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1 **Improved image quality for x-ray CT imaging of gel**
2 **dosimeters**

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23

24 **Abstract**

25 **Purpose:** This study provides a simple method for improving precision of x-ray computed
26 tomography (CT) scans of irradiated polymer gel dosimetry. The noise affecting CT scans of
27 irradiated gels has been an impediment to the use of clinical CT scanners for gel dosimetry
28 studies.

29 **Method:** In this study, it is shown that multiple scans of a single PAGAT gel dosimeter can
30 be used to extrapolate a 'zero-scan' image which displays a similar level of precision to an
31 image obtained by averaging multiple CT images, without the compromised dose
32 measurement resulting from the exposure of the gel to radiation from the CT scanner.

33 **Results:** When extrapolating the zero-scan image, it is shown that exponential and simple
34 linear fits to the relationship between Hounsfield unit and scan number, for each pixel in the
35 image, provides an accurate indication of gel density.

36 **Conclusions:** It is expected that this work will be utilised in the analysis of three-dimensional
37 gel volumes irradiated using complex radiotherapy treatments.

38 Key words: Gel dosimeter, gel dosimetry, CT imaging, SNR, Zero-scan image, radiotherapy,
39 polymer gel

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50 I. INTRODUCTION

51

52 Gel dosimeters, consisting of a radiation sensitive material infused in a 3D gel matrix, are
53 increasingly being investigated for radiotherapy dose verification and quality assurance.¹

54 When a volume of gel is irradiated, the radiation sensitive material undergoes a measurable
55 change in the magnetic relaxation, density and optical density which is directly related to the
56 radiation dose received, potentially providing a high-resolution three-dimensional
57 measurement of the dose absorbed by the gel.²⁻²¹

58 An important consideration for any dosimeter prior to clinical use is the spatial resolution and
59 the accuracy in the measurement of the absorbed dose and many authors have investigated
60 gel dosimetry as a solution.²⁻⁷

61 One of the challenges in gel dosimetry is the extraction of the dose information once the gel
62 has been irradiated. Various techniques have been employed for gel dose readout including
63 magnetic resonance imaging (MRI)¹¹⁻¹³, optical CT scanning (OCT)¹⁵⁻¹⁶; X-ray computed
64 tomography (CT)²⁰⁻²¹, and ultrasound¹⁶. In MRI imaging of polymer gel dosimeters the spin-
65 spin relaxation rate (R_2) is used to determine the radiation induced polymerization
66 corresponding to the absorbed dose.¹¹⁻¹³ One issue with MRI imaging is that artefacts are
67 significant issues affecting the accuracy of the gel dosimeters, which requires careful
68 selection of scanning parameters to ensure accuracy.¹⁴

69 OCT has been demonstrated as viable readout technique for polymer gel dosimeters due to a
70 post-irradiation change in optical density.¹⁵⁻¹⁶; however this technique is susceptible to
71 artefacts due to refraction of light¹⁸⁻¹⁹. Ultrasound imaging¹⁶ utilizes changes in acoustic
72 speed of propagation, absorption and attenuation which vary with radiation induced
73 polymerization.

74 CT has been employed to exploit post irradiation changes in linear attenuation coefficient in
75 polymer gel dosimeters.²⁰⁻²⁶ The availability of CT scanners in radiotherapy centres makes
76 this imaging technique attractive as a routine technique for imaging of gel dosimeters.
77 However, the small changes in gel density arising from radiation exposure means that this
78 technique suffers from a low signal to noise ratio (SNR). Attempts to reduce stochastic noise
79 by averaging several CT images result in an additional dosing of the gel.^{20,21} An alternative
80 approach for the reduction of noise in CT imaging of polymer gel dosimeters has been the
81 application of image processing techniques.^{24,27,28}

82 The aim of the current work is to investigate the feasibility of a simple image analysis
83 technique whereby data from multiple scans is used to provide a hypothetical ‘zero-scan’
84 image representing the irradiated gel prior to CT scanning. A simple example of a normoxic
85 polymer gel irradiated to a range of doses is used to establish that this method is capable of
86 appreciably improving CT image quality.

87 **II. METHODS AND MATERIALS**

88 **II.A. Gel Preparation and Irradiation**

89 A PAGAT gel dosimeter was prepared as described by Venning *et al.*²⁹ with 8 mM of Tetrakis
90 (Hydroxymethyl) Phosphonium Chloride (THPC) for improved stability.³⁰ The gel dosimeter
91 was prepared under normal atmospheric conditions and poured into a cylindrical Polyethylene
92 terephthalate (PET) container of 10 cm height and 5 cm radius. It was then stored at 4°C for 24
93 hrs before irradiation. The gel dosimeter was irradiated with three small (1.5 cm x 1.5 cm)
94 fields of 118, 233 and 384 cGy parallel to the central axis of the container with a Varian
95 linear accelerator using a 6 MV photon beam at 600 MU/min. A further 686 cGy was
96 delivered using a fourth test field, close to the centre of the container. The area of this high-
97 dose field was reduced to 1.0 cm x 1.0 cm to minimise possible scatter into the other test
98 regions of the gel.

