

Investigation of Temperature Dependence of Polymer Gels for Use with Scanning Magnetic Resonance Imaging

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Abstract

Polymer gels are three-dimensional dosimetric tools. The purpose of the present study was to investigate the temperature dependence of polymer gels during scanning Magnetic Resonance Imaging. Prepared gels were irradiated with a 6MV X-ray beam at intensities ranging from 0 to 20 Gy in order to investigate their dose- R_2 and dose- R_1 responses. Irradiated gels were evaluated from 1.5-T magnetic resonance R_2 and R_1 images for each 5°C change in temperature from 5°C to 41°C, and then the four-field box technique irradiation plan was used to deliver a total dose of 4 Gy using the same beam weight in each direction to the prepared gels. The profile of the dose map generated from the four-field irradiated gel data at 20°C was then compared with the planned data. The dose- R_2 response curve was linear up to 20 Gy at 20°C, with a slope of 1.17 Gy⁻¹·s⁻¹. The slopes of the fitted curves of the dose- R_2 decreased as gel temperature increased. The slopes of the dose- R_1 curves were more parallel than the slopes of the dose- R_2 curves between 5 and 41°C. The difference in the full width of half maximum of the gel profile data obtained using the four-field box technique at 20°C and the planned data were below 5% on average. The dose map from the irradiated gels obtained using the dose- R_2 curve was the same as that from the planned data under the same temperature conditions. Measurement of difference between various temperatures is significant with dose accuracy. It is suitable to evaluate the gel dosimeter under the thermal equilibrium condition, MRI room temperature from the point of view of the stability of the irradiated gels.

Keywords

Polymer Gel Dosimetry, Temperature Dependence, MRI Scanning

1. Introduction

Polymer gels provide new three-dimensional (3D) dosimetric tools that hold promise for the 3D measurement of 3D doses during clinical radiotherapy, thus enhancing quality assurance. Radiotherapy is complex and requires precise monitoring [1]-[6]. Measurements using an ion chambers are precise, but an ion chamber is a point-detector, and thus is not suited to 3D dosimetry.

The clinical use of current polymer gel dosimeters faces several problems, including the temperature stability of polymer gels under irradiation and during dosimetric evaluation using Magnetic Resonance Imaging (MRI). Polymer gel dosimetry evaluated using MRI is conducted in low temperature environments because the gels melt at temperatures over approximately 25°C, providing poor spatial information and inaccurate dosimetric results. Precise measurements of polymer gels in clinical settings are thus needed in order to determine the temperature dependence of polymer gels.

MRI is commonly used to evaluate the dose received by irradiated polymer gels. The spin-spin relaxation rate ($R_2 = 1/T_2$, s^{-1}) and spin-lattice relaxation rate ($R_1 = 1/T_1$, s^{-1}) provide the degree of polymerization of the irradiated gels and the radiation dose. This study investigated the dependence of R_2 on the dose received by polymer gels exhibiting a higher dose response than R_1 . The fundamental properties of the temperature dependence of R_1 of polymer gels remain unknown. This study was designed to investigate the fundamental temperature effects on R_1 and R_2 by investigating the differences in temperature properties between the dose and R_1 , and the dose and the R_2 calibration curve.

Several clinical irradiation studies using polymer gels have been conducted to date [6]-[13], but fundamental investigations of the temperature dependence of these gels are needed prior to the clinical application of gel dosimetry. To determine the feasibility of polymer gel dosimetry in clinical radiotherapy, we attempted to investigate the temperature-dependent properties of the polymer gels under simulated clinical irradiation conditions using a Radiation Treatment Planning System (RTPS).

2. Materials and Methods

2.1. Gel Preparation

BANG-3-type (Bis, Acrylamide, Nitrogen and Gelatin) polymer gels (BANG3PRO; MGS Research, Inc., Guilford, CT), and PAGAT (Polyacrylamide Gel and THPC) gels [14] were prepared. The BANG-3 type polymer gels were prepared using a BANG kit. The unmodified gel melted at 55°C; thus, several additives were used [15]. PAGAT gels were prepared using 89% w/w water, 3% acryla-

mide, 3% N,N'-methylenebisacrylamide (Bis), 5% gelatin (300 bloom) and 5 mM tetrakis (hydroxymethyl) phosphonium chloride (THPC). Gelatin was added to water, followed by heating at 50°C on a hot plate/magnetic stirrer. After the solution became clear, it was cooled to 45°C and Bis was added. After the Bis dissolved completely, THPC was added.

