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Can fully iterative reconstruction technique enable routine abdominal CT at less than 1 mSv?

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ARTICLEINFO	A B S T R A C T	
A R T I C L E I N F O Keywords: Iterative reconstruction Forward projected model-based Filtered back projection Sub-milli-Sievert Radiation dose CT abdomen	<i>Objective:</i> We assessed the effect of the forward projected model-based reconstruction technique (FIRST) on lesion detection of routine abdomen CT at < 1 mSv. <i>Materials and methods:</i> Thirty-seven adult patients gave written informed consent for acquisition of low-dose CT (LDCT) immediately after their clinically-indicated, standard of care dose (SDCT), routine abdomen CT on a 640-slice MDCT (Aquillion One, Canon Medical System). The LDCT series were reconstructed with FIRST (at STD (Standard) and STR (Strong) levels), and SDCT series with filtered back projection (FBP). Two radiologists assessed lesions in LD-FBP and FIRST images followed by SDCT images. Then, SDCT and LDCT were compared for presence of artifacts in a randomized and blinded fashion. Patient demographics, size and radiation dose descriptors (CTDIvol, DLP) were recorded. Descriptive statistics and inter-observer variability were calculated for data analysis. <i>Results:</i> Mean CTDIvol for SDCT and LDCT were 13 ± 4.7 mGy and 2.2 ± 0.8 mGy, respectively. There were 46 true positive lesions detected on SDCT. Radiologists detected 38/46 lesions on LD-FIRST-STD compared to 26/46 lesions on LD-FIRST-STD were seen in patients with BMI > 25.8 kg/m ² . Diagnostic confidence for lesion assessment was optimal in LD-FIRST-STD setting in most patients regardless of their size. The inter-observer agreement (kappa-value) for overall image quality were 0.98 and 0.84 for LD-FIRST-STD and STR levels, respectively. <i>Conclusion:</i> FIRST enabled optimal lesion detection in routine abdomen CT at less than 1 mSv radiation dose in patients with body mass less than ≤ 25.8 kg/m ² .	

1. Introduction

With increasing use of CT in modern medical practice, concerns have been raised over the associated radiation doses. CT vendors have introduced hardware technologies and image reconstruction methods to enable radiation dose optimization while maintaining an acceptable level of diagnostic quality. With the conventional filtered back projection and image-based iterative reconstruction methods, a decrease in CT radiation dose can impair the diagnostic confidence particularly for evaluation of low contrast organs or lesions such as in the abdomen and brain. Although a "smoother" or "softer" reconstruction kernels reduce the image noise in low dose CT, they impair spatial resolution [1]. Iterative reconstruction (IR) techniques for reconstructing CT images were described soon after the invention of CT scanners, as early as in 1973, but their application was hampered due to slow and low computational power [2]. With improvements in the computation power and speed, thanks to the video gaming industry, most CT vendors introduced commercial IR techniques to enable radiation dose reduction without sacrificing diagnostic confidence compared to the conventional filtered back projection techniques [3,4].

IR techniques can be classified into the hybrid IR (HIR) and the model-based IR (MBIR) techniques [1,5]. The MBIR employs models for image acquisition, statistics, and system geometry, whereas the HIR combines the analytical and iterative approaches to reduce the image

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noise [6]. Most CT vendors have developed diverse IR approaches. GE Healthcare (Waukesha, Wis.) introduced HIR named adaptive statistical IR (ASIR) in the year 2008. In the same year, Siemens Healthineers (Forchheim, Germany) commercialized HIR called image reconstruction in image space (IRIS) [4,7]. At about the same time, iDose (Philips Healthcare, Best, The Netherlands) and adaptive iterative dose reduction (AIDR-3D, Canon Medical Systems Corporation, Tochigi, Japan) were introduced [3,8]. In subsequent years, other IR methods were introduced including Veo and ASiR V (GE Healthcare), Safire and Admire (Siemens), IMR (Philips) and FIRST (Canon) [9–11].

