

1 Strategies to reduce radiation dose in cardiac PET/CT

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32 **Abstract**

33 **Background:** Our aim was to investigate CT dose reduction strategies on a hybrid PET/CT scanner  
34 for cardiac applications.

35 **Materials:** Image quality and does estimate of different CT scanning protocols for CT coronary  
36 angiography (CTCA) and CT-based attenuation correction for PET imaging were investigated.  
37 Fifteen patients underwent CTCA, perfusion PET imaging at rest, under stress, and FDG PET for  
38 myocardial viability. These patients were divided into three groups based on the CTCA technique  
39 performed: retrospectively gated helical (RGH), ECG tube current modulation (ETCM) and  
40 prospective gated axial (PGA) acquisitions. All emission images were corrected for photon  
41 attenuation using CT images obtained using default setting and an ultra low dose CT (ULDCT)  
42 scan.

43 **Results:** Radiation dose of RGH technique was  $22.2\pm 4.0$  mSv. It was reduced to  $10.95\pm 0.82$  mSv  
44 and  $4.13\pm 0.31$  mSv when using ETCM and PGA techniques, respectively. Radiation dose in CT  
45 transmission scan was reduced by 96.5% (from  $4.53\pm 0.5$  mSv to  $0.16\pm 0.01$  mSv) when applying  
46 ULDCT as compared to the default CT. No significant difference in terms of image quality was  
47 found among various protocols.

48 **Conclusion:** The proposed CT scanning strategies, i.e. ETCM or PGA for CTCA, ULDCT for PET  
49 attenuation correction, could reduce radiation dose up to 47% without degrading imaging quality in  
50 an integrated cardiac PET/CT coronary artery examination.

## 51 **Introduction**

52           Coronary artery disease (CAD) is the leading cause of death around the world. For the  
53 diagnosis of CAD, medical imaging emerges as a powerful method over the past decade. PET  
54 imaging is one of the well-established tools for the evaluation of ischemic heart, blood flow  
55 quantification, myocardial perfusion and viability [1, 2]. Clinical protocols usually include cardiac  
56 PET with  $^{13}\text{N}$ -ammonia and  $^{18}\text{F}$ -FDG to assess myocardial perfusion and viability, respectively. On  
57 the other hand, cardiac CT has enabled not only the detection and quantification of coronary artery  
58 calcification by using CT calcium scoring, but also the grading of coronary stenosis through  
59 contrast-enhanced CT coronary angiography (CTCA). PET/CT is a hybrid imaging instrumentation  
60 that can perform these examinations in one patient exam session (Fig. 1) [3-5]. Moreover, the CT  
61 image can be used to correct PET scans for photon attenuation [6]. So the integrated system offers  
62 the abilities to decrease overall scanning time and improve localization of vessels or  
63 regions-of-interest in cardiac imaging. However, radiation dose of these cardiac PET/CT  
64 examinations shown in Fig. 1 can be up to 37.67 mSv, while the proportion of CT radiation dose is  
65 about 75.0 % (28.27 mSv) [2, 7]. The rationale of our study was to investigate CT dose reduction  
66 strategies on a hybrid PET/CT scanner for cardiac applications.

67

## 68 **Materials and Methods**

69

### 70 *Patient population*

71 A total of 15 consecutive patients (9 male, 6 female; mean age  $50.8 \pm 8.4$  years; range 39-68  
72 years) were enrolled for this cardiac PET/CT study between November 2009 and May 2010. All  
73 patients underwent the cardiac PET/CT scanning protocol as shown in Fig.1. Patients with prior  
74 allergic reaction to iodinated contrast media or atrial fibrillation were excluded. This study was  
75 institutional review board-approved, and written informed consent was obtained for all subjects.

