ADVANCES IN CARDIAC IMAGING (U. JOSEPH SCHOEPF, SECTION EDITOR)

The Role of Iterative Reconstruction Techniques in Cardiovascular CT

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Published online: 14 September 2013 © Springer Science+Business Media New York 2013

Abstract Iterative reconstruction (IR) techniques for cardiovascular computed tomography (CT) have enjoyed a resurgence of interest in recent years as computer power has increased enough to enable reasonably timely reconstructions. The major purported benefit of current IR techniques involves image noise reduction, which both provides improved image quality and enables radiation dose reductions. Several widely available products have been released by the major CT vendors that vary in their underlying techniques but, according to the current literature, give similar results. Future algorithms should both refine current IR techniques and expand the role of IR to additional cardiovascular CT applications. This review examines the technical basis of IR, the IR products available commercially, the current data on IR in cardiovascular CT, and the future directions of the field.

Keywords Iterative reconstruction · Radiation dose · Cardiac CT angiography · Image quality · Artifacts

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Introduction

The development of computed tomography (CT) for medical imaging has had a profound impact on the practice of medicine over the past four decades. Technical advances in CT continue to develop at a remarkable pace. Nowhere is this more apparent than in cardiovascular CT. The inherent problems associated with imaging small, moving structures limited the role of cardiovascular CT for several decades. However, the past 20 years have seen a host of new developments that have brought cardiovascular CT into routine clinical practice. These began with the rise of cardiovascular electron beam CT in the mid 1990s and the subsequent development of multi detector-row CT and ECG-gating techniques just before the new millennium [1•, 2, 3]. Clinicians and industry continue to promote hardware and software solutions with the fundamental objectives of improving image quality (e.g. improving temporal and spatial resolution; reducing artifacts), expanding clinical applications (e.g. myocardial perfusion imaging; left ventricular functional analysis), and reducing radiation dose. This review examines the current and future role of one such advance-iterative reconstruction (IR) techniques.

IR technology, although attracting much attention in CT applications in recent years, is not a recent development. Initial CT efforts in the early 1970s used IR techniques to create the first images from CT projectional data [4], and emission tomography has used iterative techniques since the 1960s [5]. IR, however, necessitates complex mathematical models and performs multiple reconstructions in the creation of images (see below), and, as CT resolution improved, increased computational demands rendered IR too slow for clinical use. Faster analytical algorithms, most prominently filtered back projection (FBP), became the dominant reconstruction methods, and IR was relegated to

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research applications. The renewed interest in IR comes as the exponential growth in computer power has started to meet the requirements for practical clinical use.

The current role of IR in cardiovascular CT is evolving. The fundamental technical advantage of IR is improved image quality, mainly via noise reduction but also as a result of artifact suppression. Several specific situations in particular may benefit, including examinations involving obese individuals, which are traditionally limited by high noise secondary to quantum mottle, and evaluation of coronary stents and heavily calcified vessels, which suffer from beam-hardening artifacts and suboptimum spatial resolution. Importantly, image noise is inversely related to radiation dose; therefore, improved noise characteristics have the potential to reduce the radiation dose associated with CT while retaining acceptable levels of image quality. The potential for significant dose reduction has been described as the main clinical advantage of IR, especially given current concerns from both the public and medical communities regarding radiation from diagnostic imaging [6]. In addition, researchers are currently developing new algorithms with the objectives of both improving the traditional strengths of IR and providing new advantages and applications of the technique.

Technical Basis

The mathematics underlying IR techniques are beyond the scope of this review. The reader is referred to Beister et al. [7] and similar articles for a more detailed technical discussion. Nevertheless, a conceptual understanding of IR techniques is necessary to appreciate their benefits, potential applications, and vendor-specific differences. The latter point deserves emphasis—"IR" is a fairly nonspecific term, and there are significant variations in the techniques, advantages, and disadvantages of the currently available commercial products.

Basic Data Acquisition

CT data are acquired by transmitting a fan beam of photons through the body to an array of detectors; to acquire a complete dataset this must be performed over multiple angles (at least 180°) around the long axis of the patient. Because the beam passes through the patient, the individual data points measured by each detector represent the sum of the attenuation of all tissues through which the beam has passed. All reconstruction methods create stacks of twodimensional (2D) images from this raw projection data. 2D image matrices are made up of an array of pixels, with each pixel corresponding to a specific area within the imaged subject and assigned an attenuation value. Differences between reconstruction techniques involve determining how this attenuation value is assigned in the final image.

Reconstruction with Traditional Analytical Methods

Traditional analytical reconstruction methods, for example FBP, gather all of the individual data points for a given detector element (the line integrals representing the total attenuation of the beam as it takes a radial path through the patient) and project that information back along the radial path, dividing the total attenuation evenly across all pixels in the path. This is repeated as the tube(s) and detectors rotate around the patient, and the combined attenuation values provided from each of these back projections are summed for each pixel, resulting in the final image. Mathematical filters are applied to the data before back projection to modify image noise and resolution. For simplicity, total available image noise reduction and spatial resolution can be regarded as fixed, and filters provide the optimum balance between resolution and noise for a given application (each at the expense of the other).