The recently introduced Forward projected model-based Iterative Reconstruction SoluTion (FIRST) deploys a three-dimensional (3D) image reconstruction approach based on a forward and statistical model in the projection data to enhance the spatial resolution and reduce the image noise [12]. Compared to AIDR-3D, the FIRST optimizes image quality jointly in sinogram and image spaces [13]. The FIRST images are subjected to a regularization step to adapt the images based on organ specificity such as for abdomen, heart, lung, and bone. These features are then combined and undergo several iterations before converging to a final image. To our best knowledge, prior studies have not assessed the FIRST for enabling sub-milli-Sievert abdomen CT [14]. We hypothesized that FIRST can enable radiation dose reduction while maintaining diagnostic confidence in low radiation dose abdomen CT. To test this hypothesis, we assessed the effect of the FIRST on lesion detection in routine abdomen CT performed at low dose (LDCT at < 1 mSv).

2. Material and methods

2.1. Approvals and disclosures

The Human Research Committee of the Institutional Review Board approved our prospective multi-institutional study. The study was compliant with the Health Insurance Portability and Accountability Act (HIPAA). A study coauthor, MKK received research grants from Canon Medical Systems USA, Inc. and Siemens Healthineers. YH and EA are employees of Canon Medical Systems USA, Inc. All study coauthors had unrestricted access to the study data and the manuscript.

2.2. Patients

We prospectively enrolled 37 consecutive adult patients (17 women and 20 men; mean age = 63 ± 14 years) undergoing routine contrastenhanced abdomen CT examinations for clinically indicated reasons. The most common clinical indications were abdominal pain, cancer diagnosis and staging, and unexplained loss of weight. All participating patients gave written informed consent after understanding the risks associated with additional radiation dose. All patients met the following inclusion criteria: hemodynamic stability, age > 56 years, body mass index (BMI) $\leq 32 \text{ kg/m}^2$, understand English, ability to follow breathhold instructions, and hold breath for at least 5 s. We excluded patients undergoing urgent CT, patients with hemodynamic instability, shortness of breath, and inability to understand or follow scan instructions from the study. We recorded weight and height of all patients to estimate their body mass index (BMI).

2.3. Scanning techniques

All patients underwent standard-of-care routine abdomen CT (SDCT) in a supine position on a 640-slice multidetector-row CT scanner (Aquilion One, Canon Medical System Corporation). Patient's arms were placed over their head as per the standard of care procedure. After positioning the patients on the scan table, anteroposterior and lateral planning radiographs were acquired at 120 kV and 20 mA. To avoid motion related artifacts, patients were instructed to avoid any voluntary motion and follow breath-hold instruction for both SDCT and

LDCT acquisitions.

The scan parameters for SDCT included helical scan mode, 80×0.5 mm detector configuration, 0.5-second gantry rotation time, beam pitch of 0.9–1.1:1, 120 kV, and automatic exposure control (^{Sure}Exposure 3D, Canon Medical System Corporation). Scan length extended from the top of the diaphragm to the pubic symphysis. All patients received 80–100 mL of intravenous non-ionic iodinated contrast agent (Iopamidol 370 mg%, Bracco Diagnostics, Princeton, NJ) at a rate of 2–3 mL/s via antecubital vein. Scanning was initiated at a fixed delay of 60 s to obtain images in the portal venous phase.

All scan parameters with exception of tube current and scan length were kept identical between the LDCT and SDCT acquisitions. LDCT images were acquired through the upper abdomen from the top of the liver to its inferior tip. As opposed to the SDCT, we used fixed tube current so that additional dose length product (DLP) from the research LDCT images was less than 65 mGy.cm (to maintain an estimated effective dose < 1 mSv). Thus, most CT exams were performed between 30–50 mA s to attain DLP of 65 mGy.cm for LDCT images. Use of fixed tube current for acquiring LDCT has been reported in prior publications as well [15,16]. Both CT Dose Index volume (CTDIvol) and DLP were recorded for all SDCT and LDCT image series. We planned the acquisition of the LDCT image series before acquiring SDCT so that the time interval between the LDCT and SDCT series was less than 5–10 seconds.

2.4. Image reconstructions

Following acquisition of the SDCT and LDCT, projection data was de-identified and exported offline for image reconstruction. Images were reconstructed at 5 mm section thickness with 5 mm section overlap. Both SDCT and LDCT data were reconstructed to obtain conventional FBP images (SD-FBP and LD-FBP) and FIRST images at standard (SD-STD and LD-STD) and strong (SD-STR and SD-STR) strengths. The approximate time for reconstructing FIRST images varied between 5–7 minutes. Mean HU and image noise (standard deviation of mean HU) were recorded by drawing a region of interest (1-1.5cm²) each on right hepatic lobe and main portal vein. These values were used to calculate signal-to-noise (SNR) and contrast-to-noise (CNR) ratios were calculated (Figs. 1–3).