76

### 77 *PET/CT Acquisition Protocols*

78 All studies were performed on a PET/CT scanner (Discovery<sup>TM</sup> VCT, Germany, GE). Data  
79 acquisitions in the CT and PET studies were performed with a matrix of  $512 \times 512 \times 64$  and  
80  $128 \times 128 \times 47$ , respectively. Patients with heart rate (HR)  $>70$  bpm prior to CTCA were administered  
81 40 mg of propranolol orally after the scout scans and returned for CTCA after their HR decreased to  
82  $<70$  bpm. For those enrolled 15 patients, 5 patients (HR =  $60 \pm 8$  bpm, height =  $1.6 \pm 0.16$  m, weight =  
83  $65.5 \pm 7.78$  kg) underwent CTCA with retrospectively gated helical (RGH) acquisitions, and 5  
84 patients (HR =  $59 \pm 8$  bpm, height =  $1.6 \pm 0.08$  m, weight =  $63.0 \pm 9.85$  kg) underwent CTCA with  
85 ECG tube current modulation (ETCM) acquisitions, 5 patients (HR =  $59 \pm 6$  bpm, height =  $1.6 \pm 0.03$   
86 m, weight =  $66.2 \pm 9.65$  kg) underwent CTCA with prospectively gated axial (PGA) acquisitions.  
87 Based on the weight of the patients, 70-90 mL of a nonionic contrast medium (Optiray 350, Tyco

88 Healthcare, Montreal, Quebec, Canada) was injected at a flow rate of 5 mL/s, followed by a 25 mL  
89 bolus of saline at the same rate using a dual-head injector (Stellant D; Medrad, Warrendale, PA,  
90 USA). PET imaging examinations include  $^{13}\text{N}$ -labeled  $\text{NH}_3$  rest-stress perfusion PET and  $^{18}\text{F}$ -FDG  
91 viability scan. All patients received 700-900 MBq and 298-458 MBq injections of  $^{13}\text{N}$ -labeled  $\text{NH}_3$   
92 and  $^{18}\text{F}$ -FDG via the peripheral vein before the start of serial transaxial tomographic imaging of the  
93 heart, respectively. To compensate effects of photon attenuation in PET imaging, two CT scanning  
94 protocols were applied: the default setting and the ultra low-dose CT (ULDCT). Table 1 lists the  
95 parameters of CT scanning protocols for coronary angiography and CT-based attenuation  
96 correction.

97

#### 98 *CT Data Analysis*

99 Coronary arteries were classified into 15 segments according to the scheme proposed by the  
100 American Heart Association [8] and the intermediate artery was designated segment 16, if present.  
101 Images were analyzed and graded randomly by 2 independent cardiovascular radiologists, each with  
102 >5 years experience. Images included transverse source images, (curved) multi-planar reformations,  
103 thin-slab maximum intensity projections, and volume-rendering mode, and were presented to the  
104 observers to identify coronary image quality. After the optimal reconstruction interval was  
105 determined, we used motion artifact as the figure-of-merit for assessing image quality. We  
106 performed semi-quantitative analysis by using a 4-point ranking scale (1 = no motion artifacts; 2 =  
107 mild blurring; 3 = moderate blurring without structure discontinuity; 4 = severe artifacts and

108 doubling). For any disagreement in data assessment, the 2 readers reviewed the data together until  
109 consensus was obtained. The image quality at the best reconstruction interval from RGH, ETCM  
110 and PGA patients was then compared with Wilcoxon signed ranks test (NCSS version 2007,  
111 NCSS).

112

### 113 *PET Data Analysis*

114 The mutual information and the correlation coefficient were computed to characterize  
115 similarity between PET images corrected for photon attenuation using CT images obtained with  
116 default setting and ULDCT. Mutual information (MI) was applied to estimate the non-linear  
117 intensity distribution between two sets of images. The random variables  $X$  and  $Y$  are defined as sum  
118 of all grey value pairs at corresponding positions between two sets of images. Their intensity value  
119 at a certain coordinate in the images is the joint outcome of a random experiment. The MI between  
120  $X$  and  $Y$ , denoted as  $I(X, Y)$  is defined as equation (1):

$$121 \quad I(X, Y) = \sum_{x \in X} \sum_{y \in Y} p(x, y) \log \frac{p(x, y)}{p(x)p(y)} \quad (1)$$

122 where  $p(x)$  is the histogram of  $X$ ,  $p(y)$  is the histogram of  $Y$ , and  $p(x,y)$  is the joint histogram of  $X$   
123 and  $Y$ . The larger  $I(X,Y)$ , the more similar two images are [9,10].

