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Title	Synthesized effective atomic numbers for commercially available dual-energy CT
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Relation	



1	Technical note: Synthesized effective atomic numbers for commercially available dual-energy CT
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17	Keyword: effective atomic numbers, dual-energy CT, monochromatic CT number, beam-hardening
18	Conflict of Interest Notification: none
19	
20	ABSTRACT

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23tissue characteristic phantom and contrast material of varying iodine concentrations using single-source fast kilovoltage 24switching dual-energy CT (DECT) scanner. 25Methods: A newly developed multi energy tissue characterisation CT phantom and an acrylic phantom with various iodine 26concentrations of were scanned using single-source fast kilovoltage switching DECT (GE-DECT) scanner. The difference 27between the measured and theoretical values of Zeff were evaluated. Additionally, the difference and coefficient of variation 28(CV) values of the theoretical and measured values were compared with values obtained with the Canon-DECT scanner that 29was analysed in our previous study. 30 **Results:** The average  $Z_{eff}$  difference in the Multi-energy phantom was within 4.5%. The average difference of the theoretical and measured Zeff values for the acrylic phantom with variation of iodine concentration was within 3.3%. Compared to the 31 32results for the single-source Canon-DECT scanner used in our previous study, the average difference and CV of the theoretical 33 and measured Zeff values obtained with the GE-DECT scanner were markedly smaller. 34 Conclusions: The accuracy of the synthesized Zeff values with GE-DECT had a good agreement with the theoretical Zeff values 35for the Multi-Energy phantom. The GE-DECT could reduce the noise and the accuracy of the Zeff values than that with Canon-36 DECT for the varying iodine concentrations of contrast medium. 3738Advances in knowledge: The accuracy and precision of the Zeff values of the contrast medium with the GE-DECT could be 39 sufficient with human equivalent materials.

**Purpose:** The objective of this study was to assess synthesized effective atomic number ( $Z_{eff}$ ) values with a new developed

# 41 Introduction

42	Dual-energy computed tomography (DECT) enables direct calculation of the effective atomic number ( $Z_{eff}$ ), the
43	monochromatic energy CT number, and electron density on pixel by pixel basis <sup>1,2)</sup> . The beam hardening artefact can be reduced
44	by DECT and DECT provide more quantitatively accurate attenuation measurements <sup>3-5</sup> ). Additionally, DECT can estimate
45	iodine content in tissues by using the iodine map <sup>6,7</sup> ). In clinic, DECT has been applied to bone removal, and the automatic
46	characterization of stone compositions <sup>8-10</sup> .
47	Various commercial DECT scanners are available that can acquire CT datasets at two different energies: a single-
48	source dual-energy scanner with fast kilovoltage switching; a dual-source, dual-energy scanner; a single-source CT scanner
49	that switches kilovoltages between gantry rotation; and a single-source, dual-energy scanner with two detector layers. In the
50	current study, the dual-energy scanner with fast kilovoltage switching is used. The advantages of DECT with fast kilovoltage
51	switching is temporal registration between two-different energy datasets, that are acquired simultaneously.
52	Mitchell et al. evaluated the accuracy of the Zeff values that were calculated with fast kilovoltage switching with a
53	single detector layer with a GE Discovery CT750 DECT scanner (GE Healthcare, Princeton, NJ, USA) <sup>11)</sup> . They investigated
54	the accuracies of the synthesized effective atomic number and monochromatic images maps. Recently, a new DECT system,
55	Revolution HD CT(GE-DECT) scanner (GE Healthcare, Milwaukee, WI), has been developed. This scanner is expected to
56	improve the accuracy of the Zeff values compared to Discovery CT750 HD scanner (GE Healthcare, Milwaukee, WI). It is able
57	to be expected to improve the accuracy of the $Z_{eff}$ values using the Revolution HD CT.
58	DECT has the advantage that it can create the iodine maps image <sup>12</sup> ). The lesion target and normal tissue delineation,
59	extraction of the blood vasculature could be achieved by quantification of the iodine concentration in cancers. An iodine
60	distribution map is a promising tool for predicting the tumor response after treatment for cancers. Lee et al. showed that there

