# Unsupervised PET Reconstruction from a **Bayesian Perspective**

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Abstract—Positron emission tomography (PET) reconstruction has become an ill-posed inverse problem due to low-count projection data, and a robust algorithm is urgently required to improve imaging quality. Recently, the deep image prior (DIP) has drawn much attention and has been successfully applied in several image restoration tasks, such as denoising and inpainting, since it does not need any labels (reference image). However, overfitting is a vital defect of this framework. Hence, many methods have been proposed to mitigate this problem, and DeepRED is a typical representation that combines DIP and regularization by denoising (RED). In this article, we leverage DeepRED from a Bayesian perspective to reconstruct PET images from a single corrupted sinogram without any supervised or auxiliary information. In contrast to the conventional denoisers customarily used in RED, a DnCNN-like denoiser, which can add an adaptive constraint to DIP and facilitate the computation of derivation, is employed. Moreover, to further enhance the regularization, Gaussian noise is injected into the gradient updates, deriving a Markov chain Monte Carlo (MCMC) sampler. Experimental studies on brain and whole-body datasets demonstrate that our proposed method can achieve better performance in terms of qualitative and quantitative results compared to several classic and state-of-the-art methods.

Index Terms—PET image reconstruction, unsupervised learning, deep image prior, regularization by denoising, **Bayesian inference** 

#### I. INTRODUCTION

**P**OSITRON emission tomography (PET) is an important imaging modality in the fail of imaging modality in the field of nuclear medicine that can reflect the molecular-level activities of various organs and tissues in vivo by injecting specific radioactive tracers. Since the amount of injected radiotracer in current protocols will increase the potential risks of radiation exposure for the human body, reducing the dose is of clinical significance. However, simply reducing the injected dose can result in low-count PET projection data (sinogram), which leads to undesired noise and artifacts in the reconstructed images, further negatively affecting the subsequent clinical diagnosis. How to reconstruct high-quality images has always been a popular topic, and

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extensive efforts have been devoted over the past decades. These algorithms can be roughly classified into three groups: 1) analytical method; 2) image postprocessing; and 3) modelbased iterative reconstruction (MBIR).

The most famous analytical method is filtered backprojection (FBP) [1], which applies a convolution filter to remove blurring and relies on the central slice theorem. However, FBP is famously susceptible to measurement noise.

A simple solution for this problem is the postprocessing method. This kind of method directly processes the reconstructed PET images with image restoration algorithms, such as nonlocal mean (NLM) [2], BM3D [3] and multiscale transform [4]. Although these methods have no requirement to access the raw data and have a relatively fast processing speed, the latent information in the sinogram is ignored, and the results have potential risks of being oversmoothed.

To tackle the disadvantages of analytical and postprocessing methods, MBIR methods have become mainstream in the past two decades. Maximum-likelihood expectation maximization (ML-EM) [5] is a classic iterative method that maximizes the log-likelihood of projection data. However, such an iterative reconstruction method without regularization formed by specific prior knowledge about images or sinograms may increase the noise after a certain number of iterations. As a result, integrating MBIR with specific regularization is desirable to suppress noise efficiently. Numerous different types of regularization have been proposed. Some typical methods include total variation [6]– [9], edge-preserving filtering [10], [11], dictionary learning [12], kernel methods [13] and anatomical priors [14]– [16]. Despite promising results achieved by these methods, heavy computational cost and laborious parameter adjustment severely hinder their clinical applications. Moreover, since the construct of the regularization term is based on specific prior knowledge, it is difficult to find a universal regularization term, which can be easily adapted to different cases.

Over the past decade, deep neural networks have dramatically improved the state of the art in different fields of computer vision, e.g., object detection [17], image recognition [18], image segmentation [19], and even medical imaging. For instance, convolutional neural networks (CNNs) and generative adversarial networks (GANs) have been applied in CT and MRI reconstruction [20]-[25], which have shown competitive performance to traditional methods. Recent studies on PET imaging reconstruction based on deep learning can be roughly divided into three groups. 1) Directly learn the mapping from

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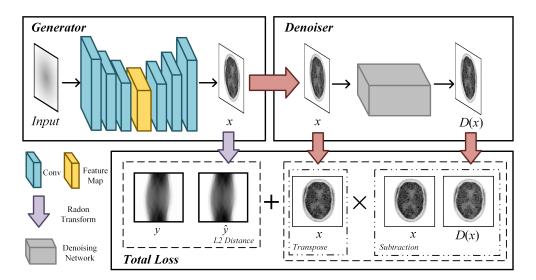


