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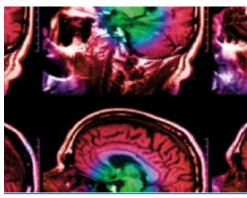


TOPICAL REVIEW • OPEN ACCESS

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TOPICAL REVIEW

Photon-counting x-ray detectors for CT

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Abstract

The introduction of photon-counting detectors is expected to be the next major breakthrough in clinical x-ray computed tomography (CT). During the last decade, there has been considerable research activity in the field of photon-counting CT, in terms of both hardware development and theoretical understanding of the factors affecting image quality. In this article, we review the recent progress in this field with the intent of highlighting the relationship between detector design considerations and the resulting image quality. We discuss detector design choices such as converter material, pixel size, and readout electronics design, and then elucidate their impact on detector performance in terms of dose efficiency, spatial resolution, and energy resolution. Furthermore, we give an overview of data processing, reconstruction methods and metrics of imaging performance; outline clinical applications; and discuss potential future developments.

1. Introduction

1.1. A brief history of photon-counting detectors

To count photons is the most intuitive approach for detecting x-rays, and if it were not for technical challenges, photon-counting detectors (PCDs) would have been standard from the beginning of radiology. Early on, gas detectors were commonplace, and Geiger–Müller devices counted individual interactions of ionizing radiation. The Nobel-prize-awarded multi-wire proportional chamber (Charpak 1997) combined photon counting with spatial resolution. The development was driven by fundamental physics research, but photon-counting gas detectors were briefly used for imaging in a Paris hospital (Dubousset *et al* 2007). In nuclear imaging, photon counting was used from the very beginning for the Anger camera as well as for the first PET system. In this case, a scintillator, typically NaI or CsI, converted the incident gamma rays into visible light that was detected by photosensitive devices. For x-ray imaging the challenges for photon counting are much harder than for nuclear imaging. The average energy of the photons is only around 70 keV as compared to 140 keV for SPECT and 511 keV for PET. Moreover, the x-ray fluence rate for a computed tomography (CT) scan can be up to $10^9 \text{ mm}^{-2} \text{ s}^{-1}$, while for nuclear imaging it is as low as 100 mm⁻² s⁻¹, putting much higher constraints on fast pulse processing for x-ray imaging.

The first photon-counting imaging system approved by the U.S. Food and Drug Administration was the Sectra MicroDose Mammography (Åslund *et al* 2007) in 2011, with around one thousand installations worldwide for breast-cancer screening and diagnosis. The first full-field photon-counting CT prototype was evaluated in the clinic in 2007 based on a CdZnTe detector, and though limited in the count rate it could accommodate, it still produced material-specific images of high quality (Benjaminov *et al* 2008). There are currently at least four photon-counting CT systems under evaluation. Three of these, including one mobile head CT system, utilize cadmium-based sensors (Yu *et al* 2016c, Si-Mohamed *et al* 2017a, Han-soo 2017), and one uses silicon-based sensors (da Silva *et al* 2019). In particular, one cadmium-based system has generated a large number of publications (Yu *et al* 2016a, 2016c, Symons *et al* 2018a, 2018b, 2017b, Pourmorteza *et al* 2016, 2017). Furthermore, a CdTe-based photon-counting system limited to breast CT (Kalender *et al* 2017) has been evaluated on patients. We are now at a crossroads to emerging clinical systems

within the next few years, and the different solutions will likely continue to compete based on imaging performance, reliability, and cost.

1.2. The clinical case for photon-counting CT

Despite the considerable improvements that CT technology has undergone in nearly half a century of existence, there are numerous areas where the improvements promised by PCDs can provide solutions to clinical problems (Willemink *et al* 2018). Their higher spatial resolution can improve visualization of lung nodules (Kopp *et al* 2018); small bone details, such as in temporal bone imaging; small blood vessels, such as coronary arteries where calcium blooming is a problem; or implants such as stents (Symons *et al* 2018a, Mannil *et al* 2018, Sigovan *et al* 2019). The improved contrast-to-noise ratio (CNR), in particular for contrast-enhanced imaging (Gutjahr *et al* 2016), can be used, for example, to improve the visualization of blood vessels (Symons *et al* 2018b) or contrast-enhancing tumors. PCDs have also been demonstrated to improve differentiation between grey and white matter in the brain (Pourmorteza *et al* 2017). Furthermore, the ability to generate material-selective images has applications in neuroradiology, such as distinguishing intracranial hemorrhage from calcification (Hu *et al* 2016) and iodinated contrast (Phan *et al* 2012), for which dual-energy CT has been shown to be effective. The potential for improved material-selective imaging with PCDs may also offer better visualization of atherosclerotic plaques by improving the ability to separate materials such as lipid, calcium, and iodine (Alessio and MacDonald 2013, Boussel *et al* 2014).

Another promising aspect is improved low-dose imaging. The health hazard posed by low-dose ionizing radiation is a contested question, but pediatric patients undergoing CT examinations with cumulative doses of more than 50–60 mGy have been shown to exhibit an increased risk of brain cancer and leukemia (Pearce *et al* 2012). In spite of this, CT is often the imaging alternative with best benefit versus risk for the patient, and the dose in x-ray imaging should therefore be kept as low as reasonably achievable. By rejecting electronic noise, PCDs can allow new imaging protocols with lower doses than what is possible today. This fact can have important implications for imaging of children and adolescents, who are more radiation-sensitive than adults, and for lung-cancer screening, where it is particularly important to minimize the dose since the examination is performed on large numbers of healthy individuals (Symons *et al* 2016).

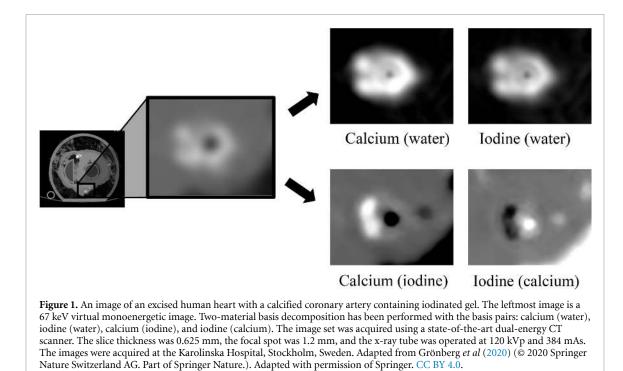
The energy-resolving capabilities of PCDs can be used to reduce artifacts due to beam hardening or metal implants (Nasirudin *et al* 2015). They can thus provide improved diagnostic quality when imaging regions with weak attenuation differences surrounded by dense bony structures, such as in the brain (Pourmorteza *et al* 2017). Another potential application is found in the emerging field of radiomics, where large numbers of quantitative radiomic features are extracted from medical images and analyzed to find patterns indicative of diagnosis. These radiomic features are currently dependent on acquisition parameters, which limits reproducibility (Berenguer *et al* 2018). The potential of photon-counting CT to generate quantitative material maps can mitigate this variability. Improved tissue characterization also shows potential for improving radiotherapy planning (Simard *et al* 2019).

Among the more speculative proposed applications of photon-counting CT are the use of new contrast agents. Gadolinium, which is used in MRI, is an attractive candidate for photon-counting CT because of the location of its K-edge in the central part of the diagnostic energy range (Roessl *et al* 2011). However, it is still unclear whether gadolinium contrast agents can generate sufficient contrast at patient-safe concentrations. Dual- or triple-contrast agent protocols, where iodine, gadolinium, and potentially also bismuth are administered at different time points to show different phases of contrast uptake in a single scan, have also been proposed (Symons *et al* 2017a, 2017b, Dangelmaier *et al* 2018). Further, there is ongoing research into the development of new nanoparticle-based contrast agents containing heavy elements such as gold, tantalum, or ytterbium (Cormode *et al* 2017, Kim *et al* 2018, Lambert *et al* 2018).

As an example of the clinical use of a high-resolution spectral PCD, an excised human heart with a calcified coronary artery containing an iodinated gel was imaged using a state-of-the-art dual-energy CT scanner (figure 1) and a prototype silicon-based spectral PCD (figure 2). The imaging task was to visualize the artery and separate calcified regions from iodine. The dual-energy image was decomposed into several material basis pairs: calcium (water), iodine (water), calcium (iodine) and iodine (calcium), where the basis not shown is within the parenthesis. The spectral PCD images (figure 2) were decomposed into three bases: water, iodine, and calcium. An overlay image was formed using the basis images in which calcium and iodine were colored red and green, respectively. The high resolution combined with the simultaneous spectral acquisition facilitated by the multi-energy-bin PCD shows potential to improve the diagnostic quality of the CT images.

1.3. Outline

The purpose of this article is to summarize and discuss recent insights into the characteristic features of PCDs and how they relate to the imaging performance of the CT system. Section 2 outlines the theoretical



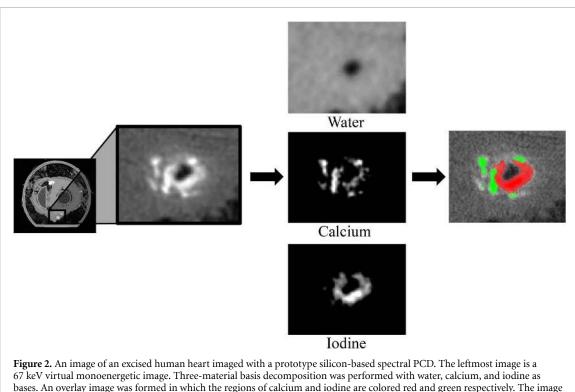


Figure 2. An image of an excised human heart imaged with a prototype silicon-based spectral PCD. The leftmost image is a 67 keV virtual monoenergetic image. Three-material basis decomposition was performed with water, calcium, and iodine as bases. An overlay image was formed in which the regions of calcium and iodine are colored red and green respectively. The image was acquired at 120 kVp using a focal spot 0.4 mm, and reconstructed with a 0.625 mm slice thickness. The dose was matched to that of the dual-energy image in figure 1. Adapted from Grönberg *et al* (2020) (© 2020 Springer Nature Switzerland AG. Part of Springer Nature.). Adapted with permission of Springer. CC BY 4.0.

benefits of PCDs over energy-integrating detectors (EIDs). Section 3 outlines the different design choices that must be made when developing a PCD. Section 4 discusses how image quality is affected by the detector properties. Section 5 describes the steps needed to generate images from the measured data. Section 6 describes different metrics of imaging performance that can be used to guide the system design. Finally, in section 7, we discuss what these insights mean for the future of photon-counting CT.

3

2. Benefits of PCDs

2.1. Weighting of photons

In the photon-detection process and/or image formation, different weight can be given to photons of different energy, affecting both the contrast and the noise of the signal.

The contrast between two projection measurements depends on the energy of the transmitted x-ray photons. Generally, low-energy photons carry more contrast information than high-energy photons. Also, if two projection measurements have different material composition, the dependency of contrast on photon energy is elevated, and increased contrast can be obtained by increasing the weight given to photons that carry more contrast information. Weighting some photons more than others increases the variance relative to the mean value, thus reducing the signal-to-noise ratio (SNR). There is a fundamental trade-off between the increase of contrast and the reduction of the SNR, and there is an optimal set of photon weights that maximizes the CNR for a given imaging task (Schmidt 2009). The weighting of photons will often be performed indirectly via the material-decomposition process (see section 5.2), where it can be seen as the utilization of the contrast and noise of the signal to obtain a material thickness estimation with minimum variance.

An EID weights each detected photon by its energy, thus adding additional weight to high-energy photons, for which the contrast is lower. In addition, the non-uniform weighting of photons results in a reduction of the SNR. The reduction of the SNR² of an EID due to the weighting of the photons is commonly referred to as the Swank factor (Swank 1973), and it is given by

$$\mathbf{I}_{\rm EID} = \frac{\left[\int_0^\infty \epsilon S(\epsilon) d\epsilon\right]^2}{\int_0^\infty \epsilon^2 S(\epsilon) d\epsilon},\tag{1}$$

where ε is signal amplitude and $S(\varepsilon)$ is the normalized distribution of signal amplitudes registered by the detector. The magnitude of the Swank factor depends on the distribution of signal amplitudes; the more the amplitudes are distributed over different energies, the lower the Swank factor. The distribution of signal amplitudes in turn depends on the spectrum from the x-ray tube, the attenuation of the imaged object, and the response of the detector.

Ideal purely PCDs intrinsically weight all photons equally (one photon, one count). Counting photons therefore gives relatively more weight to low-energy photons compared to energy integrating, resulting in a higher contrast, in particular for low absorbing materials.⁴ PCDs also avoid the negative effect of the Swank factor. In addition, spectrally resolving PCDs allow giving different weight to photons of different energy as a part of the signal processing, and the weighting can be, for example, tailored to maximize the CNR.

As an illustration of the effect of photon weighting, consider the task of separating between two projection measurements with a typical 120 kVp x-ray tube spectrum for which the x-rays pass through (a) 10 cm water and (b) 9 cm water plus 1 cm of 10 mg ml⁻¹ iodine-water solution, respectively. In this example, we simulate an ideal PCD that detects all photons and registers the counts in two energy bins, s_{low} and s_{high} , separated by a threshold at 50 keV. The results are compared to a simulation of an ideal EID that integrates the energy for all transmitted photons for the same imaging task. In order to optimize the CNR for the PCD, a weighted sum of the two energy bins is formed:

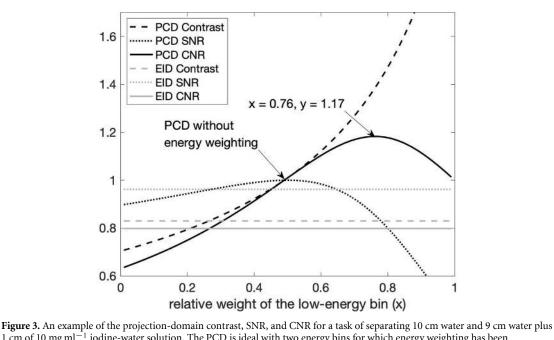
$$s = xs_{\text{low}} + (1 - x)s_{\text{high}},\tag{2}$$

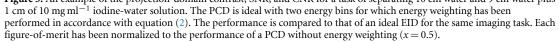
where $0 \le x \le 1$, where x = 0.5 corresponds to the PCD without energy weighting. As figures-of-merit we evaluate the following: the contrast between the two projections, defined as $C = (s_A - s_B)/s_A$; the signal-to-noise ratio, defined as $SNR = s_A/\sigma(s_A)$, where σ is the standard deviation; and finally, the contrast-to-noise ratio, defined as $CNR = C \times SNR = (s_A - s_B)/\sigma(s_A)$.

The contrast, SNR, and CNR for the PCD are plotted versus *x* in figure 3, and the performance of the EID is included for reference. The SNR of the PCD obtains its maximum at x = 0.5, i.e. equal weight to both energy bins. The contrast, on the other hand, increases monotonically as more weight is given to the low-energy bin (higher *x*). The CNR, which is the product of the contrast and the SNR, reaches a maximum of 1.17 for x = 0.76. The corresponding relative CNR², which is proportional to x-ray dose, is 1.37. In other words, for this imaging case, it is motivated to sacrifice SNR in order to gain contrast, and thus achieve a superior CNR.

For the simulated ideal EID (gray lines in figure 3), on the other hand, the suboptimal energy weighting results in a relative contrast of 0.82 (compared to the PCD without energy weighting), and a relative SNR of

⁴ Materials with low absorption transmit relatively more low-energy photons.





0.96 (in accordance with the Swank factor). In total, the sub-optimal energy weighting of the EID results in a relative CNR of 0.79, or, equivalently, a relative CNR^2 of 0.63, which is to be compared with 1.37 for the PCD with optimal energy weights. In other words, the ideal spectral PCD with optimal energy weights is, for this particular imaging task, 2.17 times more dose efficient compared to the ideal EID. It can be noted that the performance of the EID is similar to that of the spectral PCD with x = 0.27.

2.2. Material-specific imaging

Another benefit of energy-resolving PCDs is that they allow measuring the composition of the imaged object through a process known as material decomposition. This method is based on the fact that the linear attenuation coefficient $\mu(E)$ of any material in the human body can be well approximated by a linear combination of a small number of basis functions $f_j(E)$: $\mu(E) = \sum_{j=1}^{N_m} a_j f_j(E)$, where the coefficients a_i are referred to as basis coefficients (Alvarez and Macovski 1976). Since there are two dominant physical processes contributing to x-ray attenuation in the diagnostic CT energy range, namely photoelectric absorption and Compton scattering, $N_m = 2$ basis functions are commonly assumed to be sufficient for substances containing only light elements, such as human tissues. In addition to these two, one additional basis function must be included for each heavy element whose attenuation coefficient contains a K-edge discontinuity within the diagnostic energy range (Roessl and Proksa 2007, Schlomka *et al* 2008).

This low dimensionality means that it is possible to completely characterize the energy dependence of the linear attenuation coefficient at every point in the imaged volume with a small number of energy bins: at least two bins are needed for non-enhanced imaging, and three bins with a contrast agent. Compared to the dual-energy systems in clinical use today, the capability to measure the amounts of three or more basis materials is one of the benefits of PCDs, along with other advantages such as the reduction of spectral overlap, the absence of spatial mismatch between the energy-bin images, and the ability to obtain spectral information also in the peripheral parts of the image as opposed to dual-source systems, where one detector has a limited field of view due to geometric constraints.

From the energy-resolved measured data, the material composition at each point in the imaged volume can be inferred through material decomposition. This process, which can take place before, during, or after the image reconstruction, (see section 5.2) results in a set of reconstructed basis images. These basis images show the distribution within the imaged object of each of the selected basis materials, e.g. water, calcium, and iodine. Once such a set of basis images has been generated, it is straightforward to obtain distribution maps of other substances (e.g. muscle and fat) through a simple linear transformation. Another possibility is to use the basis material maps to generate images of the x-ray attenuation coefficient at each energy (i.e. virtual monoenergetic images; see Leng *et al* (2017)), which are free of beam-hardening artifacts (see section 5.4).

A further possibility is to form a linear combination of the basis images to generate an image that is optimal for maximizing detectability for a specific imaging task. It turns out that this methodology gives a detectability for an optimal linear observer that is equal to the detectability obtained from optimal weighting of the original energy-bin images, provided that photon statistics are sufficient (Alvarez 2010, Persson *et al* 2018b). In other words, as long as the detector is operating far enough above the fluence level that causes photon starvation, the material-decomposition process preserves the information available in the measured data. Since material decomposition also has the theoretical ability to remove beam hardening completely, displaying weighted basis images makes it possible to avoid the trade-off between beam-hardening artifacts and image detectability that is observed when generating weighted sums of the original bin images (Shikhaliev 2005).

Since a larger number of estimated basis components makes the material decomposition process more ill-conditioned, the benefit of adding each new basis material diminishes with an increasing number of basis functions used in the decomposition. At the same time, using a detector with a large number of energy bins makes it possible to choose a number of basis functions that is optimal for a given task. By using four or more energy bins, it is thus possible, in principle, to quantify several contrast agents independently.