61	is a possibility to distinguish between different cancers by quantifying iodine concentration <sup>13</sup> ). But, before the iodine
62	distribution map can be used clinically, it is necessary to understand the accuracy of iodine quantitation with DECT. Our
63	previous study investigated the accuracy of the estimated $Z_{\text{eff}}$ values for varying iodine concentrations of contrast medium
64	(CM) compared with the theoretical Z <sub>eff</sub> values using a single-source CT that switches voltages between gantry rotations, as
65	implemented in Canon Aquilion ONE <sup>TM</sup> DECT scanner (Canon-DECT) (Canon Medical Systems Corporation, Ōtawara-shi,
66	Japan). In the current study we found that the average difference between the theoretical and estimated $Z_{eff}$ values for the CM
67	was within 11.2% <sup>14)</sup> .
68	The aim of the current study was to assess a new developed phantom with inserts of tissue material that replicates
69	expected Hounsfield unit (HU) dependencies from low energy to high energy using the GE-DECT. Moreover, the accuracy of
70	the synthesized Z <sub>eff</sub> values with contrast material of varying iodine concentrations using the GE-DECT were evaluated, and the
71	comparison of the $Z_{eff}$ values of the GE-DECT and the Canon-DECT was performed.
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# 82 Methods and Materials

83 A) Data acquisition:

84	The current study used following different DECT scanners: a) the Revolution DECT scanner (GE Healthcare, Princeton,
85	NJ, USA) which will be referred to as GE-DECT, and b) the Canon Aquilion ONE <sup>TM</sup> (Canon Medical Systems
86	Corporation, Ōtawara-shi, Japan) which will be referred to as Canon-DECT. The scan data was obtained from previous
87	study <sup>13</sup> ). The GE-DECT scanned at 80 and 140 kV tube voltages and exposures of 560 mA were used. The other scanning
88	parameters were field of view (FOV) of 360 mm, slice thickness (ST) of 0.5 mm, and a rotation time (RT) of 1.0 s. The
89	72 middle slices of a total of 80 slices was analysed. The Canon-DECT was scanned at tube voltages of 135 and 80 kV
90	using the volume scanning method. The exposures were 800 and 200 mA, and the time taken to switch the tube voltage
91	between 135 and 80 kV was 0.4 s. The other parameters were FOV of 400 mm, ST of 0.5 mm, and a RT of 1.0 s. The $Z_{eff}$
92	was reconstructed from the scanned DECT image. Also, the 70 keV monoenergetic CT image was reconstructed from the
93	scanned DECT image to evaluate the accuracy of the iodine concentration.
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- 95 B) Phantom:

Two phantom were scanned: 1) A Multi-Energy phantoms with inserts of varying iodine and calcium concentrations (Sun Nuclear, Middleton, WI, USA) (Figure 1a), and 2) an in-house developed acrylic phantom with inserts syringes filled with different iodine concentrations (Figure 1b). The size of the acrylic phantom is 32 cm Ø and 6 cm height. The syringes filled with CM (Omnipaque 300, GE Healthcare, Prnceton, NJ, USA) diluted water to predetermined iodine concentrations of 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 20, 40, 60, 90, and 130 mg iodine per ml. Here, the syringes filled with

- the CM used in the current study were created in our previous study <sup>13</sup>). The syringe was not emptied and refilled between 101 the current and previous studies. 102
- Multi-Energy CT phantom can improve material decomposition in clinical, such as distinguishing calcification from 103 iodinated contrast and blood from calcification<sup>15</sup>. However, the maximum concentration of the CM in the Multi-Energy 104 phantom is 15 mg/ml. Jang et al. reported that Lipiodol, which is used in trans-arterial chemoembolization (TACE), had 105 a larger CT number, and its value was over 2000 HU at maximum <sup>16</sup>). Our previous study showed the correlation of the 106 CT number and the concentration of the CM. A high concentration of the CM at over 20 mg/ml has been used for TACE. 107108 Thus, the current study evaluates the Zeff values for high concentrations of the CM at over 20 mg/ml with the in-house 109 developed CM phantom.
  - (a)



