Original paper

The optimal balance between quality and efficiency in proton radiography imaging technique at various proton beam energies: A Monte Carlo study

A.K. Biegun^{a,*}, M.-J. van Goethem^{a,b}, E.R. van der Graaf^a, M. van Beuzekom^c, E.N. Koffeman^c, T. Nakaji^d, J. Takatsu^d, J. Visser^c, S. Brandenburg^a

^a KVI-Center for Advanced Radiation Technology, University of Groningen, Zernikelaan 25, 9747AA Groningen, The Netherlands

^b Department of Radiation Oncology, University Medical Center Groningen, University of Groningen, Hanzeplein 1, 9700 RB Groningen, The Netherlands

^c National Institute for Subatomic Physics (Nikhef), Science Park 105, 1098 XG Amsterdam, The Netherlands

^d Department of Radiation Oncology, Graduate School of Medicine, Osaka University, 2-2 D10 Yamadaoka, Suita, Osaka 565-0871, Japan

ARTICLE INFO

Article history:

Received 16 January 2017

Received in Revised form 4 August 2017

Accepted 9 August 2017

Available online 30 August 2017

Keywords:

Proton radiography

Proton scattering angle

Proton treatment plan

ABSTRACT

Proton radiography is a novel imaging modality that allows direct measurement of the proton energy loss in various tissues. Currently, due to the conversion of so-called Hounsfield units from X-ray Computed Tomography (CT) into relative proton stopping powers (RPSP), the uncertainties of RPSP are 3–5% or higher, which need to be minimized down to 1% to make the proton treatment plans more accurate.

In this work, we simulated a proton radiography system, with position-sensitive detectors (PSDs) and a residual energy detector (RED). The simulations were built using Geant4, a Monte Carlo simulation toolkit. A phantom, consisting of several materials was placed between the PSDs of various Water Equivalent Thicknesses (WET), corresponding to an ideal detector, a gaseous detector, silicon and plastic scintillator detectors. The energy loss radiograph and the scattering angle distributions of the protons were studied for proton beam energies of 150 MeV, 190 MeV and 230 MeV. To improve the image quality deteriorated by the multiple Coulomb scattering (MCS), protons with small angles were selected. Two ways of calculating a scattering angle were considered using the proton's *direction* and *position*.

A scattering angle cut of 8.7 mrad was applied giving an optimal balance between quality and efficiency of the radiographic image. For the three proton beam energies, the number of protons used in image reconstruction with the *direction* method was half the number of protons kept using the *position* method.

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1. Introduction

The quality of cancer treatment with protons critically depends on the accurate determination of proton stopping powers (PSP) of the tissues traversed by the protons. Proton treatment planning is based on stopping powers derived from CT images leading to systematic uncertainties of 3–5% and even up to 10% in regions containing bone [1–4]. This may result in a lower dose being deposited in parts of the tumor and/or an overdose in healthy tissues.

The use of proton radiography, which gives direct access to PSPs, can reduce the uncertainty in the translation of the X-ray CT image into a map of PSPs.

Several strategies for proton radiography and its 3D extension proton Computed Tomography (pCT) are being explored by various laboratories [5–10]. Measuring the energy loss and scattering angle of an individual proton passing through the patient is the best way to obtain a sufficiently accurate radiograph of the phantom (patient). For these measurements, the initial proton beam energy should be higher than that chosen for patient treatment to ensure that the protons leave the patient's body. From the entry and exit points and angles of each proton, combined with the deposited energy, the most likely path can be calculated for each proton. This, together with already available CT data, can improve the accuracy of the PSP prediction, and thereby improve the quality of the proton treatment plan.

At KVI-CART we performed proton radiography (pRG) experiments using a novel prototype of a gaseous PSD (Timepix-based Time Projection Chamber, TPC), developed by Nikhef [8,11]. In these experiments, we used a proton beam energy of 150 MeV to

* Corresponding author.

E-mail address: a.k.biegun@rug.nl (A.K. Biegun).

irradiate a phantom. The experiments were simulated with Geant4. The comparison of both experimental and simulated data was in excellent agreement, which was a motivation for further studies [8].

The challenge in pRG imaging arises from the MCS of the protons traversing different materials causing blurring of the radiography image. Thus, to improve the image quality and identify each material in the phantom, cuts for the proton scattering angle have to be tuned carefully.

