# Neural Network PET Reconstruction using Scattered Data in Energy-dependent Sinograms

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- Keywords: Positron Emission Tomography, Medical Imaging, Neural Network, Deep Learning, Scattered Coincidences, Direct Reconstruction.
- PET image reconstruction largely relies on pre-reconstruction data correction, which may add noise and Abstract remove information. This loss is particularly notable when correcting for scattered coincidences, which are useful for image reconstruction, though algorithmic scatter reconstructions require a detector energy resolution that exceeds the current state-of-the-art. Preliminary research has demonstrated the feasibility of using convolutional neural networks (CNNs) to reconstruct images directly from sinogram data. We have extended this approach to reconstruct images from data containing scattered coincidences. Monte Carlo simulations were performed to simulate PET data from digital phantoms. Data were modeled using 15% FWHM energy resolution detectors. Energy-dependent sinograms (EDSs), containing true and scattered coincidences, were constructed from the data. After data augmentation, 210,000 sinograms were obtained. A CNN was trained on the EDS-activity pairs for image reconstruction. A second network was trained on sinograms containing only photopeak coincidences. Images were also reconstructed using FBP, and MLEM approaches. The EDS trained network outperformed the photopeak trained network, with a higher mean structural similarity index ( $0.69 \pm .05$  vs.  $0.63 \pm .05$ ) and lower average mean square error ( $0.16 \pm .04$  vs. 0.20 $\pm$  .04). Our work demonstrates that CNNs have the potential to extract useful information from scattered coincidences, even for data containing significant energy uncertainties.

# **1** INTRODUCTION

Typical positron emission tomography (PET) image reconstruction techniques make corrections to the raw data prior to image reconstruction. These include corrections for dead time, attenuation, random and scattered coincidences, and normalization. Such corrections are imperfect. As such, they introduce noise and can remove valuable data (Cherry, 2012; Bai & Asma, 2016). The former is evident from the decreased noise equivalent counting rate (NECR) that results from scatter and random correction (Cherry, 2012, p. 340). Additionally, any alteration of the data destroys its Poisson nature (Bai & Asma, 2016, p. 266).

Scattered coincidences, in particular, contain information that is beneficial for image reconstruction. Such information may increase sensitivity due to a lower energy window threshold (Conti et al., 2012), and when used in image reconstruction, has been shown to improve contrast recovery and decrease noise (Sun & Pistorius, 2013a, 2013b). Furthermore, scattered coincidences contain information about the electron density of the scattering medium, thereby allowing attenuation maps to be estimated from scatter data (Berker et al., 2014; Brusaferri et al., 2020; Sun et al., 2015).

To date, scatter reconstruction studies have primarily focused on iterative approaches. These take advantage of the physics of Compton scattering to constrain annihilation positions to within areas or volumes in a 2-D or 3-D image space. Both time-offlight (TOF) (Conti et al., 2012) and non-TOF (Conti et al., 2012; Sun, 2016; Sun et al., 2015; Sun & Pistorius, 2013a, 2013b) methods have been implemented. Though most approaches have focused on coincidences where only one photon is scattered, it has also been demonstrated that even when both photons are scattered, this is sufficient to constrain the annihilation position (Sun, 2016).

In the absence of high-resolution TOF information, detector energy resolution must be

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sufficient to extract useful spatial information (Conti et al., 2012). Given current detector energy resolutions, coincidence timing resolution (CTR) values must be significantly better than 58-80ps, which is challenging to achieve (Meikle et al., 2021, p. 22).

## 1.1 Deep Learning in Image Reconstruction

Deep-learning (DL) networks have been applied to PET imaging. To date, most published studies focus on networks working entirely in image-space, taking as an input an image, and producing another image as output. A common application is generating attenuation-corrected (AC) PET images from nonattenuation corrected (NAC) PET images (Dong et al., 2019, 2020; Shiri et al., 2019). These approaches do not make use of scatter. Another PET image-space application is the so-called "super-resolution" problem, wherein a network attempts to derive a highresolution image from a low-resolution one. Song et al. (Song et al., 2020) provide a particularly thorough example of this technique. Lastly, CNNs have been employed to denoise images (Tian et al., 2019). A comprehensive list of PET image-space studies can be found in (Lee, 2021).