99 **II.B. x-ray CT imaging**

100 One day after irradiation, the gel was imaged using a GE Lightspeed RT 4 CT scanner. The
101 CT scans used an x-ray tube load of 300 mA with 1s rotation, beam energy of 120 kVp, 5
102 mm slice thickness, image size of 512x512 and 25 cm field of view. The CT dose from this
103 protocol is estimated as 83.4 mGy per scan at the scan centre, based on the ImPACT
104 CTDI_{100(soft tissue)} (ImPACT CT Patient Dosimetry Calculator V1.0, ImPACT, London, UK).
105 The gel was scanned 360 times at a single slice location while placed in a cylindrical water
106 tank similar to that described by Trapp *et al.*²¹ The gel was positioned in the tank such that
107 the CT scanning plane was orthogonal to the radiation beam direction. All scans consisted of
108 one slice only, with no couch motion, providing a transverse image of the phantom showing
109 all four irradiated regions at a depth of 4 cm from the surface onto which the radiation was
110 incident. An additional set of 60 scans of the tank were obtained, with the gel removed such
111 that the tank only contained water.

112 **II.C. Image Analysis**

113 The CT images, in DICOM format, were imported into Matlab (version 7.8.0.342, The
114 MathWorks, Natick, MA, USA). The 60 images of the water phantom were averaged to
115 reduce random noise.²¹ This averaged water image was then subtracted from each of the gel
116 images, to remove CT artefacts from the gel images, as described by Trapp *et al.*²¹

117 Three composite data sets were then obtained from the processed CT images, using the
118 following method. For each pixel, the Hounsfield unit (HU) was acquired sequentially from
119 the series of 360 processed CT images to form a data set of 360 data points for each gel
120 voxel. For each dataset linear, quadratic (a least squares fit with the inbuilt *Polyfit* Matlab
121 function) and exponential fits (using the *ezfit* function from the *ezyfit* curve fitting toolbox
122 based on Matlab's built-in *FMINSEARCH* function (Nelder-Mead method)) were applied to

123 the data. Thus, estimates of the relationship between scan number and HU for each voxel
124 were obtained using Eq. (1-3):

$$125 \quad HU(i, j) = L_1(i, j)N + L_0(i, j) \quad (1)$$

$$126 \quad HU(i, j) = Q_2(i, j)N^2 + Q_1(i, j)N + Q_0(i, j) \quad (2)$$

$$127 \quad HU(i, j) = E_0(i, j) + (A(i, j) * e^{-N/B(i, j)}) \quad (3)$$

128 where N is the scan number and the arrays $L_n(i, j)$, $Q_n(i, j)$ and $E_n(i, j)$, $B_n(i, j)$ and $A_n(i, j)$ are
129 free parameters for the linear, quadratic and exponential fits respectively, evaluated for each
130 pixel (i,j). A new image was then constructed, referred to here as the zero-scan image,
131 whereby each pixel in the new image is the intercept of the fit for the corresponding pixel in
132 the processed CT images, i.e. the zero-scan images are maps of $L_0(i, j)$, $Q_0(i, j)$, $E_0(i, j)$ and
133 $A(i, j)$. The rationale behind this technique is that the intercept of the fitted data most closely
134 matches the properties of the gel dosimeter immediately before CT scanning commences.

135 In each of these three new images, four regions of interest (ROIs) were selected; each
136 consisting of 121 pixels centred at the location of the 118, 233 and 384 and 686 cGy fields.
137 One ROI was also selected corresponding to an un-irradiated region in the gel container. To
138 provide measurements of the signal and noise in the images, the mean HU value and the
139 standard deviation were calculated for these selected ROIs.

140 An analysis was also carried out to find out the optimum number of CT scans required for
141 reconstructing the zero-scan image. Starting from the first 50 images and subsequently
142 adding images up to 300 the average percentage error was calculated for each group of CT
143 images. As a first step the linear reconstructed image using all the 360 scans was considered
144 as the standard image. For each of the four ROI's described above the mean HU value was
145 calculated and compared with the standard image, and the percentage error was calculated for