- 110
- Figure 1 (a) Multi-Energy phantom, (b) Acrylic phantom with variation of iodine concentration of CM. 111
- 112
- Theoretical Zeff value: 113C)
- 114The theoretical Zeff values for the Multi-energy phantom and acrylic phantom with CM were calculated using Mayneord's
- equation <sup>17</sup>): 115
- 116

117 
$$Z_{\rm eff} = \sqrt[2.94]{\sum_{i=1}^{n} a_i Z_i^{2.94}}$$
(1)

- 119 where  $Z_i$  is the atomic number and  $a_i$  is the fractional of the electrons in the *i*-th element in the mixture to the total number 120 of electrons. The material composition information is used rereleased by the manufacturer.
- 121
- 122 D) Measured Z<sub>eff</sub> value:

The  $Z_{eff}$  image reconstructed by GE and Canon DECT scanners was analyzed using the ImageJ (National Institutes of Health, Bethesda, MD, USA). The effect of the beam hardening was evaluated by measuring the centre and peripheral region as shown in Figure 2(a). The syringe was filled with the water only. A circular region of interest (ROI) for each image was drawn within 0.8 cm area in the syringe. The mean (M) and standard deviation (SD) of the  $Z_{eff}$  values within a circular ROIs for each slice were measured. The average of the M and SD for 72 slices were evaluated. For the evaluation of the CM, the M, SD, and coefficient of variation (CV) of the  $Z_{eff}$  values in the syringes were evaluated. The CV is the ratio of the standard deviation to the average  $Z_{eff}$  values in the different pixels of the ROI, as follows.

130 CV = SD/M (2)

The average of M, SD, and CV of the  $Z_{eff}$  values for 72 slices were evaluated for the syringe with the CM. The ROI for each image were drawn within 0.8 cm area in the syringe. At low concentration of the CM, the mean  $Z_{eff}$  value is smaller, thus the effect of the SD might be larger, relatively. Thus, the CV was used to evaluate the variation in the images at low and high concentrations of the CM. The proportionality of contrast enhancement to iodine concentration is near constant within 15 mg/ml<sup>-18</sup>. Thus, the current study assumed that the correlation of the CT number and iodine concentration was fitted to a linear function. The concentration was calculated from the CT number with a linear function, which was compared with the iodine concentration which we defined. Thus, the mean and SD of the CT numbers at 70 keV image reconstructed from the GE-DECT in the syringe with the iodine concentration within 10 mg/ml were evaluated. For the Multi-energy phantom, the method of the measurement was the same with the CM. A circular ROI for each image was drawn within 0.8 cm area in the material inserts. The M and SD of the  $Z_{eff}$  values within a circular ROIs for each slice were measured. The average of the M and SD for 72 slices were evaluated.



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Figure 2. (a) Method of measurement with the acrylic phantom by the beam hardening effect. The distance of the center of the ROI and peripheral of the ROI was 13 cm. The mean and SD were measured by creating a circular ROI with 0.8 cm. (b) Method of measurement with the acrylic phantom that inserted the syringes filled with CM that the diameter is 1 cm in a syringe that the diameter was 1.5 cm. The mean and SD were measured by creating a circular ROI with 0.8 cm diameter in the syringe.

150 E) Evaluation:

151 In the current study, the following items were investigated. i) The accuracy of the  $Z_{eff}$  values in the Multi-Energy phantom. 152 ii) The accuracy of the  $Z_{eff}$  values for the CM phantom. The measured  $Z_{eff}$  values were compared with theoretical values 153 and the relative average differences were calculated. The accuracy of the  $Z_{eff}$  values with the GE-DECT was compared 154 with the Canon-DECT. iii) The difference of the CV values between the GE-DECT and the Canon-DECT.

#### 157 **Results**

158 A) Accuracy of the iodine concentration:



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Figure 3 The average of M and SD of the  $Z_{eff}$  values at iodine concentrations of 0–10 mg/ml for 72 slices. The fitting was performed with linear function.