There is an alternative approach. So-called *direct* reconstruction approaches perform a domain transform to reconstruct images directly from binned data. These typically employ convolutional neural networks (CNNs). Compared to algorithmic or combined algorithmic/DL approaches, certain advantages are present. First, direct reconstruction is generally simpler. Second, DL approaches offer a possible solution to the ill-posedness of the PET inverse problem (Bai & Asma, 2016, p. 269). This solution takes the form of a non-linear filter that approximates regularization or image smoothing and is learned directly from the data. This may be thought of as similar to maximum a-posteriori (MAP) estimation where, from among images of similar likelihood values, one is chosen that is most probable given the prior (Bai & Asma, 2016, p. 270). As DL approaches are data-driven, the prior may be conceptualized as being learned implicitly from thousands of training examples.

Another probable advantage of direct reconstruction is the implicit incorporation of pointspread functions (PSFs). For iterative reconstruction methods, incorporating PSFs has been shown to improve image quality (Tong et al., 2010). A DL network may implicitly learn a spatially variant PSF from the data, thereby improving image quality. In earlier work on direct reconstruction (Häggström et al., 2019; Liu et al., 2019; Zhu et al., 2018), scatter was not employed. In the present context, the primary advantage of using direct reconstruction is—as we demonstrate—the ability of CNNs to make use of low-energy resolution scatter data to improve image quality.

# 2 METHODS

#### 2.1 Simulation Parameters

Twelve XCAT digital phantoms (Segars et al., 2010) were obtained, ranging in sex, weight, race and age. Each phantom was modified ten times by varying the activity uptake ratios, organ scaling and rotation, yielding a total of **120** phantoms. From each phantom, 250 transverse slices were obtained, spaced every 3 mm. Thus, 30,000 2-D activity distributions (images) were generated. Slice dimensions were 40 x 40 cm (71 x 71 pixels).

Monte Carlo simulations were performed on the XCAT phantoms. The Geant4 Application for Tomographic Emission (GATE) (Jan et al., 2004) was used to simulate positron emission and annihilation, photon propagation, and detection of coincidences. A three-dimensional cylindrical array of detectors was created around the phantoms. The dimensions of the crystal detectors were 3 mm x 3 mm x 20 mm. The height and radius of the cylinder were set to 750 mm and 150 mm, respectively, and the coincidence window was set to 10 ns.

The GATE source code was modified so that annihilation and scattered photon propagation were constrained to within the transverse polar plane. As this study focused on 2-D image reconstruction, doing so reduced the number of photons that needed to be simulated and thus reduced computation time. Although the simulated acquisition was performed in 3D mode, constraining the photons had the effect of simulating 2D acquisition.

The detector energy resolution was set to 15% FWHM in order to represent realistic PET photon detectors.

#### 2.2 Data Acquisition

Data were recorded in list mode. Approximately  $10^6$  counts were recorded per slice. Scatter fractions ranged from 30% to 50%, depending upon the size of the slice (torso, head, etc.). Three bins were established according to photon energy: 456 - 506 keV (bin 1), 478 - 528 keV (bin 2), and 486 - 536 keV (bin 3). Where,

for a single coincidence, both photon energies fell into bin 1, the coincidence was assigned to bin 1. Where one photon energy fell into bin 1, and the other fell into bin 2, the coincidence was assigned to bin 2. Similarly, where one photon energy fell into bin 1, and the other fell into bin 3, the coincidence was assigned to bin 3. Coincidences that could be placed into two bins, due to the overlapping energy ranges, were placed into both. Therefore, the binning of the data was accomplished via a sliding window.