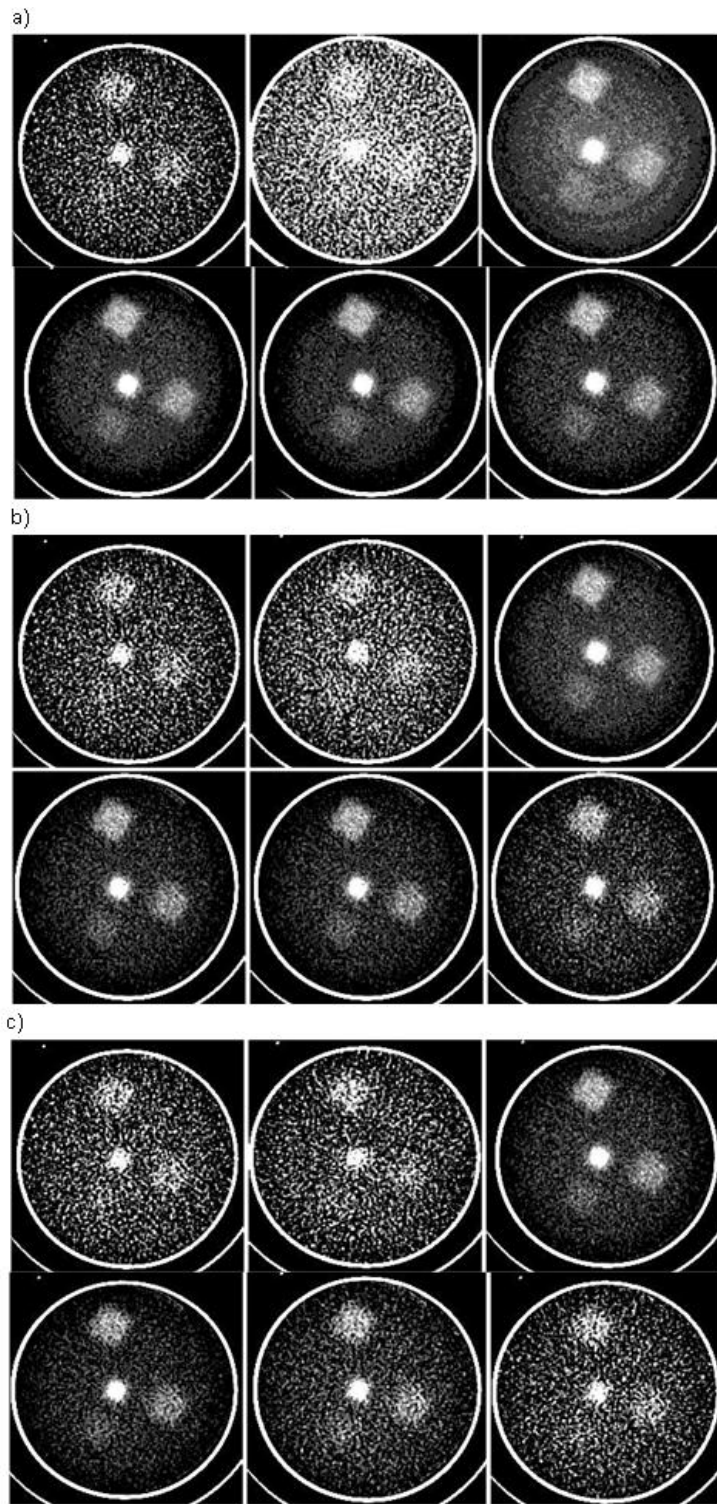
146 each ROI. In the final step all the individual % errors in the four ROI's were averaged out
147 and a single averaged percentage error was calculated for each group of CT images.

148 **III. RESULTS**

149 **III.A. Qualitative**

150 In Fig. 1 three sets of zero-scan images are shown, constructed from sets of all 360 images,
151 the first 50 images, and the first 16 CT images respectively. The relatively poor signal to
152 noise affecting un-processed CT scans of dosimetric gels is apparent in Fig. 1(a) top left and
153 top middle panels , which show the first and last of the 360 CT images of the gel dosimeter
154 series (with the averaged water image subtracted). Both images are windowed to the same
155 pixel values and an overall increase in HU is evident, above noise, in the overall lighter
156 appearance of Image 360. The source of this increase in HU is the gel density increase due to
157 the radiation dose delivered during the CT imaging. Fig. 1(a) top right panel shows an
158 average of all of the CT images of the gel dosimeter, and a reduction of noise compared to
159 the single images can clearly be seen. Lower left, middle and right panels show the zero-scan
160 images created from exponential, linear and quadratic fits to the data as described in Section
161 II.C. A reduction of noise compared to the upper panels is clearly evident.

162 Fig. 1(b) and 1(c) show results where only the first 50 and first 16 images of the dataset
163 respectively were used to create the zero-scan image. As fewer images are used an increase in
164 noise in the zero-scan image is clearly visible when a quadratic fit is used and less apparent
165 when exponential and linear fits are used.



166

167

168 FIG. 1. a) Top left-First CT image, top center-360th CT image, Top-right-Averaged CT image. Lower left-

169 Zero-scan image from exponential fit, Lower middle- Zero-scan image from linear fit, Lower right- Zero-scan

170 image from quadratic fit. b) Top left-First CT image, top center-50th CT image, Top-right-Averaged CT image.