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168 B) Reproducibility of the measured Z<sub>eff</sub> values and effect of the beam hardening:

169 Figure 4(a) shows the  $Z_{eff}$  values in the centre and peripheral region. The maximum difference of the  $Z_{eff}$  values in

- 170 the centre region and peripheral region was 0.01. The beam hardening effect is significantly smaller than the SD of the  $Z_{eff}$
- 171 values. Figure 4(b) shows the reproducibility of the measurement Z<sub>eff</sub> value. The difference of the Z<sub>eff</sub> values for three scans was
- $172 \qquad \text{within the SD of the } Z_{\text{eff}} \text{ values}.$



Figure 4 (a) The average of the M of the measurement  $Z_{eff}$  values in the center and peripheral region for 72 slices. The error bar represents the average of the SD of the measurement  $Z_{eff}$  values for 72 slices. (b) Reproducibility of the measurement  $Z_{eff}$  value for three scans. The error bar represents the SD of the measurement  $Z_{eff}$  value for three scans.

Figure 5 (a) represents the theoretical Zeff values and the average M and SD of the measured Zeff values, and Figure 179180 5 (b) represents the deviation between the theoretical Zeff values and the measured Zeff values in the Multi-Energy phantom. As shown in Figure 5 (b), the difference of the Zeff values for all material inserts were within 5.1%. The average and SD of the 181 182difference of theoretical and measurement Zeff values for all material inserts were 2.5% and 1.4%. Figure 6 (a) represents the 183theoretical Zeff values and the average M and SD of the measured Zeff values, Figure 6 (b) represents the deviation between the 184theoretical Zeff values and the measured Zeff values in the acrylic phantom. The Zeff values were larger at higher concentration of the CM. The maximum standard deviation of the CM was within 0.1. As shown in Figure 6 (b), the maximum difference 185186 was within 0.6%. At the low concentration of the CM within 10 mg/ml, the deviation between the theoretical Zeff values and 187 the average measured Zeff values were scattered. Figure 7 represents the difference of the Zeff values in the CM scanned by the 188 GE-DECT that was measured in the current study and the Canon-DECT that was found in our previous study. The average





 $200 \qquad \text{Figure 5} (a) \text{ The theoretical } Z_{\text{eff}} \text{ values and the average measured } Z_{\text{eff}} \text{ values. Error bars represent standard deviation of the average} \\$ 











209 Figure 7 The difference of the theoretical and measured Z<sub>eff</sub> values with the GE-DECT and the Canon-DECT in the acrylic phantom

210 with variation of iodine concentration of CM.

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Figure 8 The difference of the CV of the Z<sub>eff</sub> values with the GE-DECT and the Canon-DECT in the acrylic phantom with variation

215 of iodine concentration of CM.

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### 217 **Discussion**

218 This study evaluated the accuracy of  $Z_{eff}$  values in tissue equivalent materials and the CM. The past study reported 219 the accuracy of the  $Z_{eff}$  values with various DECT scanner types and various tissue-equivalent phantoms. The past study 220 reported the accuracy of the  $Z_{eff}$  values with various DECT scanner types and various tissue equivalent phantoms. Mitchell, et

221	al. investigated the accuracy of the $Z_{eff}$ values estimated from DECT scans acquired with a Discovery CT/50 DECT scanner;
222	they found the Zeff values of the Catphan phantom and tissue characterization were within 15% 10). In the current study,
223	Revolution HD CT was used. The accuracy of the $Z_{eff}$ values in tissue equivalent phantom was within 4.5%. The average and
224	SD of the difference of $Z_{eff}$ values in tissue equivalent phantom was within 2.5% and 1.4%, respectively. The Revolution CT
225	has enabled increasing 20% energy separation between the high and low energies by improving generator hardware enabling
226	faster kV to rise and fall times by comparing with Discovery CT750 HD <sup>19</sup> . Thus, the beam hardening artefact and the noise
227	could be reduced <sup>20)</sup> . Material discrimination could be accurate by increasing spectral separation <sup>16)</sup> . This contributed to the
228	improved accuracy of the $Z_{eff}$ values using the Revolution HD CT. Moreover, our previous study evaluated $Z_{eff}$ values in raw-
229	data based reconstruction image with the Canon-DECT implicated by Canon for the tissue equivalent phantom <sup>13</sup> ). The accuracy
230	except of the lung inserts were within 8.4%. The Canon-DECT was scanned with 135 kV and 80 kV, thus the higher kV energy
231	was lower than the GE-DECT implicated by GE Healthcare. This could potentially result in an increased spectral separation
232	that contribute to the reduced noise and better material discrimination.
000	