Prepared gels were poured into polyethylene-terephthalate (PET) vials and containers. Vials were 45 × 30 × 30 mm³ and were used for dose-R₂, R₁ calibration. Containers were 177 × 74 × 74 mm³ and were used to measure clinical irradiations by 3D dose distribution. Gels in the PET vials and containers were stored wrapped in aluminum foil in a refrigerator at 4°C until irradiation, as the gels melt at high temperature.

2.2. Design of Gel Phantom for Specific Treatment Plan and Simulated Clinical Irradiation Studies

Figure 1 shows a gel phantom using PAGAT gel for clinical irradiation. The phantom was 200 × 140 × 125 mm³, made of Styrofoam, and housed the gel container. The phantom was filled with water and then scanned in A-helical scan mode using an X-ray CT device (Aquilion LB; Toshiba Medical Systems, Tochigi, Japan) during the treatment plan. CT exposure conditions were as follows: tube voltage, 120 kV; tube current, 400 mA; exposure time, 0.5 s; slice thickness, 3.0 mm; and imaging field of view, 550 × 500 mm². The number of slices per CT image was 70.

Treatment plans were prepared using the RTPS system (Pinnacle³; Philips

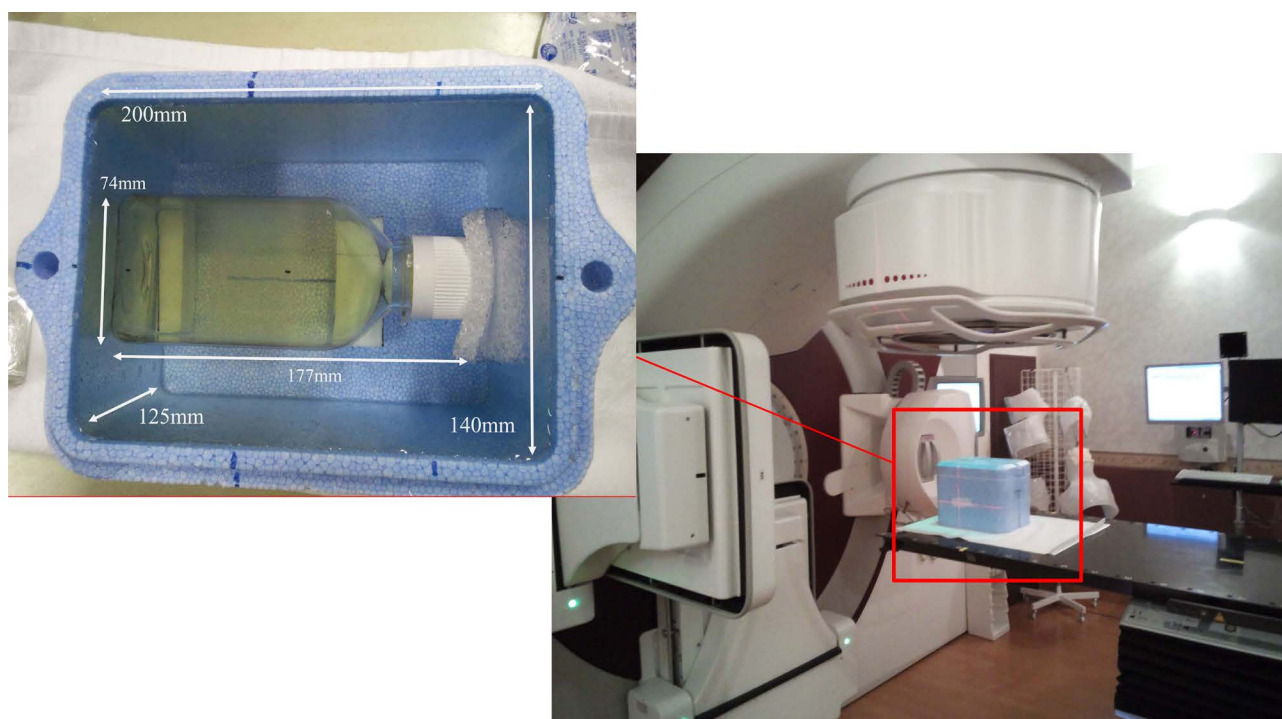


Figure 1. The photograph of gel phantom of the size 200 × 140 × 125 mm³ is shown. The PAGAT gel container was set on the gel phantom in the center position. The size of the PAGAT gel container is 177 × 74 × 74 mm³ (including neck).

Healthcare, Andover, MA) and used a four-field box irradiation technique (**Figure 2**). The four-field box technique (crossfire) irradiation plan prescribed a total dose of 4 Gy and the same beam weight in each direction. The four beam irradiation calculations assumed the treatment of esophageal cancer and determined the dosimetric results of the polymer gel.

2.3. Irradiation Using Photon Beams for Dose Calibration and Clinical Irradiation

The prepared BANG-3-type gels and PAGAT gels were irradiated with a 6 MV X-ray beam on a linear accelerator (ELEKTA Synergy; ELEKTA, Stockholm, Sweden) at Tsukuba Medical Center Hospital.

Plan dose map (left, blue) was exported from RTPS and dose map from R_2 image (right, red) was calculated from in-house program. This figure was shown that crossfire 4 beams irradiation calculations supposed that treatment of a esophageal cancer and dosimetric result of polymer gel using 6 MV photon beam. The profiles between plan dose and dose map from R_2 were compared.

First, the polymer gels in PET vials were irradiated with no collimator at the isocenter of a $300 \times 300 \times 300 \text{ mm}^3$ water tank to calibrate the dose versus R_2 and R_1 from 0 to 20 Gy in the beam axis. After irradiation, gels were stored in a