171 Lower left- Zero-scan image from exponential fit, Lower middle- Zero-scan image from linear fit, Lower left-

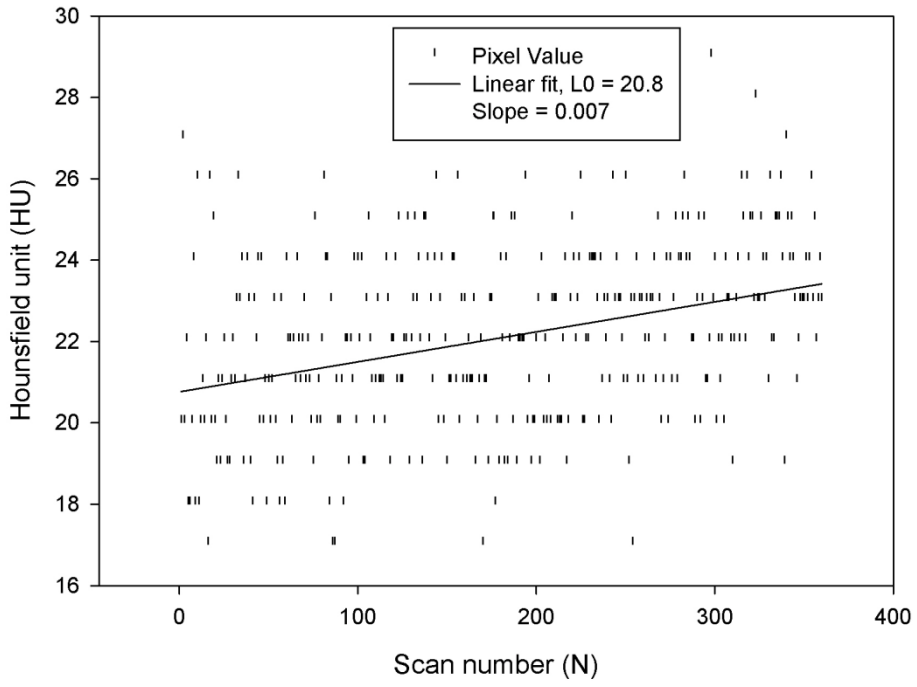
172 Zero-scan image from quadratic fit. c) Top left-First CT image, top center-16th CT image, Top-right-Averaged

173 CT image. Lower left- Zero-scan image from exponential fit, Lower middle- Zero-scan image from linear fit,
174 Lower left- Zero-scan image from quadratic fit. All images are windowed to the range 19-24 HU. Profiles
175 through the reconstructed images have been included as supplementary material.

176 **III.B. Quantitative**

177 Figs. 2(a) and (b) show examples of the variation in the HU values of two individual pixels
178 throughout the acquisition of the 360 CT images. A linear fit to the data for the pixel that
179 received 686 cGy (Fig. 2(a) has a steeper gradient (0.007 ± 0.001)) than the linear fit to the
180 data for the non-irradiated pixel (0.004 ± 0.001) as shown in Fig. 2(b). This suggests that the
181 un-irradiated gel is less sensitive to the additional dose increments delivered during the
182 scanning process, possibly due to inhibition of the low-dose response caused by the residual
183 presence of oxygen within the non-irradiated gel, as described by DeDeene *et al.*^{31,32}
184 Moreover, variations in manufacturing conditions may lead to different concentrations of
185 residual oxygen between batches leading to a variations in response to the CT dose utilized in
186 this technique. Therefore, this technique remains suitable only for relative dosimetry unless
187 an internal absolute calibration is undertaken.⁸⁻¹⁰

188 (a)

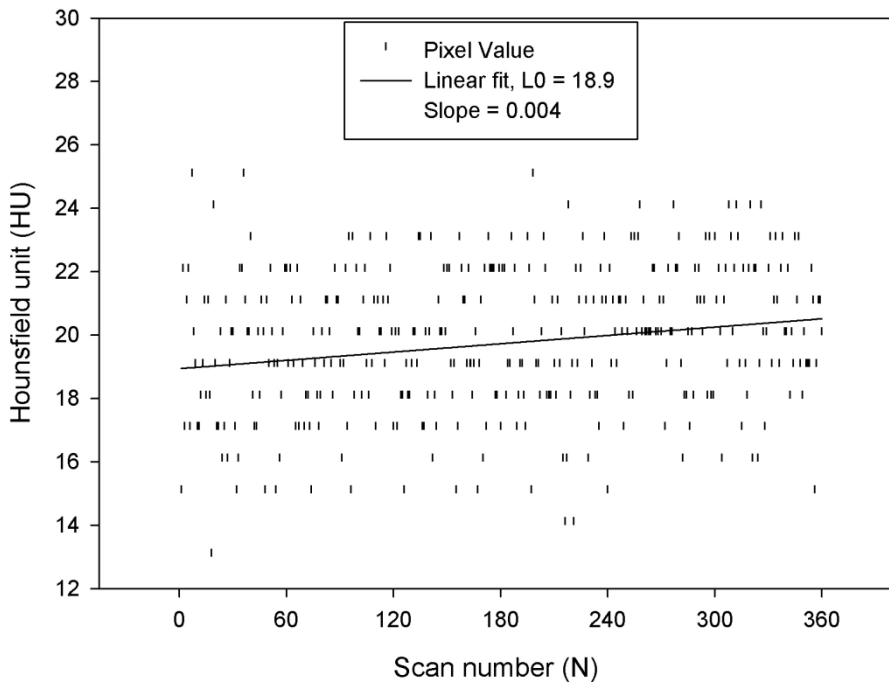


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(b)