Furthermore, it is still an open research question whether the two-basis approximation is exact down to the precision that photon-counting CT scanners are able to measure. Multiple studies indicate that the two-material approximation is insufficient for performing high-precision measurements of the attenuation coefficient (Williamson *et al* 2006, Bornefalk 2012b, Alvarez 2013). This suggests that it should be possible to separate more than two components (e.g. water, calcium and iron) from each other when imaging unenhanced human tissue. Measuring the concentration of three materials can be desirable when imaging, for example, vulnerable plaques (Wang *et al* 2011b) or liver iron overload (Luo *et al* 2015).

However, even if three-material decomposition of unenhanced tissue turns out to be feasible, it is likely that the high sensitivity to noise in three-basis decomposition will limit its application to large-area tasks (Alvarez 2013), unless prior assumptions about mass or volume preservation are included (section 5.2). The two-basis approximation is also inaccurate at sharp edges in the object, which get a unique spectral signature due to the non-linear partial volume effect (Glover and Pelc 1980), a phenomenon that can be used to obtain subpixel spatial information from spectral measurements (Persson *et al* 2018a).

2.3. Spatial resolution

One of the main advantages of the PCD is the improved spatial resolution compared to conventional detectors. PCDs that have been designed for full-body clinical CT generally have pixel sizes ranging from 0.225 to 0.5 mm at the detector; this is smaller than conventional EIDs today, which have pixel sizes generally on the order of 1 mm (Leng *et al* 2016, Persson *et al* 2014, Shefer *et al* 2013, da Silva *et al* 2019). Recent developments have, however, enabled improvements of the spatial resolution of EIDs reaching approximately 0.5 mm pixel pitch at the detector (Yanagawa *et al* 2018).

The spatial resolution of EIDs has been held back by several practical limitations. To mitigate pixel crosstalk, the detectors are made up of scintillator crystals that have to be diced into pixels, and on the edges of each pixel, there is a reflecting material which keeps the secondary light from escaping the pixel (Shefer *et al* 2013). The finite thickness of the reflectors creates dead area between the pixels, reducing the geometric efficiency. If the pixels are made smaller, the dead area takes up a larger fraction of the detector area. Semiconductor detectors, on the other hand, do not emit any secondary light and therefore require no reflectors. Instead, the detector consists of a continuous piece of semiconductor material, and these direct conversion detectors are pixelated by charge-collecting electrodes. The pixels can therefore be made smaller without losing geometric efficiency.⁵

Also, due to the nature of the energy-integrating signal formation, there is a small contribution of integrated electronic noise for each detector channel. If the pixels are made smaller, the number of detector channels per detector area increases, as does the total amount of integrated noise. A PCD, on the other hand, can adapt the lowest energy threshold to the noise floor for each channel and avoid counting noise, even for very small pixels.

Several approaches have been suggested for improving the spatial resolution of scintillator detectors, such as using an attenuating grid to obtain a smaller pixel aperture (Flohr *et al* 2007). However, this comes at the expense of a substantial reduction in dose efficiency. It has also been proposed to use high-resolution flat-panel detectors for CT (Gupta *et al* 2006). However, there are some remaining issues that have to be addressed in order to make the technology feasible, including the characteristics of the scintillator material

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⁵ At the expense of an increased fraction of charge sharing; see section 3.2.4.

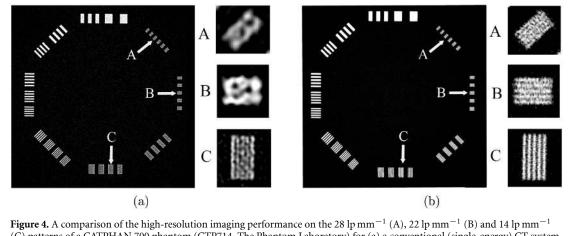


Figure 4. A comparison of the high-resolution imaging performance on the 28 lp mm⁻¹ (A), 22 lp mm⁻¹ (B) and 14 lp mm⁻¹ (C) patterns of a CATPHAN 700 phantom (CTP714, The Phantom Laboratory) for (a) a conventional (single-energy) CT system with 0.7 mm focal spot, (b) a prototype silicon-based PCD with 0.4 mm focal spot. Both images were acquired with axial mode at 120 kVp with comparable dose and reconstructed with a high-resolution kernel. Adapted with permission from da Silva *et al* (2019). © 2019 Society of Photo-Optical Instrumentation Engineers (SPIE).

(e.g. speed, afterglow, and lag), object scatter corrections due to removal of the anti-scatter grid, and the dynamic range of the detector.

The benefit of the improvement in spatial resolution made possible by PCDs has been demonstrated on multiple occasions (Leng *et al* 2018, Bartlett *et al* 2019, Symons *et al* 2018a, von Spiczak *et al* 2018). Figure 4 shows an example of the high-resolution performance of a conventional CT system and a silicon-based full-body PCD prototype. Details about the prototype system can be found in da Silva *et al* (2019). In this example, the PCD can resolve approximately twice as many line pairs per millimeter as the conventional CT system design. For example, if the spectral separation comes from the source, it is common to use a larger focal-spot size to facilitate a sufficient tube output. Spectral PCD systems, on the other hand, will facilitate simultaneous spectral and high-resolution imaging. Figure 5 shows a comparison of the spatial resolution of a state-of-the-art dual-energy system and a silicon-based full-body PCD prototype for imaging the structures of the inner ear.

In addition to the improved spatial resolution in the transaxial and longitudinal direction caused by the smaller pixel size, PCDs also have an advantage over EIDs in terms of the angular spatial resolution. The reason for this is twofold: First, PCDs do not suffer from scintillator afterglow which can lead to lag between consecutive measurements in EIDs if the sampling rate is high. Second, the electronic noise level in EIDs tends to increase for higher sampling rate, so that increasing the angular resolution leads to a penalty in dose efficiency. PCDs, on the other hand, are able to reject the electronic noise as will be discussed in section 2.4 and can therefore avoid this dose penalty. This allows higher sampling rates, provided that enough bandwidth is available to read out the data (Sjölin and Danielsson 2017). Potentially, the faster sampling rates could also be used to detect rapid motion.

With an increased spatial resolution of the detector comes a risk of introducing aliasing artifacts in the reconstructed image. This can be avoided by maintaining sufficient radial and angular sampling rates. In the radial direction, the sampling rate is automatically increased since the sampling interval (pixel center-to-center distance) is reduced. In the angular direction, on the other hand, the sampling rate (number of views per revolution) must by increased in order to avoid visible angular blur and aliasing artifacts.

As a final remark on spatial resolution, we note that the detector pixel size and the angular sampling rate are not the only factors impacting the spatial resolution of the imaging system. In order to fully utilize the increased spatial resolution of the detector, appropriate adjustments must be made to the focal spot size, image reconstruction grid, and reconstruction algorithms. All of these aspects significantly impact the CT imaging chain.

2.4. Low-dose imaging

A further application of photon-counting imaging is low-dose imaging. There are two reasons why PCDs are able to provide superior performance for low-dose imaging tasks compared to energy-integrating detectors.

The first reason is the overall improvement in image quality achievable with PCDs, as described in previous sections. This includes CNR improvement through better utilization of the energy information in the detected x-ray beam and improvement in visualization of small objects. Since the detectability of a

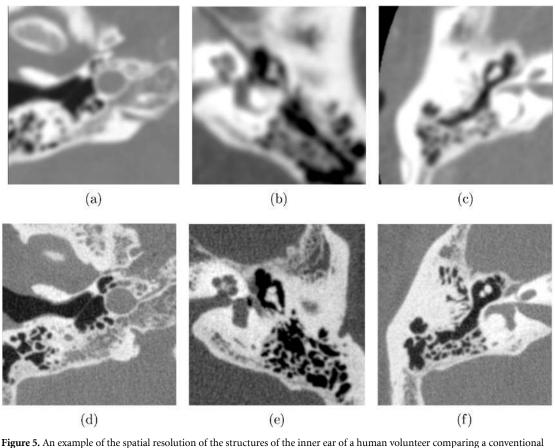


Figure 5. An example of the spatial resolution of the structures of the inner ear of a human volunteer comparing a conventional dual-energy CT system (top row), and a silicon-based full-body PCD prototype (bottom row). The images were dose-matched, and both images were acquired at 120 kVp and reconstructed at 67 keV monoenergy. The dual-energy images were acquired using 1.2 mm focal spot, and the PCD images were acquired using a 0.6 mm focal spot. The dual energy images were acquired at the Karolinska Hospital, Stockholm, Sweden. The PCD prototype is described in described in da Silva *et al* (2019).

feature in the image increases with dose, these potential improvements in image quality at equal dose can be traded for a lower dose by keeping the image detectability fixed and lowering the tube current instead.

Second, PCDs have a particular advantage for imaging with low dose levels, namely their ability to reduce the impact of electronic noise. PCDs use a threshold to separate real counts from noise, and by setting the threshold high enough above the noise floor, the electronic noise can be rejected. In this case, the only degradation caused by electronic noise is a small broadening of the energy-response function due to random fluctuations in the pulse-height measurements. EIDs, on the other hand, measure the total x-ray energy deposited during a certain time interval, and the electronic noise will be included in the measurement as a random additive term. The magnitude of this term is relatively constant, whereas the quantum noise is proportional to the incident fluence rate; therefore the electronic noise can go more or less unnoticed at high doses but becomes prohibitive at low doses or for large patients (Duan *et al* 2013). On the other hand, the detective quantum efficiency (DQE) (see section 6.3) of a PCD is, theoretically, constant down to the zero-flux limit. The relative advantage of photon-counting CT compared to energy-integrating CT is therefore larger at low dose levels (Yu *et al* 2016a). PCDs may therefore allow new low-dose imaging protocols at dose levels where electronic noise is prohibitive with current state-of-the-art scanners, with potential applications in pediatric imaging and lung cancer screening (Symons *et al* 2016).

3. Detector design considerations

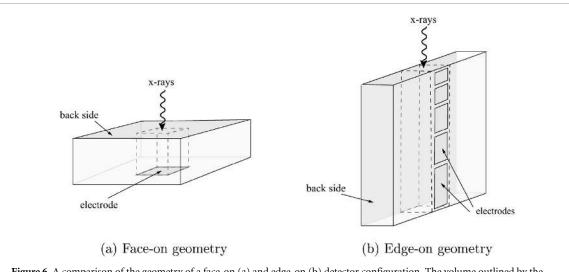
3.1. General functionality of a PCD

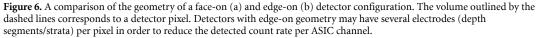
PCDs for x-ray CT are so-called direct-conversion detectors, wherein the x-ray photons are converted directly into an electric signal as opposed to first being converted to visible light. PCDs for CT generally consist of semiconductor sensors with an applied bias voltage. An interacting photon creates a cloud of charge carriers, creating a signal which is processed and registered by an application-specific integrated circuit (ASIC). Each detected photon interaction results in a count in an energy bin corresponding to the energy deposited in the interaction.

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Table 1. Comparison of properties of cadmium-based and silicon-based detectors. Compared to a CdTe/CZT detector, constructing a full-area silicon detector system is more complex because of the edge-on sensor geometry. FWHM, full width at half maximum.

Detector material	CdTe/CZT	Silicon
Preferred orientation	Face-on	Edge-on
System construction complexity	Less complex	More complex
Typical lowest threshold	20–25 keV	5–10 keV
Typical energy resolution (FWHM)	5–10 keV	3.5–5.4 keV
K-fluorescence escape probability	High	Low
Compton-scatter probability	Low	High
Mobility-lifetime product	Low	High
Amount of polarization at high fluence rate	High	Low
Material processing technology	Under development	Mature





Despite many years of development, PCDs have only recently approached the performance necessary for being used in clinical CT scanners. The main challenges are achieving good performance at high count rate, obtaining good spectral fidelity, and manufacturing a full-field detector with sufficiently low density of imperfections at a competitive cost. In the following, we examine the different detector design considerations that need to be accounted for in order to optimize detector performance.

3.2. The detector material

There are currently two main converter material candidates: cadmium (zinc) telluride (CdTe or CZT) and silicon (Si). Both candidates have pros and cons, some of which we will mention here, and their properties are summarized in table 1. More comprehensive accounts for the properties of the detector material and the functionality of the ASICs can be found in the literature (Ballabriga *et al* 2016).

3.2.1. Detector physics

One of the main differences between CdTe/CZT and Si detectors is the relative x-ray stopping power. CdTe has a high linear attenuation coefficient, requiring only about 1.7 mm to stop 95% of the x-rays in a 120 kVp spectrum filtered by 30 cm of water. Silicon, on the other hand, has a relatively low atomic number and requires roughly 55 mm to stop the same fraction of x-rays under the same conditions. As it is not feasible to make the semiconductor wafers much thicker than a few millimeters, the silicon wafers are mounted *edge-on* with respect to the incoming x-rays (Bornefalk and Danielsson 2010). This way, the effective depth of the detector is determined not by the thickness of the wafer, but by the length of the wafer, allowing the detector to have as long an absorption length as necessary. See figure 6 for an illustration of the face-on (CdTe/CZT) and edge-on (Si) geometries.

Another difference between the detector materials is the prevalence of the different types of x-ray interactions that occur in the material. In materials containing heavy elements (high-Z materials), such as CdTe/CZT, the main interaction mechanism is photoelectric absorption; Compton and Rayleigh scattering accounts for only a few percent of the total absorption. Silicon, on the other hand, has a high probability of

Interaction	Si	CZT
Total	0.94	0.94
Photoelectric	0.28	0.87
Compton	0.56	0.04
Rayleigh	0.10	0.04

Table 2. The distribution of primary interactions in a 50 mm deep Si detector and a 1.6 mm deep CZT detector for a 120 kVp spectrum attenuated by 30 cm of water.

Compton interactions, which dominate over the photoelectric effect for photons with energy higher than 48 keV. An example of the distribution of primary interactions in a 50 mm Si detector and a 1.6 mm CZT detector is shown in table 2. The distribution of interactions is computed for a typical 120 kVp x-ray tube spectrum that has been filtered through 30 cm of water (Cranley *et al* 1997, Hubbell and Seltzer 2004).

However, keep in mind that it is not straightforward to assess the signal quality based on the distribution of primary interactions in the detector material. For example, in the event of a Rayleigh scattering, there is a high likelihood that the photon will scatter with a small angle and interact again within the same detector pixel. In the same way, the charge from a photoelectric interaction can be shared between two pixels, distorting the measured energy and resulting in a double count. A Compton-scattered photon deposits parts of its energy in the first interaction and continues in a new direction. If a Compton-scattered photon is detected at least once, it adds to the detection efficiency of the detector. However, the spectral information is largely lost. If the Compton-scattered photon is detected more than once, then the signal is degraded due to count multiplicity (see section 4.2).

For high-Z detector materials, there is a high probability of K-fluorescence emission, namely, a photoelectric interaction with a high-Z element (e.g. Cd and Te) results in the emission of K-shell characteristic x-rays (Shikhaliev *et al* 2009). For CZT, approximately 70% of all photoelectric absorptions result in the emission of characteristic x-rays (Krause 1979). The energies of the emitted characteristic x-rays lie between 23 and 27 keV, and the mean free path in the detector material is approximately 120 μ m. The emission and reabsorption of characteristic x-rays result in a skewing of the energy spectrum (see section 4.4), a reduction of the spatial resolution (see section 4.3), and count multiplicity, which reduces the quantum efficiency of the detector (see section 4.2).

As illustrated in this section, the detection mechanisms of PCDs are complex. In the end, what matters for the signal quality is (1) the number of photons that are registered at least once (section 4.1), (2) the count multiplicity (section 4.2), (3) the spatial distribution of the counts (section 4.3), and (4) the associated spectral response (section 4.4). To get a full picture of the imaging performance, these effects must be evaluated simultaneously (section 6.3).

3.2.2. Collection of charge carriers

When a photon interacts in the semiconductor detector material, a cloud of charge carriers (electron–hole pairs) is created. The number of charge carriers is proportional to the energy deposited in the interaction. The electrons and holes drift in opposite directions through the detector material by an applied high-voltage bias (typically 150–1000 V, depending on the material and the material thickness), and the electrons and holes are collected by the electrodes and the back side, or vice versa (Fang *et al* 2018).

The signal in the electrode does not result from the collection of the charge itself, but rather from the movement of the charge in the electric field in the detector bulk (Hamel and Paquet 1996). When the charged particles move in the electric field, an electrical current is induced on the electrode. Therefore, the particles that are collected on the back side also contribute to the signal. How much the movement of particles induces a signal on the electrode depends on the strength of the so-called weighting potential (Xu *et al* 2011). By using very small pixels, the weighting field is strong only very close to the electrode, and only particles moving there contribute significantly to the signal. This is the so-called *small-pixel effect*(Barrett and Myers 2003), and it is used to minimize the signal from slowly moving holes in CdTe/CZT.

It is possible to collect either the holes or the electrons at the electrode. For detectors with low hole mobility and a high risk of hole trapping, such as for CdTe/CZT, it is advantageous to collect the electrons. For silicon detectors, the two types of charge carriers have similar mobility and there is a low risk of trapping, and both options are therefore feasible. For CdTe/CZT, it is desirable to drift the holes as short a distance as possible due to the low mobility, and since they are collected at the back side, the back side of the sensor should preferably face the x-rays.

3.2.3. Sensor thickness

The sensor thickness plays a part in determining the average charge collection time and the charge cloud diameter, and therefore the amount of charge sharing. Thicker sensors also degrade spectral response due to

increased probability of charge-carrier recombination during the drift through the wafer. For face-on detectors, such as CdTe/CZT, the sensor thickness determines the total absorption efficiency of the detector, and typical thicknesses range from 0.9 to 3 mm (Barber *et al* 2013, Taguchi and Iwanczyk 2013). For edge-on detectors, the wafer thickness determines the pixel pitch in one direction (the pixel pitch in the other direction is determined by the width of the charge-collecting electrodes; see figure 6). A typical value of sensor thickness for edge-on Si detectors is 0.5 mm (Xu *et al* 2013a).

3.2.4. Charge sharing

Charge sharing affects the quantum efficiency of the detector (since a photon can be lost if the charge is split, so that both pulses are below the lowest energy threshold), the spectral response, and the DQE via count multiplicity.

Factors that affect the prevalence of charge sharing include the distribution of energies deposited in the detector material (higher deposited energies create more charge sharing); the size of the charge cloud when it reaches the electrode (affected by charge mobility, thickness of material, bias voltage, and location of interaction); the pixel size (smaller pixels result in more charge sharing); the number of neighboring pixels (four for face-on, and two for edge-on); and the lowest threshold (a high lowest-energy threshold reduces the rate of double counts). Furthermore, charge sharing can be reduced by using an attenuating anti-scatter grid located over the pixel boundaries (Tkaczyk *et al* 2009). Many approaches for modeling the effects of crosstalk between detector pixels and energy bins have been investigated (Taguchi *et al* 2016, Faby *et al* 2016).

3.2.5. Electronic noise

The electronic noise on an ASIC channel is affected by many factors, including the capacitance connected to the ASIC channel input, the sensor leakage current, the temperature, and the properties of the ASIC analog channel (Xu *et al* 2013b). The electronic noise in a PCD affects the energy resolution and the quantum efficiency of the detector by defining the lowest possible threshold setting without counting noise.