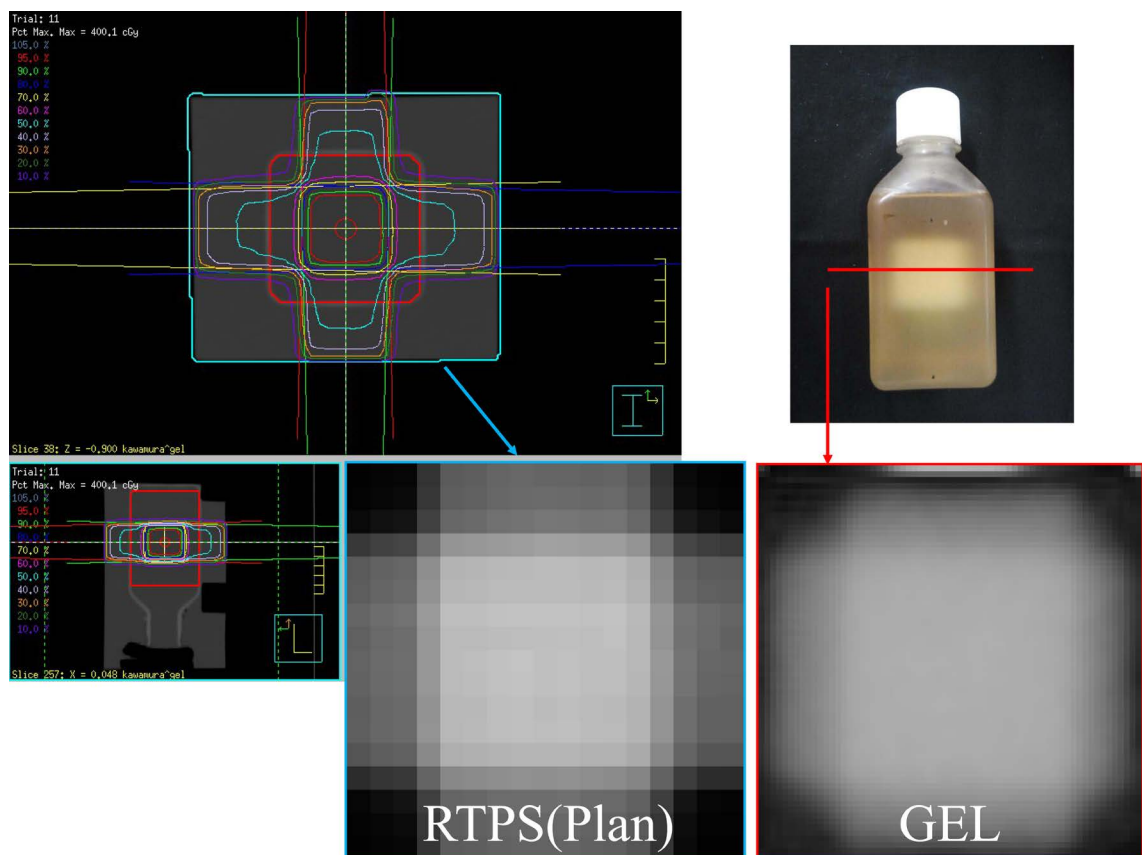


Figure 2. Treatment plan image was shown that four-field box technique. The sky blue box of the upper image was surrounded a gel phantom and the red box of all images was surrounded a gel container within fabricated polymer gels. Treatment plan was made by acquired X-ray CT images.

refrigerator at 4°C until MRI scanning.

Second, gel phantoms were irradiated with 6 MV X-ray beams at the isocenter in order to simulate clinical irradiation. The average dose rate was 300 MU/min (**Figure 1**).

2.4. R_1 and R_2 Measurements Using MRI

MRI measurements for dose evaluation of the gels were performed on a 1.5-T Siemens AVANTO 1 day after irradiation. Irradiated gels were positioned in a quadrature (QD) coil for scanning.

For R_2 measurements, gels were imaged using a multi-echo fast spin echo pulse sequence. Ten echoes of the sequence were used (echo time: TE = 15 to 150 ms; echo time interval, 15 ms). For each scan, a repetition time (TR) of 1000 ms was used, with a 1 mm² resolution (field of view (FOV) = 192 mm, matrix number = 192 × 192) using 5-mm-thick planes. The obtained data were used to calculate T_2 images using the MapIt program (Siemens) [16].

For R_1 measurements, gels were imaged using a 3D volumetric interpolated breath-hold examination (3D-VIBE) sequence (TE = 1.62 ms, TR = 15 ms). For each scan, a resolution of 0.5 mm² was used (FOV = 192 mm, matrix number = 384 × 384) using 5-mm-thick planes, the same as for R_2 . R_1 and R_2 images were created to invert these pixel values using the original program.