In this paper we present the analysis of two different proton scattering angle calculations, based on *direction* and *position* of a proton, and its effect on energy loss radiographs and statistics. The analysis was performed for detectors with different WET, these include: an ideal massless detector, a gaseous detector, silicon (Si) and plastic scintillator detectors. The proton energies chosen for the simulations were 150 MeV, 190 MeV and 230 MeV.

2. Material and methods

2.1. Monte Carlo simulations

The Monte Carlo (MC) simulations discussed in this paper were performed with Geant4 version 9.6.p03 [12]. A scheme of the simulation setup implemented in our code is sketched in Fig. 1.

Each of $5.0 \cdot 10^7$ generated protons passed through the two PSDs before and after the phantom, and it was stopped in the RED. The proton beam field was generated at a distance of 100 cm from the PSD1 with the size of $6 \text{ cm} \times 10 \text{ cm}$, in x and y direction, respectively (Fig. 1). No divergence of the beam was applied. Simulations were performed for the WET of each of the PSDs, thus mimicking the various detector types that have been used in pRG/pCT systems with tracking detectors. The list of studied PSD detectors with their specifications is shown in Table 1. Different PSD types, their materials and thicknesses were taken from

setups that were used in proton radiography systems described in the literature [10]. The thickness of the gaseous detector was taken based on the system that we previously used in our experiments at KVI-CART [8], the thickness of the Si detector was taken from the PRIMA I experiment [7], and the thickness of the plastic scintillator was taken from the proton radiography setup used at the Paul Scherrer Institute [13]. The RED in all simulations was filled with BaF2, similarly to what was used during our experiments at KVI-CART.

The proton beam energies used in the simulations are $E_{\text{beam}} = 150 \text{ MeV}$, 190 MeV and 230 MeV . The energy loss of a proton in the phantom, ΔE , was calculated as a difference between the known proton beam energy, E_{beam} , and the residual energy, E_{RED} , detected in RED:

$$\Delta E = E_{\text{beam}} - E_{\text{RED}} \quad (1)$$

The reference physics list used in our simulations was *QGSP_BIC_EMY*, which has been suggested by Cirrone et al. [14] to obtain a high level of accuracy. This physics list has been optimized for hadron radiotherapy simulations.

2.2. Phantom

The phantom is composed of a block of polymethyl methacrylate (PMMA), in which six cylinders of various sizes and materials, including tissue surrogates, were inserted (Fig. 2, left).

Three of the cylinders contain only one tissue surrogate material, such as: liver (In1), breast (In3) and lung (In4) and the other three inserts simulate an air cavity (In2), a lung tumour (In5) and dental fillings (In6). The diameters of the implemented cylinders are 8, 10, 12, 15, and 20 mm for In3, In2, In6, In1, and In4 and In5, respectively. Additionally, the topmost cylinder (In6) is inserted at an angle of 60 degrees with respect to the z-axis (beam direction).

Fig. 1. Proton radiography setup implemented in the Geant4 simulations. The momentum vectors of an individual proton from a scattered beam before and after the phantom ($\vec{p}_1$ and $\vec{p}_2$ respectively) are shown. The positions of a proton at PSD1 (x_1, y_1, z_1) and PSD2 (x_2, y_2, z_2) are also marked (red dots). All dimensions are in cm unless specified otherwise. (For interpretation of the references to colour in this figure legend, the reader is referred to the web version of this article.)

Table 1
PSD detector specifications used in the Geant4 MC simulations. The number of PSD detectors, material thickness and WET are shown for the PSD detector placed before and after the phantom.

PSD detector type	Number of PSD detectors	Material	Physical density (g/cm ³)	Material thickness (mm)	WET (mm)
Ideal	1	Air	0.0012	0.001	–
Gaseous (TPC)[8]	1	Isobutene C ₄ H ₁₀	0.0025	30.0	0.394
Silicon[7]	2	Silicon	2.330	0.4	0.752
Plastic scintillator[13]	2	Bicron BCF12	1.032	4.0	4.106

Fig. 2. (Left) Scheme of the phantom ($x = 5.4$ cm, $y = 9.4$ cm) with six inserts of various sizes and contents. The thickness of the phantom in the beam direction (z) is 6.0 cm. Lengths (L , z -axis) of materials in various inserts (In1, In2, ..., In6) are expressed in mm. Diameters (D) of metal implants (Al, Ti, Cu) and an air cavity (Air) in the insert with cortical bone (In6) are also shown in mm. (Right) Table with physical densities of materials that are present in the phantom and are implemented in the Geant4 simulations. Materials marked * represent Gammex tissue surrogates [15].

