

## Mini Review

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# Double-Low Dose Protocol of Computed Tomography Pulmonary Angiography (CTPA) in the Diagnosis of Pulmonary Embolism: A Feasible Approach for Reduction of Both Contrast Medium and Radiation Doses

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### ABSTRACT

This article discusses a hot topic on double low-dose protocol of computed tomography pulmonary angiography (CTPA) in the diagnosis of pulmonary embolism, with a special focus on a recent paper published in the British Journal of Radiology about the feasibility of this technique. Three aspects will be discussed in this review: First, both kVp and contrast medium can be reduced without affecting image quality when compared to the standard CTPA protocol; second, a low-pitch protocol is comparable to the high-pitch spiral image acquisition with similar image quality achieved, but at lower radiation dose; and finally, the double low-dose CTPA protocol can achieve diagnostic images in patients with body mass index up to 35 kg/m<sup>2</sup>.

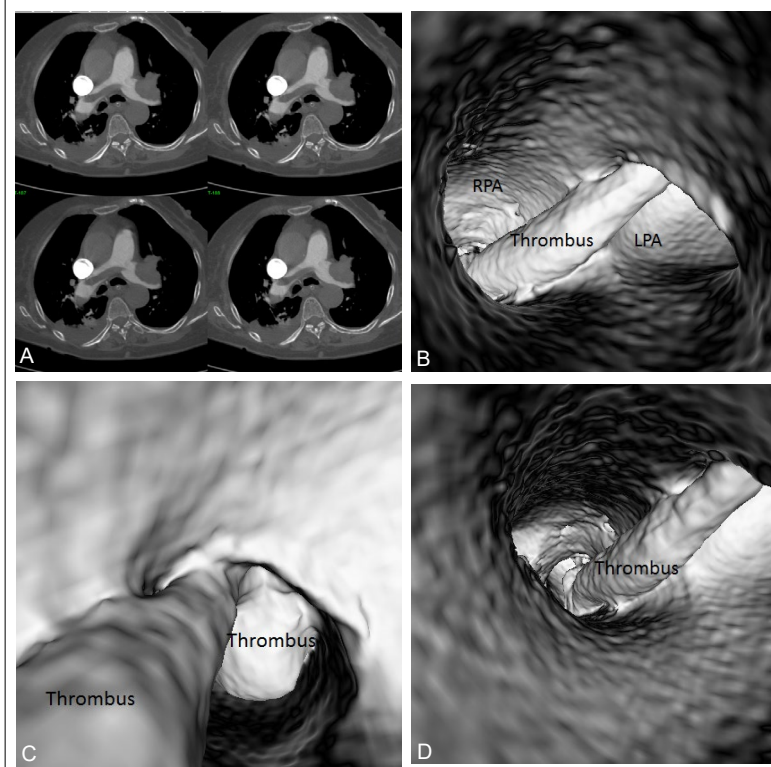
**KEYWORDS:** Computed tomography pulmonary angiography; Contrast medium; Optimization; Image quality; Pulmonary embolism; Radiation dose.

**ABBREVIATIONS:** CTPA: Computed Tomography Pulmonary Angiography; PE: Pulmonary Embolism; VIE: Virtual Intravascular Endoscopy; CIN: Contrast-Induced Nephropathy; BMI: Body Mass Index; CNR: Contrast-to-noise ratio.

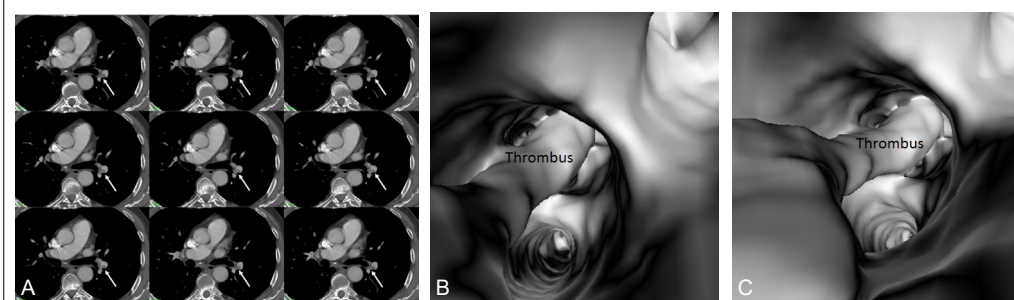
Computed tomography pulmonary angiography (CTPA) has become the method of choice for the diagnostic assessment of patients with suspected pulmonary embolism due to technological advancements in CT imaging.<sup>1-8</sup> Modern CT scanners with superior spatial and temporal resolution enable detection of segmental and subsegmental thrombus in the pulmonary arteries with high accuracy.<sup>9,10</sup> In addition to 2D axial images, CTPA allows for generation of different image reconstructions including unique intraluminal views of the thrombus in the pulmonary arteries (Figures 1 and 2).<sup>9</sup> Increased detection of pulmonary embolism seems to be associated with increased use of CTPA, in particular in the emergency department.<sup>11-17</sup>