2.5. Qualitative assessment

de-identified SDCT $(n = 37 \times 3 = 111)$ A11 and LDCT $(n = 37 \times 3 = 111)$ were viewed on a standalone image viewing software (ClearCanvas workstation, ClearCanvas Inc., Toronto, Canada). Images were assessed in soft tissue window settings (window level = 40, window width = 400); the radiologists were permitted to adjust window settings according to their preference. Two radiologists (RS with 6 years and RDK with 4 years of experience) independently assessed all six CT image series. For each patient, a single image series was displayed at a time. The LDCT images series were displayed prior to the SDCT image series. The interpreting radiologists did not take part in patient recruitment, scanning, data de-identification, or image reconstruction. Each image series was separately evaluated for lesion detection, conspicuity, size, and number.

2.6. Statistical analysis

Data were recorded and analyzed in Microsoft EXCEL (Microsoft Inc., Redmond, Wash.). Mean and standard deviations were calculated for the dose descriptors (CTDIvol and DLP). Wilcoxon signed rank and Cohen's kappa tests were performed to compare qualitative evaluation and to assess interobserver agreement. A p-value of ≤ 0.05 was considered statistically significant. For Cohen's Kappa analysis, a kappa of > 0.6 was termed as a good agreement, 0.2-0.4 as fair agreement and < 0.2 as poor agreement.



Fig. 1. A 68-year-old woman (BMI 32 kg/m²) underwent contrast-enhanced abdomen CT at SDCT (CTDIvol 22 mGy) and LDCT (2.8 mGy). All SDCT and LDCT images demonstrate a necrotic hepatic lesion (white arrow) in the right hepatic lobe. LD-FIRST images had lower noise and better diagnostic performance than the LD-FBP image.

3. Results

The mean BMI of the recruited patients was $26.2 \pm 3 \text{ kg/m}^2$. None of the patients had metallic hardware within the scanned field of view

of the LDCT. Respective CTDIvol and DLP for the SDCT were 13 \pm 4.7 mGy (range, 11–16 mGy) and 549 \pm 262 mGy.cm (range, 471–679 mGy.cm). Corresponding values for LDCT were 2.2 \pm 0.8 mGy (2–3 mGy) and 49 \pm 14 mGy.cm (39–60 mGy.cm),



Fig. 2. A 62-year-old man (BMI 23 kg/m²) underwent contrast-enhanced CT of the abdomen at SDCT (CTDIvol 11 mGy) and LDCT (2.8 mGy) images. All SDCT images demonstrate low attenuation hepatic lesions likely cysts in right hepatic lobe. LDCT images detected all lesions except a small cyst (arrow).



Fig. 3. A 69-year-old man (BMI 31 kg/m²) underwent contrast-enhanced CT of the abdomen at SDCT (CTDIvol 16 mGy) and LDCT (1.7 mGy). All SDCT and LDCT images demonstrated cholelithiasis (arrow). LD-FIRST-STD image had lower noise and better diagnostic performance than the unacceptable LD-FBP image.

Table 1

Mean (\pm standard deviation) of SNR and CNR for different image reconstruction methods.

Image series	SNR: mean ± standard deviation		CNR: mean \pm standarrd deviation
	Liver	Portal Vein	
SD-FBP LD-FBP LD-FIRST-STD LD-FIRST-STR	9 ± 2 10 ± 3 2 ± 1 16 ± 5	$ \begin{array}{r} 11 \pm 3 \\ 13 \pm 4 \\ 2 \pm 0.7 \\ 20 \pm 7 \end{array} $	3 ± 3 4 ± 3 0.4 ± 0.5 5 ± 3

respectively. There was a significant statistical difference between the dose descriptors for the SDCT and LDCT (p < 0.0001). The SNR and CNR are summarized in Table 1.