2.3. Calculations of proton scattering angles

In proton radiography the images are blurred because of MCS. Thus, it is important to find the best way to reduce the impact of MCS on the extracted proton energy loss radiographic image to minimize the blurring and, consequently, to improve the accuracy of the energy loss map. The scattering angle of a proton was calculated using both the *direction* and the *positions* of a proton before and after traversing the phantom, according to the following formulas:

$$\phi_{12}^{\vec{p}}(\text{rad}) = \cos^{-1} \frac{\vec{p}_1 \vec{p}_2}{|\vec{p}_1| |\vec{p}_2|} \quad (2)$$

where: $\vec{p}_1$ (p_{x1} , p_{y1} , p_{z1}) and $\vec{p}_2$ (p_{x2} , p_{y2} , p_{z2}) are the proton *directions* defined with momentum vectors at PSD1 and PSD2, respectively (Fig. 1).

$$\phi_{12}^{\text{pos}}(\text{rad}) = \tan^{-1} \frac{\sqrt{(x_2 - x_1)^2 + (y_2 - y_1)^2}}{(z_2 - z_1)} \quad (3)$$

where: x_1 , y_1 , z_1 and x_2 , y_2 , z_2 represent the proton *positions* at PSD1 and PSD2, respectively. This represents a “zero order” approximation.

3. Results and discussion

Fig. 3 shows the proton scattering angle distributions determined from the proton *direction*, $\theta_{12}^{\vec{p}}$ (Fig. 3a) and the proton *position*, θ_{12}^{pos} (Fig. 3b), respectively, according to (2) and (3). The angular distributions are shown for $E_{\text{beam}} = 150$ MeV. The PSD1 and PSD2 were filled with water and the corresponding WET is shown in Table 1. The scattering angle determined from the proton *direction* (Fig. 3a) shows that a significant number of protons travel in the forward direction, with a minimum scattering, while the distributions based on the proton *position* (Fig. 3b) is broader.

Various cuts on the scattering angle were applied in order to have a sharper image and to better distinguish the various insertions in the object, as discussed in our previous study [9]. The best trade-off between sharp transitions of various materials in the phantom and the statistics was found for scattering angles smaller than 8.7 mrad. Therefore, this scattering angle was also used in this analysis.

Fig. 4 shows the percentage of protons surviving the 8.7 mrad cut, when proton *direction* (Fig. 4a) and *position* (Fig. 4b) are used to make the cut. Using the proton *position* to calculate scattering angles θ_{12}^{pos} , more protons, by a factor of two, are accepted to build the radiographic image. Based on the results from [9] we verified

Fig. 3. Scattering angle distributions using proton direction (a) and position (b) according to formulas (2) and (3), respectively. The distributions were calculated for PSD1 and PSD2 with various WET (Table 1) with proton beam energy of 150 MeV.

Fig. 4. Accepted numbers of protons for a proton beam energy of 150 MeV, 190 MeV and 230 MeV after applying an angular cut of 8.7 mrad, using both proton direction (a) and position (b).

Fig. 5. Proton energy loss radiographs of the phantom obtained with 150 MeV protons scattering up to $\theta_{12}^{\text{pos}} < 8.7$ mrad. Both PSD1 and PSD2 are considered to be (a) ideal detectors (1 μm thickness, filled with air), (b) WET of a gaseous detector, (c) WET of a silicon detector and (d) WET of a plastic scintillator. The black dashed line in (a) at X = 1.15 cm shows the position of the projection in Fig. 6.

that the results for $E_{\text{beam}} = 150$ MeV are reproducible. In addition, the number of protons surviving the cut for higher proton beam energies of 190 MeV and 230 MeV were also determined and shown in Fig. 4.

Using the proton *position*, the number of accepted protons is nearly 50% and increases to 65% and 75% for $E_{\text{beam}} = 150$ MeV, 190 MeV and 230 MeV, respectively (Fig. 4b), while using proton *direction* (Fig. 4a), these numbers correspond to 25%, 31% and 37% for $E_{\text{beam}} = 150$ MeV, 190 MeV and 230 MeV. Thus, the statistics shows that it is better to use the proton *position* method to calculate the proton scattering angle.