Analytical reconstruction methods are fast and fairly robust in many situations; however, there are multiple problems with the technique. In the most basic terms, analytical reconstructions use the measured signal as if all data were perfect. The photon beam is assumed to be a straight ray that arises from a single point on the surface of the anode, terminates at a point at the center of the detector, and interacts with a point in the center of each voxel. Realworld geometric considerations, such as focal spot size, anode heel effect, the three-dimensional (3D) interaction of the beam with the voxel, and the 2D interaction of the beam with the detector, are ignored. X-ray spectra are assumed to be monoenergetic, and nonlinear effects along the assumed ray, for example scatter and beam-hardening, are not considered. Perhaps most importantly, analytical reconstruction methods do not account for image noise that results from Poisson statistical variations in photon numbers across the image plane.

IR

The fundamental differences between analytical and IR can be described as follows: whereas analytical reconstruction methods perform a single reconstruction assuming all data are perfect, IR techniques perform repetitive reconstructions applying mathematical models to account for known imperfections in the projection data. Each "iteration" is slightly modified from the previous until a predefined criterion is met. As noted above, "IR" is a nonspecific term, and there is substantial variation in the technical details of commercially available IR products beyond the preceding generalization. The advantages and disadvantages of specific methods are largely dependent on the type and scale of mathematical modeling applied to iterations. Several basic classification schemes have been described that—while imperfect—may be useful. The most prevalent of these include classification based on:

- 1 the general type of algorithm used; and
- 2 the data (i.e. raw data or image data) used in the reconstruction process.

Most IR products in current clinical use apply models of photon counting statistics to reduce image noise: There are random variations in the number of photons striking the detector across the image plane, with the distribution of photon number across a given area described by Poisson statistics. Noise is represented by the relative standard deviation of photon number, which is inversely proportional to the number of photons striking the detector. Practically speaking, this explains why increasing tube current, which increases the number of X-rays transmitted, reduces image noise. By correlating the noise patterns with the signal (photon number) at the detector, statistical IR methods are able to selectively eliminate image noise with each successive iteration (Fig. 1). Statistics-based methods also may employ regularization techniques, the most common of which are "smoothing" algorithms that limit the allowable difference in attenuation between adjacent voxels.

More advanced "model-based" IR (MBIR) products go beyond statistical modeling. Both geometric (e.g. the area of the anode, interaction of the photon beam with the voxel and detector, anode heel effect, etc.) and physical (e.g. the X-ray spectra, scattered photons, beam-hardening, etc.) models can be applied. These models are used to predict the volumetric image, with the objective of approximating the actual image as closely as possible. The predicted image is forward-projected to create an artificial raw data set that is then compared with the actual raw data set. The predicted data are corrected on the basis of the actual data, and this correction is back projected to create an updated image. The process is repeated until a specific criterion is fulfilled—it can be performed for a fixed number of iterations, until the difference between the predicted and actual data reaches a predefined threshold, or until indicators of image quality reach a specified level [7].

IR methods may also be classified on the basis of the data that are reconstructed. IR performed only in the image/slice domain first reconstructs an image using FBP; this is then forward-projected using photon statistics to selectively remove noise across successive iterations. Techniques using both image and projection/raw data first reconstruct an image using FBP. Rather than directly modifying the real data as before, the actual raw data are compared with an artificial raw data set that was created by use of Poisson statistics-based noise models. Successive iterations again reduce image noise until the real and predicted data converge. Finally, IR techniques utilizing the projection data only are akin to the previously described MBIR methods. Complex statistical, geometric, and physical models attempt to predict the projection data. Predicted raw data are compared with actual raw data and modified over successive iterations [7, 8..].

The major advantage of statistical IR techniques involves noise reduction without a corresponding decrease in spatial resolution. The lower the signal to noise ratio (SNR), the more impact IR can have on image quality. Situations in which the SNR is inherently lower, for example evaluation of large patients and high-resolution acquisition utilizing small voxels (e.g. cardiac imaging), are particularly well-suited to IR applications (Fig. 2). The noise reduction properties of IR also enable reduction of radiation dose without unacceptable sacrifices in image quality. MBIR provides incremental improvements in image quality by reducing a variety of artifacts caused by invalid assumptions and physical and geometric imperfections in the transmission data. Streak artifact, beamhardening, motion, and scatter effects can be reduced.