Coincidences in bin 1, centered about 511 keV, were more likely true coincidences, relative to those in bins 2 and 3. Coincidences in bin 2 were likely due to a singly scattered coincidence. The likely scattered photons in this bin had energies centered about 503 keV, which corresponds to a 10-degree Compton scatter. Bin 3 had a central energy of 481 keV, which corresponds to a 20-degree Compton scatter. The binning of the data therefore structures it approximately according to scattering angle, binning coincidences mainly where only a single photon is which constitute most scattered. scattered coincidences (Conti et al., 2012). Due to the energy uncertainty of the detectors, however, some coincidences may have been improperly binned.

#### 2.3 Sinogram Construction

Two categories of sinograms were constructed from the binned data: *energy-dependent sinograms* (EDSs) and *photopeak sinograms*. Photopeak sinograms were created from bin 1 data only. To construct the EDSs, individual sinograms were created from the data in each bin and then combined into a single 3-D array with dimensions s,  $\phi$  and bin number, where s and  $\phi$ are the polar coordinates of a 2-D sinogram.

An attenuation correction that assumed that the simulated phantoms were composed entirely of water was performed for both types of sinograms. The average total distance travelled in matter for the photons in each scattered coincidence was calculated for all possible singly scattered travel paths. Only the section of each path which intersected with the volume of the phantom counted towards the average. This average distance, together with the Compton cross-section for water, was used to calculate the average attenuation coefficient for each scattered coincidence. Future studies will take an attenuation map as an additional input to the network to account for variations in electron density.

As doubly scattered photons were confined to the transaxial plane by the alteration to the GATE source code, an *attenuated* coincidence (that is, one in which more than one scatter event occurred) could still be

binned, so long as the scattered photon energy was not below 456 keV (the threshold for bin 3). While the coincidence would then effectively be counted twice, we relied on the network to learn corrections for this. However, it would also be possible to employ a more sophisticated energy-dependent attenuation correction factor, which depends on a restricted Compton cross-section, to deal with this complication (Sun, 2016, p. 58).

No dead time, random, arc, or normalization corrections were performed; we relied on the network to learn these.

Data augmentation was performed to efficiently increase the number of training examples. Each 2-D activity image was randomly flipped and rotated, with the corresponding sinograms modified appropriately. This was done six times for every image-sinogram pair, yielding a total of 210,000 image-sinogram pairs.

## 2.4 Network and Training

A detailed representation of the network we employed is depicted in Fig. 1. Similar to (Häggström et al., 2019), we implemented an encoder-decoder architecture, but as in (Whiteley et al., 2020) we used fully connected (dense) layers in the center of the network. These ensured that each pixel in the image space had as its receptive field the entire input sinogram, which was desirable due to the presence of scatter in the data.

Two different networks (which employed the same network architecture) were trained on the two different types of sinograms and are referred to as the *EDS network* and the *photopeak network*. Therefore, the dimensions of the input data—and thus also the dimensions of the first set of filters—varied depending upon the sinogram type. EDSs were of size 101 x 180 x 3 (the last digit comes from the three energy bins), whereas photopeak sinograms were of size 101 x 180 x 1.

The encoding portion of the network began with a convolutional layer followed by a ReLu activation. Next, a max-pooling layer contracted the data. This was followed by two more convolutional + ReLu layers. The data were then reshaped (flattened), passed through two dense layers, and then was reshaped into an 83 x 83 array. Lastly, the decoding portion of the network was composed of four convolutional + ReLu layers and one convolutional + softmax layer. The output was an image of size 71 x 71 pixels. Due to the final layer of the network being a softmax layer, each 2D activity image in the training and test sets was normalized so that the total activity per slice equaled 1.

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Figure 1: The network takes either energy-dependent sinograms (EDSs) or photopeak sinograms as input. The photopeak sinograms are 2-D arrays composed of 101 x 108 elements. By contrast, the EDSs have a  $3^{rd}$  dimension which contains the scattered coincidences, arranged according to the energy of the scattered component. Thus, the filter size of the first layer is different depending upon the input type. Multiple convolutional layers, as well as dense layers in the middle of the network, ensure that the receptive field for every pixel in the output image comprises the entire input sinogram. This guarantees that the full scatter data is utilized.