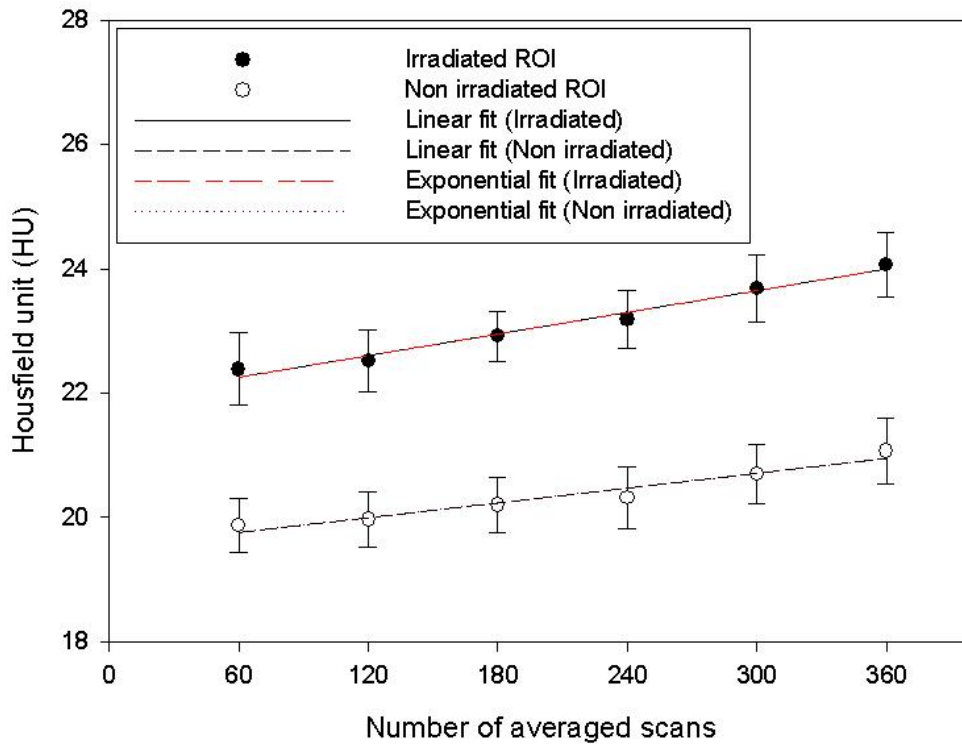


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(c)



195

196 FIG. 2. (a) HU value of a single pixel inside the irradiated ROI (686 cGy), over the 360 images. (b) HU value of
 197 a single pixel outside the irradiated ROIs, over the 360 gel images c) Averaged HU values within irradiated and
 198 non irradiated ROIs (121 pixels) with linear and exponential fits. The error bars represent the mean standard
 199 deviation within the ROI.

200 Fig. 2(c) shows the mean pixel value of an ROI of 121 pixels calculated in the regions
 201 corresponding to irradiated and non irradiated portions of the gel, and further averaged over
 202 increasing numbers of images starting with the first raw CT/zero image. The data in Fig. 2(c)
 203 has been fitted with linear and exponential fits. Results from statistical analysis of the
 204 exponential and linear fits using 360 images are shown in Table I, which suggests that the
 205 linear and exponential fits yield similar results. Although only the data for a single pixel
 206 within the 686 cGy field is shown, similar results are noted for other pixels in this and other
 207 fields.

208

209

210

211 TABLE I. Analysis of linear and exponential fits

Fit type	Equation	Coefficients(95 % confidence bounds)	SSE	R ²
Linear Model	$f(x) = p1 * x + p2$	P1 = 0.005804 (0.00463, 0.006979)	0.04509	0.9792
		P2 = 21.91 (21.63, 22.18)		
Exponential Model	$f(x) = a * \exp(-x / b) + c$	a = -5632 (- 2.246e+007, 2.246e+007)	0.04511	0.9792
		b = 9.701e+005 (-3.867e+009, 3.867e+009)		
		c = 5654 (- 2.244e+007, 2.244e+007)		

212

213 Fig. 3(a-c) shows the mean HU in each ROI in the zero-scan images of the gel, with error
 214 bars representing +/- one standard deviation from the mean for the three data sets (i.e. 360, 50
 215 and 16 images). Since the first CT image in the series is the closest in value to the irradiated
 216 gel dosimeter prior to imaging, it is used for comparison with the other images.