Fig. 1 Standard FBP (a) and iterative (b) reconstructions at the same level of the ascending aorta. Image noise expressed as the standard deviation of the attenuation (HU) in the region of interest was significantly lower in images reconstructed using IR (SAFIRE; Siemens) (*circle* in b) than in those reconstructed using FBP (*circle* in a)

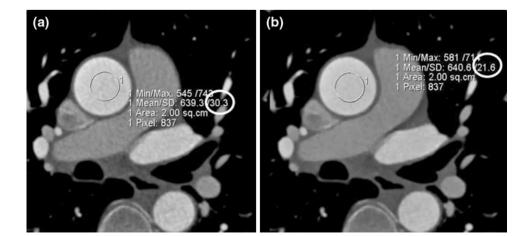
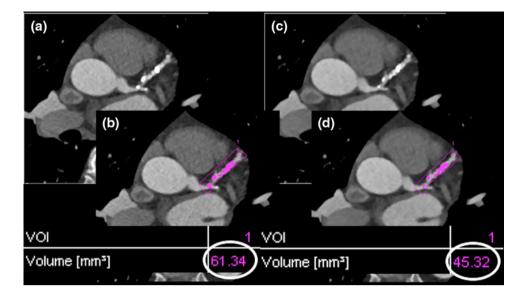


Fig. 2 Transverse **a** FBP and **b** IR (IRIS; Siemens) images at the level of the aortic root show extensive calcified plaque burden in the proximal left anterior descending coronary artery. Threshold-dependent volumetry of calcifications (*purple*) resulted in a measured volume of 61.34 mm³ on FBP reconstructions (**c**) and of 45.32 mm³ on IRs (**d**) (Color figure online)



The main disadvantage of current IR techniques is the increased computational effort. Whereas this is negligible in most statistics-based methods, MBIR in the raw data domain can still take as long as 10–90 min per reconstruction [9]. Computational power continues to increase; however, the increased complexity of mathematical models and expansion of complex acquisition techniques, for example dualsource and dual-energy CT, have somewhat mitigated improvements in reconstruction time [7]. The quality of IR is also dependent on the robustness of the applied algorithms. Improper models and overcorrection can result in reduced image quality and novel artifacts. There is potential to lose information when data are manipulated in the projection domain. Finally, IR results in an alteration in the expected image appearance, which has been described as "waxy" or "plastic" [9]. The psychophysical and cognitive effects of this altered image appearance are difficult to quantify and may affect interpretation. Vendors have responded by providing different iteration "strengths" or enabling variable blending of IR with FBP images.

Currently Available IR Products

Adaptive Statistical IR

Adaptive statistical IR (ASIR; GE Healthcare, Waukesha, WI, USA) was the first modern CT IR product released, in 2008. For each axial image, ASIR creates an image noise map and a traditional FBP reconstruction. Attenuation variances between pixels in the FBP image are examined, and the ASIR algorithm identifies variances that are statistically unlikely, i.e. attenuation fluctuations more likely secondary to image noise than actual anatomic structures. The image noise map is then used to modify the FBP image on the basis of ideal noise modeling. This process is repeated until the modified FBP image and ideal pixel values converge. ASIR enables selective blending with FBP projections, with the most commonly reported blending ratios using 40–60 % ASIR [8••, 10] (Fig. 3).

IR in Image Space

IR in image space (IRIS; Siemens Healthcare, Forchheim, Germany) also uses statistics-based noise modeling; however, the entire process is carried out in the image domain. A nonlinear algorithm is applied to each area of an axial FBP image to remove attenuation variances that are likely to be because of noise. In general, a predefined and fixed number of iterations are performed [11] (Fig. 4).

Sinogram Affirmed IR

The second IR product released by Siemens, sinogram affirmed IR (SAFIRE), models noise on the basis of raw data. Attenuation variances are removed on the basis of this model and the resulting image is compared with, and combined with, the initial FBP image; the process is then repeated. SAFIRE most commonly uses five iterations; however, the number or "strength" of iterations can be modified [12] (Fig. 5).

Adaptive Iterative Dose Reduction

Adaptive iterative dose reduction (AIDR, Toshiba Medical Systems, Otawara, Japan) was the first IR method introduced by Toshiba; similar to IRIS, AIDR performs denoising of the image data on the basis of photon counting statistics. AIDR 3D was subsequently released with several major differences. Both electronic and quantum noise are selectively removed in the projection domain, by both both noise and anatomic modeling in image reconstruction. noise filtering and modeling. An additional model accounts for the physical properties of the CT system at the time of acquisition. These two processes are combined and subsequent iterations compare the original FBP and the modified reconstructions in the image domain. Anisotropic diffusion is used to ensure that high-frequency structures representing fine details are preserved [13, 14]. Put simply, anisotropic diffusion models the shapes of the anatomic area being examined and provides shape-adaptive smoothing. For example, the system might detect high variation between two adjacent pixels within the left ventricular cavity. This is unexpected and would be attributed to noise and selectively eliminated. However, high variation between pixels at the myocardial-left ventricular border would be expected; this border would be left unchanged. Similar to other methods, iterations are repeated until a specific criterion is met. AIDR images are then blended with FBP images. Modifications of blending ratios are not available on current commercial products.

