Low-dose computed tomography with 4th-generation iterative reconstruction algorithm in assessment of oncologic patients

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Abstract

AIM
To compare radiation dose and image quality of low-dose computed tomography (CT) protocol combined with hybrid-iterative reconstruction algorithm with standard-dose CT examinations for follow-up of oncologic patients.

METHODS
Fifty-one patients with known malignant diseases which underwent, during clinical follow-up, both standard-dose and low-dose whole-body CT scans were enrolled. Low-dose CT was performed on 256-row scanner, with 120 kV and automated mA modulation, and iterative reconstruction algorithm. Standard-dose CT was performed on 16-rows scanner, with 120 kV, 200-400 mAs (depending on patient weight). We evaluated density values and signal-to-noise ratio, along with image noise (SD), sharpness and diagnostic quality with 4-point scale.
RESULTS
Density values in liver, spleen and aorta were higher in low-dose images (liver 112.55 HU vs 103.90 HU,  \( P < 0.001 \)), as SD values in liver and spleen (liver 16.81 vs 14.41). Volumetric-Computed-Tomographic-Dose-Index (CTDIvol) and Dose-Length-Product (DLP) were significantly lower in low-dose CT as compared to standard-dose (DLP 1025.6 mgY/cm vs 1429.2 mgY/cm,  \( P < 0.001 \)) with overall dose reduction of 28.9%. Qualitative analysis did not reveal significant differences in image noise and diagnostic quality.

CONCLUSION
Automatic tube-current modulation combined with hybrid-iterative algorithm allows radiation dose reduction of 28.9% without loss of diagnostic quality, being useful in reducing dose exposure in oncologic patients.

Key words: Computed tomography; Low-dose computed tomography; Tube current modulation; Oncologic imaging; Radiation dose

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Core tip: Introduction of new generation of multidetector computed tomography (MDCT) scanner allowed thin-collimation scanning and high spatial resolution, and reducing at same time the delivered radiation dose to patients by using new iterative reconstruction algorithm. This new mathematical model approach permits to reduce the radiation dose, especially in patients who undergo serial follow-up study for oncologic (staging and restaging) purpose. On these basis in our study we evaluated radiation dose and image quality of CT examinations in a population of oncologic patients undergoing follow-up examinations with a new generation MDCT scanner (256-rows) using automatic modulation of tube current and iterative reconstruction algorithm (DoseRight system).

INTRODUCTION
Since the advent of computed tomography (CT) and, more recently, the development of multidetector computed tomography (MDCT) technique, the number of CT scans have increased significantly[1,2]; according to National Council on Radiation Protection and Measurements (NCRP) report 160, in 2006 United States population was exposed to more than seven times ionizing radiation from medical procedures than in the early 1980’s, and CT contributes to nearly one-half of all this radiation imaging exposure[3,4]. This high number of CT examinations have raised concern because of the potential risk of radiation-induced malignancy[5].

In the clinical management of patients with malignant diseases CT, with other cross-sectional imaging techniques such as MRI and FDG-PET/CT, has a major role for initial diagnosis of the disease, for staging, and during follow-up to monitor response to treatment and evaluate disease remission or relapse[6]. CT of the chest, abdomen and pelvis has the ability to obtain a global evaluation of the patient and to depict both primary tumor and metastasis in most cases, more quickly, safely and accurately than other more invasive or less sensitive imaging techniques. On these bases, oncologic patients typically undergo multiple CT investigations during their clinical follow-up, and risks derived from repeated scans and exposure to ionizing radiation should be balanced against the benefits of diagnostic imaging: CT examinations must have a correct clinical justification, and every CT scan must be technically optimized to keep radiation doses as low as possible while providing requested diagnostic information[7,8].

To optimize radiation dose in CT various technological strategies can be applied: These include active management of individual acquisition parameters (number of phases, section thickness, peak voltage, tube current-time product, pitch) or using an automated exposure control system[9]. Recently, iterative reconstruction algorithms have been introduced to reduce image noise, allowing further reduction in radiation dose beyond levels previously achievable with filtered back projection reconstruction (FBP)[10]. iDose\(^{4}\) is a hybrid iterative algorithm that is able to reduce noise on both the raw and image data[11,12]. The purpose of our study was to evaluate the radiation dose and image quality of CT examinations in a population of oncologic patients undergoing follow-up examinations comparing a new generation MDCT scanner (256-rows), using automatic modulation of tube current and iterative reconstruction algorithm (DoseRight system and iDose\(^{4}\)), with a 16-MDCT scanner using standard dose protocol and FBP.

MATERIALS AND METHODS

Patients population
Institutional research board approval was obtained for this retrospective study with waiver of the requirement for written consent.