124 Correlation coefficient (CC) was applied to calculate the intensity relationship point by point  
125 between two image sets. The CC value represents the linear information about intensity difference  
126 between two imaging sets of  $n$  voxels in each. The equation defining the correlation coefficient is:

127 
$$CC = \frac{S_{u,v}}{S_u * S_v} = \frac{\sum_{i=1}^n (u_i - \bar{u}) * (v_i - \bar{v})}{\sqrt{\sum_{i=1}^n (u_i - \bar{u})^2} * \sqrt{\sum_{i=1}^n (v_i - \bar{v})^2}} \quad (2)$$

128 where  $S_u$  is the standard deviation of object  $u$ ,  $S_v$  is the standard deviation of object  $v$ ,  $S_{u,v}$  is  
129 the covariance of object  $u$  and  $v$ ,  $CC$  is the correlation coefficient of  $u$  and  $v$ , which is between -1  
130 and 1.

131

### 132 *Radiation Dose*

133 The volume CT dose index (CTDI<sub>vol</sub>) was displayed after each scan on the scanner's console.  
134 To obtain the effective dose, the CTDI<sub>vol</sub> was multiplied by the scan length to get the dose-length  
135 product (DLP), an indicator of the integrated radiation dose of an entire CT examination. Values of  
136 DLP were converted into effective dose using a conversion factor of 0.017 mSv · mGy<sup>-1</sup> · cm<sup>-1</sup> for  
137 adult chest CT [11].

138

139 **Result**

140 A total of 220 segments were evaluated in the 15 patients. Twelve segments were missing  
141 because of anatomic variants and 8 segments were too small to visualize. When the best  
142 reconstruction interval was used, images without motion artifacts (score 1) were obtained in 40 of  
143 the 76 segments for RGH group, 38 of the 74 segments for ETCM group and 38 of the 70 segments  
144 for PGA group. No statistical significance was seen among image quality obtained from the three  
145 CTCA groups. Fig. 2 shows curved multiplanar reconstruction images of the right coronary artery,  
146 reconstructed from RGH, ETCM and PGA acquisitions.

147 For PET acquisitions of  $^{13}\text{N-NH}_3$  for perfusion (rest/stress) and FDG for viability with two  
148 different CT attenuation corrections, we got  $0.878\pm 0.041$ ,  $0.764\pm 0.037$ ,  $0.833\pm 0.037$  for the mutual  
149 information and  $0.976\pm 0.006$ ,  $0.974\pm 0.009$ ,  $0.982\pm 0.005$  for the correlation coefficient, respectively.  
150 These results indicate that the PET data sets corrected for photon attenuation using two different CT  
151 acquisition protocols provide similar image quality and information. Fig. 3 shows CT images for  
152 CT-based attenuation correction and corresponding FDG-PET images with attenuation correction  
153 using these CT images. Image noise is more noticeable in CT image obtained using ULDCT  
154 because of the reduced photon flux. However, no significant difference is observed between the two  
155 sets of PET images.

156 The effective dose of patients underwent a cardiac PET/CT examination are show in Fig. 4.  
157 Radiation dose of RGH technique was  $22.2\pm 4.01$  mSv. It was reduced to  $10.95\pm 0.82$  mSv and  
158  $4.13\pm 0.31$  mSv when using ETCM and PGA techniques, respectively, leading to a dose reduction of



159 50-83%. Radiation dose in CT transmission scan was reduced by 96.5% from  $4.53 \pm 0.5$  mSv to  
160  $0.16 \pm 0.01$  mSv when applying ULDCT protocol. When CT images for attenuation correction were  
161 obtained using ULDCT, radiation dose of the whole cardiac PET/CT examination using ETCM and  
162 PGA for CTCA could be reduced by 29.8-47.9%.