iDose

iDose, released by Philips Healthcare (Eindhoven, the Netherlands), can be compared with AIDR 3D in that it uses First, the projection data are examined, areas with very low photon counts (i.e. noisy areas) are identified, and noise is selectively eliminated. Again, an anisotropic diffusion process is utilized to penalize noise while maintaining true anatomic edges. The process then moves to the image domain, where noise subtraction is performed using an estimated map of image noise so that edges are maintained. This "hybrid" (i.e. noise and anatomic modeling) method is purported to enable greater IR strength, resulting in less noise, without resulting in the waxy or overcorrected images that resulted from earlier generations of IR techniques. Seven different IR levels can be set (L1–L7) that correspond to increasing levels of noise reduction, ranging from a noise reduction factor of 0.89 (89 % noise compared with the equivalent FBP reconstruction) for L1 up to 0.45 for L7. These noise reduction factors are not arbitrary values; rather, they correspond to the noise increases that would result from increased levels of dose reduction, from 20 % dose reduction for L1 to 80 % reduction for L7 [15] (Fig. 1).

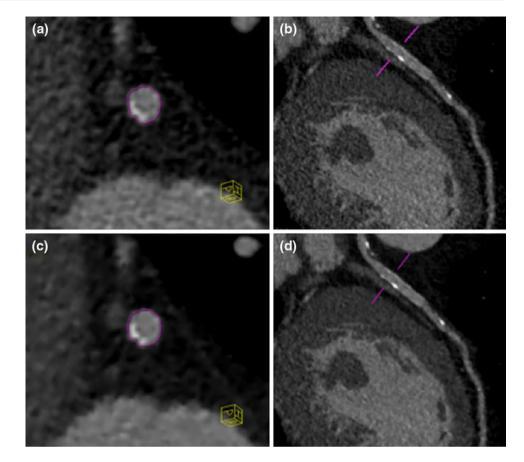
MBIR/Veo

MBIR, later released under the trade name Veo (GE Healthcare, Waukesha, WI, USA), is the only fully model-

Fig. 3 CT topography of an obese patient (a) with corresponding transverse reconstructions of a coronary CT angiography using FBP (b) and IR (IRIS; Siemens) (c). Image noise expressed as the standard deviation of the attenuation (HU) in the region of interest was significantly lower in images reconstructed using IR than in those reconstructed using FBP-18.9 versus 39.3 HU



Fig. 4 Cardiac CT angiography (cCTA) study of a patient with an implanted coronary artery stent. Images are displayed as automatically generated curved multiplanar reformat along the vessel centerline (*right figures*; b and d) and as cross-sections perpendicular to the centerline (*left figures*; a and c). The *lower figures* (c and d) show images reconstructed using IR (SAFIRE; Siemens) compared with those reconstructed using FBP (a and b)



based reconstruction product currently available. It uses complex statistical, optical, geometric, and physical modeling that attempts to model the X-ray beam from anode to detector (see above). There are many potential benefits to the technique; but the complexity of the method is its major limitation, with reconstruction times ranging from 10 to 90 min [9].

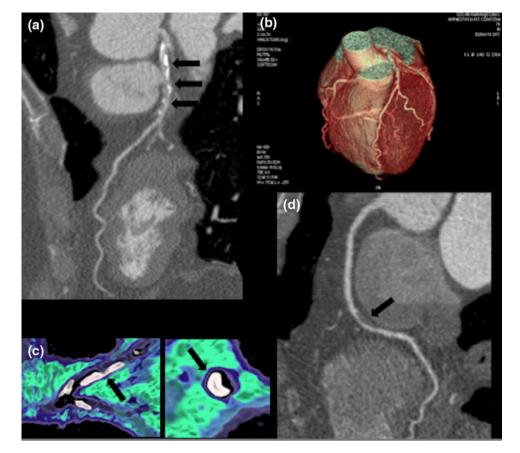
Current Applications and Evidence

Noise and Radiation Dose Reduction

The primary role of IR techniques in current cardiovascular CT practice is, via noise reduction, to enable radiation dose reduction without sacrificing image quality. Tube voltage and tube current are related to both radiation dose and image noise. Low voltage and/or low current acquisition procedures have become increasingly popular as dose-reduction strategies, but at the expense of increased noise. Numerous studies have shown that IR can improve objective noise, SNR, contrast-to-noise ratio (CNR), and subjective image quality (SIQ) compared with FBP reconstructions of the same projection data; furthermore, low-dose acquisition using IR can maintain the image quality of higher-dose acquisition using FBP.