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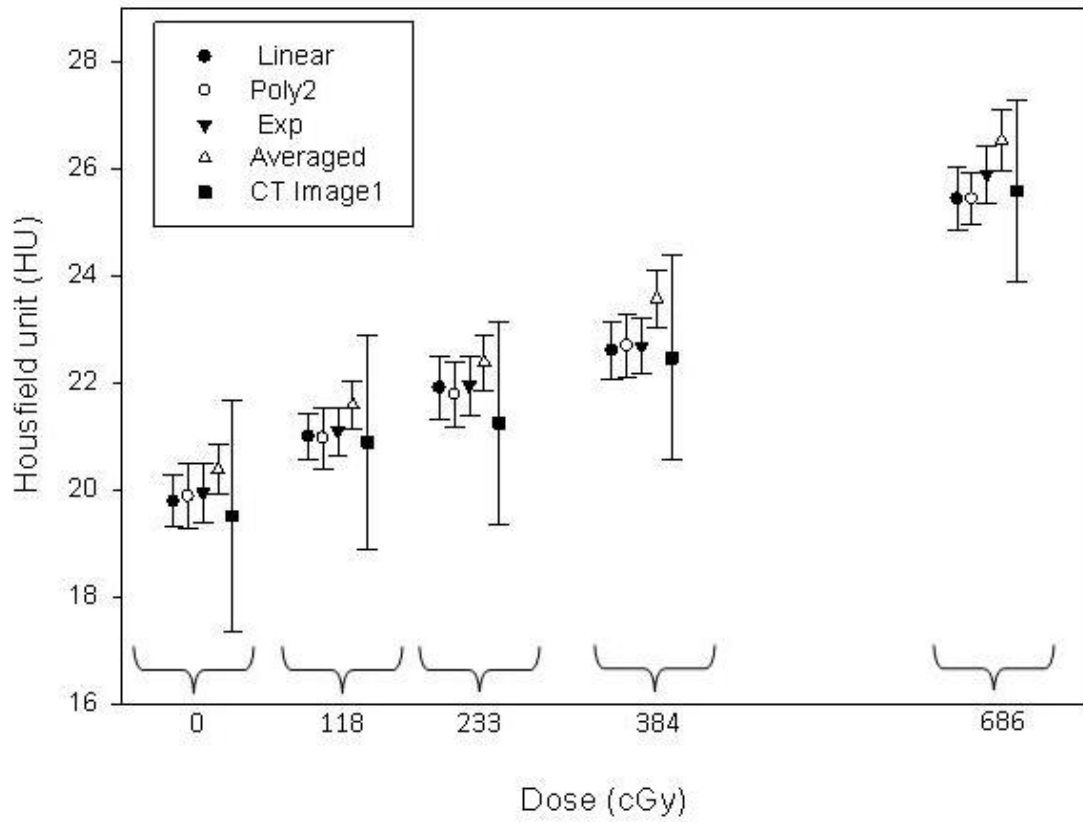
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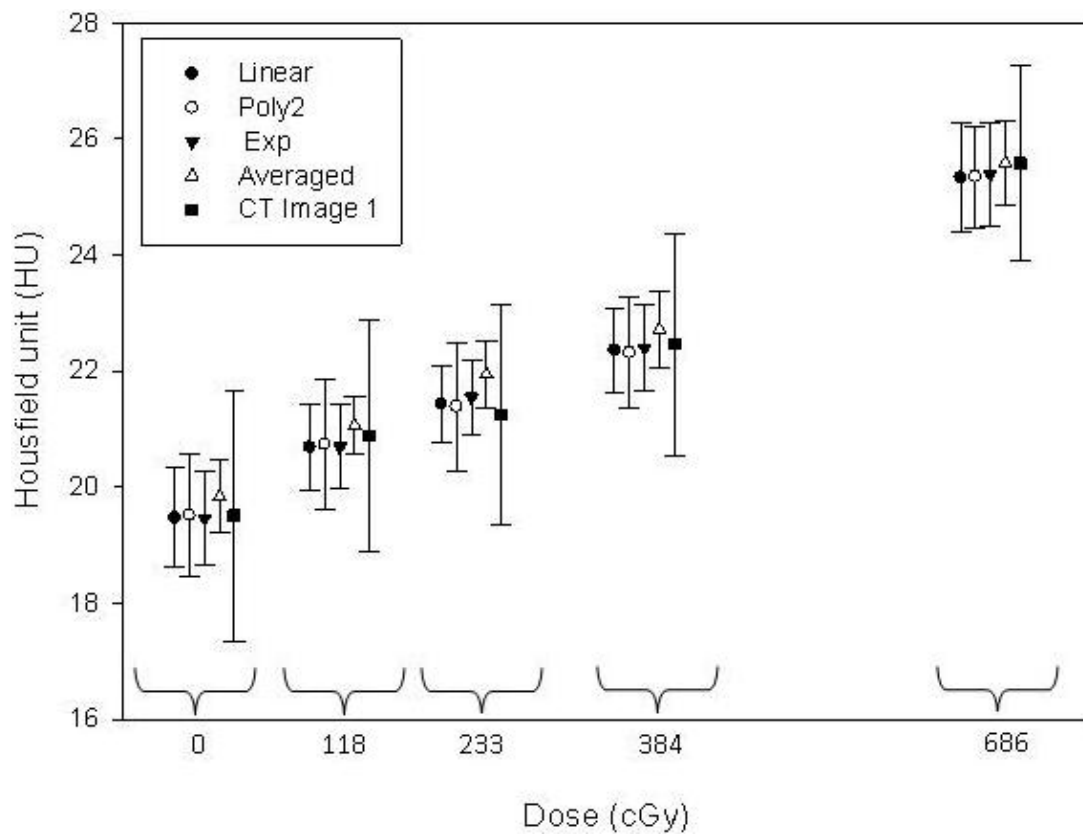
a)