Fig. 1. Overview of our proposed model. The detailed generator and denoiser are shown in Fig. 2 and Fig. 3, respectively.

the sinogram to the corresponding PET image. For example, DeepPET [26] was proposed to employ the sinogram as input and outputs PET images directly based on encoder-decoder network architecture. Furthermore, Hu et al. [27] used an improved Wasserstein generative adversarial network aided by perceptual loss to reconstruct PET images from sinograms, which alleviated the problem of oversmoothness and detail loss that usually appeared in plain networks. 2) Learn the mapping from low-dose or low-count images to their corresponding full-dose or full-count images [28]-[31], which can be treated as image restoration or enhancement tasks. 3) Iterative reconstruction networks, which combine a neural network with MBIR by unrolling the iterative algorithm or substituting the regularization term with a learning-based prior. Typically, inspired by the kernel method, Gong et al. [32] embedded a trained network into a constrained optimization problem and solved it using the alternating direction method of multipliers (ADMM). Mehranian et al. [33] proposed a forwardbackward splitting algorithm to integrate deep learning into the framework of MAP reconstruction, which is composed of regularization, expectation-maximum and weighted fusion. BCD-Net [34] is another iterative neural network constructed by unfolding a block coordinate descent MBIR algorithm, showing significant improvement in extreme low count PET reconstruction [35]. All these abovementioned methods are supervised and require large amounts of samples to train a proper network. However, it is not always feasible in clinical practice, especially for pilot clinical trials, to obtain such an amount of training samples, especially for paired low-dose or low-count data and their corresponding normal-dose or normal-count data.

To overcome this obstacle, recently, a trend of incorporating the well-known unsupervised learning model, called deep image prior (DIP) [36], has come to the fore. DIP demonstrates that CNNs without pretraining can capture lowlevel statistics intrinsically. CNNs' ability to solve inverse problems in the field of image processing, such as denoising and super resolution, is due to the network architecture itself instead of learning. Gong et al. [37] integrated DIP into PET reconstruction while optimizing a Poisson distributionbased log-likelihood function. Yokota et al. [38] incorporated DIP with non-negative matrix factorization for dynamic PET reconstruction. Under the framework of DIP, Cui et al. employed the patient's CT/MR images as the auxiliary input of conditional GAN to improve the performance of the original GAN [39]. Nonetheless, without early stopping or an appropriate stop criterion, DIP is apt to overfit. Designing a proper stop criterion is the key point to ease this problem. Mataev et al. [40] proposed the DeepRED framework, merging the concept of regularization by denoising (RED) [41] into DIP by introducing an explicit prior. Cheng et al. [42] provided a novel Bayesian view of DIP by conducting posterior sampling based on stochastic gradient Langevin dynamics (SGLD) [43]. DeepRED and SGLD were applied to PET reconstruction in [44] and [45], respectively. However, DeepRED usually takes BM3D or NLM as its denoiser, which suffers from two main disadvantages. First, traditional methods have limited ability in representation learning as they can only capture specific features such as self-similarity. Second, for most traditional denoisers, the calculation of derivatives for the denoising function would be a daunting task [40], which may jeopardize the integration of DeepRED with neural networks.

In this paper, we propose a PET reconstruction model combining the DeepRED framework with approximate Bayesian inference. Our model reconstructs PET images from a single sinogram without any anatomic prior, e.g., corresponding CT or MR images. In particular, the traditional denoiser commonly used in DeepRED is substituted with a learnable denoiser. As a result, the proposed model includes two CNNs: one is a generator that takes arbitrary input and outputs the reconstructed image, and the other plays the role of denoiser in DeepRED. Additionally, the denoiser learns to impose a task-based constraint on DIP and obtains adaptivity that does not exist in traditional denoisers. On the other hand, our denoiser is differentiable, which makes backpropagation possible to optimize the loss function. We also considered

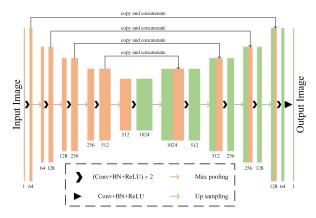


Fig. 2. Architecture of the generator network in our proposed model.

SGLD. Specifically, Gaussian noise is injected into the gradient updates, which derives a Markov chain Monte Carlo (MCMC) sampler from stochastic gradient descent (SGD), further alleviating overfitting from a Bayesian perspective. The main contributions of this work are summarized as follows: 1) an unsupervised end-to-end PET image reconstruction model under the framework of DeepRED, which does not need paired samples, is proposed. 2) A learnable denoiser embedded in DeepRED is used, which adds force to DIP adaptively and facilitates the calculation of the gradient. 3) DeepRED is combined with SGLD to further improve the regularization performance.