There are limitations in the currently available evidence. Most studies use objective and subjective measures of image quality; however, only limited data are available comparing the actual diagnostic accuracy of FBP and IR techniques. There are few studies that compare specific IR products with each other. Most IR applications have modifiable settings involving the iterative strength or proportion of IR blending with FBP; however, the optimum settings for each product and situation are not well-defined. There are limited data on the impact of IR on clinical workflow. Finally, the radiation dose reductions reported in many studies are simulated or estimated (e.g. by removing half of the projection data or adding artificial noise to the images) rather than actual.

With these limitations in mind, the current evidence shows that IR techniques may have extraordinary dosereduction potential in a variety of clinical scenarios. Selected studies are summarized in Table 1. We can summarize the trends as follows: IR results in reduced noise with no to minimal effects on attenuation (signal) compared with FBP reconstructions of the same data, resulting in improved SNR and CNR. This usually, but not uniformly, results in improvements in SIQ and vessel assessment. In general, noise reduction is proportional to the iterative strength or proportion of IR blending; however, most studies reveal that SIQ does not follow the same Fig. 5 cCTA study of a 70 year old patient (BMI 24) using a prospective gated scan and ASiR (Adaptive Statistical IR; GE) image reconstruction. Despite a dose-lengthy product (DLP) of only 37 mGy-cm (0.5 mSV), excellent image quality was achieved. a, d Curved multiplanar reformation of the left anterior descending coronary artery (LAD) and the right coronary artery (RCA). Black arrows indicate severe lumen narrowing caused by mixedplaques (a) and noncalcified plaque (d). b 3D volume rendering from left anterior oblique perspective. c Corresponding virtual angioscopy with severe calcified lesions (black arrow). (Images courtesy of GE Health Care.)



pattern. As iterative strength/proportion increases, SIQ first improves, peaks, but then begins to diminish, because of overcorrection and/or the introduction of new artifacts, and/or an increasingly unfamiliar subjective image appearance may evolve. Most studies that compare actual or simulated low-dose acquisitions reconstructed with IR to higher-dose acquisitions reconstructed with FBP demonstrated no significant differences or persistent improvements in SIO with the use of IR [12, 14, 16-26, 27, 28-33]. Several real-world observational studies have recorded cardiovascular CT dosimetry data after implementation of IR methods and likewise shown significant dose reductions without unacceptable sacrifices in SIO [18, 31, 34•]. The few studies that have assessed the diagnostic accuracy of IR using intravascular ultrasound (for coronary artery plaque characterization) or invasive coronary angiography (for coronary artery stenosis detection and quantification) have reported equivalent or improved accuracy compared with FBP [12, 25, 26, 35, 36].

These findings are valid for all currently available IR products except MBIR, for which experiences are extremely limited. ASIR has resulted in improved image quality at a dose reduction of 25 % compared with FBP [37] and equivalent image quality and diagnostic accuracy with dose reductions up to 72 % [25]. Observational studies after

implementation of ASIR have demonstrated 44-54 % reductions in effective dose [18, 34•]. AIDR has shown improved subjective and objective image quality with simulated 50 % dose reduction compared with full-dose FBP reconstructions of the same data [38], and one observational study demonstrated dose savings of 22 % without compromising quality [31]. Low-dose acquisitions using IRIS have demonstrated improved image quality compared with routine acquisitions using FBP with dose savings up to 62 % [27•]. Likewise, SAFIRE has shown equivalent image quality as FBP at simulated dose reductions of 50 % [39] and improved image quality with simulated dose reductions of 50-80 % [12, 29]. Both fixed and adaptive-dose acquisition procedures using iDOSE have shown that radiation reductions of 55-63 % are allowable with improved or equivalent image quality compared with higher-dose FBP [15, 20, 40]. Preliminary investigations using MBIR have shown favorable data on diagnostic accuracy [36], demonstrating improved image quality compared with ASIR in one study [41]; however, the evidence for MBIR in cardiovascular CT is currently limited to ex-vivo studies.

The optimum method for dose reduction using IR is not well-defined. Most of the more impressive dose reductions were demonstrated in simulated or fixed-dose reductions