163 **Discussion**

164 Radiation dose is becoming a major issue for cardiac imaging. The current PET/CT growth  
165 leads to the volume of cardiac diagnostic procedures involving the use of ionizing radiation – within  
166 both emission and transmission scanning – increased considerably. Due to improved accuracy, the  
167 number of cardiac PET/CT scans is growing rapidly [2,12]. A number of techniques can be used to  
168 minimize dose from integrated cardiac PET/CT examination. For CTCA, PGA should be employed  
169 when it is expected that multiple reconstructions at different positions of the cardiac cycle will not  
170 be necessary for diagnosis. This is generally the case for patients with regular rhythm, little or no  
171 ectopy, and well-controlled HR after administration of beta-blockers such as metoprolol.  
172 Beta-blockers play an important role in dose reduction in addition to improving image quality by  
173 decreasing coronary artery velocity.

174 Another important consideration is the optimization of tube current and voltage. In modern  
175 PET imaging, CT has replaced Ge-68 for the transmission scan. However, the drawback of the  
176 helical CT technology is the higher radiation dose to patients. Effective dose increases linearly with  
177 tube current [13], and therefore tube current should be minimized to the lowest level yet still  
178 providing acceptable image quality for performing attenuation correction on PET images. Therefore,  
179 we use 10 mA tube current for ULDCCT in our study. The low tube current causes low photon flux  
180 thus leading to low signal-to-noise ratio in the images. However, these CT images are mainly for  
181 attenuation correction purpose thus diagnostic image quality is not needed. Our results of MI and  
182 CC showed that there is actually no significant difference between the PET images with attenuation

183 correction using default CT and ULDCT. Thus, the diagnostic accuracy of PET images corrected  
184 using ULDCT should not be affected. Pascal Koepfli *et al.*[14] once compared the usage of  
185 conventional Ge-68 external source and CT with different tube currents (120 mA, 80 mA, 40 mA  
186 and 10 mA) as the transmission scans for PET myocardial perfusion scans. Their study shows that  
187 for the assessment of qualitative and quantitative myocardial blood flow (MBF) with a hybrid  
188 PET/CT scanner, the use of CT with a tube current of 10 mA, instead of Ge-68, can provide  
189 accurate results for attenuation correction in PET images. However, the effective doses for the CT  
190 using tube current of 80 mA and 10 mA are 0.38 mSv and 0.05 mSv respectively. Thus, the  
191 radiation dose can be lowed by 8-fold by using tube current of 10 mA as compared to 80 mA  
192 without compromising diagnostic information. On the other hand, Michael Souvatzoglou *et al.*[15]  
193 assessed the image quality and radiation dose for 3 different CT scans: (1) LDCT(low dose CT) :  
194 120 kV , 26 mA; (2) SCT(slow CT) : 120 kV , 99 mA; (3) ULDCT(ultra low dose CT) : 80 kV ,  
195 13 mA for the attenuation correction in cardiac PET/CT scans. Their results showed that all 3 types  
196 of CT can provide PET images with good image quality and the ULDCT showed the least radiation  
197 dose. It is thus feasible to use 80 kV, a value that is lower than that used in the clinical setting, for  
198 CT attenuation correction purpose for dose reduction. Although using ULDCT as attenuation  
199 correction does not affect the diagnosis in the PET images, our results showed that there are still  
200 some minor discrepancies between ULDCT and default CT in terms of MI and CC analysis. It is  
201 probably because CT images are usually a snap shot of a respiration cycle, while PET scans are  
202 results of average of respiration cycles. This temporal difference between the scans often introduces

203 misalignment artifacts in PET images. These artifacts will cause registration errors and affect the  
204 quantitative results.

205 In conclusion, this study demonstrated that radiation dose in cardiac PET/CT can be  
206 significantly reduced without degrading imaging quality. The proposed imaging protocol, with  
207 radiation dose reduction up to 47.9% for an integrated cardiac PET/CT coronary artery examination,  
208 should be able to provide unique insight into diagnosis and management of coronary heart disease.