Silicon sensors need to be sensitive to low-energy Compton interactions in order to obtain a high detection efficiency, and it is therefore important to be able to set the lowest-energy threshold as low as possible. This is possible only if the electronic noise in the ASIC is low. In comparison, the noise requirements for an ASIC used with a CdTe/CZT sensor are less stringent, since most of the useful events deposit an energy of 25 keV or more.

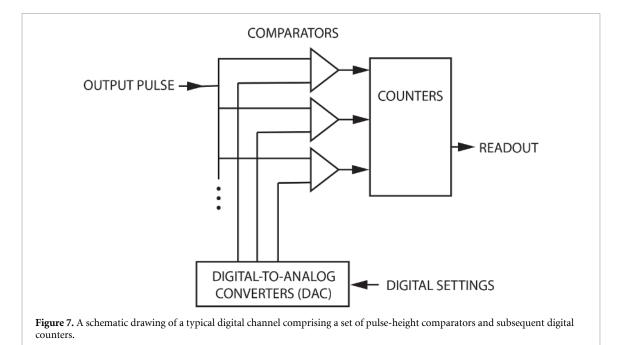
3.2.6. Depth segmentation

A sensor in the edge-on geometry can divide the pixels into depth segments/strata, effectively dividing the count rate by the number of segments (Liu *et al* 2016). This capability is important if the count-rate tolerance is an important factor in the total detector performance. In addition, the depth segmentation introduces a redundancy in the pixel channels that improves reliability.

3.2.7. Imperfections in the detector material

One drawback of cadmium telluride is its relatively high density of imperfections, which can act as traps or recombination centers for electrons and holes (Bolotnikov *et al* 2005). This is detrimental to the detector performance in two ways: First, the build-up of trapped charges in the semiconductor gives rise to an electric field that causes the charge-collection efficiency to degrade or even break down at high photon fluence rates, a phenomenon known as polarization (Siffert *et al* 1976, Bale and Szeles 2008). Second, the amount of charge that contributes to the measured signal becomes dependent on the interaction location, resulting in degraded energy resolution in the form of tailing in the energy spectrum (Xu *et al* 2011). This tailing can be reduced by making the detector pixels small relative to the wafer thickness, thereby decreasing the sensitivity of the measured signal to the hole motion, and therefore also hole trapping, through the small-pixel effect (Barrett *et al* 1995). At the same time, using too-small electrodes can result in loss of energy resolution, in addition to the spectral degradation caused by charge sharing, due to incomplete charge collection in the case when the surface of the material is not a perfect dielectric and there is a slight surface conductivity (Bolotnikov *et al* 1999).

In contrast, silicon sensors can be manufactured with a very low density of imperfections. Since this material also has a higher charge-carrier mobility compared to CdTe, the mobility-lifetime products for electrons and holes in silicon are approximately 3–4 orders of magnitude larger than those of CdTe (Fang *et al* 2018), meaning that most charge carriers live long enough to contribute to the signal. This leads to negligible charge tailing compared to CdTe/CZT, apart from the Compton interactions (Xu *et al* 2013a), and ensures that polarization is much less of a problem.



3.2.8. Other factors

Other important factors related to the detector material are production reliability, stability over time, and production cost. In the end, these factors may be important to the adoption of PCDs in the clinic.

3.3. Signal processing

3.3.1. The analog channel

The induced current on the electrode is convolved with the transfer function of the ASIC analog processing, which in general terms consists of a charge-sensitive amplifier and a shaping filter. The main purpose of the analog processing is to produce an output pulse with a pulse height that is proportional to the integral of the induced current, which in turn is proportional to the charge collected on the electrode, and to do so with a high signal-to-noise ratio.

One of the main parameters of the analog channel is the shaping time, which determines the temporal width of the ASIC filter kernel. For high count-rate applications, the shaping time should be kept as short as possible in order to avoid pulse pileup. However, the shaping time needs to be long enough to ensure that the height of the output signal is proportional to the energy. If the shaping time is too short, then the amplitude of the output signal will depend on the length of the input signal (a longer input signal resulting in a lower amplitude and vice versa), and the energy resolution is adversely impacted. A long shaping time also generally leads to a decreased relative noise level, since the input signal is low-pass filtered more heavily by the shaper. Hence there is a trade-off between pileup tolerance, noise level, and energy resolution, which has been addressed by, for example, using dual shapers (Sundberg *et al* 2018).

3.3.2. The digital channel

The ASIC channel on a multi-bin PCD has several pulse-height comparators (thresholds) that are used to identify the arrival of a new photon pulse and to estimate the energy of the photon interaction (see figure 7). Each comparator compares the amplitude of the output signal from the analog channel to a programmable reference voltage supplied by a digital-to-analog converter. The comparator returns a one or a zero depending on whether the signal exceeded the reference voltage or not. The photon counts are categorized into a set of counters (energy bins), generally one for each comparator.

The number of energy thresholds varies between different photon-counting ASICs (Taguchi and Iwanczyk 2013). In order to perform a two-basis material decomposition, it is necessary to have only two energy thresholds. However, the performance of the material decomposition depends on the position of the thresholds, and the optimal selection of the thresholds is task dependent. Having more thresholds ensures that a close-to-optimal CNR can be obtained for all imaging tasks in a scan (Shikhaliev 2008, Zheng *et al* 2020), and that close to the minimum possible noise can be achieved in material-specific images (Alvarez 2011, Faby *et al* 2015). Also, more thresholds can improve the effectiveness of corrections, such as charge-sharing or pileup correction (see section 3.3.2).

The way the digital part of the ASIC analyzes the output from the comparators is commonly referred to as the *counting mode*, and the exact implementation differs between ASICs. The counting modes are generally categorized as either *paralyzable* or *non-paralyzable*, referring to the behavior of the channel under heavy pulse pileup (section 4.5).

For a non-paralyzable detector, when a photon pulse crosses the lowest threshold, a counter is incremented, and a *dead time* is initiated during which no other photons are counted. After the dead time, the channel can count again. The dead time is selected such that the signal induced by the longest input pulses are below the lowest energy threshold at the end of the dead time. With increasing number of detected photons per second on the channel, the output from the non-paralyzable channel approaches a fixed number (the maximum number of dead times that fit within a single readout interval).

In a paralyzable detector, the dead time is extended until the signal drops below the threshold. At high flux rates, pileup can cause the signal level to exceed the threshold level for longer periods, resulting in increased dead time. Paralyzable behavior occurs, for example, if the detector is configured to count the number of times the input signal crosses each threshold, known as the threshold-crossing frequency. In presence of severe pileup, the input signal does not fall below the highest threshold, and the threshold-crossing frequency therefore drops to zero.

It is also possible to implement other counting modes, such as detection of local maxima (peak sample and hold) using the comparators, and Hsieh and Pelc (2016) have shown that more sophisticated counting modes can substantially improve the performance of the detector.

3.3.3. Charge summing and anti-coincidence logic

Charge sharing and the emission/reabsorption of characteristic x-rays has a significant negative impact on the imaging performance of the detector (Schlomka *et al* 2008, Xu *et al* 2011, Shikhaliev *et al* 2009).

The corrupted signal can be partially restored by means of interpixel communication (Koenig *et al* 2014, Ji *et al* 2018). One way of correcting the signal is to use *analog charge summing* (e.g. the Medipix3RX ASIC) (Ballabriga *et al* 2007, Koenig *et al* 2013, Nilsson *et al* 2007). A circuit on the ASIC sums the charge in overlapping clusters of, for example, 2×2 pixels prior to comparison with the energy thresholds, and the photon count is allocated to the pixel that registered the largest collected charge. The analog charge-summing mode obtains an energy resolution corresponding to the increased pixel size, while keeping the spatial resolution defined by the native pixel size. However, the detector reduces its capability to cope with high count rates by a factor equal to the number of pixels for which the charge is summed.

Digital anti-coincidence logic has been implemented in, for example, silicon strip detectors developed for mammography (Fredenberg *et al* 2010b). The scheme identifies double-counting events where the pulses in two neighboring pixels cross over the lowest energy threshold simultaneously, and it keeps the first detected pulse (which generally is larger) and disregards the second pulse. This method improves the noise properties and the spatial resolution by removing double counting, but does not improve the spectral imaging capability since no energy correction is made.

A spectral version of the digital anti-coincidence logic, referred to as *digital charge summing*, has been evaluated in simulation (Hsieh and Sjolin 2018). For the evaluated imaging cases, and for a detector with only two energy thresholds, the digital charge summing achieved roughly half the benefit (improved dose efficiency) compared to the analog charge summing. Digital charge summing has the potential to be much faster than its analog counterpart, since the digital charge summing needs only a short coincidence window to register if two events are simultaneous; otherwise, the channel operates as normal. The probability of false coincidences (i.e. two events in neighboring pixels randomly occurring within the coincidence window) is therefore low, but not negligible.

Further, it has also been suggested that, instead of correcting the charge-sharing events, the coincidences can be registered in so-called coincidence counters, read out from the ASIC and handled in post-processing (Hsieh 2020).

4. Effects impacting image quality

The effect of x-ray scatter, characteristic fluorescence reabsorption, and charge sharing between pixels is described by three processes: degradation of the spatial resolution, increase of noise, and degradation of the spectral information.

4.1. Quantum efficiency

The key figure of merit for the dose efficiency of an x-ray detector is the quantum efficiency, which describes how many of the incoming photons are detected. For the quantum efficiency, it does not matter if the photon

is detected in a photoelectric event or a Compton-scattering event, or if it is affected by K-fluorescence or charge sharing. As long as the photon is detected at least once, it contributes to the detection efficiency.

Naturally, the thickness of the detector material impacts the quantum efficiency. However, thicker detectors can affect other important aspects of the detector performance. For face-on detectors, a thicker material affects the charge collection time, and therefore the energy response, via an increase of charge trapping/recombination and charge sharing. Edge-on detectors do not suffer from the same downside to increasing the depth of the detector, apart from the increase in detector material and the resulting more stringent demands on alignment precision, since the distance the charge travels remains the same.

The position of the lowest threshold can have a large impact on quantum efficiency by determining how many low-amplitude input pulses are detected. Any process of spectral degradation that causes some of the photon pulses to become lower than the lowest threshold leads to a reduction of the DQE. For Si, this effect can be caused by charge sharing and Compton scattering. For CdTe/CZT, this effect can, in addition, be caused by K-fluorescence emission/reabsorption and the trapping/recombination of charge carriers. Photon-induced pulses dropping below the lowest energy threshold as a consequence of charge sharing can lead to regions of insensitivity at the boundaries between pixels (Tlustos *et al* 2006). The lowest energy threshold also determines the number of double counts to a large extent by inclusion or exclusion of low-energy charge sharing and K-fluorescence reabsorption. For Si, the lowest threshold is often set at around 5–15 keV in order to include many Compton interactions (Bornefalk and Danielsson 2010, Persson *et al* 2014), and for CdTe/CZT, the lowest energy threshold is often assumed to lie at 20–25 keV (Xu *et al* 2011, Shikhaliev 2009, Yu *et al* 2016a). In practice, it may not be optimal to reject electronic noise completely, but instead to operate the detector with the threshold at a low enough level where a small number of false counts are generated by the noise floor, as long as the extra noise is outweighed by the benefit of detecting more primary detection events (Rajbhandary and Pelc 2018).

The semiconductor wafer is not necessarily sensitive in the entire bulk of the detector. For CdTe/CZT, there can be a dead layer near the back side of the sensor which is insensitive to x-rays (Matsumoto *et al* 2000, Moralles *et al* 2007),⁶ and for Si, there is generally an inactive guard ring around the wafer protecting against leakage current. The dead layers can have a large impact on the performance of the detector, in particular for CdTe/CZT since the detector material is highly attenuating. For example, a 20 μ m dead layer on a CdTe/CZT detector reduces the quantum efficiency by approximately 9%, whereas a 200 μ m guard ring on a Si detector reduces the quantum efficiency by approximately 1.5% (for a 120 kVp spectrum attenuated by 10 cm of water). A dead layer on CdTe/CZT detectors can also form as a consequence of high x-ray flux (Du *et al* 2002). Furthermore, there is a dead layer close to the pixels, where an interaction gives rise to charges that only move through part of the weighting field and therefore generate a reduced signal (Boucher 2013).

4.2. Count multiplicity

As a consequence of scatter, K-fluorescence, or charge sharing, a photon can be counted more than once. Count multiplicity gives extra weight to a fraction of the photons, resulting in a reduction of the zero-frequency DQE and noise correlations between neighboring pixels (Michel *et al* 2006).

As a consequence of count multiplicity, the zero-frequency DQE is reduced by a factor

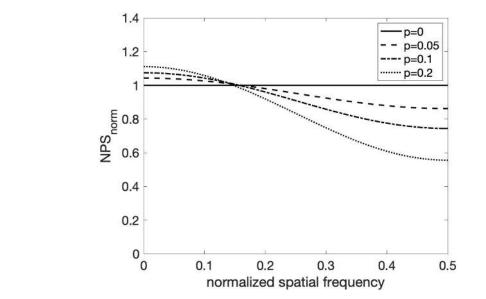
$$I_{\text{multiplicity}} = \frac{\left(\sum_{n=1}^{\infty} nr_n\right)^2}{\sum_{n=1}^{\infty} n^2 r_n},\tag{3}$$

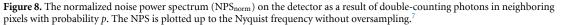
where r_n is the fraction of detected photons that are counted *n* times. Comparing equation (3) to the Swank factor of an EID (1), we can see that the expressions are identical apart from that the signal amplitude (ε) in equation (1) is exchanged for the multiplicity (*n*) in equation (3).

Counting photons in more than one pixel results in pixel-to-pixel correlations and a non-white noise power spectrum (NPS) (Xu *et al* 2014). The detector NPS can be obtained by taking the discrete Fourier transform (DFT) of the auto-covariance of the counts measured by the detector array (Cunningham 2000). In the special case for which a fraction p of the λ photons that are counted in a pixel are counted also in the neighboring pixels (p/2 to the left and p/2 to the right), the auto-covariance is given by

$$\operatorname{Cov}[N_0, N_i] = \begin{cases} \lambda(1+p), & \text{for } i = 0\\ \lambda p, & \text{for } |i| = 1\\ 0, & \text{for } |i| > 1, \end{cases}$$
(4)

⁶ The x-rays generally enter through the back side.





where N_i is the number of counts in the *i*th pixel. To illustrate the effect of the pixel-to-pixel correlations on the noise, consider the NPS normalized by the square of the mean number of counts in each pixel:

$$NPS_{norm}(u) = \frac{NPS(u)}{\lambda^2 (1+p)^2},$$
(5)

where NPS(u) = DFT[Cov[N_0, N_i]], and u is spatial frequency. Examples of NPS_{norm} for different values of p are shown in figure 8 (the curves have been normalized by $1/\lambda$ to obtain unity NPS_{norm} for p = 0, and the spatial frequency axis is normalized by the sampling frequency). The increase of NPS_{norm} at the zero frequency is inversely proportional to the decrease of the zero-frequency DQE given by (3):

NPS_{norm}
$$(u = 0) = 1/I_{\text{multiplicity}} = \frac{(1-p)+4n}{((1-p)+2n)^2}.$$
 (6)

An important observation is that the pixel-to-pixel correlations lead to a higher number of events detected in each pixel; each pixel registers $\lambda(1+p)$ photons, as opposed to only λ . This can also be seen as the integral of the normalized NPS reduces as p increases. When the pixels are considered together, however, it becomes clear that the quantum efficiency has indeed been reduced, as indicated the increase of the zero-frequency NPS. This property of correlated noise must be considered when, for example, evaluating the detector's CNR performance.

4.3. Spatial resolution

The spatial resolution of the PCD is determined, first and foremost, by the center-to-center distance between the electrodes on the semiconductor surface. For edge-on detectors (e.g. silicon), the thickness of the semiconductor wafer determines the spatial resolution in one of the dimensions.

Charge sharing, K-fluorescence reabsorption, and Compton scattering degrade the spatial resolution somewhat by causing events to be detected in positions other than where they first interacted with the detector. This effect leads to a blur of the point-spread function (PSF), and therefore a degradation of the modulation transfer function (MTF). These effects can partially be corrected using anti-coincidence logic or similar methods.

Charge sharing, unless corrected, also limits the smallest feasible pixel size indirectly due to the increase in count multiplicity and the degradation of the energy response of the detector. If pixels that are too small are used, the detector will lose much of its spectral capabilities and some of its dose efficiency (Rajbhandary *et al* 2018).

The spatial resolution of the detector is not constant over the range of detected energies. High-energy interactions are predominantly registered close to the center of the pixel, since charge sharing occurs at the

⁷With oversampling (e.g. quarter-pixel offset of the detector), the NPS is mirrored around the native Nyquist frequency.

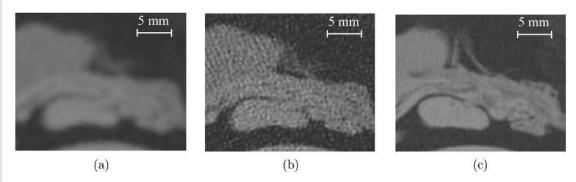


Figure 9. An example of the benefits of a native high-resolution image acquisition, here when imaging tissue of an excised human heart (same as that shown in figure 2). Images (a) and (b) were acquired with a conventional CT system with 0.7 mm focal spot, and reconstructed with a soft kernel and a bone kernel, respectively. Image (c) was acquired by a prototype silicon-based PCD with roughly half the native pixel size and a 0.4 mm focal spot, and reconstructed with a high-resolution filter kernel. The images were acquired at comparable dose and slice thickness (0.625 mm). The native high-resolution acquisition of the PCD has less noise in a high-resolution image, which improves the visibility of the low-contrast high-resolution structures within the tissue.

pixel boundaries, giving them a more narrow PSF, whereas low-energy interactions can be registered over the full extent of the pixel (Stierstorfer *et al* 2019). This implies that the detector MTF has an energy dependence, and the spatial resolution can be enhanced by giving more weight to the high-energy bins at the cost of increased noise in the reconstructed image. In the case that monoenergetic images are formed from material basis maps, the spatial resolution will therefore depend on the choice of monoenergetic energy. In order to have a full characterization of the spatial resolution of the detector, the full energy range should be evaluated.