Between August 2013 and April 2015, 259 adult patients (> 18 years old) with known malignant diseases (32 lung, 9 colorectal, 3 ovarian, 3 stomach, 2 uterine, 1 non Hodgkin lymphoma and 1 testicular) underwent contrast enhanced chest-abdomen-pelvis CT examination in a single venous phase during their clinical follow-up on a 256-MDCT scanner with a protocol implemented in our institution from August 2013 (with automatic modulation of current tube and iDose\(^{4}\) reconstruction algorithm). These examinations were identified with a retrospective...
review of CT studies archived with PACS system (AGFA Diagnostic Software, Impax, version 6.4.0.3125; Agfa, Mortsel; Belgium). In this group we selected patients which had undergone also standard-dose contrast enhanced CT scan on a 16-MDCT in a different time. Fifty-one patients were finally selected as our study group and were categorized into four groups according to their weight (kg): 41-60 kg (group A), 61-80 kg (group B), 81-90 kg (group C) and > 90 kg (group D) (Table 1).

Table 1  Descriptive table of weight groups in the patients population (n = 51)

| Weight groups | No. of patients | M/F | Mean ± SD (yr) |
|---------------|----------------|-----|----------------|
| Group A (41-60 kg) | 8 | 3/5 | 70.5 ± 8.6 |
| Group B (61-80 kg) | 25 | 9/16 | 67.6 ± 11.5 |
| Group C (81-90 kg) | 13 | 11/2 | 71.8 ± 7.4 |
| Group D (> 90 kg) | 5 | 4/1 | 59.2 ± 10.1 |
| All patients | 51 | 34/18 | 68.3 ± 10.4 |

M: Male; F: Female.

All patients had undergone thorax-abdomen-pelvis CT examinations both on a 16-rows MDCT scanner (Brilliance, Philips Medical Systems, Eindhoven, The Netherlands) and on a 256-rows MDCT (iCT, Philips).

For both examinations, CT data were acquired after the intravenous bolus injection of non-ionic iodinated contrast material (Xenetix 350; Guerbet, Aulnay, France), injected using a 18-gauge catheter positioned into the antecubital vein at a rate of 3.5 mL/s, with image acquisition during portal venous phase (55-70 s after the initiation of the contrast bolus), in accordance with an institutionally defined protocol. The volume of contrast agent was calculated on the basis of the patient’s body weight, with total dose ranging from 80 to 130 mL, and it was followed by a saline flush of 50 mL of NaCl at 3.5 mL/s. The patients were instructed to hold their breath during scanning. All studies were started from the lung apices and proceeded in a cephalocaudal direction until ischial tuberosity, to include chest, abdomen and pelvis of the patients.

The technical parameters for scanning included: (1) for 16-rows CT scanner: 120 kVp, mAs depending on patients’ weight (41-60 kg: 200 mAs; 61-80 kg: 300 mAs; 81-90 kg: 350 mAs; > 90 kg: 400 mAs), section thickness 2 mm, pitch 0.813, 0.75 s rotation time, display field of view (FOV) depending on the patient’s physique (median values of 350), beam collimation 6 mm × 0.625 mm (to reduce the overranging and to improve the Z-DOM modulation performance). The Brilliance iCT scanner can control radiation exposure with advanced dose reduction tools, such as X-ray tube current automatic modulation system (the automatic current selection ACS, which automatically suggests tube current settings according to estimated patient diameter in the scan region, and the Z-axis dose modulation system Z-DOM, that modulates mAs along the patient longitudinal axis using the attenuation profile estimated from the Surview), SmartShape and IntelliBeam shaping filters, the Eclipse asymmetric collimator for over-ranging reduction, NanoPanel detectors and ClearRay 2D anti-scatter grid.

The 16-MDCT images were reconstructed using a standard FBP algorithm with a standard soft-tissue kernel, while the 256-MDCT images were reconstructed with hybrid iterative reconstruction algorithm (iDose): iDose Level 3 was chosen.

At the end of every examinations, the volumetric computed tomographic dose index (CTDIvol) and the dose-length product (DLP) were provided by the scanners in the dosimetric report.

CT values (HU) and standard deviation of CT values (SD) were also measured by placing one ROI in the liver and the spleen as: SNR = HUROI/SDROI, where HUROI is the mean CT value in Hounsfield units of the tissue and SDROI the standard deviation of CT values in the same ROI.

Statistical analysis
Continuous variables (HU, SD, SNR, DLP, CTDIvol, patients weight) were presented as means ± SD. The Wilcoxon signed-rank test for paired samples or paired Student t-test were used, where appropriate, to compare values of continuous variables between standard dose protocol images and lower dose protocol images. To evaluate differences in qualitative analysis between the two protocols the Wilcoxon signed-rank test for paired samples was applied. A P-value < 0.05 was considered statistically significant.