Ref.	IR method	IR acquisition and reconstruction procedure	FBP acquisition and reconstruction procedure	Quantitative image quality	Qualitative image quality	Diagnostic accuracy	Radiation dose reduction	Notes
Pontone et al. [25]	ASIR	Adaptive procedure based on BMI	Fixed-dose procedure	No difference in SNR or CNR		No significant difference on a per- segment or per- patient basis	72 % reduction with IR (2.1 vs. 7.5 mSv)	
Leipsic et al. [37]	ASIR 30 %	Adaptive procedure based on predefined allowable noise index of 25	Adaptive procedure based on predefined allowable noise index of 21	Improved noise, and SNR with IR	Improved subjective image quality with IR		25 % reduction with IR	Included noncontrast chest CTs
Leipsic et al. [21]	ASIR at 20, 40, 60, 80, and 100 %	Low-dose cCTA	Intrapatient comparison	Reduced noise with increased level of IR; no difference in attenuation	40 and 60 % ASIR had higher image quality and increased interpretable coronary artery segments			
Leipsic et al. [34•]	ASIR (40 %)	Individually selected; partially BMI-based	Individually selected; partially BMI- based	No difference in attenuation, noise, or SNR	No difference in interpretability per coronary artery or per patient		44 % reduction with IR (2.3 vs. 4.1 mSv).27 % reduction with IR after adjusting for scan settings	"Real-world" observational study
Gosling et al. [18]	ASIR	Individually selected; partially BMI-based	Individually selected; partially BMI- based				54 % reduction with IR (2.5 vs. 5.4 mSv)	"Real-world" observational study
Stolzman et al. [36]	ASIR and MBIR	Ex-vivo CT and IVUS of coronary atteries	Intrapatient comparison			No significant difference in accuracy or reader reliability between FBP, ASIR, or MBIR		
Shen et al. [42]	ASIR (40 %)	Adaptive procedure with target noise 35 HU	Adaptive procedure with target noise 35 HU	No difference in SNR			31 % reduction with IR (9.9 vs. 6.8 mSv)	
Scheffel et al. [41]	ASIR and MBIR	Ex-vivo CT of hearts	Intrapatient comparison	MBIR had lowest noise, followed by ASIR and then FBP. CNR highest with MBIR	MBIR had higher percentage of coronary cross-sections rated "excellent" (26 %) compared with ASIR (4 %) and FBP (13 %)			
Fuchs et al. [57]	ASIR at 20, 40, 60, 80, and 100 %	Routine acquisition	Intrapatient comparison	No difference in assessment of plaque volume, plaque components, vessel measurements, or stenosis measurements				

262

Table 1 con	continued							
Ref.	IR method	IR acquisition and reconstruction procedure	FBP acquisition and reconstruction procedure	Quantitative image quality	Qualitative image quality	Diagnostic accuracy	Radiation dose reduction	Notes
Yoo et al. [14]	AIDR 3-D	640-slice CT with lowest possible voltage; determined by automatic exposure control	Intrapatient comparison	IR had lower noise, higher SNR and CNR	Subjective image quality and mid and distal coronary artery interpretability improved with IR			
Tomizawa et al. [31]	AIDR	Individually selected	Individually selected	No difference in noise, SNR, or CNR	No difference in subjective image quality		22 % reduction with IR	"Real-world" observational study
Tatsugami et al. [30]	AIDR	320-slice single heartbeat cCTA	Intrapatient comparison	IR had lower noise, higher SNR	Subjective image quality higher with IR			
Chen et al. [38]	AIDR 3-D	Routine acquisition; reconstructed simulating 50 % dose reduction	Intrapatient comparison; reconstructed at full dose and simulating 50 % dose reduction	SNR and CNR similar to slightly higher with IR at 50 % dose	Subjective image quality higher with IR at 50 % dose		Simulated 50 % dose reduction with IR	
Renker et al. [27•]	IRIS	80 or 100 kVp acquisition	120 kVp acquisition	IR had lower noise	Subjective image quality higher with IR		62 % reduction with IRIS	
Renker et al. [26]	IRIS	Routine acquisition	Intrapatient comparison	IR had lower noise	Subjective image quality higher with IR	Improved measures of per-segment diagnostic accuracy with IR (overall diagnostic accuracy 95.9 vs. 91.8 %)		
Park et al. [24]	IRIS	100 kV; 200–320 mAs based on BMI	100–120 kV based on BMI; 320 mAs	IR had lower noise, higher SNR. No difference in spatial resolution	No difference in subjective image quality		40-51 % radiation dose reduction with IR, depending on BMI	
Bittencourt et al. [16]	IRIS	Routine acquisition	Intrapatient comparison	IR had lower noise, higher SNR.	No difference in subjective image quality of number of evaluable segments per patient			
Wang et al. [39]	SAFIRE	Routine acquisition; reconstructed simulating 50 % dose reduction	Intrapatient comparison; reconstructed at full dose	No difference in noise or SNQ	No difference in subjective image quality		49 % estimated dosereduction (4 vs.7.9 mSv)	
Takx et al. [29]	SAFIRE	Routine acquisition; reconstructed simulating 80 % dose reduction	Standard acquisition; reconstructed at full dose and simulating 80 % dose reduction	IR at 80 % dose reduction had lower noise, higher SNR and CNR than FBP at full dose	IR at 80 % dose reduction had higher subjective image quality		80 % simulated dose reduction	
Schuhbaeck et al. [28]	SAFIRE	Ultra-low-dose acquisition (80 kVp; 50 mAs; average effective dose 0.06 mSv)	Intrapatient comparison	IR had lower noise. No difference in SNR or CNR	IR had higher subjective image quality			