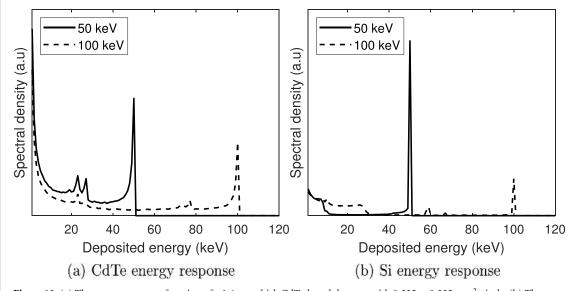
4.3.1. The spatial resolution and noise trade-off

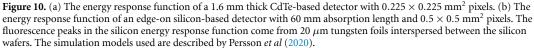
A common misconception when it comes to high-resolution detectors is that having small pixels comes with a noise penalty. Indeed, the noise in each detector pixel will increase if the pixels are smaller, since fewer photons are detected in each pixel. However, high-resolution image acquisition improves the resolution-noise trade-off in the reconstructed image. That is, for the same image noise, the high-resolution system will have better spatial resolution at equal dose, and conversely, for the same spatial resolution, the high-resolution system will have lower noise. The literature predicts noise variance reductions between 14% and 83% from a $2\times$ increase in native resolution (Baek *et al* 2013, Kachelrieß and Kalender 2005). The highest noise reductions are observed when reconstructing high-resolution images. For low-resolution images (e.g. for soft-tissue imaging) the benefit is reduced. In addition, the high-resolution system can use reconstruction kernels with higher cut-off frequency without introducing aliasing artifacts than a system with lower native resolution. An example of the benefit of a native high-resolution detector is shown in figure 9, in which the two images to the left are acquired using a conventional CT system reconstructed with a soft kernel (a) and a bone kernel (b), and the right-most image (c) was acquired using a prototype PCD, with roughly half the pixel size, and reconstructed using a high-resolution kernel. The native high-resolution acquisition of the PCD improves the spatial resolution and noise trade-off, and allows reconstructing high-resolution images with less noise and aliasing artifacts.

4.4. Energy response

In contrast to properties like detection efficiency and spatial resolution, which are important for all detectors, the ability to measure the energy distribution of the incident spectrum is unique to energy-resolving detectors. The precision with which each deposited energy is measured is partly determined by the number of energy thresholds (see section 3.3.2). At the same time, the number of thresholds is not the only factor that affects the spectral imaging performance.

The energy-resolving capability of PCDs is also determined by non-ideal effects in the detector materials and the readout electronics that give a non-ideal energy response, i.e. cause the registered signal to deviate from the expected amplitude. In a hypothetical ideal PCD, each photon gives rise to an electrical signal with amplitude proportional to the incident photon energy. In CdTe-based and CZT-based detectors (figure 10(a)), K-fluorescence gives rise to two secondary clusters of peaks in the detected spectrum distinct from the photopeak: fluorescence peaks at the fluorescence energy (23 and 26 keV for Cd, and 27 and 31 keV for Te), (Thompson *et al* 2009) and K-escape peaks at the original energy minus the fluorescence energy (Shikhaliev *et al* 2009, Xu *et al* 2011). The response of a silicon-based detector (figure 10(b)), on the other hand, exhibits a large fraction of Compton interactions at low energies and reproduces the photoelectric part of the spectrum with diminished magnitude, but it is otherwise not distorted in the way characteristic of the





response from a CdTe-/CZT-based detector (Bornefalk and Danielsson 2010). In both CdTe/CZT and Si detectors, charge sharing causes spectral tailing, which is a background of charge-shared counts of relatively constant magnitude (figure 10) (Xu *et al* 2011, Liu *et al* 2014). Spectral tailing can also be caused by incomplete charge collection (section 3.2.7).

In addition, electronic noise in the readout electronics causes further spectral degradation by adding a random error to the registered amplitude of each pulse. In practice, this characteristic has the effect of convolving the detected spectrum with a Gaussian kernel. Typical photopeak widths reported for detectors designed for high-flux x-ray imaging are 3.5–5.4 keV full width at half maximum (FWHM) for Si (Xu *et al* 2013a) and 5–10 keV FWHM for CdTe/CZT (Iwanczyk *et al* 2009, Brambilla *et al* 2013, Barber *et al* 2015). Spectral response models incorporating physical effects and electronic properties have been published for silicon (Liu *et al* 2015a, 2015b) and CdTe/CZT (Schlomka *et al* 2008, Cammin *et al* 2014, 2016, Taguchi *et al* 2016, 2018b).

The effect of the spectral distortion is that it degrades the performance of the detector for tasks that have a strong energy-dependent component. While the performance for density-imaging tasks, such as detection of non-enhancing tumors, is only weakly affected by non-ideal energy resolution (Schmidt 2010), imaging tasks involving calcium and iodine are more strongly affected (Rajbhandary *et al* 2018). Good energy-resolving capability is especially important for material quantification tasks, since these cannot be performed at all with a non-energy-resolving CT. The impact of the detector energy response on material quantification performance has been investigated by several authors (Roessl and Herrmann 2009, Wang and Pelc 2012, Rajbhandary *et al* 2018).

Apart from the detector energy response, the incident spectrum shape is also an important factor affecting spectral imaging performance, and multiple studies have investigated how performance can be optimized by varying the x-ray tube voltage (kVp) and prepatient filtering, such as with a K-edge filter (Wang and Pelc 2012, Chen *et al* 2015, Atak and Shikhaliev 2015, 2016). To understand how the incident spectrum shape interacts with the detector response function, note that misregistration of photon energies does not degrade spectral imaging performance if it is possible to correct for the spectral distortion and thereby recover the original spectrum. Instead, degradation of imaging performance occurs only when there is *overlap* between detected energies originating from different incident energies, so that it is impossible to recover the incident spectrum without ambiguity (Persson *et al* 2018b). Figure 11 illustrates how K-fluorescence can give rise to spectral overlap, depending on the incident energy spectrum distribution. K-fluorescence and charge sharing give rise to spectral tailing that overlaps with incident energies present in the true spectrum, which exacerbates the harmfulness of these effects on the spectral imaging performance. On the other hand, Compton scatter events in a silicon-based detector do not overlap much with the spectrum of photoelectric events. Instead, their detrimental effect on energy-resolving imaging stems from the fact that one energy of a Compton event can correspond to a range of incident energies. The Compton

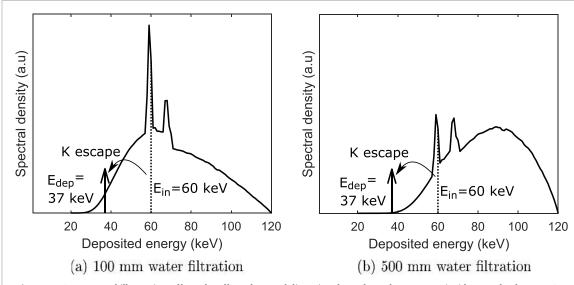


Figure 11. A conceptual illustration of how the effect of spectral distortion depends on the spectrum incident on the detector. A photon with the original incident energy $E_{in} = 60$ keV loses 23 keV through the escape of a K-fluorescence event and deposits $E_{dep} = 37$ keV. (a) For a spectrum filtered through 100 mm of water, the deposited energy overlaps with the incident spectrum, and the original energy cannot be recovered after detection. (b) For a spectrum filtered through 500 mm of water, the deposited energy does not overlap with the spectrum, and the original energy can be recovered by adding 23 keV to the measured energy. Note that this is a simplified example, and that a more realistic model should also include spectral overlap due to charge sharing.

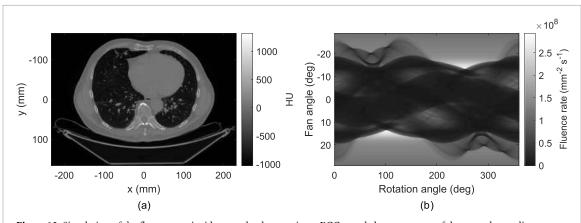
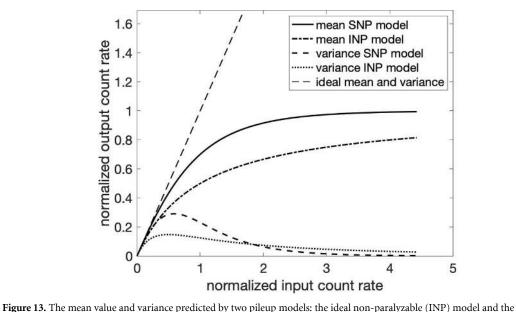


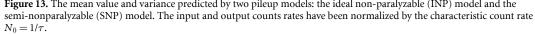
Figure 12. Simulation of the fluence rate incident on the detector in an ECG-gated chest scan, one of the most demanding imaging cases in terms of detector count-rate performance. A tube voltage of 120 kVp and a fixed tube current of 550 mA was assumed, representing the high-current phase of the ECG-gated protocol. (a) Chest CT image used for the simulation, with attenuation values in Hounsfield units (HU). (b) Incident fluence rate on the detector, after transmission through the patient. The highest fluence rates occur near the skin line and behind the lungs. The simulation model is described in Persson *et al* (2016).

events are therefore less useful than the photoelectric events for determining the incident energy distribution, but they do not confound the spectral information provided by the photoelectric events.

4.5. Pulse pileup

Photons that interact within a short time period 'pile' onto each other, resulting in count loss, distortion of the Poisson counting process, and degradation of the energy information. In order for a PCD to be feasible for clinical CT, the detector has to be able to cope with very high fluence rates that can exceed $3.5 \times 10^8 \text{ mm}^{-2} \text{ s}^{-1}$ at the detector (Persson *et al* 2016). Figure 12 shows how the photon fluence rate is distributed on the detector for an imaging protocol that is demanding in terms of high-count-rate capability. The effects of pulse pileup can be mitigated by decreasing the pixel size, but the benefit is limited since small pixels are more prone to charge sharing, which causes double counting and spectral degradation (Hsieh *et al* 2018). The mathematical description of pulse processing by an electronic channel is known as a dead-time problem, and these problems have been studied extensively for several decades and have applications in many scientific fields (Müller 1973, Carloni *et al* 1970, Wielopolski and Gardner 1976). Still, analysis of the effects of pulse pileup in spectral PCDs, and how to account or correct for this effect, is an active topic of investigation today.





4.5.1. Statistical degradation

In the presence of pileup, the output count process is no longer Poisson distributed. A simple, but commonly used, model of the photon-counting process is the so-called ideal non-paralyzable (INP) model (Tenney 1984). The INP model predicts the statistics and spectral distortion under two simplifying conditions: (1) there is a fixed dead time after each photon detection during which all additional events are lost, and (2) the registered energy in the case of pileup is the sum of the energy of the associated photon interactions. The mean and variance of the output count process Y for the INP model is given by

$$\mu_{\rm Y} = \frac{\lambda}{1+\lambda\tau}, \quad \sigma_{\rm Y}^2 = \frac{\lambda\tau}{(1+\lambda\tau)^3},\tag{7}$$

where λ is the input count rate and τ is the dead time. The mean value and the variance of the output count rate for a non-paralyzable detector with non-zero pulse length, the so-called semi-nonparalyzable (SNP) model (Xu *et al* 2011), have been derived analytically (Grönberg *et al* 2018) and are given by

$$u_{\rm Y} = \frac{\lambda}{e^{-\lambda\tau_{\rm s}} + \lambda\tau}, \quad \sigma_{\rm Y}^2 = \frac{\lambda e^{-\lambda\tau_{\rm s}}(2 - e^{-\lambda\tau_{\rm s}})}{(e^{-\lambda\tau_{\rm s}} + \lambda\tau)^3},\tag{8}$$

where λ is the input count rate, τ is the dead time, and τ_s is the average time at the end of the dead time during which, if a photon interacts, a new dead time is initiated directly after the first dead time. The length of τ_s depends on the length and shape of the photon-induced pulses and is often a large fraction of τ (0.77%–0.86% in Liu *et al* (2016)). Examples of the mean value and the variance predicted by the INP and the SNP model with $\tau_s = 0.8$ as a function of count rate is plotted in figure 13. The input and output count rates have been normalized by the characteristic count rate $N_0 = 1/\tau$. In the ideal case, without pileup and with Poisson statistics, the output count rate is equal to the input count rate, and the variance is equal to the mean value.

Pileup introduces a non-linear response to variations in the flux rate, and the statistics start to deviate from Poisson behavior. The effect of pileup on image quality is therefore not straightforward to assess, and looking at, for example, count-rate linearity tells only half the story. One way to incorporate both the effect of non-Poisson statistics and the non-linearity of the output count rate is to consider the CNR for the task of differentiating between two similar projection measurements with a small difference δ in photon flux. A flux-dependent doseefficiency can then be defined as the ratio of the CNR² with and without pileup:

$$DE(\lambda) = \frac{CNR_{\text{with pileup}}^2}{CNR_{\text{without pileup}}^2} = \frac{(\mu_Y(\lambda+\delta) - \mu_Y(\lambda))^2 / \sigma_Y^2(\lambda)}{(\lambda+\delta-\lambda)^2/\lambda} = \frac{(\mu_Y(\lambda+\delta) - \mu_Y(\lambda))^2}{\delta^2 \sigma_Y^2(\lambda)/\lambda}.$$
(9)

An example of the dose efficiency as a function of count rate is plotted in figure 14(a), and the CNR² versus the thickness of attenuating material (here water) is shown for different maximum count rate (N_0 , $2N_0$, and

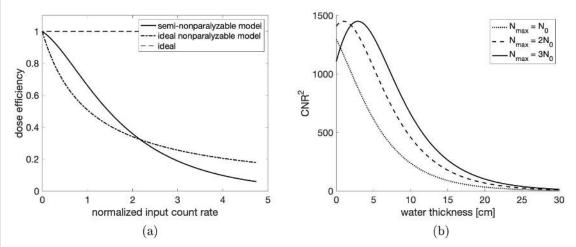


Figure 14. (a) The dose efficiency for a low-contrast imaging task as a function of the input count rate (normalized with N_0), comparing the ideal non-paralyzable model and the semi-nonparalyzable. (b) The CNR² estimated by the semi-nonparalyzable model versus the thickness of water attenuation for three different input count rates of the unattenuated beam (N_{max}). Even though the input count rate is high when there is little water in the beam path, and the DQE is reduced by pileup, the CNR² is still relatively high since pileup mainly affects high-flux projections, for which the SNR is high.

 $3N_0$) in figure 14(b). As an example of a maximum input count rate, consider a case for which the maximum x-ray flux on the detector is 3.5×10^8 mm⁻² s⁻¹ (Persson *et al* 2016), and the detector has 0.25×0.25 mm² pixels and a 100 ns dead time; then the maximum input count rate is $2.2N_0$

 $(3.5 \times 10^8 \text{ mm}^{-2} \text{s}^{-1} \times 0.25^2 \text{ mm}^2 \times 100 \times 10^{-9} \text{ s} = 2.2)$. Even though the dose efficiency is reduced for the higher count rates, the CNR² is still high since pileup mainly affects projections that have high statistics.

The effect of the statistical degradation from pileup in the final CT image is limited by several factors. First, the central rays, where the x-ray flux is generally low due to patient attenuation, contribute much more to the image than the peripheral rays due to the nature of tomographic image reconstruction. Second, the x-ray flux at the periphery of the patient is lowered by the bowtie filter of the x-ray tube. Also, pileup increases the noise for high-flux projections, for which the SNR is high (see figure 14(b). Combined, these effects help mitigate the impact of statistical degradation from pulse pileup.

4.5.2. Spectral distortion

Pulse pileup introduces errors in the registered photon energies, resulting in a distortion of the energy spectrum. The consequence is a reduction in the spectral separation between energy bins, and thus a reduction of the basis-material-decomposition performance.

The effect of spectral degradation from pileup occurs earlier than the loss of dose efficiency (equation (9)), which can, for example, be seen by observing that the response of the individual energy bins deviates from linear behavior at much lower count rates than the total number of counts (Fredenberg *et al* 2010b). It has also been shown that pulse pileup affects the imaging performance more for spectral detection tasks than for density imaging tasks (Wang *et al* 2011a).

The spectral distortion depends on, for example, the photon-counting mode, the dead time (if applicable), the pulse shape in the ASIC, and the input spectrum. Many approaches for modeling the spectral effects of pulse pileup can be found in the literature (Taguchi *et al* 2010, 2011, Wang *et al* 2011a, Cammin *et al* 2014, Roessl *et al* 2016). Other approaches for handling the effects of pileup include empirically estimating parameters to model the non-ideal effects (Zimmerman and Schmidt 2015, Alvarez 2011). Also, methods for correcting the effects of pileup after read-out (e.g. by using neural networks) have been investigated (Touch *et al* 2016, Feng *et al* 2018).

Some authors have also suggested using the so-called *pileup trigger* method to help mitigating spectral pileup (Kappler *et al* 2011). This method uses a threshold that is placed above the highest energy of the input spectrum to identify if a photon has been subject to pileup.

5. Image formation

5.1. Energy bin images

The most straightforward ways of generating reconstructed images from spectral CT data are either summing the counts in all energy bins before reconstruction, thereby generating a conventional-looking CT image, or reconstructing a CT image from the measured data in each energy bin (Yu *et al* 2016c). In

comparison with material decomposition, reconstructing bin images has the advantage of being easy to implement since it does not require accurate models of the source and detector. On the other hand, this method has the drawback of not being able to correct for the effect of beam hardening. The bin images can be reconstructed with filtered backprojection or with an iterative algorithm similar to those used for single-energy CT (Salehjahromi *et al* 2018). By summing the images with carefully selected weight factors, the optimal CNR for a specific imaging task can then be attained (Schmidt 2009).

Another option is to display the images individually, which has the benefit of showing the attenuation in a narrow energy range, but leads to high noise since only a subset of the photons are used to form the image. This trade-off between precise spectral information and noise can be mitigated by applying postprocessing algorithms that act jointly on the different bin images and exploit similarities between them to reduce noise. An example is the HYPR-LR algorithm, in which low spatial frequencies from the individual bin images is combined with high spatial frequencies from a full-spectrum image, thereby generating images combining spectral information with low noise (Leng *et al* 2011). This type of bin image noise reduction can also be incorporated in an iterative reconstruction algorithm, such as the spectral prior image constrained compressed sensing (SPICCS) algorithm, which uses total variation regularization to penalize structural differences between the bin images and a full-spectrum image (Yu *et al* 2016b).

Other, more sophisticated regularization methods that exploit redundancy between the bin images have also been studied, including total nuclear variation (Rigie and La Rivière 2015), total generalized variation (Niu *et al* 2018b), and dictionary learning (Zhang *et al* 2017). Denoising based on non-local means filtering where the broad-spectrum image is used to weight pixels according to similarity was studied by Li *et al* (2017) and Zhang *et al* (2016 combined non-local-means-like regularization in the spectral dimension with SPICCS. Block-matching, where patches with similar structure in different parts of the image are identified in the broad-spectrum image and used to denoise each other (Harrison *et al* 2017) have also been demonstrated. The observation that most of the image pixels tend to be spanned by a small number of basis materials, whereas k-edge elements are often confined to localized regions, can be exploited for noise reduction as shown by Gao *et al* (2011). There are also algorithms exploiting the low dimensionality of spectral CT images in combination with SPICCS (Zeng *et al* 2020) or patch-based denoising (Niu *et al* 2018a). Finally, a new and promising research direction is the application of convolutional neural networks for bin image denoising (Clark and Badea 2020).

5.2. Material decomposition

A standalone material-decomposition step can be performed before the reconstruction (in projection space) or after the reconstruction (in image space). Another possibility is to perform reconstruction and material decomposition simultaneously, as discussed in section 5.3. Image-space decomposition is straightforward to implement, since it can take the form of a simple linear transformation of the energy-selective reconstructed images (Persson *et al* 2014, Symons *et al* 2017a). However, this does not compensate for the non-linearity of the x-ray attenuation and therefore does not remove beam-hardening artifacts completely. Iterative image-space decomposition has been proposed as a way of solving this problem (Heismann and Balda 2009).