209

210 **Reference**

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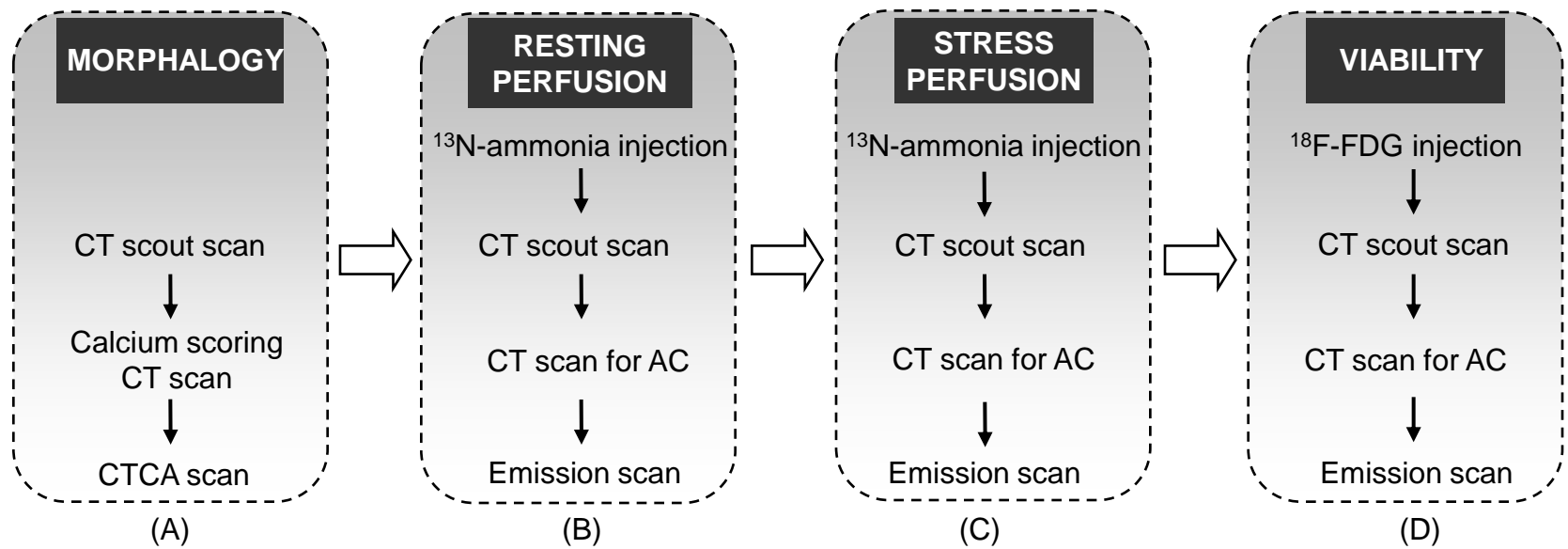


Fig.1 Flowchart of cardiac PET/CT scanning procedures, including cardiac CT imaging (A), cardiac perfusion PET imaging at rest (B) and under stress (C), and FDG PET for myocardial viability (D).