223

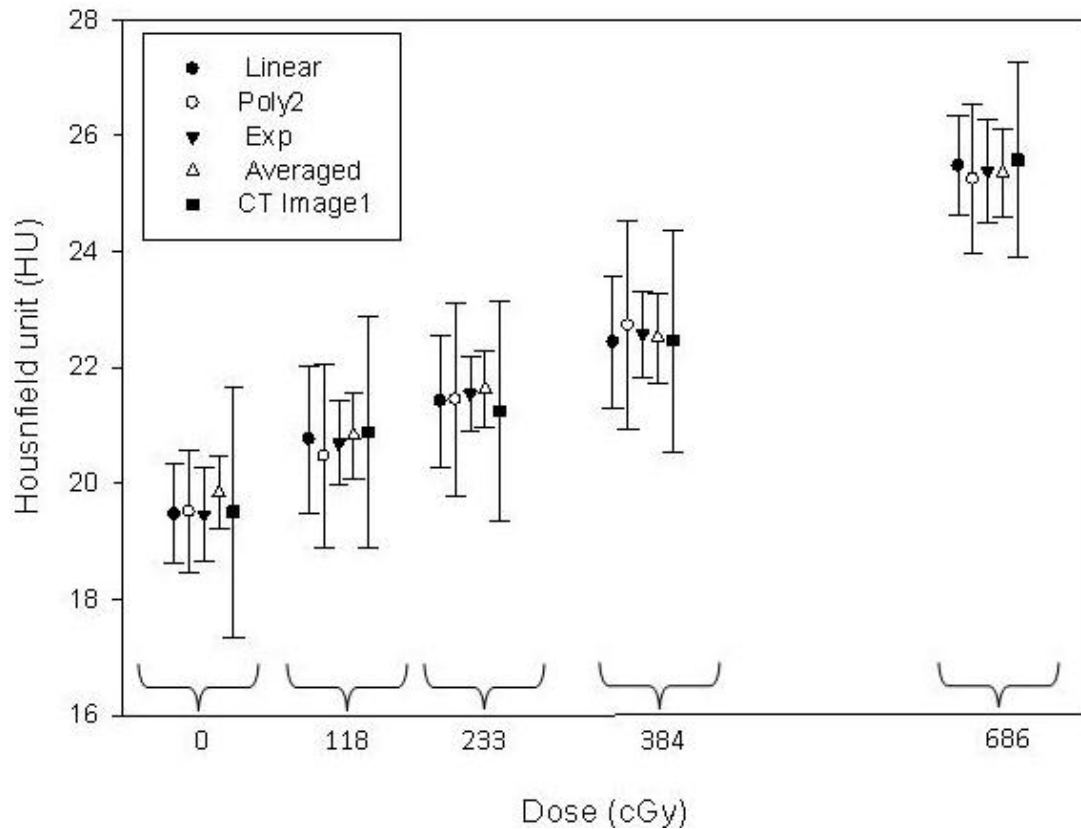
224

b)



225

226



228

229 FIG.3. Plot of mean HU versus dose in cGy for all types of fit. 'CT # 1' refers to the first gel image. 'CT
 230 Averaged' refers to the average of all gel CT images. On the x-axis the data points were shifted for visual clarity
 231 a) using 360 images b) using 50 images c) using 16 images. Error bars represent one standard deviation of the
 232 pixel values and therefore 68% confidence interval.

233 Fig. 3(a) shows that when all CT images are averaged together stochastic noise is reduced (as
 234 indicated by the smaller error bars on the data from the averaged scans), but the resulting
 235 mean HU values in the ROIs are consistently higher than the mean CT numbers from the
 236 corresponding ROIs in the first CT image. This suggests that although the averaging of CT
 237 images may produce more precise images, the accuracy suffers. In fact, for almost all ROIs,
 238 the mean HU value in the first CT image falls outside the error bars of the averaged image,
 239 indicating the severity of the inaccuracy of the data from the averaged image.

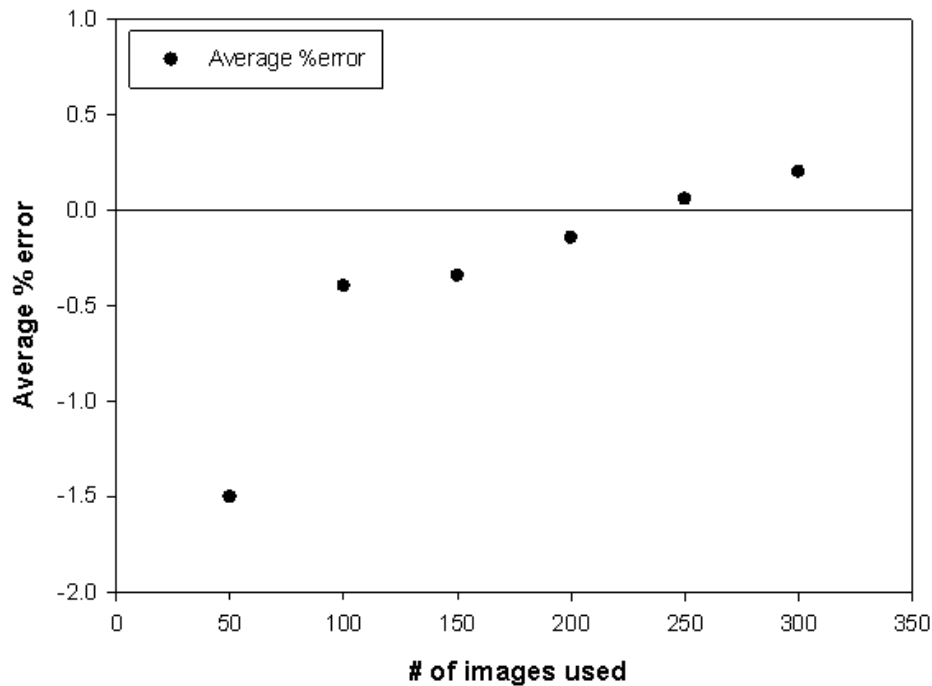
240 Fig. 3(a) shows that when using 360 CT images to create the zero-scan image, all of the
 241 fitting functions produce mean HU values in each ROI which are of a close match to the

242 values from the first CT image. In fact, the mean ROI values from CT image 1 falls within
243 the error bars of all corresponding zero-scan images.