Cohen’s kappa was used to evaluate agreement between the two readers. Analysis was performed with commercially available statistical software (SPSS Statistics 17.0, Chicago, IL).

RESULTS
Radiation dose
Mean values of obtained DLP and CTDIvol were significantly lower with lower-dose protocol in comparison with standard dose protocol (P < 0.001): For lower-dose CT, DLP and CTDIvol were respectively 1025.6 ± 446.4 mGy cm, 1025.6 ± 446.4 mGy cm, 1025.6 ± 446.4 mGy cm.
370.9 mGy*cm and 15.4 ± 5.2 mGy, compared with 1429.2 ± 297.7 mGy*cm and 21.4 ± 4.0 mGy for standard-dose protocol. The low-dose protocol provided a mean DLP reduction of 28.9% compared to standard-dose protocol (Figure 1).

The average weight was 74.1 ± 14.9 kg for low-dose scans (range 41-114 kg) and 73.9 ± 15.0 kg for standard dose scans (range 43-114 kg) (P = 0.705). By dividing patients into four groups of weight, we obtained a higher rate of DLP reduction in patients in groups A, B and C with a statistically significant difference between the two protocols; in patients with high weight (> 90 kg, group D), for values of DLP and CTDIvol only few differences between the two protocols were recorded, not statistically significant, with a radiation dose reduction of 2.5% (Table 2).

**Qualitative analysis**

Qualitative analysis results are shown in Table 3. There was a good inter-reader agreement, as shown by k Cohen values. There weren’t significant differences in the qualitative evaluation of image noise and diagnostic quality for both readers (Figure 2), and of image sharpness for one reader. The other reader assigned to low-dose images superior grades for sharpness, with a significant difference (P = 0.012).

**Quantitative analysis**

CT values of density (HU) measured within abdominal aortic lumen and liver and spleen parenchyma were significantly higher using low-dose CT with iDose® (P < 0.001) (Table 4). The mean liver and spleen parenchymal noise (SD) was higher with low-dose protocol as well, while SD of abdominal subcutaneous fat was higher but without reaching statistical significance. The measured noise varied according to the weight of the patients, with higher values of SD in patients with higher weight; values of SD in the liver parenchyma with low-dose protocol vs standard protocol were respectively 14.62 ± 1.80 vs 11.62 ± 2.03 in group A, 16.66 ± 1.45 vs 13.10 ± 2.18 in group B, 18.03 ± 1.98 vs 16.26 ± 1.54 in group C and 17.91 ± 2.24 vs 20.59 ± 3.28 in group D.

SNR values, calculated as SNR = HU_{ROI}/SD_{ROI}, were lower in low-dose images, reaching a significant difference within the liver parenchyma (6.94 ± 1.32 vs 7.80 ± 2.30, P = 0.002) and without statistical
**DISCUSSION**

In the latest years many studies investigating the potential of radiation dose reduction by applying different iterative reconstruction algorithms have been published for abdomen, chest, head, coronary and chest angiography, and they showed significant dose reduction while maintaining, or sometimes improving, image quality[@12-20].

Arapakis et al[@13] addressed the effect of iterative algorithm on radiation dose and image quality of chest-abdomen-pelvis (CAP) CT scans. They applied iDose® hybrid iterative reconstruction algorithm in a group of 84 patients and compared images to those obtained with “old standard” protocol with filtered back projection reconstruction algorithm in a group of 99 patients, obtaining an overall 46.5% decrease in effective dose with lower image noise and higher values of SNR and CNR; in their study, the greatest dose reduction was recorded in patients with lower weight[@13].

Karpitschka et al[@14] retrospectively evaluated 40 patients which underwent CT scans for staging of malignancy with both a standard-dose (tube current time product 250 mAs and FBP reconstruction) and a reduced-dose CT scan (150 mAs and with Iterative Reconstruction between two protocols in group C and higher in group D (Table 3).
in Image Space IRIS), obtaining a greater than 45% dose reduction at maintained image quality; the authors recommend the use of IR in oncological patients in order to reduce radiation dose to patients. Moreover, for abdominal CT scans, Prakash et al. showed a reduction of radiation dose by 25% using weight-based adjustment of automatic exposure control technique and Adaptive Statistical Iterative Reconstruction (ASIR) in comparison with FBP reconstructed scans, while May et al. obtained a 50% reduction in abdominal CT by using IRIS.