There were no differences between the number and types of lesions detected on the SD-FBP and both the SD-FIRST series (p > 0.9) (Fig. 1-3). The average lesion size was $13 \pm 10 \text{ mm}$ (range = 3-32 mm). Both radiologists detected 55 lesions on the SDCT images which included low attenuation liver lesions (n = 23), renal cysts (n = 17), adrenal nodules (n = 6), low attenuation lesions in the pancreas (n = 4), cholelithiasis (n = 2), retroperitoneal fat stranding with thickening (n = 1), ascites with mesenteric fat stranding (n = 1), and retroperitoneal lymphadenopathy (n = 1).

On the LD-FBP images, the radiologist 1 (R1) detected 39 lesions that included low attenuation liver lesions (n = 13), renal cysts (n = 16), adrenal nodules (n = 5), low attenuation lesions in the pancreas (n = 1), cholelithiasis (n = 1), retroperitoneal fat stranding with thickening (n = 1), ascites with mesenteric fat stranding (n = 1), and retroperitoneal lymphadenopathy (n = 1). On these images, the radiologist 2 (R2) detected 35 lesions that included low attenuation liver lesions (n = 13), renal cysts (n = 12), adrenal nodules (n = 4), low attenuation lesions in the pancreas (n = 3) cholelithiasis (n = 2), and retroperitoneal lymphadenopathy (n = 1).

Both radiologists detected additional lesions on the LD-FIRST-STD images that were not documented for the LD-FBP images. These included lesions that included low attenuation liver lesions (n = 2–4), adrenal nodule (n = 1), low attenuation lesions in the pancreas (n = 2), cholelithiasis (n = 1), retroperitoneal fat stranding with thickening (n = 1), and ascites with mesenteric fat stranding (n = 1). Patients (BMI > 25.8 kg/m²) with missed lesions on the LD-FIRST-STD images relative to the SDCT images were larger than the patients without missed lesions (p = 0.01).

All lesions on LD-FBP images were also seen on the LD-FIRST-STD and STR settings. However, radiologists could not detect three lesions on the LD-FIRST-STR that were seen on the LD-FIRST-STD; these

Table 2

Number of lesions detected by each radiologist on SDCT and different LDCT image series.

-		
Image reconstruction methods	R1 (number of detected lesions)	R2 (number of detected lesions)
SD-FBP LD-FBP LD-FIRST-STD LD-FIRST-STR	55 39 49 46	55 35 43 40

included low attenuation liver lesion (n = 1), low attenuation lesions in the pancreas (n = 1) cholelithiasis (n = 1), and adrenal nodule. Table 2 summarizes the lesion detection by each radiologist.

The number of lesions detected on different SDCT image series were identical.

There were no major artifacts on any of the SDCT or LDCT image series regardless of the reconstruction technique. The inter-observer agreement (kappa-value) were 0.98 and 0.84 for LD-FIRSTSTD and STR levels, respectively.

4. Discussion

FIRST algorithm has been evaluated for radiation dose reduction in patients with lung nodules [14,17] and those undergoing CT angiography of the abdomen [12] or coronary arteries [13,18-20]. Our radiation doses (2.2 \pm 0.8 mGy) were substantially lower than the prior abdominal CT angiography study with FIRST (7 \pm 2.8 mGy) [12]. Despite the differences in radiation doses between our study and that of Wu et al [12], we noticed an improved lesion detection with the LD-FIRST images as compared to the LD-FBP images. Both radiologists detected more lesions on LD-FIRST images than on LD-FBP images. The additional lesions on the LD-FIRST images included additional liver, adrenal and pancreatic lesions. Both radiologists preferred LD-FIRST images at standard strength over the strong setting since the latter images were "too smooth" for evaluation of major organs such as the liver, pancreas, and kidneys. Other studies have also reported the "too smooth" appearance with different IR techniques at higher strengths of noise reduction [16].

Previous studies have described use of IR techniques to enable radiation dose reduction below 1 mSv for abdomen CT [21–26]. However, most successful studies with sub-mSv radiation doses using IR techniques were performed for evaluation of kidney stones and CT colonography [22–26]. Kidney stones are high contrast findings are not as much affected by high image noise at sub-mSv radiation doses [22]. Air filled lumen also provides a much higher contrast to the colonic wall and polyps, which enables uncompromised evaluation at sub-mSv