Fig. 5 shows radiographic images of the phantom reconstructed based on θ_{12}^{pos} of a proton that passed through all the detectors and that scattered up to 8.7 mrad. The proton energy loss, ΔE , is calculated using (1).

Fig. 5a represents the energy loss radiograph when both PSDs were filled with air (ideal detectors), while Fig. 5b, c and d represent PSDs filled with WET of TPC, Si and plastic scintillators (Table 1). Fig. 5a also indicates with the position of the dashed line, located at $X = 1.15$ cm, for which the projections in Fig. 6a are plotted. The reconstructed energy loss radiographs for PSDs filled with air and WET of TPC are very comparable, Fig. 5a and b, with all inserts visible. When PSDs are filled with Si (Fig. 5c), a part of the radiographic image is blurred for inserts with cortical bone, liver and breast (see Fig. 2 for the reference), while these three inserts are hardly recognizable when PSDs are filled with plastic scintillator; the radiographic image in Fig. 5d is blurred the most.

Fig. 6a demonstrates the differences in the proton energy loss profiles through selected material of the phantom for various WET of PSDs (Table 1) and the transitions between materials.

ΔE profiles of the radiographic image were calculated in correspondence of the dashed line in Fig. 5a and were obtained with 150 MeV protons scattering up to 8.7 mrad (θ_{12}^{pos}). It can be

observed that the transition between various phantom materials is least steep for the WET of plastic scintillators. This is expected, because the WET of plastic scintillators is the largest (Table 1). For both the WET of TPC and Si, sharp transitions between materials are visible (Fig. 6a). The thickness of gas is significantly larger than the one of the Si and only 0.4 mm of Si has been considered in the simulations for each PSD. Furthermore, no visible difference in sharpness between PSDs filled with air (ideal detector) and the WET of TPC is observed, which is expected because the WET of TPC is very small (<0.4 mm). For higher proton beam energies, 190 MeV and 230 MeV, the differences in ΔE profiles between various PSD materials are smaller, as expected due to decreasing MCS of protons with increasing proton beam energy.

Fig. 6b presents the energy loss ratio (ΔE ratio) between the energy loss measured with TPC, Si, plastic scintillator and air, for 150 MeV protons. The ΔE ratio for TPC and Si is comparable and the fluctuation is between 0.95 and 1.1 (15%), with the average value of 1.02 (determined by visual inspection), whereas the fluctuation for plastic scintillator is larger, between 0.97 and 1.31 (34%), with an average value of 1.12.

Similar ΔE profiles and ΔE ratios were obtained for the same cut of scattered protons of 8.7 mrad, where the scattering angle θ_{12}^{pos} was calculated using the proton *direction* (formula 2). The fluctuation of the profiles and ratios were observed to be larger than in Fig. 6a and b, with the average value of 1.03 for protons with energy of 150 MeV, and can be seen in Table 2. Furthermore, both ΔE profiles and ΔE ratios were obtained for higher proton beam energies of 190 MeV and 230 MeV (not shown) and similar trend was observed. The average values of ΔE ratios are smaller when scattering angle is calculated using the proton *position* than the proton *direction* (1.01 for θ_{12}^{pos} vs. 1.03 for θ_{12}^{dir} , for proton beam energies of 190 MeV and 230 MeV, respectively). All the average values of ΔE ratio fluctuations are shown in Table 2.

Fig. 6. (a) ΔE profiles through selected material of the phantom obtained with 150 MeV protons scattering with an angle θ_{12}^{pos} up to 8.7 mrad for different WET (Table 1). (b) ΔE ratio for 150 MeV protons and θ_{12}^{pos} up to 8.7 mrad for different WET.

Table 2

Average values of ΔE ratio fluctuations obtained for a proton beam energy of 150 MeV, 190 MeV and 230 MeV after applying an angular cut of 8.7 mrad, using both proton *direction* (θ_{12}^{dir}) and *position* (θ_{12}^{pos}).

ΔE ratio (average)	θ_{12}^{dir}			θ_{12}^{pos}		
	150 MeV	190 MeV	230 MeV	150 MeV	190 MeV	230 MeV
Gaseous (TPC)	1.03	1.03	1.03	1.02	1.01	1.01
Silicon	1.03	1.03	1.03	1.02	1.01	1.01
Plastic scintillator	1.12	1.07	1.04	1.12	1.06	1.04

4. Conclusions

In this paper we present the results of a study on proton radiography, an imaging technique aiming at a direct (and therefore more accurate) calculation of the PSP of an object under study (patient).