The Styrofoam container described earlier was set in the QD coil and filled with water at 5.0°C. Irradiated gel samples in vials and in containers were individually set in the center of the Styrofoam container and the water temperature in the container was raised from 5.0°C to 41.0°C at 5.0°C /min.

2.5. R_2 , R_1 versus Dose Linearity of the Polymer Gel Calibration Curve

The obtained data were used to calculate R_2 images using the MapIt program. Two-dimensional (2D) R_1 and R_2 images were constructed from the T_1 and T_2 images using an in-house program. Data points for the dose R_2 , R_1 characteristic curve were obtained by averaging the R_2 values from the region of interest (ROI) in the polymer gel.

2.6. Comparison between the RTPS Data and the R_2 and R_1 Images

The acquired R_2 images were converted into dose images using calibration data from the dose- R_2 curve. Dose images from the R_2 images of the irradiated gels in the center profile were compared with the calculated plan data from the RTPS with four-box irradiation fields from the simulated clinical situation (**Figure 2**).

3. Results

3.1. Temperature Dependency of R_2 versus Dose Linearity of the Polymer Gel Calibration Curve

Figure 3 shows the R_2 values of the BANG polymer gels as a function of photon

dose between 0 and 20 Gy. Each R_2 calibration curve was fit to a straight line for doses below 20 Gy. The fitted straight line for photon dosimetry at 20°C had a gradient of $1.17 \text{ Gy}^{-1}\cdot\text{s}^{-1}$ and an intercept of 3.87 s^{-1} , and the coefficient of correlation was 0.998.

