Reliability of virtual non-contrast computed tomography angiography: comparing it with the real deal

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Abstract

Background: Computed tomographic angiography (CTA) may require a non-contrast enhanced dataset for the diagnostic workup. By using dual-energy acquisition, it is possible to obtain a virtual non-contrast-enhanced (VNC) dataset, thereby possibly eliminating the non-contrast acquisition and reducing the radiation dose.

Purpose: To compare image quality of VNC images reconstructed from arterial phase dual-energy CTA to true non-contrast (TNC) images, and to assess whether VNC images were of sufficient quality to replace TNC images.

Material and methods: Thirty consecutive patients with suspected abdominal aortic aneurysm, aortic dissection, or subacute control after EVAR/TEVAR were examined with dual-energy CT (DECT). The examination protocol included a single-energy TNC, DECT arterial phase (80 kV/Sn140 kV), and single-energy in venous phase of the aorta. A VNC dataset was obtained from the DE acquisition from arterial phase scans. Mean attenuation and image noise were measured within regions of interest at three levels in the aorta in TNC and VNC images. Comparison of the TNC and VNC images for artefacts was made side-by-side. Subjective evaluation included overall image quality on a 4-grade scale, and quantitative analysis of algorithm-induced artefacts by two experienced readers.

Results: For all cases, the aortic attenuation was significantly higher at VNC than at TNC. Image noise measured quantitatively was also significantly higher at VNC than at TNC. Subjective image quality was lower for VNC (mean = 3.1 for VNC, 3.7 = for TNC) but there were no cases rated non-diagnostic.

Conclusion: VNC images based on arterial phase CTA have significantly higher mean attenuation and higher noise levels compared to TNC.

Keywords

Computed tomography, CT, dual-energy CT, virtual non-contrast, CT angiography, attenuation values

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Introduction

Computed tomography angiography (CTA) is commonly used for imaging of acute and chronic aortic disease. Endovascular aneurysm repair (EVAR) may require life-long follow-up by CTA to diagnose complications such as endoleaks or stentgraft migrations. For EVAR follow-up, a triple phase examination protocol including non-contrast medium (unenhanced) phase and contrast medium (enhanced) early and delayed venous phase acquisitions is the mainstay for followup (1,2), inducing a substantial radiation dose to the patients, although there is no generally accepted ¹Department of Clinical Sciences, Lund University, Lund, Sweden
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Creative Commons Non Commercial CC BY-NC: This article is distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 License (http:// www.creativecommons.org/licenses/by-nc/4.0/) which permits non-commercial use, reproduction and distribution of the work without further permission provided the original work is attributed as specified on the SAGE and Open Access page (https://us.sagepub.com/en-us/nam/open-access-at-sage). consensus as to which imaging strategy would be most advantageous. One method to reduce radiation dose would be to use a two-phase (unenhanced and splitbolus mixed arterial and venous phase) protocol instead of a three-phase (unenhanced, arterial and venous phase) protocol. Javor et al. has used a single acquisition with split contrast bolus with timing acquisition in both venous and arterial phases when replacing TNC images with VNC images (3).

Dual-energy CT (DECT) utilizes the different attenuation properties between materials or tissues for different X-ray energies. The differences are more pronounced in materials with high atomic numbers, such as iodine and calcium, due to their strong photoelectric effect causing high attenuation at lower photon energies. Water and muscle have similar attenuation at different photon energies and can therefore function as reference materials (4,5). The DE technique has several applications for clinical imaging and can allow reduction of the iodine contrast medium dose, as well as the radiation dose (6,7).

Imaging of arteries may benefit greatly from DECT. Using contrast-phase acquisition with two different X-ray tube peak voltages, where the lower peak voltage enables greater conspicuity of vessel attenuation, contrast medium can be reduced significantly with preserved image quality (8,9). The differences in atomic number between calcium and iodine make them readily separable with use of DE techniques. Using post-processing algorithms, it is possible to identify and remove bone structures, and also identify and remove the iodine content, and thereby obtain a reconstructed image comparative to an unenhanced dataset, virtual non-contrast (VNC) images, without having to do true non-contrast (TNC) acquisitions (10,11).

It has not been well established if the VNC is of acceptable quality to replace the TNC in aortic CTA (12).

The purpose of this study was to assess whether VNC images reconstructed from arterial phase DECTA were of sufficient quality to replace TNC images to reduce the radiation dose to the patients.