[25,26]. Conversely, routine abdomen CT is a more commonly performed procedure which entails evaluation of low-contrast solid organs and lesions in the liver, pancreas, and kidneys. In 2012, Padole et al reported that IR techniques from four different CT vendors (other than the vendor assessed in our study) could not provide acceptable image quality in non-obese patients undergoing sub-mSv routine abdomen CT [21]. Although LD-FIRST images outperformed LD-FBP in terms of lesion detection, both radiologists missed several lesions on both LD-FBP and FIRST images they reported on SDCT images series. However, in contradiction to prior studies, there were no lesions on the sub-mSv abdomen CT images reconstructed with FIRST in patients with $BMI \leq$ 25.8 kg/m^2 . Interestingly, O'Neill et al have also reported that sub-mSy abdomen CT images reconstructed with ASIR technique enabled diagnostic interpretation in non-obese patients with a BMI $< 25.2 \text{ kg/m}^2$ [26]. The study tested visibility of only bowel findings related to Crohn's disease on LDCT of the abdomen [26] as opposed the smaller and lower contrast lesions on our study at similar radiation doses.

Implications of our study include feasibility of deploying sub-mSv routine abdomen CT protocol with FIRST in non-obese, non-overweight patients with BMI $\leq 25.8 \text{ kg/m}^2$. When using FIRST technique for image reconstruction, a standard setting provides better lesion delineation and retention of image texture as opposed to the strong setting. In overweight patients (BMI > 25.8 kg/m^2), sub-mSv CT for routine abdomen is not possible with the FBP or the FIRST. Missed lesions at sub-mSv CT include liver metastases and pancreatic lesions which have serious diagnostic implications. Future sub-mSv studies should target high contrast situations in the abdomen for evaluation of abdominal vasculature, colonography and kidney stones.

Our study has limitations. Because of the prospective nature of our study, associated additional radiation dose from acquisition of LDCT image series, and stringent inclusion and exclusion criteria, our study sample size is not large (n = 37 patients). We did not perform quantitative assessment of FBP or FIRST images since the conventional parameters such as image noise (standard deviation of CT numbers), signal-to-noise ratio, and contrast-to-noise ratio, do not provide an accurate assessment of image quality or diagnostic acceptability of iterative reconstruction techniques. As expected from any CT examination with two or more post-contrast acquisition, there was a short delay between the acquisition of SDCT and LDCT images. We addressed this limitation with planning of LDCT image series before acquisition of SDCT. Due to the portal venous scanning for the SDCT and LDCT image series, the minimal time difference (up to 10 s) did not have a notice-able effect on the evaluation of these two series.

We excluded obese patients since LDCT images for routine abdomen in such patients were unlikely to attain the required diagnostic performance with either FBP or FIRST reconstruction techniques. Suboptimal performance of both FBP and FIRST images in the larger patients included in our study confirmed our decision. Our study focused on routine abdomen protocol for LDCT, and therefore, our results should not apply to other abdominal CT protocols where radiation doses are lower than for routine abdomen CT. For example, abdominal CT for renal colic and colonography are performed at a lower dose than routine abdomen CT. Also, it is unlikely that radiation doses (CTDIvol 2.2 ± 0.4 mGy) used in our study can provide sufficient information for liver and pancreatic protocols where a preponderance of low-contrast lesions require lower noise and better image quality than needed for the routine abdomen CT images. Another limitation of our study pertains to the use of fixed tube current for acquiring LDCT images as opposed to automatic exposure control for the SDCT images. We used fixed tube current for LDCT because it is easy to apply to reach a targeted radiation dose (DLP 65 mGy.cm; estimated effective dose just under 1 mSv). Also, because for such LDCT the tube current with automatic exposure control over a short length of acquisition in the upper abdomen will not change substantially from slice to slice or within each slice depending on the patient size.

reconstruction technique (FIRST) enabled optimal lesion detection in routine abdomen CT at less than 1 mSv radiation dose in patients with body mass less than $\leq 25.8 \text{ kg/m}^2$. The standard setting of FIRST outperforms stronger setting in terms of lesion detection and image texture. Compared to our standard of care, routine abdomen CT protocol, the assessed low dose CT FIRST images can enable a radiation dose reduction of up to 85%.

Conflict of interest

A study coauthor, MKK received research grants from Canon Medical Systems USA, Inc. and Siemens Healthineers. YH and EA are employees of Canon Medical Systems USA, Inc. All study coauthors had unrestricted access to the study data and the manuscript.

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In summary, recently introduced forward-projected, model-based

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