Although there are no guidelines available about the minimum acceptable yield of CTPA, it is generally agreed that a diagnostic yield of CTPA less than 10% indicates overuse of CTPA as a diagnostic tool.<sup>18,19</sup> Sharma and Lucas recently reported their single center experience of CTPA in the diagnosis of pulmonary embolism over a period of 8 years. Authors found a direct correlation between increased number of CTPA scans and percentage of positive pulmonary embolism (the positive diagnostic yield ranges from 12% to 28.1%), suggesting that the use of CTPA is clinically appropriate.<sup>11</sup> Mountain and colleagues in their multi-center study showed similar findings with association of increasing use of CTPA with increased rates of

**Figure 1:** CT Pulmonary Angiography Shows Multiple Pulmonary Emboli in an 85-Year-Old man. A: 2D Axial Images Show Large Thrombus Formation in the Right and Left Main Pulmonary Arteries with Filling Defect Observed. B: 3D Virtual Intravascular Endoscopy (VIE) Shows Large Thrombus Involving Left Pulmonary Artery (LPA) and Right Pulmonary Artery (RPA). C: Close VIE Visualization of the Thrombus in the LPA. D: Close VIE View of the Thrombus Extends to the RPA.



**Figure 2:** CT Pulmonary Angiography Shows Pulmonary Embolism in a 65-Year-Old Man. A: 2D Axial Images Demonstrate Thrombus in the Left Lower Lobar Artery (arrows). B and C: 3D Virtual Intravascular Endoscopy Offers Intraluminal Views of the Thrombus in the Left Lower Lobar Artery in Relation to the Surrounding Branch Artery.



pulmonary embolism diagnosis. Their study involved 14 clinical sites (15 emergency departments) across Australia and New Zealand consisting of more than 7000 CTPA scans with >94% performed on  $\geq 64$ -slice CT. The overall diagnostic yield of CTPA at these 14 clinical sites was 14.5% (range: 9.3 to 25.3%) with significant variations in the diagnostic yield among the clinical sites. Of these clinical centers, four sites were found to have significantly lower yield which is less than the acceptable rate of 15.3%. Despite the yield variation in this multi-center study, increased use of CTPA is significantly correlated with pulmonary embolism (PE) diagnosis, thus, justifying its clinical value.<sup>17</sup>

The increased use of CTPA raises concerns about subjecting patients to excessive radiation exposure and contrast-induced nephropathy (CIN).<sup>7,13,20</sup> CT is a well-recognized modality with high radiation dose, although significant dose reduction has been achieved in recent years with implementation of many dose-reduction strategies. This has been widely reported in cardiovascular CT imaging, such as coronary CT angiography,<sup>21-24</sup> abdominal aortic imaging,<sup>25,26</sup> and CTPA.<sup>27-30</sup> Similarly, CIN has been drawing more attention in the recent literature due to routine use of contrast medium injection during CT scans including CTPA. Thus, reduction of contrast medium volume is equally

important as reduction of radiation dose, which is shown in some studies.<sup>24,31</sup> This leads to the currently recommended protocol of double low-dose CTPA protocol, which has been addressed in a recent study supported by other relevant reports.