In our study, values of DLP obtained using a low-dose protocol with iDose iterative algorithm were, on average, 28.9% lower compared to our standard dose with FBP reconstruction. Despite higher levels of quantitative noise (as demonstrated by SD values) within liver and spleen parenchyma, however, qualitative analysis didn’t reveal significant differences in overall image noise and diagnostic quality when compared with standard-dose images in the same patients (Table 2). These results and lower rate of dose reduction in comparison with other CAP studies may be correlated to a different level of strength of the iterative reconstruction of iDose applied in our institution (L3) which determines less noise reduction and, with a fixed noise index, can be associated with higher levels of tube current and radiation dose.

The measured noise and SNR in low-dose images varied according to the weight of the patients, with higher values of SD and lower values of SNR in patients with higher weight (Table 3). These data were confirmed by qualitative analysis, which showed increasing levels of subjective noise by increasing patients weight, but without significant differences in image sharpness and diagnostic quality between the four groups (Table 2). In our patient group with weight greater than 90 kg (group D), values of SD within liver and spleen parenchyma and abdominal fat were the highest, with worst values of SNR; however, compared to standard-dose images, these values of SNR were higher indicating a better image contrast, although the difference was not statistically significant.

In group D the DLP reduction rate was substantially lower than those in the other patient groups (2.5% vs 29.6%, 33.5% and 29.9%). With automatic tube current modulation, tube current is automatically adjusted to the X-ray attenuation of the patient to keep the radiation dose as low as possible while maintaining a constant image noise level and specified image quality as in a previously defined reference image. In patients with large body habitus, to maintain a constant image quality, there is a risk of high radiation doses when this technique is applied in abdominal MDCT. Our results could be explained then by the higher tube current needed and used in this group of patients to fulfill the fixed noise level and desired final image quality. In heavy patients, the optimal noise indexes and image quality should be adjusted to patient habitus, considering that subjective image quality in abdomen CT of these patients is usually higher because of the amount of fat deposition around the abdominal organ that improve tissue contrast.

There were several limitations in our study. First of all, it was a retrospective study and acquisition of CT paired studies in our patients were not realized on the same scanner; despite many technical parameters were identical between the two protocols, some of them and the scanners were different and it may have introduced some bias. Because of the interval time between studies, which was remarkable in a small number of patients, and the retrospective nature of the study, we couldn’t compare conspicuity or detection rate of the primary tumor and secondary lesions between the two protocols, which should be addressed in further prospective studies.

In conclusion, in our sample of oncologic patients, automatic modulation of tube current and iDose reconstruction algorithm allowed a mean radiation dose reduction of 28.9%, without significant loss of subjective diagnostic quality, and this protocol could be useful in reducing dose exposure in patients with malignancy which undergo a high number of chest-abdomen-pelvis CT scan during their clinical follow-up.
preserving the underlying edges associated with true anatomy or pathology.

Applications
In this study the authors compared radiation dose and image quality of CT examinations in a population of oncologic patients undergoing follow-up examinations with a new generation MDCT scanner (256-rows) using automatic modulation of current tube and iterative reconstruction algorithm (DoseRight system and Dose4) and with a 16-MDCT scanner with standard dose protocol and FBP. The importance of this work relies on the fact that the results confirm the high diagnostic quality and the important radiation dose sparing of whole body CT scan examination obtained with a low-dose setting protocol combined with the new state of art iterative reconstruction algorithm in comparison with a standard-dose protocol. Moreover in this manuscript the authors compared and commented the results with those of previous literature on this field by using different vendor approach.

Terminology
MDCT: Multidetector row-CT are new generation of CT scanner with high number of detector, which allow to obtain high spatial resolution images with thinner collimation; FBP: Filtered back projection (FBP) has been the industry standard for CT image reconstruction for decades, representing a very fast and fairly robust method to reconstruct the raw data obtained from routinely CT scan acquisition; Hybrid Iterative Reconstruction Algorithms: In the literature, the term hybrid IR usually refers to algorithms that mainly decrease image noise by iterative methods. IR approaches are not new and were, in fact, the initially proposed method for data reconstruction in the early days of CT technology during the 1970s. However, due to its mathematically demanding properties and the large amount of data in CT imaging, until recently IR has not been practical for clinical purposes. The current evidence on the clinical implementation of IR into CT protocols shows substantial promise for major improvements in image quality, chiefly noise reduction-with subsequent radiation dose reduction-and artifact suppression; iDose*: Dose4 is a fourth-generation hybrid IR algorithm introduced by Philips Healthcare. With this algorithm the noise can be controlled for high spatial resolution reconstructions, hence providing high-quality, low-contrast, and spatial resolution within the same image, when radiologist work with low dose approach; through an iterative mathematical process, the noisy data are penalized and edges are preserved.

Peer-review
This is a very interesting attempt to achieve lower radiation dose in follow-up CT of oncologic patients with parallel comparison of thorax-abdomen-pelvis CT with 4th generation hybrid iterative reconstruction algorithm and standard dose examination.

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