Material and Methods

Patient population

Thirty consecutive patients (15 men, 15 women, median age = 65 years) examined by DECT for suspected acute aortic disease (25 patients) or complications after abdominal EVAR (five patients) were included in the study. The study was approved by the local ethics committee (#2014/811). Patients with a body mass index (BMI) > 35 kg/m² were excluded because of the presumed insufficient transmission of 80 peak kilo voltage (kVp) quanta. Patients with impaired renal

function (glomerular filtration rate <45 mL/min) were examined with a reduced contrast medium dose. Patient characteristics are shown in Table 1.

CT parameters and radiation dose

All examinations were performed on a dual source CT scanner (Siemens Somatom Definition Flash, Siemens Healthcare, Forchheim, Germany) equipped with two X-ray tubes mounted in the gantry at an angle of 95° and two 64-channel detectors. The maximum fields of view (FOV) were 50 cm and 33 cm, respectively. The TNC series were acquired with a tube potential of 120 kVp and a quality reference tube current of 140 mAs. The VNC series were acquired with tube A potential of 80 kVp and tube B of Sn140 kVp. The quality reference tube current values were 167 mAs and 83 mAs for tube A and tube B, respectively (Table 2). Automatic exposure control (CareDose 4DTM, Siemens Healthcare, Forchheim, Germany) was used to adept the tube current to variations in patient attenuation, both between different patients and within any given patient.

 Table 1. Basic characteristics of the study patients.

Variables	
Patients (n)	30
Age (years)	65 (37–89)
Weight (kg)	77 (53–126)
Height (cm)	171 (158–193)
Body mass index (kg/m ²)	26 (19–34)
Plasma creatinine (µmol/L)	81 (46–122)

Median values (2.5 and 97.5 percentiles) are given unless otherwise stated.

 Table 2. CT scanning parameters of true non-contrast (TNC)

 acquisition and virtual non-contrast (VNC) reconstructions.

Single-energy non-contrast phase for TNC	Dual-energy early arterial contrast phase for VNC
140	167/83
120	80/Sn140
0.85	0.85
0.5	0.33
128×0.6	128×0.6
38.4	38.4
130f	D26f
512×512	512×512
5	5
	non-contrast phase for TNC 140 120 0.85 0.5 128 × 0.6 38.4 130f 512 × 512

The examination was performed in the supine position in cranio-caudal direction at breath-hold from the thoracic aperture to the groins. All scanning parameters regarding the non-unenhanced phase for TNC analyses and the early arterial enhanced phase for VNC analyses are listed in Table 2. Individual contrast media volume and injection rates were calculated based on contrast media dose per kg body weight (maximum dose weight = 80 kg) at a dose of 340 mg I/kg and median volume of 80 mL (range = 38-99 mL). The early arterial-phase scan was acquired after initiation of the intravenous injection of contrast media (Omnipaque 300 mg I/mL, GE Healthcare, Oslo, Norway) with a fixed injection time of 15s followed by a 50-mL saline chaser at 4 mL/s. Bolus-tracking with a threshold of 120 HU at the level of the renal arteries was used.

For both acquisitions, effective dose (ED) was calculated from the dose-length product (DLP) registered by the CT scanner and multiplied by the mean of the ED/DLP conversion factor for chest/abdomen/ pelvis ($0.186 \text{ mSv}/(\text{mGy} \times \text{cm})$) based on the IRCP 103 tissue weighting factors (13). The effective dose of a biphasic unenhanced and enhanced protocol was compared with a single-phase protocol, i.e. the arterial phase only.

True and virtual non-contrast image reconstruction

TNC images were reconstructed with a slice thickness/ increment of 5/5 mm. The VNC dataset was obtained by a patented algorithm based on the dataset from the two X-ray tubes with different X-ray energies. A specific dual-energy post-processing workstation with a patented algorithm (Syngo MMWP version VA 20; Siemens Healthcare, Forchheim, Germany) was used for image analysis, generating VNC images from the arterial-phase acquisition.

Quantitative image quality analysis

Comparison of attenuation and noise between TNC and VNC images was made by drawing circular regions of interest (ROI) within the aortic lumen at the levels of the hemidiaphragms, the renal arteries, and the aortic bifurcation on a PACS workstation (IDS7, Sectra Imtec AB, Linköping, Sweden). The ROIs were made as large as possible while avoiding calcifications, plaques, and stent material. Mean attenuation and image noise (one standard deviation) in Hounsfield Units (HU) were registered.

Subjective image quality analysis

Subjective evaluation was done in two steps. First, the TNC and VNC images were evaluated side-by-side for

artifacts such as residual contrast medium, subtraction of calcifications, or stent structures. Four months later, the second evaluation was done, in which the subjective image quality was scored at a fixed window setting (level/width = 200/650 HU) in randomized order by two senior consultant interventional radiologists (JW with ten years of CTA experience and LL with nine years of CTA experience) blinded to patient information and examination protocol on a PACS workstation using the viewing and scoring software ViewDEX v2.0 (Viewer for Digital Evaluation of X-ray images) (14.15). Image quality was evaluated for overall image quality, signs of residual contrast medium, possible subtracted calcifications, and stent structures. A fourpoint scale was used for overall diagnostic quality (4 = excellent, 3 = good, 2 = moderate but sufficient for diagnosis, and 1 =non-diagnostic). The specific artefacts were scored as yes/no.