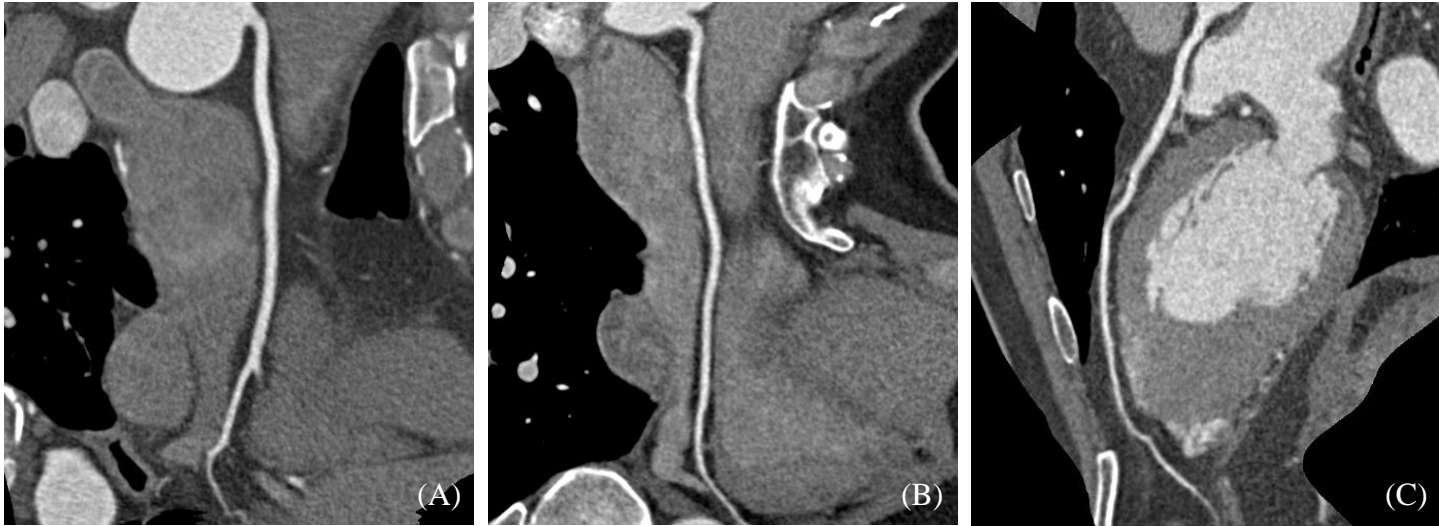


Fig. 2. Curved multiplanar reconstruction images show no motion artifact (score 1) of the RCA, reconstructed from RGH (A), ETCM (B) and PGA (C) acquisitions.

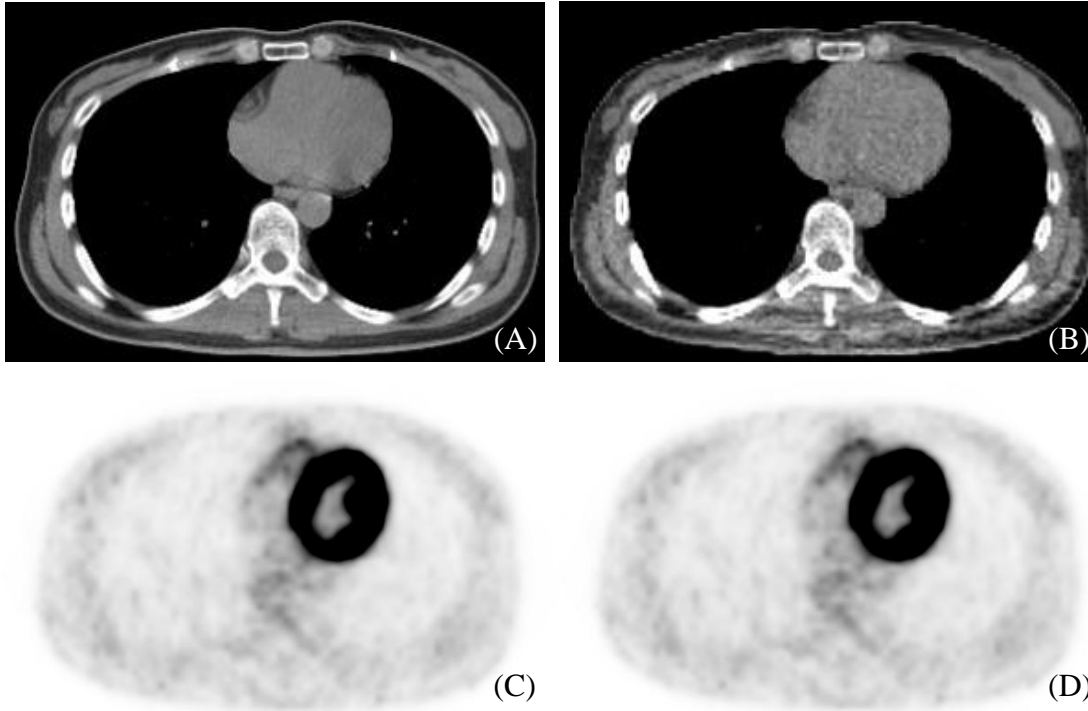


Fig. 3. CT images for CT-based attenuation correction obtained using default scan parameters (A) and ULDCT (B). FDG-PET images with attenuation correction using CT images in (A) and (B) are shown in (C) and (D), respectively.