For the acrylic phantom with the syringe filled with the CM, the beam hardening artefact was smaller and the reproducibility was significantly smaller than the SD of the  $Z_{eff}$  values in the ROI. Thus, the reliability of the measurement  $Z_{eff}$ values was sufficient. The accuracy of the  $Z_{eff}$  values was within 3.3% at the range of 0-130 mg/ml in the CM with the GE-DECT. It could be also the higher beam was used the DECT implicated by GE Healthcare, which could reduce the beam hardening artefact with the high concentration of the CM.

At the low concentration of the CM within 10 mg/ml, the CV and the deviation between the theoretical  $Z_{eff}$  values and the average measured  $Z_{eff}$  values were scattered, as shown in Figure 6(b) and Figure 8. At the low concentration of the CM,

240 the mean value of the Zeff is close to 0. Thus, the SD in the images significantly affects the deviation and CV. Moreover, the

241 CV is larger at the low concentration of the CM even if the SD is the same value between low and high concentrations of the242 CM.

243	Although the accuracy of the $Z_{eff}$ values was within 7.2% at less than 20 mg/ml in the CM, the beam hardening
244	artefact was affected at over 20 mg/ml and the maximum difference was 11.2% at 130 mg/ml with the Canon-DECT. For the
245	GE-DECT, the accuracy of the $Z_{eff}$ values was within 3.3% at the range of 0-130 mg/ml in the CM. Moreover, the CV with the
246	GE-DECT was significantly smaller than the Canon-DECT. It depends on that the SD was smaller for the GE-DECT. The
247	image reconstruction method and imaging filter, and the deviation of the high and low-kV energy were affected these
248	differences. In clinical of radiation diagnosis, the accuracy and precision of the Zeff values within 15 mg/ml are needed. From
249	above, the GE-DECT could be useful for the material decomposition.
250	In our previous study, the CM extraction method was developed, but it used only electron density and CT data <sup>20</sup> .
251	However, they did not show the accuracy of the electron density. The current study revealed that the accuracy and the
252	precision of the Zeff values were sufficient for the material decomposition. It is possible to contribute to improving the
253	estimation accuracy of the CM distribution by adding the Zeff values. The accuracy and precision were different between the
254	DECT scanner types, thus the data such as electron density and Zeff obtained from DECT should be evaluated before using

255 for the material decomposition in clinical.

## 256 Conclusion

- 257 The accuracy of the synthesized Z<sub>eff</sub> values with dual-source DECT was in good agreement with theoretical values for the
- Multi-Energy phantom. The GE-DECT could reduce the noise and improve the accuracy of the  $Z_{\rm eff}$  values compared to a
- 259 Canon-DECT for the varying iodine concentrations of CM.

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313	Figure legends
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317	the ROI and peripheral of the ROI was 13 cm. The mean and SD were measured by creating a circular ROI with 0.8 cm. (b)
318	Method of measurement with the acrylic phantom that inserted the syringes filled with CM that the diameter is 1cm in a
319	syringe that the diameter was 1.5 cm. The mean and SD were measured by creating a circular ROI with 0.8 cm diameter in
320	the syringe.
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322	Figure 3 The mean and SD for the Z <sub>eff</sub> values at iodine concentrations of 0–10 mg/ml. The fitting was performed with linear
323	function.

325	Figure 4 (a) The mean and SD of the $Z_{eff}$ values in the center and peripheral region. (b) Reproducibility of the measurement $Z_{eff}$ value
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328	Figure 5 (a) The theoretical $Z_{eff}$ values and the average measured $Z_{eff}$ values. Error bars represent standard deviation of the average
329	values. (b) The deviation between the theoretical $Z_{eff}$ values and the average measured $Z_{eff}$ values in the Multi-Energy phantom.
330	
331	Figure 6 (a) The theoretical $Z_{eff}$ values and the average measured $Z_{eff}$ values. (b) The deviation between the theoretical $Z_{eff}$ values and
332	the average measured $Z_{eff}$ values in the acrylic phantom with variation of iodine concentration of CM.
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