Statistical analysis

Statistical analysis was performed using statistical software SPSS software (version 24.0; SPSS Inc., Chicago, IL, USA). The Mann–Whitney U test was used to compare differences between the TNC and the VNC datasets regarding vascular attenuation, noise, and radiation dose. The Wilcoxon signed-rank test was used to compare subjective evaluation of image quality. A *P* value < 0.05 was considered significant. Inter-observer agreement regarding overall image quality was assessed using intra-class correlation coefficient (ICC).

Results

VNC images were compared to images from an actual unenhanced dataset (TNC) in 30 consecutive patients examined for suspected acute aortic disease or for elective EVAR follow-up. All examinations were successful without any adverse events. Although the 80-kVp dataset had a limited FOV of 33 cm, the entire aorta and the iliac arteries were within the FOV in all cases.

Radiation data

Radiation data are shown in the Table 3. The mean scanning length for the TNC acquisition was 67 cm and the calculated mean effective dose was 9.0 mSv. The mean scanning length for the arterial phase DE acquisition was 64 cm and the calculated mean effective dose was 8.4 mSv. A biphasic protocol would thus result in an average dose of 17.4 mSv. Thus, a single DE scan with VNC reconstruction would have resulted in a mean effective dose reduction of 52%.

 Table 3. Radiation parameters of true non-contrast (TNC)

 acquisition and virtual non-contrast (VNC) reconstructions.

Variables	TNC	VNC	
Scan time (s)	6 (5–7)	4 (4–7)	
Scan length (cm)	67 (55–77)	64 (55–73)	
CTDI _{vol} (mGy)	7.2 (4.3–11.3)	6.5 (5.6–9.6)	
Dose-length product (mGy \times cm)	492 (262–875)	460 (380–760)	
Effective dose (mSv)	9.0 (4.8–16.1)	8.4 (7.0–13.9)	

Median values (2.5 and 97.5 percentiles) are given unless otherwise stated.

 $\mathsf{CTDI}_{\mathsf{vol}},$ volume computed tomographic dose index; mGy, milligray; mSv, millisievert.

Table 4. Attenuation (HU) and image noise (I SD, HU) of the aortic lumen of true non-contrast (TNC) acquisition and virtual non-contrast (VNC) reconstructions.

Aortic level	TNC	VNC	P value
Hemidiaphragms			
Attenuation	38 (26–48)	80 (55-120)	<0.001
Image noise	5 (- 9)	23 (12–33)	<0.001
Renal arteries			
Attenuation	45 (36–55)	79 (52–123)	<0.001
lmage noise	14 (10–24)	22 (12–36)	<0.001
Aortic bifurcation			
Attenuation	46 (39–53)	67 (43–118)	<0.001
Image noise	14 (9–24)	22 (12–33)	<0.001

Median values (2.5 and 97.5 percentiles) are given.

HU, Hounsfield units; SD, standard deviation.

Quantitative image quality

Attenuation and image noise of the TNC and VNC images are listed in Table 4. At all aortic levels, VNC had a significant higher attenuation (P < 0.001) and image noise (P < 0.001).

When comparing VNC images to the TNC images, side-by-side artifacts in major thoracic vessels were seen in VNC images (Fig. 1, Pulsation). Removal of minor arterial wall calcifications was seen in 22 out of 30 VNC images. In all cases, the removed calcifications were in the range of 1–3 mm and located at the aortic or iliac wall (Fig. 1, Calcification). Partially removed stent structures were seen in the VNC images in two cases. The removed stent structures were seen in the VNC images in two cases. The removed stent graft, respectively (Fig. 1, Stent). In five cases, incomplete elimination of the aortic contrast medium attenuation was seen in the VNC images (Fig. 1, Contrast medium) where three of the patients were women and two were men. These five patients had

BMI of 26, 24, 33, 26, and 33 kg/m^2 . One of these patients was operated with fenestrated EVAR and four were referred to the study because of suspected aortic rupture. No obvious reason could be found during retrospective investigation.

Subjective image quality

All TNC images were rated good or excellent (the mean image quality score was 3.6 for reader 1 and 3.7 for reader 2). All VNC images were rated moderate to excellent (the mean image quality score was 3.0 for reader 1 and 3.2 for reader 2). There was a significant (P < 0.001) difference in subjective image quality between TNC and VNC datasets. For the overall image quality, the inter-observer agreement was moderate with an average ICC value of 0.743.