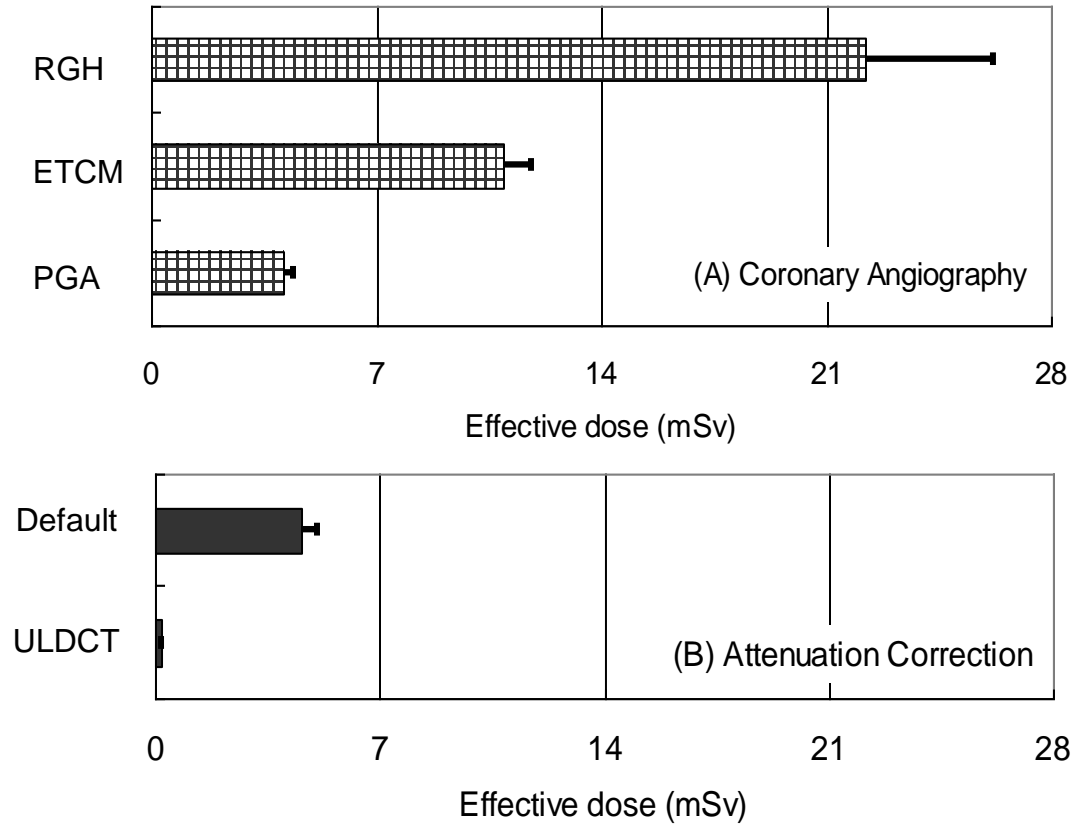


Fig. 4. Effective dose obtained from CT scans for CT coronary angiography (A) and CT-based attenuation correction (B).

Table 1. Parameters of the CT scanning protocols for coronary angiography and CT-based attenuation correction

	Coronary Angiography			Attenuation Correction	
	RGH <sup>a</sup>	ETCM <sup>b</sup>	PGA <sup>c</sup>	Default	ULDCT <sup>d</sup>
Tube voltage (kV)	120	120	120	120	80
Tube current (mA)	624.5±34.65	633±28.58	639.2±40.99	20-210 <sup>e</sup>	10
Rotation time (ms)	350	350	350	500	500
Pitch <sup>f</sup>	0.18-0.22	0.18-0.22	1	0.516	0.516
Acquisition window	0%-100%	30%-80%	75% ±5%	NA	NA

<sup>a</sup>RGH = retrospectively gated helical

<sup>b</sup>ETCM = electrocardiogram tube current modulation

<sup>c</sup>PGA = prospectively gated axial

<sup>d</sup>ULDCT = ultra low dose CT

<sup>e</sup>Variable tube current was delivered based on patient's scout image to maintain an appropriate balance between image quality and radiation dose (Smart mA).

<sup>f</sup>Pitch depends on the patient's heart rate.