A more straightforward way of eliminating beam hardening, thereby obtaining a quantitatively reliable image, is by using projection-space material decomposition, where the counts in the different energy bins are mapped to line integrals of basis coefficients before reconstruction. This method is well suited for photon-counting CT with its perfect geometric alignment between energy bins, but it can be more difficult to implement for systems exhibiting mismatch between the high- and low-energy projection rays, such as dual-source CT. In statistical terminology, the mapping from counts to line integrals is called an *estimator*. If the number of energy bins is equal to the number of basis materials, this estimator can be a simple mapping from counts to estimated line integrals, given by a polynomial fit, for example (Alvarez and Macovski 1976). If there are more energy bins than basis functions, the choice of estimator is more complicated.

There are a number of statistical properties that are desirable for such an estimator: ideally, it should be unbiased, meaning that the output should be equal to the true projected basis coefficient on average; the output should preferably have low variance; and it should be robust in the sense that it is able to perform well under a variety of conditions without generating image artifacts. For example, streak artifacts may result if the estimated basis sinograms contain anomalous measurements.

The maximum-likelihood estimator is commonly used in photon-counting CT research (Roessl and Proksa 2007, Schlomka *et al* 2008, Si-Mohamed *et al* 2018). This estimator builds on the fact that the registered counts m_i in each energy bin *i* is Poisson distributed and finds the set of projected basis coefficients A such that the x-ray beam is most likely to have passed through, given the measured counts. This estimator has the favorable property of being asymptotically unbiased and efficient for large measured count numbers, meaning that the variance approaches its theoretical minimum given by the Cramér–Rao lower bound (CRLB) (see section 6.2) (Kay 1993).

Mathematically, the maximum-likelihood estimate is calculated by minimizing the negative log-likelihood function

$$l(\mathbf{A}) = \sum_{i=1}^{N_b} \lambda_i(\mathbf{A}) - m_i \ln \lambda_i(\mathbf{A})$$
(10)

with an iterative algorithm. This equation builds on a model for the expected number of counts $\lambda_i(\mathbf{A})$ in each of the N_b energy bins, as a function of the projected basis coefficients \mathbf{A} . An example of such a model is

$$\lambda_i(\mathbf{A}) = \int S(E) w_i(E) \mathrm{e}^{-\sum_{j=1}^{N_m} A_j f_j(E)} dE$$
(11)

with N_m denoting the number of basis materials, S(E) denoting the spectral density as a function of energy E, and $w_i(E)$ denoting a weight function describing the spectral sensitivity of the energy bin to different incident photon energies. Ehn *et al* (2016) proposed a modification of this forward model that can be fitted well to a limited number of calibration measurements.

One drawback of the maximum-likelihood estimator is that it is calculated with an iterative algorithm, which is time-consuming. Other estimators have therefore been proposed with the aim of increasing speed. Alvarez proposed using a linearized, approximate forward model to obtain a linear estimator, which is then corrected using a look-up table to reduce bias (Alvarez 2011, 2016). As an extension of this method, different linearized forward models can be applied in different regions of the space of possible *A* values (Rajbhandary *et al* 2017).

Another estimator was proposed by Lee *et al* (2017a, 2017b). This estimator is a three-step method which builds on approximating the set of possible transmittance functions that can be expressed as $X(E) = e^{-\sum_{j=1}^{N_m} A_j f_j(E)}$ as a low-dimensional space. This allows the decomposition problem to be formulated as follows: (1) least-squares fitting of the transmittance X(E) that best agrees with the registered counts; (2) least-squares fitting of a combinations of basis functions $\sum_j A_j f_j(E)$ to log X(E); and (3) post-correcting for bias. In recent years, neural network estimators have also attracted attention. These are based on networks of simple processing units, called neurons, that are trained to map measured bin counts directly to the corresponding basis coefficient line integrals (Zimmerman and Schmidt 2015).

Since basis images tend to be noisier than conventional CT images, significant attention has been paid to methods of reducing basis image noise. In analogy with the SPICCS algorithm for bin image denoising, the prior-knowledge-aware iterative denoising method uses an image reconstructed from the sum of all energy bins as a prior image and penalizes structural differences between this image and basis images, after rescaling (Tao *et al* 2018). More advanced basis decomposition methods, such as block-matching methods (Wu *et al* 2019), dictionary learning (Wu *et al* 2020) or neural-network-based methods (Chen and Li 2019), are able to reduce noise by aggregating information from multiple pixels when forming the basis images.

Another way of reducing basis image noise is by enforcing a constraint, such as volume or mass conservation, in the basis decomposition (Liu *et al* 2009, Mendonça *et al* 2010, Ronaldson *et al* 2012, Malusek *et al* 2013, Lee *et al* 2014, Curtis and Roeder 2017, Ren *et al* 2019). Such a constraint provides an additional equation in addition to the x-ray measurements, allowing three-material decomposition with two energy bins or four-basis decomposition with three energy bins. Even if a sufficient number of energy bins are available, it may be desirable to include constraints in order to reduce noise in the decomposed basis images. A drawback of this approach is that the constraints build on prior assumptions about the material compositions that can occur in the body, and if these assumptions are in disagreement with reality, including the constraints will result in systematic errors in the resulting basis images (Bornefalk and Persson 2014, Yveborg *et al* 2015a). Whether the benefits of such constraints outweigh the drawbacks is therefore dependent on the specific imaging task.

5.3. Image reconstruction

The process of reconstructing photon-counting images depends on which material-decomposition algorithm is used. An ordinary single-channel reconstruction algorithm such as filtered back-projection can be used to reconstruct one image for each energy bin, allowing image-space material decomposition to be performed subsequently. Projection-space material decomposition, on the other hand, can be implemented either as a one-step or two-step inversion, depending on whether the mapping from measured counts to basis images takes place in a single processing step or whether basis sinograms are generated as an intermediate step.

In two-step inversion, projection-space material decomposition is first used to generate a set of basis sinograms, one for each basis material (Schlomka *et al* 2008). From these basis sinograms, a set of reconstructed basis images is then generated with a reconstruction algorithm such as filtered back-projection

or an iterative method. In the case of an iterative method, there is the possibility of choosing whether the image reconstruction should treat the different bases independently (Schirra *et al* 2013) or as a joint optimization problem (Sawatzky *et al* 2014). The noise in the basis images resulting from basis decomposition is anti-correlated between basis materials, since the basis decomposition algorithm is able to measure the total attenuation more accurately than it can measure the exact amount of each basis material in the beam path. By modifying the data weighting to take this anti-correlated noise structure into account, the reconstructed image noise can be reduced. At the same time, the coupling between basis materials introduced in this way increases the complexity of the reconstruction problem and can also lead to artifacts caused by crosstalk between the basis images (Persson and Grönberg 2017).

In one-step inversion, on the other hand, the mapping from registered counts to basis images takes place in one single step, eliminating the ray-wise decomposition step. This step typically takes the form of an iterative optimization, which searches for the set of basis images that best corresponds to the measured counts in each projection line and energy bin. In this case, the reconstruction necessarily treats the basis images as a joint optimization problem. The development of algorithms that optimize image quality while satisfying computational requirements is an active research topic. Several one-step algorithms have been proposed, based on primal-dual optimization (Barber *et al* 2016) or separable quadratic surrogates (Long and Fessler 2014), which can be accelerated with momentum terms (Mechlem *et al* 2018). A comparison of the computational cost of five such algorithms can be found in Mory *et al* (2018).

5.4. Image display

The result of the image reconstruction algorithm is a set of reconstructed images, one for each basis material. These can either be displayed as is (e.g. as maps of contrast agent concentration) or combined together to form a final image. Basis material decomposition is often claimed to increase image noise, and this is true in the sense that the individual basis images (e.g. a K-edge image) exhibit higher noise levels than a conventional CT image formed from the summed counts in all energy bins. On the other hand, as pointed out in section 2.2, the basis images can be combined in a weighted sum and yield the same optimal CNR as a weighted sum of the bin images (Rajbhandary and Pelc 2016). Another option is to combine the basis images with a conventional gray-scale image, which can always be reconstructed from the raw data in addition to the basis images, to form a color-overlay image combining high resolution and/or low noise with a material distribution map (Pourmorteza *et al* 2016, Symons *et al* 2017b). Thus, material decomposition does not cause any of the available information to be lost, but rather enables additional ways of presenting the image data.

One or more basis images can be used to form a virtual non-contrast image (Faby *et al* 2015). Another display technique currently used in dual-energy CT is to form virtual non-calcium images, which can suppress bone and thereby help in visualizing bone marrow (Kellock *et al* 2017). The different basis images can also be combined to form virtual monoenergetic images (Leng *et al* 2017, Symons *et al* 2018b), as is common in dual-energy scanners today. By varying the display energy, the CNR can be optimized for a given imaging task. More advanced ways of forming virtual monoenergetic images include prior-knowledge-aware iterative denoising (Tao *et al* 2019b) and neural-network-based methods (Feng *et al* 2019). Approximate material maps and virtual monoenergetic images can also be generated through image-space material decomposition from reconstructed bin images (Zhou *et al* 2019), although this approach gives less accurate results since it does not eliminate beam hardening.

Another possibility for image display is to form a synthetic Hounsfield image, which is an image with CT numbers equal to those of an energy-integrating CT image for a specified set of acquisition parameters, but with beam-hardening artifacts eliminated (Bornefalk 2012a). This may facilitate the transition from energy-integrating to photon-counting CT by providing image measurements that can be compared to existing reference values.

6. Detector performance metrics

As described in the preceding sections, the physical effects affecting detector performance can lead to trade-offs between different desirable detector properties. For example, a smaller detector pixel size gives higher spatial resolution and less sensitivity to pileup but also leads to more crosstalk between pixels, causing double counting and degraded energy resolution. When designing a PCD, it is therefore important to devise accurate metrics of detector performance in order to evaluate the effects of design decisions on image quality. Full-scale simulation of a photon-counting CT acquisition may in many cases be too time-consuming, especially if the objective is to characterize noise properties, which requires simulating many noise realizations. For this reason, a number of mathematical metrics of detector performance have been developed.

For energy-resolving PCDs, the question of how to measure detector performance is further complicated by the need to take energy resolution into account. Since energy information can be used in different ways, it also becomes useful to distinguish between different types of tasks in order to describe the detector performance. In a *detection* task, the objective is to detect if a certain feature is present or absent in the image. This feature can be a small feature, such as a bone fragment or a small blood vessel, or a large feature that is hard to distinguish from the surrounding anatomy, such as a tumor. In a material quantification or material separation task, on the other hand, the objective is to measure the composition of the imaged object. Within the framework of basis material decomposition, this is done by measuring the amount of each basis material present in every voxel in the imaged volume. This method allows making quantitative tissue characterizations, which can be important for example in bone densitometry or calcium scoring. The method also facilitates generation of material-specific maps (e.g. iodine maps) which allows distinguishing iodinated contrast agents from non-enhancing anatomical structures, or virtual non-contrast images where the contrast agent is removed and only the tissue attenuation remains. Quantification of iodine concentration is important, for example, in follow-up of cancer treatments. Separation between iodine and calcium is crucial to assess the degree of stenosis in calcified plaques, determining whether catheterized invasive diagnostics or stenting is required (Alessio and MacDonald 2013, Boussel et al 2014). It is also important in CT perfusion imaging in general and in assessment of stroke when selecting patients for thrombectomy to remove the obstruction to the blood supply.

The most important difference between detection and quantification tasks in terms of their requirements on detector performance is that quantification tasks are generally more dependent on energy resolution than detection tasks. This characteristic can be seen by considering a purely counting PCD with no energy-resolving capabilities. Such a detector can perform relatively well for most detection tasks just by detecting the difference in the number of transmitted photons caused by the presence of a feature. A non-energy-resolving detector, on the other hand, is not able to distinguish between different materials, but only to measure their total attenuation.

6.1. Contrast-to-noise ratio

A common way to compare the performance of two imaging systems is to evaluate the CNR, or perhaps more commonly the CNR², for a given imaging task. However, when comparing detector systems with different energy response, such as an EID and a PCD, the relative CNR² is highly task dependent. The relative performance depends not only on the choice of materials in the two projections for which the CNR is computed, but also on the density, or thickness, of those materials. As an example, consider figure 15, where we have compared the projection domain CNR² of a Si PCD, a Si spectral PCD with optimal energy weighting, and an EID. The two projections that were used to compute the CNR² comprised 20 cm of soft tissue and 19 cm of soft tissue plus 1 cm of an iodine/water solution with varying concentration up to 10 mg ml^{-1} . The CNR² curves reach a minimum at slightly different iodine concentrations, i.e. there is an offset between the curves. Both the PCD and the EID reach zero CNR² for some concentrations of iodine, whereas the spectral PCD never reaches zero due to the optimal energy weighting (Yveborg et al 2013). The relative performance of the photon-counting systems compared to the energy-integrating system has a strong dependence on the iodine concentration (figure 15(b)). Comparing imaging systems with a different energy response through a single relative CNR² measurement is therefore likely to be misleading. In fact, altering the imaging task can change which of the detector systems obtains the highest CNR. This also applies when comparing different photon-counting systems (e.g. silicon-based and CdTe/CZT-based detectors) since they too have different energy response. A possible solution to this problem is to make sure that the selected imaging tasks are clinically relevant and evaluate the performance for a range of imaging tasks in order to get a picture of the overall performance of the system.

6.2. Cramér-Rao lower bound

Whereas CNR indicates how much a certain feature stands out from the background relative to the noise level, another measure is needed for assessing the performance for material-specific imaging. This role is filled by the CRLB (Kay 1993). The CRLB gives a lower bound on the variance for any unbiased estimator, or a lower bound on the covariance matrix when estimating a multivariate parameter. In the context of photon-counting CT, the CRLB gives a lower bound on the covariance matrix of estimated projected basis coefficients. Roessl and Herrmann (2009) derived the CRLB for both photon-counting and EIDs and used this formula to study how the noise in basis images depends on detector design parameters. The noise level in the image domain can be obtained from the projection-space CRLB through back-projection (Roessl *et al* 2011).

It may seem like a lower bound on image variance is of limited value since it does not provide any information about the upper limit of the variance. In practice, however, the maximum-likelihood estimator

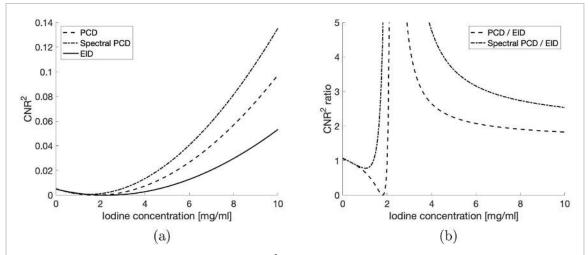


Figure 15. (a) A comparison of the projection domain CNR^2 for a photon-counting detector without energy resolution (PCD), a spectral photon-counting detector (spectral PCD) with optimal energy weighting, and an energy integrating detector (EID). The projections used to compute the CNR^2 comprised 20 cm soft tissue and 19 cm soft tissue plus 1 cm of an iodine/water solution with varying concentration. (b) The ratio of the CNR^2 curves, showing that the relative performance of the imaging systems is highly dependent on the iodine concentration.

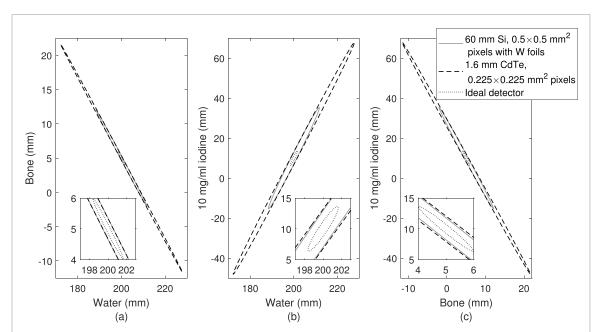


Figure 16. Illustration of CRLB for realistic simulation models of CdTe-based and silicon-based detectors and an ideal detector, in a three-basis (water/bone/iodine) decomposition. The tube voltage is 120 kVp and the object consists of 200 mm of water, 5 mm of bone and 10 mm of iodine (10 mg ml⁻¹). In each plot, the contours show the ellipse that the decomposed projected basis coefficients will fall inside with 95% probability according to the CRLB, when aggregating data from a 9×9 mm² detector segment. The CdTe-based detector is 1.6 mm thick and divided into 0.225 × 0.225 mm² pixels. The silicon sensor has an absorption depth of 60 mm, a pixel pitch of 0.5 × 0.5 mm², and 20 μ m thick tungsten scatter blockers between the wafers. The detector models are as described by Persson *et al* (2020), with the difference that the CdTe-based and the silicon-based detector here both have eight energy bins. Object scatter was taken into account assuming a one-dimensional tungsten anti-scatter grid of 25 × 0.1 mm lamellae with 1.0 mm (for Si) and 1.125 mm (for CdTe) spacing in front of the detector. The ellipses representing the silicon-based and CdTe-based detectors are more elongated compared to the ideal detector, showing that imperfect energy resolution makes it more difficult to identify the material composition.

comes close to attaining the CRLB (Roessl *et al* 2011). Figure 16 illustrates the CRLB for a three-basis decomposition task, for a CdTe-based and a silicon-based detector model.

It is important to note that the lower limit of basis projection variance is a lower limit of the variance of any unbiased estimator. If the estimator is allowed to be biased, the variance can be allowed to be lower than this limit. An extreme example would be a material-decomposition algorithm that predicts the same projected basis coefficients regardless of the measured data, which gives an estimator with zero variance and large bias. In practice, it may be desirable to use inequality constraints, such as non-negativity (Barber *et al* 2016, Long and Fessler 2014), and thereby reduce variance for certain basis configurations at the expense of a small amount of bias.

In contrast to CNR, which is relevant for detection tasks, the CRLB measures the ability of the imaging system to identify the material composition of the imaged object. The CRLB therefore provides a measure of the energy-resolving capability of a PCD, and it has been applied to investigating the relationship between pixel size and energy resolution in PCDs (Rajbhandary *et al* 2018). Faby *et al* used the CRLB to validate the statistical optimality of an image-based decomposition method, which was then used to study the effect of energy-bin correlations and compare the performance of photon-counting systems with other dual-energy CT techniques (Faby *et al* 2015, 2016). Another application of the CRLB is to estimate the covariance matrix between basis line integrals in ray-wise material decomposition in order to obtain the data weighting for the reconstruction step in a two-step material-decomposition algorithm (Schirra *et al* 2013, Sawatzky *et al* 2014).