Boos et al<sup>31</sup> in their study investigated the feasibility of using double low-dose CTPA protocol in 70 patients with suspected pulmonary embolism. They implemented a new developed scanning protocol defined as a 70-kVp simultaneous acquisition dual-source CTPA in Group A comprising 35 patients with body mass index (BMI) less than 35 kg/m<sup>2</sup>. With this protocol, the two X-ray tubes were operated at 70-kVp with a low-pitch of 0.9. Automatic tube current modulation was used with administration of 40 ml of contrast medium followed by a saline flush of 40 ml at a flow rate of 3 ml/sec. The new protocol group was compared to a control group, Group B consisting of another 35 patients with similar demographics in age, gender and BMI. In Group B, CTPA was performed with a high-pitch spiral acquisition mode with use of automatic tube current and automatic tube potential selection. Of 35 patients in Group B, 100 kVp and 120 kVp was used in 6 and 29 patients, respectively, with administration of 70 ml of contrast medium followed by the saline flush protocol as used in Group A. Images were reconstructed with a medium level of iterative reconstruction (Level 3, SAFIRE: sinogram affirmed iterative reconstruction). Quantitative assessment of image quality was based on signal-to-noise ratio (SNR) and contrast-to-noise ratio (CNR), while qualitative analysis of image quality was assessed by two independent observers using a four-point scoring scale with score of 1 indicating non-diagnostic, and score of 4 excellent image quality. No significant differences were found in both qualitative and quantitative assessment of image quality between the two groups; however, both radiation dose and contrast medium dose were significantly lower in Group A than in Group B, with corresponding reduction of 50% and 40% achieved. Details of comparative results between the two groups are shown in the Table 1.

There are three observations from Boos study that bear discussions. First, the double low-dose CTPA is feasible in acquiring images for diagnostic assessment of pulmonary embolism with use of low kVp and low contrast medium. Of various dose-reduction strategies, low kVp is highly recommended and it is widely used in many CT angiographic applications, in particular, in cardiovascular CT imaging area.<sup>32,33</sup> According to the society of cardiovascular computed tomography guidelines, selection of kVp is correlated with patient's BMI, with low kVp (100 or 80 kVp) used in patients with smaller BMI.<sup>34</sup> Reduction of kVp shows particular value in CT angiography because of the association between vessel visualization and iodine enhancement. This is confirmed in Boos' study by showing an increase in vascular CT attenuation when kVp is reduced. The CT attenuation measured in main pulmonary trunk and left lower segmental pulmonary artery was found to be significantly higher in Group A than that in Group B (414.3±149.4 HU and 416.4±139.3 vs. 259.6±69.7 and 256.0±75.0 HU,  $p<0.0001$ ) with no significant difference in signal-to-noise ratio (SNR) and contrast-to-noise-ratio (CNR) between the two groups ( $p>0.05$ ). Most of the studies tested lowering kVp value with 80 or 100 kVp on CTPA examinations while achieving diagnostic images with low radiation dose,<sup>35-40</sup> while Boos and colleagues in their study further lowered the kVp to 70 without affecting image quality. It is well known that lowering tube voltage is associated with increased image noise, but this can be compensated by iterative reconstruction, which is confirmed by Boos study.

Contrast medium has detrimental effects on renal function, thus, reduction of contrast medium during CT angiography has attracted increasing attention in the literature. Previous protocols using 80-100 ml contrast medium followed by 30-60 ml saline flush are being replaced by low volume of contrast medium, as shown in Boos and others' studies. Further lowering contrast volume to even 40 or 20 ml has also been reported in some studies Saade et al<sup>40</sup> compared two groups of patients

**Table 1:** Summary of Scanning Protocols and Results in Boos' Study.<sup>31</sup>

Parameters	Group A 35 patients	Group B 35 patients
Data acquisition mode	Simultaneous dual-source protocol	Dual-source, high-pitch helical protocol
Pitch value	0.9	2.2
kVp	70	120 kVp in 29 patients, 100 kVp in 6 patients
mAs	269±91 (range: 100-397)	144±35 (range: 82-211)
Volume of contrast medium (ml)	40	70
Body mass index (kg/m <sup>2</sup> )	26.8±3.9	26.8±4.2
Effective radiation dose (mSv)	2.0±0.6	3.9±1.1
Subjective image quality assessment	3.7±0.6	3.7±0.6
SNR <sub>Trunk</sub>	14.6±6.0	13.9±3.7
SNR <sub>LLSA</sub>	15.1±8.9	12.0±4.5
CNR <sub>Trunk</sub>	12.4±5.7	11.6±3.3
CNR <sub>LLSA</sub>	12.9±8.5	10.0±4.1