For each 2-D activity image, two sinograms were created: one energy-dependent, and one photopeak. From the 210,000 image-sinogram pairs, 180,000 photopeak sinograms and their corresponding activity distributions were used to train the photopeak network. The same number of energy-dependent sinograms (with the same 2-D activity distributions) were used to train the EDS network. In each case, this left 30,000 sinogram-image pairs for use as a validation set. As the validation sets were not used to tune network parameters, they also functioned as test sets.

Each network was trained for 150 epochs. During each epoch, the loss function of the training set and validation set were computed (Fig 2). Three loss functions were investigated for network training: Kullback-Leibler (KL), mean squared error (MSE), and Poisson. KL was chosen due to the superiority of images generated; these had less noise than those generated by a network trained with MSE and had higher contrast.

Network weights were optimized with an Adam optimizer using a learning rate equal to 10<sup>-5</sup>. Training was implemented within Python using the TensorFlow library.

#### 2.4.1 Transfer Learning

Monte Carlo simulations limit the practical size of training sets. We, therefore, investigated transfer learning as a possible method to train the network more thoroughly.

We began by training a network with sinograms calculated analytically from the activity distributions and subsequently corrupted by Poisson noise. The first 8 layers of the model were then frozen, and the remainder trained on Monte Carlo-derived data sets. However, this network underperformed relative to networks trained only on Monte Carlo data sets, even though the latter had relatively few training examples. The authors hypothesize that, as the analytic sinograms did not include energy-dependent components, initial network layers did not learn to take full advantage of the sinogram data and discarded the scattered components. If transfer



Figure 2: Training and validation losses for the EDS and photopeak networks.

learning were to be performed in future efforts, the 3rd dimension of the analytically computed sinograms should represent the energy-dependent scatter component more accurately.

## **3 RESULTS**

### 3.1 Network Performance

Once training of the EDS and photopeak networks was completed, these were tested with separate sets of 30,000 sinogram-image pairs. For comparison purposes, reconstructions of the photopeak sinograms were made using filtered back projection (FBP) and maximum likelihood expectation maximization (MLEM). The latter was terminated at 13 iterations. No additional scatter corrections were performed on the data before reconstruction. These, therefore, are illustrative and should not be taken as examples of the best possible analytic or algorithmic reconstructions. Rather, the most relevant comparison is between the EDS network and the photopeak network. Sample image reconstructions using all four methods may be found in Fig. 3.



Figure 3: Images for each reconstruction method are shown; note that the cold artifacts in images reconstructed from photopeak sinograms are absent or attenuated in the images reconstructed from EDSs.

The average mean squared error (MSE) and structural similarity (SSIM) (Zhou Wang et al., 2004) were determined from the validation set for each reconstruction method, relative to the ground truth images. Contrast values were also determined for all four methods relative to the EDS-network. This was done by averaging the intensity of a region of interest of a hot spot feature ( $I_h$ ), as well as the background area ( $I_b$ ). The contrast for a single image is then given by:  $(I-I_b)/I_b$ .

The average contrast value was computed for a 10-image subset for each reconstruction method. Then the relative contrast values for the photopeak network, MLEM and FBP were determined relative to the EDS network. Figure 4 shows the image quality metrics for each reconstruction method.

The computation times for each reconstruction method were also determined. The CNN reconstruction times were approximately 3 and 92 times faster than FBP and MLEM methods, respectively.



Figure 4: Evaluation metrics are shown for the four reconstruction methods employed. Of particular interest are the metrics for the two CNNs. The network trained on energy-dependent sinograms (EDSs) outperforms the network trained on photopeak sinograms, as evidenced by the structural similarity index metric (SSIM) and mean square error (MSE). However, for both SSIM and MSE, the difference between the metrics is less than the standard deviation of the metric values for images generated by each network; this is apparent from the error bars in the figure.

#### 3.2 Discussion

For each image metric, the mean performance of the EDS network exceeded that of the photopeak network, though the difference was within the uncertainty for all metrics. However, the uncertainties are given by the standard deviation of the metric values, and as each metric value depends on the particular image considered, comparing differences in the mean metric values with their uncertainties has limited utility. For example, it may be possible that every EDS network image has a higher SSIM than the equivalent photopeak network image, and yet the uncertainties may overlap. A more thorough analysis, to be undertaken in future work, will examine this problem in greater detail.