The MCS of protons passing through various materials blurs the energy loss radiographic image. Selecting protons with small scattering angles sharpens the edges between materials in the object, decreasing the blurring. Two ways of calculating the proton scattering angle have been studied, where either the proton *direction* ($\theta_{12}^{\vec{p}}$) or the proton *position* (θ_{12}^{pos}) was used to calculate the scattering angular distribution. When applying cuts in the radiographic image reconstruction based on θ_{12}^{pos} , the number of “accepted” protons was doubled compared to when $\theta_{12}^{\vec{p}}$ was used for all three energies, specifically: 51% vs. 25% with $E_{beam} = 150$ MeV, 65% vs. 30% with $E_{beam} = 190$ MeV, and 75% vs. 37% with $E_{beam} = 230$ MeV.

Plastic scintillators, such as PSD1 and PSD2, seem to be the most disadvantageous in the pRG/pCT imaging system, unless the highest proton beam energy of 230 MeV is used. For a more realistic representation of Si detectors, additional materials should be added to the simulations consisting of, for example, a PCB material to which the Si detector is connected. Protons from the beam pass through the PCB material undergoing MCS and causing further blurring of the radiographic image. Thus, it will make the results for PSD_{Si} less accurate than for PSD_{TPC}. Furthermore, the thickness of a single Si detector is usually taken to be at least 300 μm . As PSDs of at least two (or three in some pRG/pCT prototypes) Si detectors are placed before and after the object under study, this will increase the image blurring further. The MCS is element dependent and grows with the atomic number of the material. The most promising PSDs in a pRG/pCT system are expected to be gaseous detectors. The MCS in low-density gas is the smallest amongst the discussed materials giving the least expected blurring of the radiographic image. The readout chip in gaseous PSDs can be located at the surface perpendicular to the beam direction, such that protons will not pass through the chip material, but mostly

pass through the gas [8]. More simulations and experiments are required to study the MCS on complex phantoms and real tissues.

Acknowledgments

We would like to thank the Center for Information Technology of the University of Groningen for their support and for providing access to the Peregrine high-performance computing cluster, where the Monte Carlo simulations were performed. This work was partially supported by the Japan Society for Promotion of Science Core-to-Core Program (number 23003).

References

- [1] Schneider U, Pedroni E, Lomax A. The calibration of CT Hounsfield units for radiotherapy treatment planning. *Phys Med Biol* 1996;41:111–24.
- [2] Paganetti H. Range uncertainties in proton therapy and the role of Monte Carlo simulations. *Phys Med Biol* 2012;57:R99–R117.
- [3] Plautz T et al. 200 MeV proton radiography studies with a hand phantom using a prototype proton CT scanner. *IEEE Trans Med Imag* 2014;33(4):875–81.
- [4] Schuemann J et al. Site-specific range uncertainties caused by dose calculation algorithms for proton therapy. *Phys Med Biol* 2014;59:4007–31.
- [5] Schneider U, Pedroni E. Multiple Coulomb scattering and spatial resolution in proton radiography. *Med Phys* 1994;21:1657–63.
- [6] Ryu H et al. Density and spatial resolutions of proton radiography using a range modulation technique. *Phys Med Biol* 2008;53:5461–8.
- [7] Sipala V et al. A proton Computed Tomography system for medical applications. *JINST* 2013;8:C02021.
- [8] Biegun AK et al. Proton radiography with timepix based time projection chambers. *IEEE Trans Med Imag* 2016;35(4):1099–105.
- [9] Biegun AK et al. Proton energy and scattering angle radiographs to improve proton treatment planning: a Monte Carlo study. *JINST* 2016;11:C12015.
- [10] Poludniowski G, Allinson NM, Evans PM. Proton radiography and tomography with application to proton therapy. *Br J Radiol* 2015;88:20150134 [and references therein].
- [11] <https://www.nikhef.nl>.
- [12] Agostinelli S et al. Geant4—a simulation toolkit. *Nucl Instr Meth in Phys Res* 2003;506:250–303.
- [13] Schneider U et al. First proton radiography of an animal patient. *Med Phys* 2004;31(5):1046–51.
- [14] Cirrone GAP et al. Hadrontherapy: a Geant4-based tool for proton/ion-therapy studies. *Prog Nucl Sci Technol* 2011;2:207–12.
- [15] <http://www.gammex.com/>.