6.3. Spectral DQE (energy-dependent spatial resolution and noise)

Characterization of conventional x-ray detectors is typically done using the noise-equivalent quanta (NEQ) and DQE. These metrics are based on linear-systems theory and are typically expressed as functions of spatial frequency u. The NEQ can be calculated from the incident number of photons q, the large-area gain G, detector MTF, and NPS as follows (Cunningham 2000):

$$NEQ(\boldsymbol{u}) = \frac{q^2 G^2 MTF^2(\boldsymbol{u})}{NPS(\boldsymbol{u})}.$$
(12)

The NEQ of a detector is the number of noise quanta that an *ideal* detector, which registers every incoming photon and its exact position of incidence, would need to register in order to achieve the same detectability as the detector under consideration. This metric is dependent on the illumination conditions. For example, a detector which has an NEQ of $15\,000$ mm⁻² for a particular spatial frequency band when irradiated by 20 000 photons per mm² is 75% as efficient as an ideal detector, since the ideal detector could achieve the same detectability with only 15 000 incident photons. The DQE is obtained by normalizing the NEQ by the number of incident photons,

$$DQE(\boldsymbol{u}) = \frac{NPS(\boldsymbol{u})}{q},$$
(13)

which can be shown to be equal to the squared ideal-linear-observer detectability relative to an ideal detector, $d'(\boldsymbol{u})^2/d'_{\text{ideal}}(\boldsymbol{u})^2$ (Cunningham 2000). In the above example, the DQE is equal to 15 000/20 000 = 0.75 for the given spatial frequency band.

DQE is a useful metric of detector performance because it takes both noise and resolution into account. It is therefore a better measure of dose efficiency than CNR, which does not incorporate the effect of spatial resolution. Whereas the CNR increases by low-pass filtration of the image after acquisition, such processing leaves the DQE unchanged, reflecting the fact that low-pass filtering does not add information to the image.

PCDs with a single energy bin (i.e. non-energy-resolving detectors) can be analyzed with the traditional NEQ and DQE metrics. An early such study was made by Durst *et al* (2007); Acciavatti and Maidment (2010) compared the DQE of photon-counting and EIDs; and Stierstorfer *et al* modeled the DQE of a CdTe-based detector (Stierstorfer 2018, Stierstorfer *et al* 2019). A common way of simulating signal and noise transfer characteristics is to model the detection process as a cascade of simple processes, the so-called cascaded-systems analysis (Cunningham 2000). Tanguay *et al* (2013) developed the theory for cascaded-systems analysis of PCDs and used it to study the impact of non-ideal physical effects on detector performance (Tanguay *et al* 2015, Tanguay and Cunningham 2018). Furthermore, Xu *et al* (2014) studied a silicon-strip PCD for mammography with a cascaded model, and Ji *et al* (2018) used a cascaded model to study anti-coincidence logic in a CdTe-based detector.

To analyze the ability of PCDs to perform energy-resolved imaging, cascaded-systems analysis has also been applied to energy-resolving detectors by Tanguay and Cunningham (2018) and in the work of Taguchi *et al* (2016, 2018a, 2018b), where such a model was used to study the effect of $N \times N$ binning of pixels into macro-pixels in a CdTe-based detector. Energy-resolving silicon-strip detectors also been studied, by Fredenberg *et al* (2010a) and Chen *et al* (2016). However, fully understanding how the resulting signal transfer and noise characteristics affect image quality requires an extension of the NEQ and DQE concepts to energy-resolving detectors. Important steps in this direction were taken by Richard and Siewerdsen (2008) in the context of dual-energy radiography.

A natural extension of the NEQ and DQE metrics to energy-resolving PCDs was proposed by Persson *et al* (2018b) and will be briefly outlined here. An equivalent spatial-domain description has been described

by Rajbhandary *et al* (2019). Although it may be tempting to simply extend the NEQ to be dependent on incident photon energy in addition to spatial frequency, this definition does not work with broad-spectrum illumination since it fails to describe how different incident energies can be confounded during the detection process as described in section 4.4. Instead, NEQ needs to be defined as a function of two energy indices so that NEQ(u, E_1 , E_2) describes how much the joint presence of photons with energy E_1 and E_2 in the incident spectrum contributes to image detectability at spatial frequency u.

To compute this generalized NEQ, it is necessary to know detector characteristics. The signal transfer is described by the transfer function $H_k(\mathbf{u})$, which corresponds to the MTF but without absolute value and without the normalization to 1 at $\mathbf{u} = 0$. The noise characteristics of the sampled signal is described by the cross-spectral density W_{d^+} , essentially a frequency-dependent covariance matrix between energy bins. From these quantities, the generalized NEQ is obtained as

$$\operatorname{NEQ}(\boldsymbol{u}, E_1, E_2) = \sum_{k_1} \sum_{k_2} \overline{q}(E_1) H_{k_1}^*(\boldsymbol{u}, E_1) \left(W_{d^+}^{-1} \right)_{k_1, k_2}^*(\boldsymbol{u}) H_{k_2}(\boldsymbol{u}, E_2) \overline{q}(E_2)$$
(14)

with $\overline{q}(E)$ denoting the spectral photon density and ^{*} denoting complex conjugate.

Under the assumption that the object consists of a small set \mathcal{B} of basis materials, it is more convenient to express the generalized NEQ in matrix form through a basis transformation. The elements NEQ_{l1,l2}^{\mathcal{B}} (\boldsymbol{u}) of this NEQ matrix describe the contribution to image detectability from the addition of differential amounts of basis materials l_1 and l_2 . In addition to allowing the detectability to be calculated for detection tasks, this matrix also describes the performance for quantification tasks in a set of decomposed basis images. Namely, the matrix elements of its inverse $(NEQ^{\mathcal{B}}(\boldsymbol{u})^{-1})_{l_1,l_2}$ give, up to a scale factor, the CRLB for the covariance of basis images l_1 and l_2 as a function of frequency (Persson *et al* 2018b, equation 25). This allows calculating the squared ideal-linear-observer detectability in basis image l as

$$d_l^{\prime 2} = \int \frac{\left(\Delta \hat{\tilde{A}}_l(\boldsymbol{u})\right)^2}{\operatorname{Var}\left(\hat{\tilde{A}}_l(\boldsymbol{u})\right)} d\boldsymbol{u}$$
(15)

where $\Delta \tilde{A}_l(\boldsymbol{u})$ is the difference between target and background in the Fourier transform $\tilde{A}_l(\boldsymbol{u})$ of the basis image, and $\operatorname{Var}\left(\hat{A}_l(\boldsymbol{u})\right)$ is the variance of an efficient estimator for \tilde{A}_l .

As an illustration of equation (15), the relative squared ideal-linear-observer detectability for detection of features of different sizes in an iodine basis image is shown in figure 17 for simulation models of CdTe-based and silicon-based detectors. As shown in this figure, a system with smaller pixels, such as the simulated CdTe system, has an advantage for imaging of small features. At the same time, a system with high energy resolution, such as the simulated silicon system, has an advantage for imaging larger features in a material-selective basis image, in particular for three-basis decomposition where energy resolution is particularly important in order to separate iodine and calcium. It is also interesting to note that the CdTe detector performance exhibits a relatively weak kVp dependence, whereas the silicon detector benefits strongly from using low kVp, where the fraction of Compton scatter is low. These observations show that the spectral linear-systems framework provides important clues for how to best utilize PCDs, especially in view of the ongoing trend towards using lower kVp to save patient dose.

In analogy with equation (13), a generalized DQE matrix can also be defined by normalizing the NEQ matrix by the NEQ matrix of a detector with perfect quantum efficiency, energy resolution, and spatial resolution:

$$DQE_{l_1,l_2}^{\mathcal{B}}(\boldsymbol{u}) = \frac{NEQ_{l_1,l_2}^{\mathcal{B}}(\boldsymbol{u})}{\sqrt{NEQ_{l_1,l_1}^{\mathcal{B},ideal}NEQ_{l_2,l_2}^{\mathcal{B},ideal}}}.$$
(16)

In analogy with the conventional DQE, it can be shown that a diagonal element $DQE_{l,l}^{\mathcal{B}}$ of this matrix gives the task-specific DQE, i.e. the squared ideal-linear-observer detectability $d'_l(\boldsymbol{u})^2/d'_{ideal}(\boldsymbol{u})^2$ relative to an ideal energy-resolving detector, for detecting basis material *l*. Here, the ideal-observer detectability of a multi-channel image should be interpreted as the highest detectability that can be obtained in any image as a weighted sum of the images, where the weight factors are allowed to vary with spatial frequency (Yveborg *et al* 2015b). The generalized NEQ matrix thus encodes the performance of an energy-resolving PCD for both detection and quantification tasks. How this framework can be applied to studying PCD performance is described by Persson *et al* (2020).

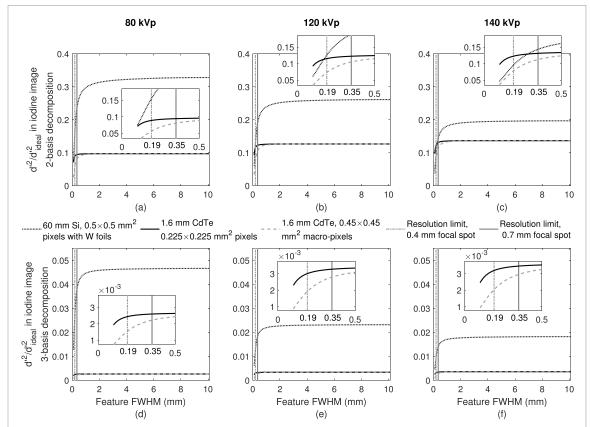


Figure 17. Simulated squared ideal-linear-observer detectability $d^{1/2}$ relative to an ideal energy-resolving detector, for realistic models of CdTe-based and silicon-based detectors. The curves show the squared detectability for Gaussian features in an iodine basis image resulting from 2-basis (iodine/water) (a)–(c) and 3-basis (iodine/bone/water) (d)–(f) decompositions, with a 200 mm thick water object. Three tube voltages were used: 80 kVp (a), (d), 120 kVp (b), (e), and 140 kVp (c), (f). The detector models and anti-scatter grid employed are the same as in figure 16, and the simulation model is described in Persson *et al* (2020). The feature size is given as full width at half maximum (FWHM) at the isocenter plane. The smaller pixel size of the CdTe-based detector gives leads to high detectability for small features, whereas the superior energy resolution of the silicon-based detector with pixels aggregated in 2 × 2 blocks, forming $0.45 \times 0.45 \text{ mm}^2$ macro-pixels. For small features, the plotted detectability should be interpreted as an upper performance limit allowed by the detector, and not as anticipated system performance, since the detectability in a real system is also affected by the size of the x-ray source. Effectively, there will be no features smaller than the smallest available focal spot size, currently $0.4 \text{ mm} \times 0.5 \text{ mm}$ (Yanagawa *et al* 2018), and this spot size is only available at a limited tube current. The vertical lines in the figure show the resolution limits in isocenter for focal spot size sof 0.4-0.7 mm (IEC 60336) corresponding to typical minimum spot sizes for state-of-the-art scanners, assuming a bimodal focal spot shape (da Silva *et al* 2019).

6.4. Quantitative image accuracy

CNR, CLRB, and DQE are metrics related to the image noise level. In addition to these noise metrics, the systematic error, or bias, in the average value in regions of the image is also an important performance measure, for example when evaluating the conspicuity of artifacts or the accuracy of quantitative image measurements. For the latter case, a relevant metric is the mean-squared error, which combines the bias with the noise variance as (Kay 1993)

$$MSE(I) = (E[I] - I^{True})^{2} + Var[I]$$
(17)

where $E[\cdot]$ and $Var[\cdot]$ denote expected value and variance, respectively, and I^{True} is the true value of the random image value *I*.

Although the standard deviation of image noise in a reconstructed image is typically several Hounsfield units, the averaging that takes place when measuring the mean CT number in a region of interest suppresses noise so that even a small systematic error can dominate the MSE. Similarly, when making quantitative measurements in a basis image, it is more important to reduce bias if the measurement region of interest is large, whereas for small regions it may be favorable to reduce noise at the expense of bias by reducing the number of basis materials (Bornefalk and Persson 2014).

One source of error is the log-normalization step transforming counts into attenuation line integrals. Since this operation is non-linear, it introduces an error that becomes larger as the dose decreases (Zhang *et al* 2018). Other sources of error are inaccuracies in the model used for material decomposition. In order to

make quantitative material composition measurements that are limited by quantum noise rather than systematic errors, it is necessary to have a detailed model of the x-ray source, the detector response, and scattered photons from the object (Bornefalk *et al* 2015). Estimating the distribution of object scatter from energy-resolved measurements is a non-trivial problem since the energy spectra of scattered and primary photons overlap in general. However, energy-resolved measurements can be helpful for scatter correction and/or measurements of the scatter distribution (Sossin *et al* 2016, 2017).

Despite the fact that more research is needed to address these challenges, measurements of K-edge contrast agents such as gold or iodine in basis images generated with prototype spectral CT systems have been shown to correlate well with the actual concentrations of these elements (Si-Mohamed *et al* 2017b, 2018).

7. Summary and outlook

After many years of technological advancements, PCDs are now on the verge of being adopted in clinical x-ray computed tomography. During the last decade in particular, there has been substantial research activity in the field of photon-counting CT, both in hardware development and in terms of improved understanding of how the resulting image quality is impacted by the detector performance. As outlined in the above sections, there is now a profound understanding of how detector design parameters (e.g. the choice of detector material and pixel size) or how choices in the signal processing chain (e.g. shaping time and threshold locations) affect the detector properties such as quantum efficiency, spatial and energy resolution, and pileup response. Likewise, there has been considerable progress towards understanding how these detector properties impact the resulting image quality, as outlined in section 6.

The challenge posed by pileup at high count rates, which was previously the main obstacle for the adoption of PCDs in clinical CT, has been largely overcome with the latest generation of detectors. In addition, there has been considerable progress in techniques for minimizing the detrimental effects of charge sharing, fluorescence, Compton scatter, and incomplete charge collection. Consequently, there are now several photon-counting CT prototypes available and used for imaging human volunteers. A first-generation photon-counting CT performs on par with conventional energy-integrating CT for unenhanced density imaging with normal dose (Symons *et al* 2016), so there is evidently room for improvement with better detectors in the future. The adoption of PCDs for CT also enables new imaging techniques that are still in the research stage, such as combining photon-counting CT with dual-energy acquisition (Faby *et al* 2015, Tao *et al* 2019a) or with phase-contrast imaging (Epple *et al* 2015).

Looking ahead, continued improvements in detector technology coupled with more accurate physics modeling and data-processing methods are expected to bring the technology closer to its full potential, leading to substantial improvements in image quality in terms of contrast and spatial resolution at lower dose, to quantitatively reliable images, and to improved material discrimination capabilities. This is anticipated to improve diagnostic performance in a diversity of application areas such as cardiovascular, neurological, thoracic, oncologic, and pediatric imaging, just to name a few. We can expect the coming decade to be even more productive than the last one for the field of photon-counting CT.

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References

Acciavatti R J and Maidment A D A 2010 A comparative analysis of OTF, NPS and DQE in energy integrating and photon counting digital x-ray detectors *Med. Phys.* **37** 6480–95

Alessio A M and MacDonald L R 2013 Quantitative material characterization from multi-energy photon counting CT *Med. Phys.* 40 031108

Alvarez R E 2010 Near optimal energy selective x-ray imaging system performance with simple detectors Med. Phys. 37 822-41

Alvarez R E 2011 Estimator for photon counting energy selective x-ray imaging with multibin pulse height analysis *Med. Phys.* **38** 2324–34

Alvarez R E 2013 Dimensionality and noise in energy selective x-ray imaging Med. Phys. 40 111909

Alvarez R E 2016 Efficient, non-iterative estimator for imaging contrast agents with spectral x-ray detectors *IEEE Trans. Med. Imaging* 35 1138–46

Alvarez R E and Macovski A 1976 Energy-selective reconstructions in x-ray computerised tomography Phys. Med. Biol. 21 733-44

Åslund M, Cederström B, Lundqvist M and Danielsson M 2007 Physical characterization of a scanning photon counting digital mammography system based on Si-strip detectors *Med. Phys.* **34** 1918–25

- Atak H and Shikhaliev P M 2015 Dual energy CT with photon counting and dual source systems: comparative evaluation *Phys. Med. Biol.* **60** 8949–75
- Atak H and Shikhaliev P M 2016 Photon counting x-ray imaging with K-edge filtered x-rays: a simulation study Med. Phys. 43 1385-400
- Baek J, Pineda A R and Pelc N J 2013 To bin or not to bin? the effect of CT system limiting resolution on noise and detectability *Phys. Med. Biol.* **58** 1433
- Bale D S and Szeles C 2008 Nature of polarization in wide-bandgap semiconductor detectors under high-flux irradiation: application to semi-insulating $Cd_{1-x}Zn_x$ Te *Phys. Rev.* B 77 035205
- Ballabriga R et al 2016 Review of hybrid pixel detector readout ASICs for spectroscopic x-ray imaging J. Instrum. 11 P01007 Ballabriga R, Campbell M, Heijne E, Llopart X and Tlustos L 2007 The medipix3 prototype, a pixel readout chip working in single
- photon counting mode with improved spectrometric performance *IEEE Trans. Nucl. Sci.* 54 1824–9 Barber R F, Sidky E Y, Schmidt T G and Pan X 2016 An algorithm for constrained one-step inversion of spectral CT data *Phys. Med. Biol.*
- 61 3784–3818
- Barber W C et al 2013 High flux energy-resolved photon-counting x-ray imaging arrays with CdTe and CdZnTe for clinical CT 3rd Int. Conf. Advancements Nuclear Instrumentation, Measurement Methods and Their Applications (ANIMMA) (New York: IEEE) (https://doi.org/10.1109/ANIMMA.2013.6728030)

Barber W, Wessel J, Nygard E and Iwanczyk J 2015 Energy dispersive CdTe and CdZnTe detectors for spectral clinical CT and NDT applications *Nucl. Instrum. Methods* A 784 531–7

Barrett H H, Eskin J and Barber H 1995 Charge transport in arrays of semiconductor gamma-ray detectors *Phys. Rev. Lett.* **75** 156 Barrett H H and Myers K J 2003 *Foundations of Image Science* (Hoboken, NJ: Wiley)

- Bartlett D J, Koo C W, Bartholmai B J, Rajendran K, Weaver J M, Halaweish A F, Leng S, McCollough C H and Fletcher J G 2019 High-resolution chest computed tomography imaging of the lungs: impact of 1024 matrix reconstruction and photon-counting detector computed tomography *Invest. Radiol.* 54 129–37
- Benjaminov O, Perlow E, Romman Z, Levinson R, Bashara B, Cohen M and Zelikovsky A 2008 Novel, energy-discriminating photon counting CT system (EDCT): first clinical evaluation—CT angiography: Carotid artery stenosis Radiological Society of North America 2008 Scientific Assembly and Annual Meeting, Chicago IL (SSA21-05)
- Berenguer R, Pastor-Juan M d R, Canales-Vázquez J, Castro-García M, Villas M V, Mansilla Legorburo F and Sabater S 2018 Radiomics of CT features may be nonreproducible and redundant: influence of CT acquisition parameters *Radiology* 288 407–15
- Bolotnikov A, Camarda G, Wright G and James R 2005 Factors limiting the performance of CdZnTe detectors *IEEE Trans. Nucl. Sci.* 52 589–98
- Bolotnikov A, Cook W, Harrison F, Wong A S, Schindler S and Eichelberger A 1999 Charge loss between contacts of CdZnTe pixel detectors *Nucl. Instrum. Methods* A 432 326–31
- Bornefalk H 2012a Synthetic hounsfield units from spectral CT data Phys. Med. Biol. 57 N83-N87
- Bornefalk H 2012b XCOM intrinsic dimensionality for low-Z elements at diagnostic energies Med. Phys. 39 654-7