Figure 4 shows the R_2 values of the BANG polymer gels as a function of dose between 5.0°C and 40°C. The fitted straight line of the data collected at 0 Gy provided a gradient of $-0.19^\circ\text{C}^{-1}\cdot\text{s}^{-1}$ and an intercept of 8.10 s^{-1} , and fitting of the data collected at 20 Gy provided a gradient of $-0.96^\circ\text{C}^{-1}\cdot\text{s}^{-1}$ and an intercept of 47.45 s^{-1} . The coefficient of correlation of the 0 Gy data was 0.894 and that of the 20 Gy data was 0.991.

3.2. Temperature Dependency of R_1 versus Dose Linearity of the Polymer Gel Calibration Curve

Figure 5 shows the R_1 values of the BANG polymer gels as a function of photon doses between 0 and 20 Gy. Each R_1 calibration curve was fit to a straight line for

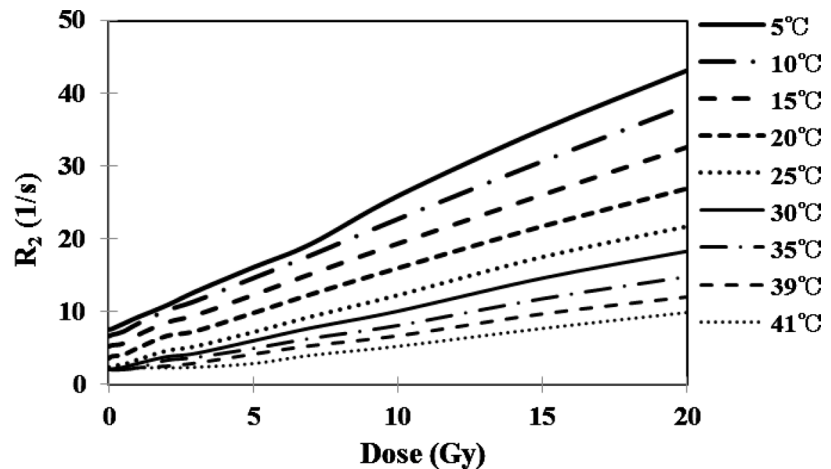


Figure 3. Dose- R_2 responses using different temperature conditions. Error bars are omitted because of overlapping bars and plots.

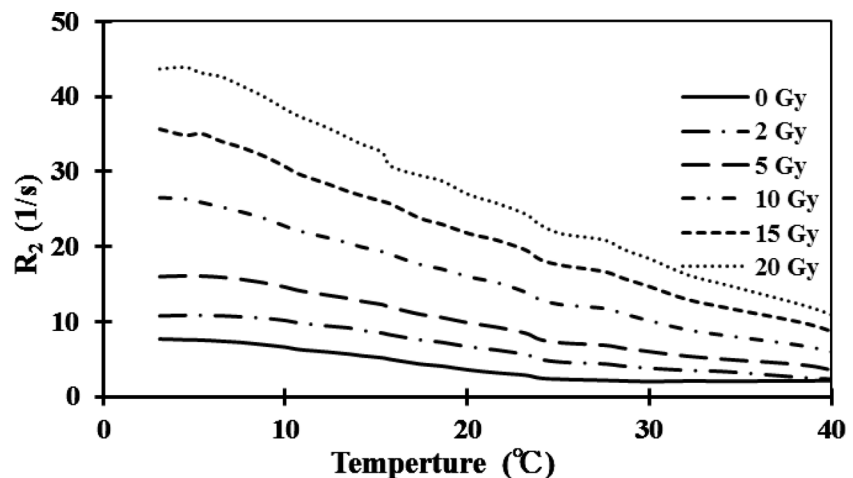


Figure 4. R_2 -temperature responses of the BANG-type gel dosimeter using different photon dose conditions. Error bars are omitted because of overlapping bars and plots.

doses below 20 Gy. Data points were obtained by averaging the R_1 values in the polymer gel. The fitted straight line for data collected at 20°C had a gradient of $0.034 \text{ Gy}^{-1}\cdot\text{s}^{-1}$ and an intercept of 0.72 s^{-1} . The coefficient of correlation was 0.962.