SNR: Signal-to-noise ratio, CNR: Contrast-to-noise ratio, LLSA: Left lower segmental pulmonary artery, Trunk: Pulmonary trunk.

undergoing CTPA with mean contrast volume being 33 ml and 29 ml, respectively. Their results showed significant improvement in the visualization of pulmonary vessels with use of low contrast medium. Another study by Lu et al<sup>30</sup> reported their experience of further lowering the contrast medium to 20 ml with tube voltage of 80 kVp and high-pitch protocol. Of 100 patients with suspected pulmonary embolism, 50 patients were scanned with the CTPA protocol of 100 kVp, pitch of 1.2 and 60 ml of contrast medium, while another 50 patients with the protocol of 80 kVp, pitch of 2.2 and 20 ml of contrast. Comparable image quality and diagnostic accuracy was found between two groups with no significant differences, while the double low-dose protocol using 20 ml contrast medium resulted in 50% radiation dose reduction.

Second, use of a low-pitch CTPA protocol is not associated with higher radiation dose when compared to the high-pitch mode. Pitch value has a direct impact on radiation dose as it is traditionally believed that higher the pitch, lower the radiation dose. However, increasing pitch to a higher level is not recommended due to increase in image noise resulting from suboptimal spatial resolution. With latest CT scanners the potentially high radiation dose associated with a low-pitch protocol could be countered by reducing the tube current time product (mAs), thus improving image quality by decreasing the image noise. This is observed in Boos' study. The mean mAs in Group A was significantly higher than that in Group B (268±91 mAs vs. 144±35 mAs,  $p < 0.0001$ ). Despite higher mAs being used in Group A, the 70 kVp protocol led to lower radiation dose with improved image quality of pulmonary arteries. High-pitch and low kVp (80 kVp) protocol has been reported in a number of studies demonstrating the further dose reduction to less than 1 mSv without compromising diagnostic image quality.<sup>30,37,38,41</sup> However, lowering kVp and increasing pitch level will increase image noise, thus may impair diagnostic performance of CTPA, hence, it is not widely implemented in clinical practice. It has been reported that better image quality was achieved with a pitch of 2.0 when compared to the CTPA protocol using a pitch of 3.0.<sup>42</sup> The mean effective dose was 2.0 mSv and 3.9 mSv for double low-dose and high-pitch CTPA protocols in Boos study, and this is higher than that reported in Lu's study which used high-pitch CTPA protocol in normal weight patients. Given the improved image quality of pulmonary vasculature, even in large patients with BMI up to 35 kg/m<sup>2</sup>, the radiation dose of 2.0 mSv is acceptable from a clinical perspective.

Finally, the double low-dose protocol of CTPA allows for acquisition of images in patients with large BMI, according to this study. BMI is one of the main factors that should be considered during CTPA as in most of the situations, kVp and mAs are adjusted based on patient's BMI because of its impact on the radiation dose and image quality. Of these reported double low-dose CTPA studies, most of them were performed in patients with normal BMI (mean value <25 kg/m<sup>2</sup>).<sup>30,35</sup> In contrast, Boos and colleagues presented diagnostic quality images in large patients as well, although they did not include extremely obese pa-

tients. The limitations of previous studies including only normal sized patients or with missing data on BMI<sup>43</sup> have been overcome by Boos' study, with findings offering additional value to the current literature.

In summary, Boos and colleagues in their study have demonstrated the feasibility of using double low-dose protocol comprising 70 kVp and 40 ml of contrast medium during CT pulmonary angiography. This protocol leads to significant reduction of radiation dose and contrast medium dose when compared to the high-pitch spiral dual-source CT pulmonary angiography. In addition, this study also confirms that low-dose CT pulmonary angiography is able to produce diagnostic images in patients with body mass index up to 35 kg/m<sup>2</sup>. Due to limited small sample size in this study, further research based on a large cohort of patients with assessment of both image quality and diagnostic accuracy is warranted.

#### CONFLICTS OF INTEREST

The authors declare that they have no conflicts of interest.

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