Differences in the quality of the reconstructed images can also be observed by visual inspection. Figure 3 depicts reconstructions of four different 2-D activity distributions using the four reconstruction methods. The photopeak network generates distinct cold artifacts that are less visible or absent in the EDS network images. Noise within constant-activity areas is also higher in the photopeak network generated images. Lastly, the contrast between proximal areas with different activity levels appears higher in the EDS network generated images. Both CNNreconstructed images share broad agreement with images reconstructed via FBP and MLEM.

Learning curves for the two CNNs are displayed in Fig. 3. The training and validation loss of the photopeak network shows slight overfitting, whereas the energy-dependent network does not. This may be due to the training set for the EDS network containing three times the amount of data as the training set for the photopeak network, whereas the number of weights—except for the first layer—was equal. This indicates that the EDS network was appropriately sized, given the size of its training set.

# 4 APPLICATIONS AND IMPROVEMENTS

#### 4.1 **3D and Total-body PET**

Improved image metrics demonstrate that including the information contained in the scatter has the potential to increase image quality, even with 15% FWHM energy resolution detectors. Given this, image reconstruction with data containing higher scatter fractions, such as occurs with 3-D acquisition or with large patients, is likely to benefit most. However, 3-D PET rebinning approaches, such as the Fourier rebinning algorithm (Defrise et al., 1997), typically only consider true coincidences. To make full use of scatter data, either fully 3-D reconstruction must be pursued, or else rebinning algorithms must be which accurately rebin scattered developed coincidences. The former approach will require a much larger CNN with many more weights, which will require many more training examples. The computational requirements for such an approach are likely to be prohibitive. The authors believe the latter approach to be more reasonable.

The highest scatter fractions occur with totalbody PET (TB-PET). Due to the increased sensitivity possible with TB-PET, reducing injected activity by a factor of 20 or more is attainable. However, if a CT scan must be performed for the purpose of attenuation correction, this offsets the benefit. Attenuation corrected emission maps may be constructed with deep learning methods, even in the absence of CT, though this may be difficult using the sparse data from low-dose scans (Meikle et al., 2021, p. 25). In such cases, using scattered data for attenuation correction and/or activity estimation looks promising, especially as scatter fractions—and thus the information contained in the scatter—are increased for total-body PET.

## 4.2 Additional Data Types

If attenuation maps are available, these have the potential to increase the utility of scatter imaging. Photons are more likely to be scattered in volumes with high electron density. Therefore, attenuation maps contain statistical information about likely scattering locations. The use of attenuation maps as prior information for determining scatter locations may therefore increase the available spatial information also increases the spatial information that can be gleaned from scattered coincidences, as demonstrated by Conti et al. (Conti et al., 2012). Therefore, networks that use attenuation or TOF data, together with scatter data, seem promising.

## 4.3 Future Work

The current study focused on a single network. Future work will explore various network configurations, including CycleGan, which often outperforms Unettype architectures. In this network type, a cycleconsistent loss is added, which penalizes projection functions for not being injective (Wang et al., 2020). Different binning schemes may also be compared. Lastly, verification of the reconstruction method using physical phantoms is necessary to guarantee that the improvements seen in simulations translate to real-world applications.

# 5 CONCLUSION

Two convolutional neural networks sharing the same network architecture were trained and tested with two different sets of data: one with coincidences where both photons fell within a  $511(\pm 25)$  keV photopeak window, and one which also included coincidences where one of the photons had a lower detected energy. The network trained with the energy-dependent scatter sinograms was observed to have a lower mean-square-error and larger structural similarity index than the network trained with only

the photopeak sinograms. Images generated by the photopeak network also contained more severe artifacts.

These results suggest that including scattered coincidences in the data has the potential to increase image quality. The authors hypothesize that by utilizing coincidences outside the photopeak energy bin, the patient dose may be lowered while maintaining the same image quality, thus improving patient care.

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