Figure 6 shows the R_1 values of the BANG polymer gels as a function of dose between 5.0°C and 40°C. The fitted straight line of the data collected at 0 Gy had a gradient of $-0.020^\circ\text{C}^{-1}\cdot\text{s}^{-1}$ and an intercept of 1.12 s^{-1} , whereas the data collected at 20 Gy had a gradient of $-0.029^\circ\text{C}^{-1}\cdot\text{s}^{-1}$ and an intercept of 1.94 s^{-1} . The coefficient of correlation of the 0 Gy data was 0.925 and that of the 20 Gy data was 0.985.

The dose uncertainties (%) of R_1 and R_2 between 5°C and 39°C are shown in **Table 1**. An increase in temperature results in large dose uncertainties of R_1 and R_2 . The temperature-related errors in dose are 8.3% for R_1 and 7.5% for R_2 . The dose-related error of the dose response results of the gel is below 8% at temperatures below 20°C.

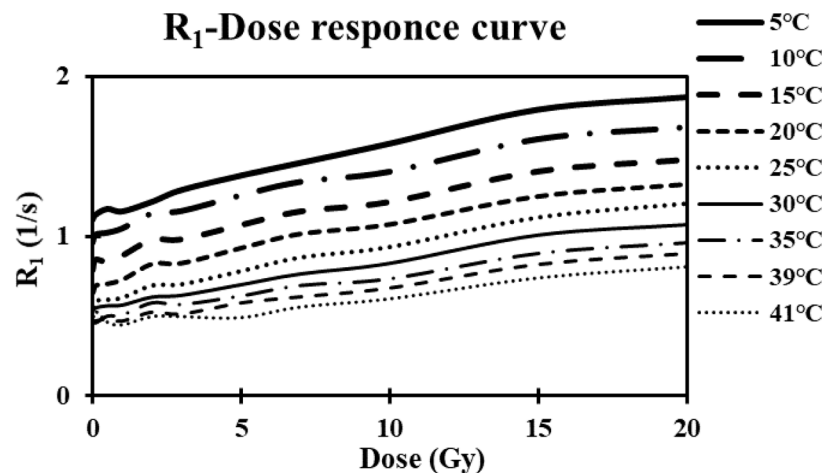


Figure 5. Dose- R_1 response curves using different temperature conditions. Error bars are omitted because of overlapping bars and plots.

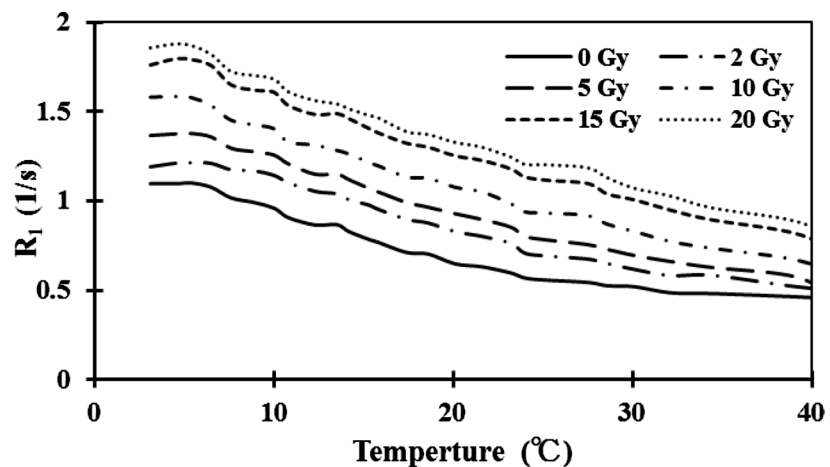


Figure 6. R_1 -temperature responses using different photon dose conditions. Error bars are omitted because of overlapping bars and plots.