Bornefalk H and Danielsson M 2010 Photon-counting spectral computed tomography using silicon strip detectors: a feasibility study *Phys. Med. Biol.* **55** 1999–2022

Bornefalk H and Persson M 2014 Theoretical comparison of the iodine quantification accuracy of two spectral CT technologies *IEEE Trans. Med. Imaging* 33 556–65

Bornefalk H, Persson M and Danielsson M 2015 Allowable forward model misspecification for accurate basis decomposition in a silicon detector based spectral CT *IEEE Trans. Med. Imaging* 34 788–95

Boucher Y 2013 Analysis of cadmium zinc telluride detector performance and characteristics for applications in gamma-ray imaging spectrometers *PhD Thesis* University of Michigan, Ann Arbor, MI

Boussel L, Coulon P, Thran A, Roessl E, Martens G, Sigovan M and Douek P 2014 Photon counting spectral CT component analysis of coronary artery atherosclerotic plaque samples *Br. J. Radiol.* **87** 20130798

Brambilla A, Ouvrier-Buffet P, Gonon G, Rinkel J, Moulin V, Boudou C and Verger L 2013 Fast CdTe and CdZnTe semiconductor detector arrays for spectroscopic x-ray imaging *IEEE Trans. Nucl. Sci.* 60 408–15

Cammin J, Kappler S G, Weidinger T and Taguchi K 2016 Evaluation of models of spectral distortions in photon-counting detectors for computed tomography *J. Med. Imaging* 3 023503

Cammin J, Xu J, Barber W C, Iwanczyk J S, Hartsough N E and Taguchi K 2014 A cascaded model of spectral distortions due to spectral response effects and pulse pileup effects in a photon-counting x-ray detector for CT *Med. Phys.* **41** 041905

Carloni F, Corberi A, Marseguerra M and Porceddu C 1970 The asymptotic method of dead-time correction in Poissonian distribution Nucl. Instrum. Methods 78 70–6

Charpak G 1997 Electronic imaging of ionizing radiation with limited avalanches in gases Rev. Mod. Phys. 65 591

Chen H, Danielsson M and Xu C 2016 Size-dependent scanning parameters (kVp and mAs) for photon-counting spectral CT system in pediatric imaging: simulation study *Phys. Med. Biol.* **61** 4105–26

Chen H, Xu C, Persson M and Danielsson M 2015 Optimization of beam quality for photon-counting spectral computed tomography in head imaging: simulation study J. Med. Imaging 2 043504

Clark D P and Badea C T 2020 Fast spectral x-ray CT reconstruction with data-adaptive, convolutional regularization *Proc. SPIE* 11312 113120A

- Cormode D P *et al* 2017 Multicolor spectral photon-counting computed tomography: *in vivo* dual contrast imaging with a high count rate scanner *Sci. Rep.* 7 4784
- Cranley K, Gilmore B J, Fogarty G W A, Desponds L and Sutton D 1997 *IPEM Report 78: Catalogue of Diagnostic X-ray Spectra and Other Data, electronic version* (York: Institute of Physics and Engineering in Medicine)

Cunningham I 2000 Applied linear-systems theory *Handbook of Medical Imaging* vol 1 Physics and Psychophysics eds R L Van Metter, J Beutel and H L Kundel (Bellingham, WA: SPIE) pp 79–160

Chen Z and Li L 2019 Robust multimaterial decomposition of spectral CT using convolutional neural networks Opt. Eng. 58 013104

- Curtis T E and Roeder R K 2017 Effects of calibration methods on quantitative material decomposition in photon-counting spectral computed tomography using a maximum a posteriori estimator *Med. Phys.* **44** 5187–97
- da Silva J, Grönberg F, Cederström B, Persson M, Sjölin M, Alagic Z, Bujila R and Danielsson M 2019 Resolution characterization of a silicon-based, photon-counting computed tomography prototype capable of patient scanning *J. Med. Imaging* **6** 043502
- Dangelmaier J *et al* 2018 Experimental feasibility of spectral photon-counting computed tomography with two contrast agents for the detection of endoleaks following endovascular aortic repair *Eur. Radiol.* 28 3318–25
- Du Y, LeBlanc J, Possin G E, Yanoff B D and Bogdanovich S 2002 Temporal response of CZT detectors under intense irradiation 2002 IEEE Nucl. Sci. Symp. Conf. Rec (New York: IEEE) pp 480–4
- Duan X, Wang J, Leng S, Schmidt B, Allmendinger T, Grant K, Flohr T and McCollough C H 2013 Electronic noise in CT detectors: Impact on image noise and artifacts AJR Am. J. Roentgenol. 201 W626–W632
- Dubousset J, Charpak G, Skalli W, Kalifa G and Lazennec J Y 2007 Système EOS : la radiographie dose efficiency la tête aux pieds face et profil simultanés à très basses doses dose efficiency radiations Revue dose efficiency Chirurgie Orthopédique et Traumatologique 93 SUP6 143-6
- Durst J, Anton G and Michel T 2007 Discriminator threshold dependency of the zero-frequency DQE of photon-counting pixel detectors *Nucl. Instrum. Methods* A 576 235–8
- Ehn S, Sellerer T, Mechlem K, Fehringer A, Epple M, Herzen J, Pfeiffer F and Noël P 2016 Basis material decomposition in spectral CT using a semi-empirical, polychromatic adaption of the beer-lambert model *Phys. Med. Biol.* **62** N1–N17
- Epple F M *et al* 2015 Phase unwrapping in spectral x-ray differential phase-contrast imaging with an energy-resolving photon-counting pixel detector *IEEE Trans. Med. Imaging* 34 816–23
- Faby S, Kuchenbecker S, Sawall S, Simons D, Schlemmer H P, Lell M and Kachelrieß M 2015 Performance of today's dual energy CT and future multi energy CT in virtual non-contrast imaging and in iodine quantification: a simulation study *Med. Phys.* 42 4349–66

Faby S, Maier J, Sawall S, Simons D, Schlemmer H P, Lell M and Kachelrieß M 2016 An efficient computational approach to model statistical correlations in photon counting x-ray detectors *Med. Phys.* **43** 3945–60

- Fang Y, Xu C, Yao Y, Pelc N, Danielsson M and Badano A 2018 Modeling charge transport in photon-counting detectors Nucl. Instrum. Methods A 899 115–21
- Feng C, Kang K and Xing Y 2019 Fully connected neural network for virtual monochromatic imaging in spectral computed tomography J. Med. Imaging 6 011006
- Feng R, Rundle D and Wang G 2018 Neural-networks-based photon-counting data correction: Pulse pileup effect arXiv:1804.10980

Flohr T, Stierstorfer K, Süss C, Schmidt B, Primak A and McCollough C H 2007 Novel ultrahigh resolution data acquisition and image reconstruction for multi-detector row ct *Med. Phys.* **34** 1712–23

- Fredenberg E, Hemmendorff M, Cederström B, åslund M and Danielsson M 2010a Contrast-enhanced spectral mammography with a photon-counting detector *Med. Phys.* **37** 2017–29
- Fredenberg E, Lundqvist M, Cederström B, Åslund M and Danielsson M 2010b Energy resolution of a photon-counting silicon strip detector *Nucl. Instrum. Methods* A 613 156–62
- Gao H, Yu H, Osher S and Wang G 2011 Multi-energy CT based on a prior rank, intensity and sparsity model (PRISM) *Inverse Probl.* 27 115012
- Glover G H and Pelc N J 1980 Nonlinear partial volume artifacts in x-ray computed tomography Med. Phys. 7 238–48
- Grönberg F, Lundberg J, Sjölin M, Persson M, Bujila R, Bornefalk H, Almqvist H, Holmin S and Danielsson M 2020 Feasibility of unconstrained three-material decomposition: imaging an excised human heart using a prototype silicon photon-counting CT detector *Eur. Radiol.* **30** 5904–12
- Grönberg F, Danielsson M and Sjölin M 2018 Count statistics of nonparalyzable photon-counting detectors with nonzero pulse length *Med. Phys.* **45** 3800–11
- Gupta R, Grasruck M, Suess C, Bartling S H, Schmidt B, Stierstorfer K, Popescu S, Brady T and Flohr T 2006 Ultra-high resolution flat-panel volume CT: fundamental principles, design architecture and system characterization *Eur. Radiol.* **16** 1191–205
- Gutjahr R *et al* 2016 Human imaging with photon counting-based computed tomography at clinical dose levels: contrast-to-noise ratio and cadaver studies *Invest. Radiol.* **51** 421–9

Hamel L A and Paquet S 1996 Charge transport and signal generation in CdTe pixel detectors *Nucl. Instrum. Methods* A 380 238–40 Han-soo L 2017 Samsung unveils next-generation radiology tech in us Korea Biomedical Review (Seoul, Korea: The Korean Doctors' Weekly Co., Ltd.) (available at: www.koreabiomed.com/news/articleView.html?idxno=1981) (Accessed: 9 Feb 2020)

- Harrison A P, Xu Z, Pourmorteza A, Bluemke D A and Mollura D J 2017 A multichannel block-matching denoising algorithm for spectral photon-counting CT images *Med. Phys.* 44 2447–52
- Heismann B and Balda M 2009 Quantitative image-based spectral reconstruction for computed tomography Med. Phys. 36 4471–85
 Hsieh S S 2020 Coincidence counters for charge sharing compensation in spectroscopic photon counting detectors IEEE Trans. Med. Imaging 39 678–87
- Hsieh S S and Pelc N J 2016 Improving pulse detection in multibin photon-counting detectors J. Med. Imaging 3 023505
- Hsieh S S, Rajbhandary P L and Pelc N J 2018 Spectral resolution and high-flux capability tradeoffs in CdTe detectors for clinical CT *Med. Phys.* 45 1433–43
- Hsieh S S and Sjolin M 2018 Digital count summing vs analog charge summing for photon counting detectors: a performance simulation study *Med. Phys.* 45 4085–93
- Hu R, Daftari Besheli L, Young J, Wu M, Pomerantz S, Lev M H and Gupta R 2016 Dual-energy head CT enables accurate distinction of intraparenchymal hemorrhage from calcification in emergency department patients *Radiology* **280** 177–83

Hubbell J and Seltzer S 2004 Tables of x-ray mass attenuation coefficients and mass energy-absorption coefficients (version 1.4) (Gaithersburg, MD: National Institute of Standards and Technology) Originally published as NISTIR 5632 (https://dx.doi.org/10.18434/T4D01F)

- Iwanczyk J S, Nygard E, Meirav O, Arenson J, Barber W C, Hartsough N E, Malakhov N and Wessel J C 2009 Photon counting energy dispersive detector arrays for x-ray imaging IEEE Trans. Nucl. Sci. 56 535–42
- Ji X, Zhang R, Chen G H and Li K 2018 Impact of anti-charge sharing on the zero-frequency detective quantum efficiency of CdTe-based photon counting detector system: cascaded systems analysis and experimental validation *Phys. Med. Biol.* **63** 095003
- Kachelrieß M and Kalender W A 2005 Presampling, algorithm factors and noise: considerations for CT in particular and for medical imaging in general Med. Phys. 32 1321–34
- Kalender W A, Kolditz D, Steiding C, Ruth V, Lück F, Rößler A C and Wenkel E 2017 Technical feasibility proof for high-resolution low-dose photon-counting CT of the breast *Eur. Radiol.* 27 1081–6

Kappler S, Hölzer S, Kraft E, Stierstorfer K and Flohr T 2011 Quantum-counting CT in the regime of count-rate paralysis: introduction of the pile-up trigger method *Proc. SPIE* **7961** 79610T

Kay S M 1993 Fundamentals of Statistical Signal Processing: Estimation Theory (Upper Saddle River, NJ: Prentice Hall PTR) Kellock T T, Nicolaou S, Kim S S Y, Al-Busaidi S, Louis L J, O'Connell T W, Ouellette H A and McLaughlin P D 2017 Detection of bone

marrow edema in nondisplaced hip fractures: Utility of a virtual noncalcium dual-energy CT application *Radiology* **284** 798–805 Kim J, Bar-Ness D, Si-Mohamed S, Coulon P, Blevis I, Douek P and Cormode D P 2018 Assessment of candidate elements for development of spectral photon-counting CT specific contrast agents *Sci. Rep.* **8** 12119

Koenig T *et al* 2013 Charge summing in spectroscopic x-ray detectors with high-Z sensors *IEEE Trans. Nucl. Sci.* 60 4713–18

Koenig T et al 2014 How spectroscopic x-ray imaging benefits from inter-pixel communication Phys. Med. Biol. 59 6195

Kopp F K et al 2018 Evaluation of a preclinical photon-counting CT prototype for pulmonary imaging Sci. Rep. 8 17386

Krause M O 1979 Atomic radiative and radiationless yields for K and L shells J. Phys. Chem. Ref. Data 8 307-27

Lambert J W et al 2018 An intravascular tantalum oxide–based CT contrast agent: preclinical evaluation emulating overweight and obese patient size Radiology 289 103–10

- Lee O, Kappler S, Polster C and Taguchi K 2017a Estimation of basis line-integrals in a spectral distortion-modeled photon counting detector using low-order polynomial approximation of x-ray transmittance *IEEE Trans. Med. Imaging* 36 560–73
- Lee O, Kappler S, Polster C and Taguchi K 2017b Estimation of basis line-integrals in a spectral distortion-modeled photon counting detector using low-rank approximation-based x-ray transmittance modeling: K-edge imaging application IEEE Trans. Med. Imaging 36 2389–403
- Lee S, Choi Y N and Kim H J 2014 Quantitative material decomposition using spectral computed tomography with an energy-resolved photon-counting detector *Phys. Med. Biol.* **59** 5457–82
- Leng S *et al* 2016 Dose-efficient ultrahigh-resolution scan mode using a photon counting detector computed tomography system *J. Med. Imaging* 3 043504
- Leng S *et al* 2018 150-μm spatial resolution using photon-counting detector computed tomography technology: technical performance and first patient images *Invest. Radiol.* **53** 655–62
- Leng S, Yu L, Wang J, Fletcher J G, Mistretta C A and McCollough C H 2011 Noise reduction in spectral CT: Reducing dose and breaking the trade-off between image noise and energy bin selection *Med. Phys.* **38** 4946–57
- Leng S, Zhou W, Yu Z, Halaweish A, Krauss B, Schmidt B, Yu L, Kappler S and McCollough C 2017 Spectral performance of a whole-body research photon counting detector CT: quantitative accuracy in derived image sets *Phys. Med. Biol.* **62** 7216–32
- Li Z, Leng S, Yu L, Manduca A and McCollough C H 2017 An effective noise reduction method for multi-energy CT images that exploit spatio-spectral features *Med. Phys.* 44 1610–23
- Liu X, Bornefalk H, Chen H, Danielsson M, Karlsson S, Persson M, Xu C and Huber B 2014 A silicon-strip detector for photon-counting spectral CT: energy resolution from 40 kev to 120 kev *IEEE Trans. Nucl. Sci.* **61** 1099–105
- Liu X, Chen H, Bornefalk H, Danielsson M, Karlsson S, Persson M, Xu C and Huber B 2015a Energy calibration of a silicon-strip detector for photon-counting spectral CT by direct usage of the x-ray tube spectrum *IEEE Trans. Nucl. Sci.* 62 68–75
- Liu X, Grönberg F, Sjölin M, Karlsson S and Danielsson M 2016 Count rate performance of a silicon-strip detector for photon-counting spectral CT Nucl. Instrum. Methods A 827 102–6
- Liu X, Persson M, Bornefalk H, Karlsson S, Xu C, Danielsson M and Huber B 2015b Spectral response model for a multibin photon-counting spectral computed tomography detector and its applications *J. Med. Imaging* 2 033502
- Liu X, Yu L, Primak A N and McCollough C H 2009 Quantitative imaging of element composition and mass fraction using dual-energy CT: three-material decomposition *Med. Phys.* **36** 1602–9
- Long Y and Fessler J A 2014 Multi-material decomposition using statistical image reconstruction for spectral CT IEEE Trans. Med. Imaging 33 1614–26
- Luo X F, Xie X Q, Cheng S, Yang Y, Yan J, Zhang H, Chai W M, Schmidt B and Yan F H 2015 Dual-energy CT for patients suspected of having liver iron overload: can virtual iron content imaging accurately quantify liver iron content? *Radiology* 277 95–103
- Malusek A, Karlsson M, Magnusson M and Carlsson G A 2013 The potential of dual-energy computed tomography for quantitative decomposition of soft tissues to water, protein and lipid in brachytherapy *Phys. Med. Biol.* **58** 771–85
- Mannil M et al 2018 Photon-counting CT: high-resolution imaging of coronary stents Invest. Radiol. 53 143-9

Matsumoto M, Yamamoto A, Honda I, Taniguchi A and Kanamori H 2000 Direct measurement of mammographic x-ray spectra using a CdZnTe detector *Med. Phys.* 27 1490–502

Mechlem K, Ehn S, Sellerer T, Braig E, Münzel D, Pfeiffer F and Noël P B 2018 Joint statistical iterative material image reconstruction for spectral computed tomography using a semi-empirical forward model *IEEE Trans. Med. Imaging* **37** 68–80

Mendonça P R S, Bhotika R, Maddah M, Thomsen B, Dutta S, Licato P E and Joshi M C 2010 Multi-material decomposition of spectral CT images *Proc. SPIE* **7622** 633–41

- Michel T *et al* 2006 A fundamental method to determine the signal-to-noise ratio (SNR) and detective quantum efficiency (DQE) for a photon counting pixel detector *Nucl. Instrum. Methods* A 568 799–802
- Moralles M, Bonifacio D, Bottaro M and Pereira M 2007 Monte carlo and least-squares methods applied in unfolding of x-ray spectra measured with cadmium telluride detectors *Nucl. Instrum. Methods* A 580 270–3
- Mory C, Sixou B, Si-Mohamed S, Boussel L and Rit S 2018 Comparison of five one-step reconstruction algorithms for spectral CT Phys. Med. Biol. 63 235001

Müller J W 1973 Dead-time problems Nucl. Instrum. Methods 112 47-57

Nasirudin R A, Mei K, Panchev P, Fehringer A, Pfeiffer F, Rummeny E J, Fiebich M and Noël P B 2015 Reduction of metal artifact in single photon-counting computed tomography by spectral-driven iterative reconstruction technique *PloS One* **10** 1–15

Nilsson H E, Norlin B, Fröjdh C and Tlustos L 2007 Charge sharing suppression using pixel-to-pixel communication in photon counting x-ray imaging systems *Nucl. Instrum. Methods* A 576 243–7

Niu S *et al* 2018b Iterative reconstruction for photon-counting CT using prior image constrained total generalized variation *Comput. Biol. Med.* **103** 167–82