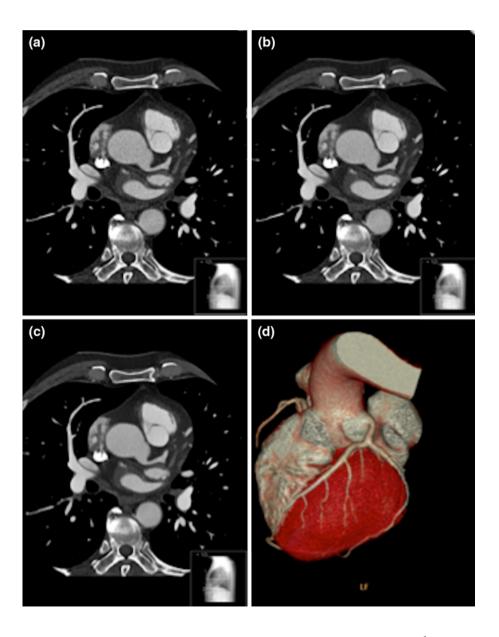
Table 1 continued	ntinued							
Ref.	IR method	IR acquisition and reconstruction procedure	FBP acquisition and reconstruction procedure	Quantitative image quality	Qualitative image quality	Diagnostic accuracy	Radiation dose reduction	Notes
Moscariello et al. [12]	SAFIRE	Routine acquisition; reconstructed simulating 50 % dose reduction	Intrapatient comparison; reconstructed at full dose	IR at 50 % dose reduction had lower noise	IR at 50 % dose reduction had higher subjective image quality	IR at 50 % dose reduction had improved accuracy in stenosis detection (overall accuracy 96.9 vs. 93.8 %)	Simulated 50 % dose reduction	
Utsonomiya et al. [33]	iDOSE (levels 3 and 7)	Routine acquisition	Intrapatient comparison	IR had higher SNR. No difference in attenuation	IR had higher subjective image quality. iDOSE 7 had improved visualization of distal vessels and number of assessable coronary artery segments			
Oda et al. [23]	iDOSE (level 7)	Low-dose (80 kVp) acquisition	Intrapatient comparison	IR had lower image noise, higher SNR	IR had higher subjective image quality			
Hou et al. [15]	iDOSE (levels 3, 4, and 5)	Tube currents of 600, 500, and 400 mAs for iDOSE levels 3, 4, and 5, respectively	Tube current of 1,000 mAs	IR at lower doses had lower noise and higher CNR than high-dose FBP.			55 % dose reduction with the 500 mAs acquisition using IR compared with FBP	500 mAs procedure using IR had the best compromise of dose reduction and image quality on ROC analysis
Hou et al. [40]	iDOSE (levels 3, 4, and 5)	Tube currents of 125, 105, 84, and 64 mAs for iDOSE levels 3, 4, 5, and 5, respectively	Tube current of 125 mAs	No difference in noise, CNR, or SNR between all groups	No difference in subjective image quality between FBP and all IR groups		63 % dose reduction with the 84 mAs acquisition using IR compared with FBP	84 mAs procedure using IR had the best compromise of dose reduction and image quality on ROC analysis
Hosch et al. [20]	iDOSE at different levels	Fixed routine acquisitions, low- dose BMI adapted acquisition, and fixed low-dose acquisitions were performed	Intrapatient comparison	BMI-adapted procedures with IR had increased SNR and CNR compared with fixed routine procedures using FBP			63 % dose reduction with BMI-adapted acquisitions compared with fixed routine acquisitions (1.2 vs. 3.2 mSv)	

that may not be valid in clinical practice or have limitations in generalization to a broader population. BMI-adaptive reconstructions using predefined acquisition settings based on patient body habitus offer a potential solution [24, 25]; other groups have proposed patient-specific adaptive-dose procedures that adjust scan settings on the basis of allowable image noise [37, 42]. Notably, specific IR products vary in their patient-specific adaptability.

The use of IR methods has also been examined specifically in pediatric populations, with similar results. Both SAFIRE and IRIS have shown improvements in image noise, CNR, SNR, and SIQ compared with equivalent-dose FBP reconstructions [19, 32]. Simulated half-dose IRIS and SAFIRE reconstructions in pediatric cardiovascular CT compare favorably with full-dose FBP reconstructions, with SAFIRE showing the greatest benefit [32].

The noise-reduction properties of IR may hold particular appeal in the evaluation of obese individuals, in whom noise is intrinsically higher secondary to reduced photon transmission. One study demonstrated a 50 % reduction in mean effective radiation dose using cardiovascular CT procedures with 100 kV tube current and SAFIRE reconstructions compared with 120 kV procedures reconstructed with FBP, without detrimental effects on image noise, SNR, CNR, or SIQ. The patient population was limited to individuals with BMI >30 kg/m² [43]. Similar results have been reported for use of ASIR for obese individuals undergoing CT examinations of the abdomen and pelvis [44].