The remainder of this paper is organized as follows. In the next section, we introduce the preliminary knowledge of DIP and DeepRED. Sections III elaborate on the proposed method. In Section IV, brain data and whole-body data experiments are conducted to evaluate our method. Finally, a discussion and conclusions are given in Section V.

## **II. PRELIMINARIES**

#### A. Deep Image Prior

DIP is a state-of-the-art unsupervised framework that can restore corrupted images without training data. DIP suggests that the unknown image to be reconstructed x can be represented using a carefully selected neural network g as  $x = g(\theta \mid z)$ , where  $\theta$  and z denote the network parameter set and a random input, respectively. Consider an inverse problem y = Px + n, where y is the measurement, P represents the sampling matrix dependent on the specific task, and n denotes additive noise, which is usually assumed to be Gaussian noise. Under the DIP framework, x can be obtained first by optimizing:

$$\theta = \arg\min_{\theta} \|P \cdot g(\theta \mid z) - y\|.$$
<sup>(1)</sup>

Once  $\hat{\theta}$  is obtained,  $\hat{x} = g(\hat{\theta} \mid z)$  is obtained.

# B. DeepRED

DeepRED was proposed to mitigate overfitting by combining DIP and regularization by denoising. The objective function of DeepRED becomes:

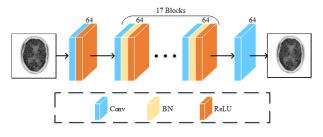


Fig. 3. Architecture of the denoiser network in our proposed model.

$$\min_{\substack{x,\theta \\ x,\theta}} \quad \|P \cdot g(\theta \mid z) - y\| + \frac{\lambda}{2} x^T (x - f(x)),$$
s.t.  $x = g(\theta \mid z).$ 
(2)

where  $f(\cdot)$  represents a denoiser such as NLM and BM3D, and  $\frac{\lambda}{2}x^T(x - f(x))$  is the so-called RED, which turns a denoiser into regularization. This framework relies on an image-adaptive Laplacian that poses cogent prior acting as a regularization term in various inverse problems [46]– [48]. This kind of regularization exhibits a better-managed optimization due to homogeneity and strong passivity on denoisers.

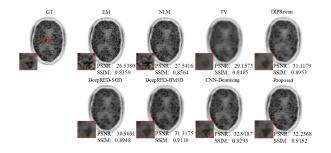


Fig. 4. Reconstructed axial brain PET image using different methods with a count level of  $1\times 10^8$  from one patient.

# III. Method

#### A. PET Reconstruction Model

The measured PET projection data  $y \in \mathbb{R}^M$  are known as sinograms, where each element  $y_i$  is the number of photons accumulated in the *i*-th line of response (LOR). For M lines of response and N pixels in original image  $x \in \mathbb{R}^N$ , a typical PET imaging model is formulated as:

$$\bar{y} = Px + r + s, \tag{3}$$

where  $\bar{y} \in \mathbb{R}^M$  is the expectation of y,  $P \in \mathbb{R}^{M \times N}$  denotes the system matrix,  $r \in \mathbb{R}^M$  and  $s \in \mathbb{R}^M$  represent the random coincidences and scattered coincidences, respectively. Assuming that  $y_i$  follows a distribution of  $p(y_i \mid x)$ , the loglikelihood for y can be written as

$$L(x \mid y) = \sum_{i=1}^{M} \log p(y_i \mid x).$$
 (4)

Under ideal conditions,  $y_i$  is modeled as an independent Poisson variable. However, because of several acquisition circumstances, such as detector systems, electronic circumstances

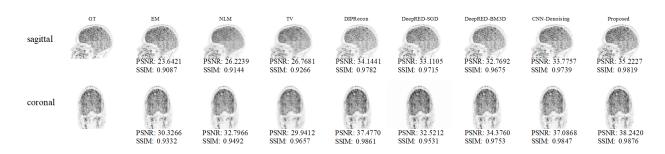


Fig. 5. Reconstructed sagittal and coronal brain PET images using different methods with a count level of 1 × 10<sup>8</sup> from another patient.

and various data preprocessing, the actual distribution often deviates from a Poisson distribution [49], [50]. Since several additive sources of error tend to form a Gaussian distribution, a Gaussian noise model is suitable for PET data. In fact, [51] shows that both Poisson and Gaussian models can be leveraged for PET reconstruction. In this article, to introduce the DeepRED framework, we adopt a Gaussian-based loglikelihood function as:

$$p(y_i \mid x) = \frac{1}{\sqrt{2\pi\sigma}} exp\{-\frac{(y_i - \bar{y}_i)^2}{2\sigma^2}\}.$$
 (5)

Assuming that the variance of each LOR is equal, we have

$$L(x \mid y) = -\|y - \bar{y}\|_{2}^{2} + C,$$
(6)

where C is a constant that can be ignored.