Table 1. Dose uncertainties (%) of R_1 and R_2 between 5°C and 39°C from the dose response results.

	TEMP (°C)	5	10	15	20	25	30	39
R_1		4.7	5.6	6.2	7.5	8.6	10.5	15.0
R_2		6.1	6.1	6.1	7.2	8.5	8.2	10.2

Dose uncertainties (%).

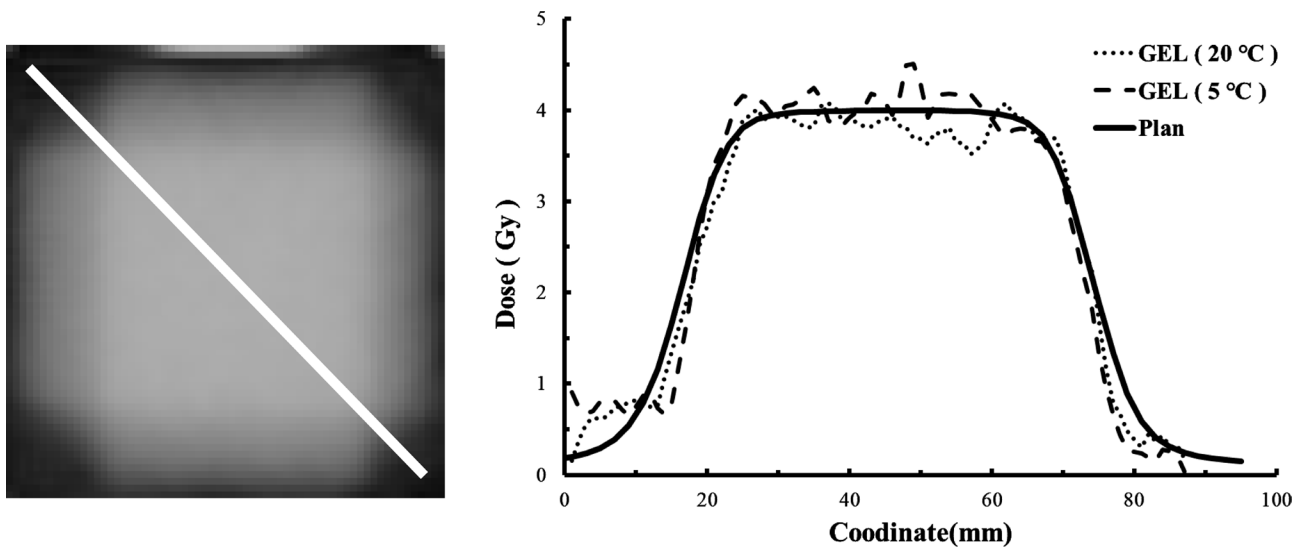


Figure 7. The dose profile (right) of the crossfire 4 beam-irradiated gels and the planned dose data. The profile of the gels is similar to that of the planned data.

3.3. Dose Profile Comparison between Dose Images Obtained Using R_2 Images of Polymer Gel and Dose Images Obtained Using RTPS

Figure 7 shows the dose profile for PAGAT polymer gel on a planned RTPS of the oblique center line of the four-field box technique and planned data for comparison. The full width at half maximum (FWHM) of the GEL (20°C), GEL (5°C), and Plan (RTPS) curves were 57.48 mm, 55.22 mm and 58.48 mm, respectively. The dose-related error of the results obtained using the gel at irradiation doses over 2 Gy at 20°C is 4.87% on average and the error at 5°C is 5.18% on average.

4. Discussion

4.1. Temperature Dependency of R_2 , R_1 versus Dose Linearity of the Polymer Gel Calibration Curve

The results confirmed dose R_1 and dose R_2 linearity between 0 to 20 Gy. The dose- R_2 gradient was steeper at low temperature, as reported previously [10] [11]. Dose- R_1 curves with respect to temperature have not previously been reported and were found to parallel the dose- R_2 gradient.