- Niu S, Yu G, Ma J and Wang J 2018a Nonlocal low-rank and sparse matrix decomposition for spectral CT reconstruction *Inverse Probl.* 34 024003
- Pearce M S *et al* 2012 Radiation exposure from CT scans in childhood and subsequent risk of leukaemia and brain tumours: a retrospective cohort study *Lancet* **380** 499–505

Persson M, Bujila R, Nowik P, Andersson H, Kull L, Andersson J, Bornefalk H and Danielsson M 2016 Upper limits of the photon fluence rate on CT detectors: case study on a commercial scanner *Med. Phys.* **43** 4398–4411

Persson M and Grönberg F 2017 Bias-variance tradeoff in anticorrelated noise reduction for spectral CT Med. Phys. 44 e242-e254

- Persson M, Holmin S, Karlsson S, Bornefalk H and Danielsson M 2018a Subpixel x-ray imaging with an energy-resolving detector J. Med. Imaging 5 013507
- Persson M, Huber B, Karlsson S, Liu X, Chen H, Xu C, Yveborg M, Bornefalk H and Danielsson M 2014 Energy-resolved CT imaging with a photon-counting silicon-strip detector *Phys. Med. Biol.* **59** 6709–27
- Persson M, Rajbhandary P L and Pelc N J 2018b A framework for performance characterization of energy-resolving photon-counting detectors Med. Phys. 45 4897–4915
- Persson M, Wang A and Pelc N 2020 Detective quantum efficiency of photon-counting CDTE and Si detectors for computed tomography: a simulation study (arXiv:2003.00390)
- Phan C, Yoo A, Hirsch J, Nogueira R and Gupta R 2012 Differentiation of hemorrhage from iodinated contrast in different intracranial compartments using dual-energy head CT *AJNR Am. J. Neuroradiol.* **33** 1088–94
- Pourmorteza A *et al* 2016 Abdominal imaging with contrast-enhanced photon-counting CT: first human experience *Radiology* 279 239–45
- Pourmorteza A, Symons R, Reich D, Bagheri M, Cork T, Kappler S, Ulzheimer S and Bluemke D 2017 Photon-counting CT of the brain: *in vivo* human results and image-quality assessment *AJNR Am. J. Neuroradiol.* **38** 2257–63
- Rajbhandary P L, Hsieh S S and Pelc N J 2017 Segmented targeted least squares estimator for material decomposition in multibin photon-counting detectors J. Med. Imaging 4 023503
- Rajbhandary P L, Hsieh S S and Pelc N J 2018 Effect of spectral degradation and spatio-energy correlation in x-ray PCD for imaging IEEE Trans. Med. Imaging 37 1910–19
- Rajbhandary P L and Pelc N J 2016 Comparison weighted energy bin vs. weighted basis material CT images Proc. 4th Int. Conf. on Image Formation in X-Ray Computed Tomography pp 327–9
- Rajbhandary P L and Pelc N J 2018 Effect of electronic noise and lowest energy threshold selection in photon counting detectors *Proc.* SPIE 10573 105734S
- Rajbhandary P L, Persson M and Pelc N J 2019 Detective efficiency of photon counting detectors with spectral degradation and crosstalk Med. Phys. 47 27–36
- Ren L, Tao S, Rajendran K, McCollough C H and Yu L 2019 Impact of prior information on material decomposition in dual- and multienergy computed tomography J. Med. Imaging 6 013503
- Richard S and Siewerdsen J H 2008 Cascaded systems analysis of noise reduction algorithms in dual-energy imaging *Med. Phys.* 35 586–601
- Rigie D S and La Rivière P J 2015 Joint reconstruction of multi-channel, spectral CT data via constrained total nuclear variation minimization *Phys. Med. Biol.* **60** 1741–62
- Roessl E, Brendel B, Engel K, Schlomka J, Thran A and Proksa R 2011 Sensitivity of photon-counting basedk-edge imaging in x-ray computed tomography *IEEE Trans. Med. Imaging* **30** 1678–90

Roessl E, Daerr H and Proksa R 2016 A fourier approach to pulse pile-up in photon-counting x-ray detectors Med. Phys. 43 1295-8

Roessl E and Herrmann C 2009 Cramér–Rao lower bound of basis image noise in multiple-energy x-ray imaging *Phys. Med. Biol.* 54 1307–18

- Roessl E and Proksa R 2007 K-edge imaging in x-ray computed tomography using multi-bin photon counting detectors *Phys. Med. Biol.* 52 4679–96
- Ronaldson J P, Zainon R, Scott N J A, Gieseg S P, Butler A P, Butler P H and Anderson N G 2012 Toward quantifying the composition of soft tissues by spectral CT with Medipix3 Med. Phys. 39 6847–57
- Salehjahromi M, Zhang Y and Yu H 2018 Comparison study of regularizations in spectral computed tomography reconstruction *Sens. Imaging* **19** 16
- Sawatzky A, Xu Q, Schirra C O and Anastasio M A 2014 Proximal ADMM for multi-channel image reconstruction in spectral x-ray CT *IEEE Trans. Med. Imaging* 33 1657–68
- Schirra C O, Roessl E, Koehler T, Brendel B, Thran A, Pan D, Anastasio M A and Proksa R 2013 Statistical reconstruction of material decomposed data in spectral CT *IEEE Trans. Med. Imaging* 32 1249–57
- Schlomka J P *et al* 2008 Experimental feasibility of multi-energy photon-counting K-edge imaging in pre-clinical computed tomography *Phys. Med. Biol.* **53** 4031–47
- Schmidt T G 2009 Optimal 'image-based' weighting for energy-resolved CT Med. Phys. 36 3018–27
- Schmidt T G 2010 CT energy weighting in the presence of scatter and limited energy resolution Med. Phys. 37 1056-67
- Shefer E et al 2013 State of the art of CT detectors and sources: a literature review Curr. Radiol. Rep. 1 76-91
- Shikhaliev P M 2005 Beam hardening artefacts in computed tomography with photon counting, charge integrating and energy weighting detectors: a simulation study *Phys. Med. Biol.* **50** 5813–27
- Shikhaliev P M 2008 Computed tomography with energy-resolved detection: a feasibility study Phys. Med. Biol. 53 1475
- Shikhaliev P M 2009 Projection x-ray imaging with photon energy weighting: experimental evaluation with a prototype detector *Phys. Med. Biol.* **54** 4971
- Shikhaliev P M, Fritz S G and Chapman J W 2009 Photon counting multienergy x-ray imaging: effect of the characteristic x rays on detector performance *Med. Phys.* **36** 5107–19
- Si-Mohamed S *et al* 2017a Review of an initial experience with an experimental spectral photon-counting computed tomography system *Nucl. Instrum. Methods* A **873** 27–35
- Si-Mohamed S *et al* 2017b Evaluation of spectral photon counting computed tomography K-edge imaging for determination of gold nanoparticle biodistribution in vivo *Nanoscale* **9** 18246–57

Si-Mohamed S et al 2018 Multicolour imaging with spectral photon-counting CT: a phantom study Eur. Radiol. Exp. 2 34

- Siffert P, Berger J, Scharager C, Cornet A, Stuck R, Bell R, Serreze H and Wald F 1976 Polarization in cadmium telluride nuclear radiation detectors *IEEE Trans. Nucl. Sci.* 23 159–70
- Sigovan M *et al* 2019 Feasibility of improving vascular imaging in the presence of metallic stents using spectral photon counting CT and K-edge imaging *Sci. Rep.* **9** 19850
- Simard M, Lapointe A, Lalonde A, Bahig H and Bouchard H 2019 The potential of photon-counting CT for quantitative contrast-enhanced imaging in radiotherapy *Phys. Med. Biol.* 64 115020
- Sjölin M and Danielsson M 2017 Compression of CT sinogram data by decimation in the view direction Med. Phys. 44 e138-e146
- Sossin A, Rebuffel V, Tabary J, Létang J, Freud N and Verger L 2017 Characterizing the behavior of scattered radiation in multi-energy x-ray imaging *Nucl. Instrum. Methods* A **850** 25–34
- Sossin A, Rebuffel V, Tabary J, Létang J M, Freud N and Verger L 2016 A novel scatter separation method for multi-energy x-ray imaging *Phys. Med. Biol.* **61** 4711–28

Stierstorfer K 2018 Modeling the frequency-dependent detective quantum efficiency of photon-counting x-ray detectors Med. Phys. 45 156–66

Stierstorfer K, Hupfer M and Köster N 2019 Modeling the DQE(f) of photon-counting detectors: impact of the pixel sensitivity profile *Phys. Med. Biol.* 64 105008

Sundberg C, Sjölin M, Wikner J, Svensson C and Danielsson M 2018 Increasing the dose efficiency in silicon photon-counting detectors utilizing dual shapers *Proc. SPIE* **10573** 105734W

Swank R K 1973 Absorption and noise in x-ray phosphors J. Appl. Phys. 44 4199–203

Symons R *et al* 2017b Dual-contrast agent photon-counting computed tomography of the heart: initial experience *Int. J. Cardiovasc. Imaging* 33 1253–61

Symons R *et al* 2018a Quarter-millimeter spectral coronary stent imaging with photon-counting CT: initial experience J. Cardiovasc. Comput. Tomogr. 12 509–15

Symons R, Cork T E, Sahbaee P, Fuld M K, Kappler S, Folio L R, Bluemke D A and Pourmorteza A 2016 Low-dose lung cancer screening with photon-counting CT: a feasibility study *Phys. Med. Biol.* **62** 202–13

- Symons R, Krauss B, Sahbaee P, Cork T E, Lakshmanan M N, Bluemke D A and Pourmorteza A 2017a Photon-counting CT for simultaneous imaging of multiple contrast agents in the abdomen: an *in vivo* study *Med. Phys.* **44** 5120–7
- Symons R, Reich D S, Bagheri M, Cork T E, Krauss B, Ulzheimer S, Kappler S, Bluemke D A and Pourmorteza A 2018b Photon-counting computed tomography for vascular imaging of the head and neck: first *in vivo* human results *Invest. Radiol.* **53** 135–42
- Taguchi K, Frey E C, Wang X, Iwanczyk J S and Barber W C 2010 An analytical model of the effects of pulse pileup on the energy spectrum recorded by energy resolved photon counting x-ray detectors *Med. Phys.* **37** 3957–69
- Taguchi K and Iwanczyk J S 2013 Vision 20/20: single photon counting x-ray detectors in medical imaging *Med. Phys.* 40 100901
 Taguchi K, Polster C, Lee O, Stierstorfer K and Kappler S 2016 Spatio-energetic cross talk in photon counting detectors: detector model and correlated poisson data generator *Med. Phys.* 43 6386–404
- Taguchi K, Stierstorfer K, Polster C, Lee O and Kappler S 2018a Spatio-energetic cross-talk in photon counting detectors: N × N binning and sub-pixel masking *Med. Phys.* 45 4822–43
- Taguchi K, Stierstorfer K, Polster C, Lee O and Kappler S 2018b Spatio-energetic cross-talk in photon counting detectors: numerical detector model (PCTK) and workflow for CT image quality assessment *Med. Phys.* **45** 1985–98
- Taguchi K, Zhang M, Frey E C, Wang X, Iwanczyk J S, Nygard E, Hartsough N E, Tsui B M and Barber W C 2011 Modeling the

performance of a photon counting x-ray detector for CT: Energy response and pulse pileup effects *Med. Phys.* **38** 1089–102 Tanguay J and Cunningham I A 2018 Cascaded systems analysis of charge sharing in cadmium telluride photon-counting x-ray detectors *Med. Phys.* **45** 1926–41

Tanguay J, Yun S, Kim H K and Cunningham I A 2013 The detective quantum efficiency of photon-counting x-ray detectors using cascaded-systems analyses *Med. Phys.*

40 041913

- Tanguay J, Yun S, Kim H K and Cunningham I A 2015 Detective quantum efficiency of photon-counting x-ray detectors *Med. Phys.* 42 491–509
- Tao A, Huang R, Tao S, Michalak G J, McCollough C H and Leng S 2019a Dual-source photon counting detector CT with a tin filter: a phantom study on iodine quantification performance *Phys. Med. Biol.* **64** 115019

Tao S, Rajendran K, McCollough C H and Leng S 2018 Material decomposition with prior knowledge aware iterative denoising (MD-PKAID) Phys. Med. Biol. 63 195003

Tao S, Rajendran K, Zhou W, Fletcher J G, McCollough C H and Leng S 2019b Improving iodine contrast to noise ratio using virtual monoenergetic imaging and prior-knowledge-aware iterative denoising (mono-PKAID) *Phys. Med. Biol.* **64** 105014

Tenney F H 1984 Idealized pulse pileup effects on energy spectra Nucl. Instrum. Methods 219 165-72

- Thompson A et al 2009 X-Ray Data Booklet (Berkeley, CA: Lawrence Berkeley National Laboratory)
 Tkaczyk J E, Du Y and Li W 2009 Method and apparatus to reduce charge sharing in pixellated energy discriminating detectors US Patent 7,486,764
- Tlustos L, Ballabriga R, Campbell M, Heijne E, Kincade K, Llopart X and Stejskal P 2006 Imaging properties of the Medipix2 system exploiting single and dual energy thresholds *IEEE Trans. Nucl. Sci.* 53 367–72

Touch M, Clark D P, Barber W and Badea C T 2016 A neural network-based method for spectral distortion correction in photon counting x-ray CT *Phys. Med. Biol.* **61** 6132

von Spiczak J *et al* 2018 Photon counting computed tomography with dedicated sharp convolution kernels: tapping the potential of a new technology for stent imaging *Invest. Radiol.* **53** 486–94

Wang A S, Harrison D, Lobastov V and Tkaczyk J E 2011a Pulse pileup statistics for energy discriminating photon counting x-ray detectors Med. Phys. 38 4265–75

- Wang A S and Pelc N J 2012 A comparison of dual kV energy integrating and energy discriminating photon counting detectors for dual energy x-ray imaging *Proc. SPIE* 8313 83130W
- Wang J, Garg N, Duan X, Liu Y, Leng S, Yu L, Ritman E L, Kantor B and McCollough C H 2011b Quantification of iron in the presence of calcium with dual-energy computed tomography (DECT) in anex vivoporcine plaque model *Phys. Med. Biol.* 56 7305–16
- Wielopolski L and Gardner R P 1976 Prediction of the pulse-height spectral distortion caused by the peak pile-up effect *Nucl. Instrum. Methods* 133 303–9
- Willemink M J, Persson M, Pourmorteza A, Pelc N J and Fleischmann D 2018 Photon-counting CT: technical principles and clinical prospects Radiology 289 293–312
- Williamson J F, Li S, Devic S, Whiting B R and Lerma F A 2006 On two-parameter models of photon cross sections: application to dual-energy CT imaging *Med. Phys.* **33** 4115–29

Wu W et al 2020 Dictionary learning based image-domain material decomposition for spectral CT Phys. Med. Biol. in press (https://doi.org/10.1088/1361-6560/aba7ce)

Wu W, Wang Q, Liu F, Zhu Y and Yu H 2019 Block matching frame based material reconstruction for spectral CT Phys. Med. Biol. 64 235011

- Xu C, Chen H, Persson M, Karlsson S, Danielsson M, Svensson C and Bornefalk H 2013a Energy resolution of a segmented silicon strip detector for photon-counting spectral CT Nucl. Instrum. Methods A 715 11–17
- Xu C, Danielsson M and Bornefalk H 2011 Evaluation of energy loss and charge sharing in cadmium telluride detectors for photon-counting computed tomography *IEEE Trans. Nucl. Sci.* 58 614–25
- Xu C, Persson M, Chen H, Karlsson S, Danielsson M, Svensson C and Bornefalk H 2013b Evaluation of a second-generation ultra-fast energy-resolved ASIC for photon-counting spectral CT *IEEE Trans. Nucl. Sci.* **60** 437–45

- Xu J, Zbijewski W, Gang G, Stayman J W, Taguchi K, Lundqvist M, Fredenberg E, Carrino J A and Siewerdsen J H 2014 Cascaded systems analysis of photon counting detectors *Med. Phys.* **41** 101907
- Yanagawa M, Hata A, Honda O, Kikuchi N, Miyata T, Uranishi A, Tsukagoshi S and Tomiyama N 2018 Subjective and objective comparisons of image quality between ultra-high-resolution CT and conventional area detector CT in phantoms and cadaveric human lungs *Eur. Radiol.* 28 5060–8
- Yu Z et al 2016c Evaluation of conventional imaging performance in a research whole-body CT system with a photon-counting detector array Phys. Med. Biol. 61 1572–95
- Yu Z, Leng S, Kappler S, Hahn K, Li Z, Halaweish A F, Henning A and McCollough C H 2016a Noise performance of low-dose CT: comparison between an energy integrating detector and a photon counting detector using a whole-body research photon counting CT scanner J. Med. Imaging 3 043503
- Yu Z, Leng S, Li Z and McCollough C H 2016b Spectral prior image constrained compressed sensing (spectral PICCS) for photon-counting computed tomography Phys. Med. Biol. 61 6707–32
- Yveborg M, Danielsson M and Bornefalk H 2015a Theoretical comparison of a dual energy system and photon counting silicon detector used for material quantification in spectral CT *IEEE Trans. Med. Imaging* 34 796–806
- Yveborg M, Persson M and Bornefalk H 2015b Optimal frequency-based weighting for spectral x-ray projection imaging IEEE Trans. Med. Imaging 34 779–87
- Yveborg M, Persson M, Crafoord J, Danielsson M and Bornefalk H 2013 Eliminated risk of iodine contrast cancellation with multibin spectral CT Phys. Med. Biol. 58 N201
- Zeng D *et al* 2020 Full-spectrum-knowledge-aware tensor model for energy-resolved CT iterative reconstruction *IEEE Trans. Med. Imaging* **39** 2831–43
- Zhang R, Cruz-Bastida J P, Gomez-Cardona D, Hayes J W, Li K and Chen G H 2018 Quantitative accuracy of CT numbers: theoretical analyses and experimental studies *Med. Phys.* 45 4519–28
- Zhang Y, Mou X, Wang G and Yu H 2017 Tensor-based dictionary learning for spectral CT reconstruction *IEEE Trans. Med. Imaging* 36 142–54
- Zhang Y, Xi Y, Yang Q, Cong W, Zhou J and Wang G 2016 Spectral CT reconstruction with image sparsity and spectral mean *IEEE Trans. Comput. Imaging* 2 510–23
- Zheng Y, Yveborg M, Grönberg F, Xu C, Su Q, Danielsson M and Persson M 2020 Robustness of optimal energy thresholds in photon-counting spectral CT *Nucl. Instrum. Methods* A **953** 163132
- Zhou W, Michalak G, Weaver J, Ferrero A, Gong H, Fetterly K A, McCollough C H and Leng S 2019 Determination of iodine detectability in different types of multiple-energy images for a photon-counting detector computed tomography system J. Med. Imaging 6 043501
- Zimmerman K C and Schmidt T G 2015 Experimental comparison of empirical material decomposition methods for spectral CT *Phys. Med. Biol.* **60** 3175–91