Discussion

This prospective study based on 30 consecutive aortic DECT scans was carried out to compare the image quality of TNC to VNC images including attenuation and noise, as well as estimating the potential reduction in radiation dose. Throughout the study, we used a standardized DECT protocol to maintain image quality at a constant level.

Several authors have reported acceptable quality VNC images as acceptable replacements for the TNC images in the abdominal aorta (16,17) while in our study we found aortic attenuation was considerably higher in all VNC images compared to TNC images throughout the acquisitions. The difference was more pronounced in the proximal part of the aorta and less pronounced in the distal part of aorta and the iliac arteries. This attenuation difference between TNC and VNC was more pronounced in our material than has previously been reported (11,12,16). There are several possible explanations to this difference. We used tube voltages of 80/Sn140 kVp compared to 100/Sn140 kVp in previous studies (12,16). Compared to the study of Toepker et al. (16), in which a total of 110 mL Iomeron 400 mg I/mL contrast medium was used, we used only 80 mL Omnipaque 300 mg I/mL contrast medium.

In the thoracic aorta, close to the heart incomplete elimination of contrast media was seen, maybe caused by the pulsating movement of the aortic and pulmonary artery walls leading to a minimal spatial difference between tube A and tube B, thereby affecting the output of the VNC algorithm. This may be of clinical relevance if mistaken for a dissection or an intramural hematoma. Five moderate ratings were caused by such artefacts in the thoracic aorta. This could possibly be avoided by using electrocardiogram (ECG) gating. In 'two patients, a partial subtraction of stents occurred

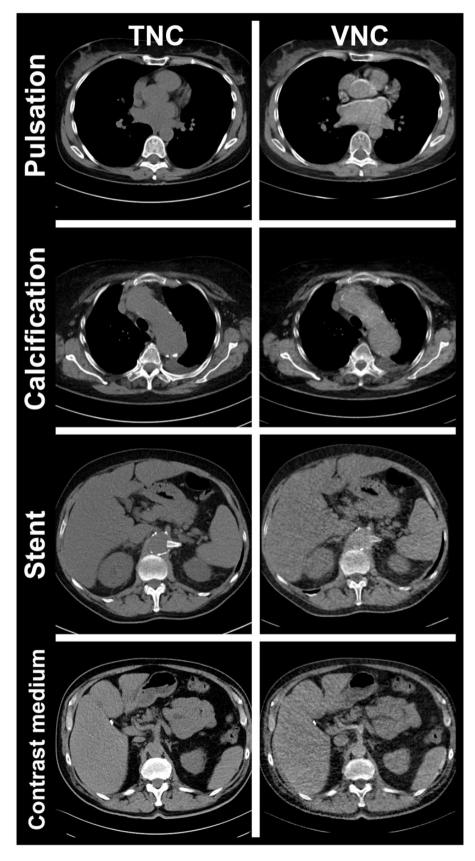


Fig. 1. Pulsation: Attenuation artifacts in main thoracic vessels in the VNC image. Calcification: Elimination of the small calcifications in the aortic wall in the TNC image. Elimination of the renal stent in the VNC image. Stent: A partly removed renal stent. Contrast medium: Incomplete elimination of the contrast media in the VNC image on the right in the abdominal aorta.

in the VNC images, resulting in diagnostic uncertainty, and rating as suboptimal image quality.

In 24 of 30 cases, some structures were removed by the VNC algorithm; in 22 cases, there were minor calcifications in the aortic wall. Subtraction of calcium from DECT data represents an in-built technical limitation of the VNC algorithm (18). This finding could represent an important limitation when the presence of calcification in aortic thrombus plays an important role in endoleak recognition. In two cases, minor parts of stents were partially subtracted which can be misinterpreted as a fracture or damage of the stent, the problem seen in some previous studies (19). One moderate rating was seen in a patient with a BMI of 35 kg/m^2 , resulting in grainy images and limited coverage of the abdominal circumference showing the 80-kVp ring all over the abdomen interfering with evaluation. Dose modulation by CareDose 4D was not fully compensated in patients with high BMI resulting in more image noise.

The present study is subjected to several limitations. The number of patients was relatively small and patient population was heterogeneous in terms of age and study indication.

In conclusion, VNC images based on arterial phase CTA have significantly higher attenuation values at the aorta and higher noise levels compared to TNC. According to our study, VNC images are not suitable as replacements for TNC images generally but can be considered when radiation dose reduction is definitely desirable or when no TNC acquisition is available.

Declaration of conflicting interests

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