneous phantoms The feasibility of range monitoring

 SOBPs irradiation of PMMA phantoms were performed using different range shifters along the beam line so that each irradiation differed from the other ones only in the proton range, with variations less than 2 mm.

Label	Material	Equivalent thickness in PMMA (mm)
a2	Aluminum	1.9
a3	PMMA	2.9
a4	PMMA	3.9
a6	PMMA	5.8

• Preliminary dosimetric measurement of each selected dose configuration was performed for accurate irradiation planning.
The experiments on homogeneous phantoms

The feasibility of range monitoring



Unfolding: preliminary results



Filter extrapolated from the data available for the Full Energy Bragg peak and mesured activity

Filtered dose: β+ activity distribution obtained from the application of the filter function to measured dose depth profiles

Resolution of air gaps in PMMA phantoms within the irradiation field

PMMA phantom with 0.5 cm Air_Gap at 2 cm depth;



•Phantom irradiation:

- Bragg peak dose: 30 Gy
- Irradiation time: 18 s;

•Beam cross sention: 2.5 cm Ø;

• Acquisition time: 20 min;



Resolution of air gaps in PMMA phantoms within the irradiation field

• Final three holes collimator: 0.5 cm Ø each;







The experiments on slab phantoms Sensitivity of the PET method



- Monoenergetic irradiation:
 - Bragg peak dose: 30 Gy;
 - Irradiation time: 18 s;
- Beam cross sention: 2.5 cm Ø;
- Acquisition time: 20 min;



The experiments on slab phantoms Sensitivity of the PET method Reconstructed isotope







rison of two dedicated "in beam" PET



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BASTEI (beta activity measurements at the Therapy with Energetic ions) is in use at GSI.

2 heads 42x21 cm2

8x8 BGO christals: 6.7x6.7x20mm3

A cylindrical PMMA phantom (7cm diameter, 7cm length) was irradiated with 3 monoenergetic 12C beams (108.53 112.60 116.57 AMeV). A square section beam of 28mm in side was adopted and a total dose of 60Gy was delivered for each energy. The acquisition time was set at ~30 minutes for both PET systems.

Preliminary

Energy	[AMeV]	108.53	112.60	116.57
Range	[mm]	21.9	23.6	25.3



Thank you for your attention