244 In Figs. 3 (b) and 3 (c), where 50 and 16 CT images respectively are used to create the zero-
245 scan image, there is an increase in noise compared to Fig. 3 (a) for the zero-scan images
246 using linear, exponential and quadratic fits. The noise in the averaged image is less as
247 compared to the fitted data however the mean value remains inaccurate indicating the
248 deposition of the dose due to averaging. The minimum dose limit of the PAGAT gel was
249 found to be 2.3 Gy for 0 Gy dose at 95% confidence interval using the approach reported by
250 Trapp *et al.*²³ with a linear fit to the data.

251 **III.C. Optimum number of scans required for reconstruction**

252 In Fig. 4 the average % error is plotted against the number of CT images used. It clearly
253 demonstrates that if less than 100 images are used then the error value increases; however the
254 error remains within 0.5% if 100 or more CT images are used in the work presented here. The
255 number of images required will naturally vary according to the uncertainty required by the
256 user together with the CT imaging parameters (for example, see Baxter *et al.*³³), and can be
257 calculated by using statistical methods.



258

259 FIG.4. Plot of average % error (ROI's compared to the linear reconstructed image) versus the number of images
 260 used in image reconstruction.

261

262 **IV. DISCUSSION**

263

264 Reducing noise in CT imaging of gel dosimeters by averaging 360 images results in an
 265 overall increase in CT number of the final image due to the gradual increase in gel density
 266 caused by the radiation dose delivered during CT scanning. This is illustrated by the
 267 comparison between Fig. 1(a), where a visible difference is seen between images from the
 268 first and last scans, and by examination of the data in Fig. 3(a).

269 Comparison of the response of the non-irradiated and irradiated regions of the gel,
 270 exemplified by the different gradients of the linear fits to the individual pixel response data in
 271 Figs. 2(a) and (b) indicates that this PAGAT gel responds differently to the incremental
 272 absorption of small radiation doses (from repeated CT scanning) depending on its degree of

273 pre-irradiation. This is additional confirmation of behaviour observed by several
274 authors.^{6,29,31,32} Some authors have shown that some polymer gel dosimeters undergo post-
275 irradiation changes at a rate which depends upon the absorbed dose, resulting in an edge
276 enhancement effect^{6,12}. In the present work the maximum delivered dose of 686 cGy is below
277 that shown to produce measurable changes in the 24 hour irradiation-to-imaging time;
278 however if this technique is used with larger doses then earlier imaging may be required to
279 ensure accuracy.

280 Fitting a function to the pixel data and using this to evaluate a zero-scan image substantially
281 reduces image noise, while providing accurate measurements (see Figs. 2 and 3). Fitting the
282 exponential fit does not result in any additional advantage in analysing CT images according
283 to the method described here as the level of extra dose delivered via CT scanning is low and
284 the gel's subsequent response shows no obvious non-linearity. When analysing CT data by
285 producing a zero-scan image, data shown in Fig. 3 suggest that the use of a simple, linear
286 fitting relationship is suitably accurate in situations where a user does not have access to
287 exponential fitting software.

288 The technique presented here can be accompanied by further techniques for noise
289 improvement including image filtering^{24,27,28}, However the application of a particular filtering
290 strategy is dependent on the nature of the dose distribution and the noise present in the
291 original CT data. Reduction of noise by averaging the CT images^{20,21} will result in
292 inaccuracies as evident from Fig. 3.

293 The data presented here represents results from specific scanning parameters on a specific
294 scanner and gel dosimeter. If parameters are varied the technique presented here will continue
295 to work if the gel dosimeter chosen changes CT number with dose. For example, using a
296 smaller slice thickness will increase the stochastic noise in each acquisition, thus increasing
297 the noise in Fig. 2, however providing a large enough sample size is acquired the fitted

298 function (and therefore intercept) will not significantly alter. Similarly, using a more sensitive
299 gel dosimeter^{34,35} may reduce the number of images required for suitable results using this
300 technique, and future work beyond the scope of this paper will refine this technique.

301 **V. CONCLUSION**

302

303 A simple method has been proposed for improving the image quality of polymer gel
304 dosimeters imaged with x-ray CT. It has been shown that a simple analysis of the increase in
305 HU with repeated imaging can be used to produce an accurate, low-noise ‘zero-scan’ image
306 of the gel. The zero-scan image prediction method described here has been shown to be
307 capable of improving the precision while maintaining the accuracy of a two-dimensional
308 single-slice CT image of a gel sample irradiated to a range of doses. Use of multi-slice or
309 cone-beam CT modalities to provide repeated three-dimensional CT scans of the gel would
310 allow this method to be applied in three dimensions, without the measured dose being
311 compromised by scattering effects from adjacent slices.

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314 dosimeters.

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322 ^{a)} Conflict of interest notification. The authors report no actual or potential conflict of interests.

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