Fig. 6 Transverse a FBP, and IRs using iDose (Philips Healthcare) at iDose level b 4 and c 7, at the level of the aortic root. A continuous decrease of image noise can be seen from images a through c. d Shows a 3D volume rendered image reconstructed using iDose level 7. (Images courtesy of Armin Huber, Department of Radiology, Klinikum rechts der Isar, Technische Universität München, Munich, Germany.)



Coronary Artery Stent Evaluation

Evaluation of coronary artery stents using cardiovascular CT has been vigorously investigated but remains a challenge because of limitations in temporal resolution, spatial resolution, and beam-hardening artifacts [45]. Recent work has examined several potential roles of IR in improving CT stent evaluation. Studies simply comparing IR to FBP reconstructions for patients with coronary artery stents have demonstrated improvements in image quality with dose reductions similar to those described above. Reductions in stent volumes [27•, 46] and image noise [27•, 46-48] have been reported, with improved in-stent visualization [46, 48] (Fig. 6). IR may also provide value in high-resolution CT stent imaging. Spatial resolution is a major limitation in stent evaluation, prompting the development of high-resolution acquisition (usually 0.23 mm spatial resolution). Reduced voxel size in these images leads to significant problems with image noise because of photon starvation; this provides the basis for IR implementation. Early studies have shown improvements in noise, blooming artifacts, in-stent visualization, and diagnostic accuracy using IR in conjunction with high-resolution reconstruction kernels and acquisitions [49-51]. Custom-built IR algorithms that specifically target blooming artifacts may provide yet another solution for CT stent evaluations [52].

Coronary Calcium Evaluation

Similar concepts provide the basis for IR use in heavily calcified vessels. Renker et al. [26] demonstrated significantly lower image noise and calcification volumes with significantly higher SIQ when using IRIS compared with FBP for patients with Agatston scores \geq 400 (Fig. 4). IR also led to significant improvements in several measures of per-segment diagnostic accuracy for detection of significant stenoses, with overall diagnostic accuracy of 95.9 % for IRIS compared with 91.8 % for FBP, using invasive coronary angiography as reference standard. Reductions in calcification volume using IR should be considered when performing noncontrast cardiovascular CT for calcium scoring; one study comparing ASIR to FBP in CT calcium scoring showed decreased noise and reduced Agatston and volumetric calcium scores when using IR compared with FBP [53]. Because calcium score risk stratification uses data from previous population-based studies that used FBP reconstructions, coronary calcium volume reduction with IR has the potential to result in diverging risk stratification and subsequent aggressiveness of risk factor modification.

Future Directions

Short-term objectives for IR should include refinement of current techniques and applications. As noted above, the optimum strategy for IR-based radiation dose reductions is not established. BMI-adaptive [24, 25] and noise-adaptive [37, 42] acquisition procedures have been proposed. In addition, several IR products enable user-specified modulation of iterative strength or proportion of blending with FBP images, and researchers should seek to evaluate the best combinations for routine clinical work. Finally, more data regarding the effects of IR on diagnostic accuracy, prognostic value, and clinical outcomes will be necessary to justify widespread clinical adoption.

Early studies have shown that IR may provide cardiovascular CT solutions beyond traditional noise and dose reductions. High-resolution acquisitions are becoming more feasible as IR tempers the negative effects of photon starvation [49–51]. Contrast material dose reductions may also be possible without prohibitive degradation of image quality [22]. Novel IR algorithms are also emerging that should expand IR applications. Beam-hardening correction algorithms have been proposed for use in cardiovascular CT myocardial perfusion imaging to reduce artifacts related to high-concentration contrast material in the descending aorta and left ventricle [54]. Preliminary data on an IR technique that imposes penalties on dense tissues has likewise suggested a role for IR in reducing blooming artifacts [52], while a novel "metal detection technique" described by Boas and Fleischmann [55] demonstrated reduced beam-hardening artifacts from metal and bone. Future algorithms can be expected to further incorporate system, geometric, and motion models [8., 56].

Conclusion

Increases in available and affordable computer power have provided the basis for (re)application of IR to cardiovascular CT. Current data suggest that commercially available IR products provide noise reduction that enables significant reduction in radiation dose without degrading image quality. Refinement of current IR techniques should stimulate increased clinical adoption, and emerging and future algorithms promise to expand the role of IR to novel cardiovascular CT applications.

Compliance with Ethics Guidelines

Conflict of Interest John W. Nance Jr and Ullrich Ebersberger declare that they have no conflict of interest. U. Joseph Schoepf receives honoraria from and is a paid consultant for Bayer and Siemens and receives payment for development of educational presentations from GE.

Human and Animal Rights and Informed Consent This article does not contain any studies with human or animal subjects performed by any of the authors.

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