Dose gradient with temperature is less pronounced in the dose R_1 curves than in the dose R_2 curves and therefore the temperature dependence has less effect

on dose R_1 linearity than on dose R_2 linearity. In contrast, the gradient of the R_1 dose linearity was 40 times smaller than that of the R_2 dose linearity. Furthermore, the overall standard deviation (SD) and coefficient of variation (calculated by dividing the R_2 or R_1 value by SD) of the R_1 images are larger than that of the R_2 images. The dose images obtained from the R_1 images included more noise when compared with the dose images obtained from the R_2 images. Therefore, the dose images obtained from the R_1 images were less precise compared with the dose images obtained from the R_2 images.

R_2 and R_1 are dependent on both the correlation time (τ_c) and the Larmor frequency (ω) from the Bloembergen-Purcell-Pound (BPP) relaxation theory [17] [18] [19]. τ_c of the gels shows an inverse correlation with their temperature: raising the temperature correspondingly decreases the τ_c of the gels. T_1 and T_2 are dependent on an inverse relationship with viscosity [17]. Here, I assumed that an increase in temperature decreased the viscosity of the gels. Therefore, the gel R_1 and R_2 values decreased as temperature increased, resulting in a larger τ_c and the gradients (Figure 4 and Figure 5) changed gradually with temperature. The change in the gradients of the dose R_2 curves was assumed to depend on both the viscosity of the gels arising from the degree of polymerization and the temperature.

4.2. Dose Profile Comparison between Dose Images Obtained Using R_2 and Dose Images Obtained Using RTPS

The dose profile is presented in Figure 7 and shows differences in the temperature dependence and a comparison of the dose profile obtained by gel dosimetry and the RTPS plan. The dose images obtained from the R_2 polymer gel data are precise regarding the process of conversion from the R_2 images to the dose image using the same temperature dose R_2 curve. Because the prepared gels melt at high temperature, it was difficult to maintain the inside of the gels at a constant temperature as required during scanning MRI. Although BANG gels have a high R_2 gradient, with changing temperature, they are less stable than PAGAT gels for the experiment to irradiated gels using RTPS. Methacrylic acid-based gel dosimeters such as BANG gel have the disadvantage of temperature dependency, in contrast to acrylic acid-based gel dosimeters such as PAGAT. BANG gels melt above 25°C, while PAGAT gels melt at about 30°C. At high and low temperatures, dose images were less precise and disagreed with the RTPS plan dose data. In this study, the dose-related error at irradiation doses over 2 Gy at 20°C was less than the error at 5°C.

The difference in the FWHM of the dose profiles between the gels at 20°C and the RTPS plan dose data was 0.51 mm, whereas at 5°C, the difference was 1.8 mm, despite the dose error of the dose response at low temperature being smaller than at high temperature, as shown in Table 1. The inhomogeneity related to the temperature of the irradiated gels in the containers may have caused the high dose error at low temperature. In addition, it is difficult to maintain a constant low temperature inside the gel phantom during MRI scanning and thus the

temperature inhomogeneity inside the large phantom rose during exposure to the scanning RF pulse [20]. The temperature inhomogeneity resulting from RF exposure in a high field MRI scanner is large. Accordingly, dose evaluation using high field MRI scanning gives rise to data with large uncertainties. Dosimetric evaluation at MRI room temperature is required due to the stability of the scans during scanning MRI. Thus, it is necessary to maintain a constant at temperature inside the gels for precise measurements.

Future work will be aimed at clinical applications and will assess the temperature-related dose error using γ analysis, dose differences, and distance-to-agreement measurements.

5. Conclusions

This study revealed the temperature dependency of polymer gel dosimeters during scanning MRI. Dose- R_1 linearity and dose- R_2 linearity were shown between 0 to 20 Gy. These results indicate that temperature dependency has a greater effect on the gradient of the dose- R_2 curves than the gradient of the dose- R_1 curves.

Although the gradients of dose- R_1 curves are more constant than the gradients of dose- R_2 curves, the R_1 dose images have more noises because of smaller dose gradients.

Comparison of the FWHM in the dose profile of dose images obtained using polymer gels at 20°C and using the RTPS plan data showed the difference to be below 5%.

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