#### B. The Proposed Reconstruction Model

In the proposed method, the reconstructed image x is represented by:

$$x = G(\theta_G \mid z), \tag{7}$$

where G denotes a generative network. Specifically, in this work, U-Net is employed as the backbone of G.  $\theta_G$  represents the parameter set of G, and z is the input of G. In contrast to [37], in which other modality images, such as MRI or CT, are selected as z, we simply set  $z = P^T y$  to keep our method more robust. By introducing a negative log-likelihood as a fidelity term into DeepRED, the objective function is obtained as

$$\min_{x} \|y - Px - r - s\|_{2}^{2} + \frac{\lambda}{2} x^{T} (x - f(x)).$$
(8)

In the original DeepRED [40], which uses NLM or BM3D as a denoiser, the derivative of  $f(\cdot)$  with respect to x is difficult to calculate. To circumvent this obstacle, ADMM is employed. However, ADMM introduces additional parameters to be tuned. Furthermore, traditional denoisers have demonstrated limited performance compared to recently proposed deep learning-based models. To further improve the denoising performance, we replace traditional denoisers with a DnCNNlike network [52] and denote it as D. Then, (8) can be rewritten as

$$\min_{\theta_G, \theta_D} \|y - PG(\theta_G \mid z) - r - s\|_2^2 + \frac{\lambda}{2} G(\theta_G \mid z)^T (G(\theta_G \mid z) - D(\theta_D \mid G(\theta_G \mid z))),$$
(9)

where  $\theta_D$  represents the parameter set of D. Then, we have two networks D and G in our model and will train them in an end-to-end manner.

Although RED can mitigate overfitting to a certain degree, the optimization of (9) still leads to point estimation, which is apt to cause overfitting. To further relieve the overfitting and construct an overall regularization, we treat this problem from a Bayesian perspective. In Bayesian inference, a suitable prior distribution  $p(\theta_G)$  is added to the parameter set  $\theta_G$ , and the final estimation  $\hat{G}$  is formulated by marginalizing the posterior:  $\hat{G} = \int G(\theta_G \mid z)p(\theta_G \mid y)d\theta_G$ . A common solution to this posterior average is MCMC. However, MCMC converges more slowly than backpropagation while optimizing the network parameters. To handle this problem, SGLD was proposed for generating an MCMC sampler by injecting a proper amount of Gaussian noise in the updating step. SGLD formulates the updating equation as:

$$\theta_{t+1} = \theta_t + \frac{\epsilon}{2} (-\nabla_{\theta_G} obj(\theta_G) + \nabla_{\theta_G} p(\theta_G)) + \eta_t, \quad (10)$$

where  $\eta_t \sim \mathcal{N}(0, \epsilon)$  is the injected Gaussian noise,  $\epsilon$  is the step size, and  $obj(\cdot)$  denotes the objective function.  $\nabla_{\theta_G} obj(\theta_G)$ and  $\nabla_{\theta_G} p(\theta_G)$  represent the derivatives of  $obj(\theta_G)$  and  $p(\theta_G)$ , respectively, with respect to  $\theta_G$ . The first term on the righthand side of (10) is the standard SGD updating, where  $-\nabla_{\theta_G} obj(\theta_G)$  can be considered a likelihood function, and  $\nabla_{\theta_G} p(\theta_G)$  can be treated as a prior, such as the L1 or L2 penalty. Theoretically,  $\epsilon$  converges to zero. Since SGLD with diminishing step size converges weakly to a posterior distribution [53], a constant rather than diminishing step size is commonly used in practice [54]-[56]. In this work, a constant step size specific to different images and count levels is adopted. The randomness comes from two sources in (10): one is the injected Gaussian noise, and the other is the noise in the stochastic gradient. In the early phase, the stochastic gradient takes control over the updating equation, leading SGLD to behave like SGD. In the later phase, the injected Gaussian noise dominates, and the algorithm steps into the Langevin dynamics phase, allowing estimation over the posterior phase.

Fig. 1 shows the overview of our proposed model. y is the measured sinogram, which is the only data we have in our algorithm.  $P^T y$  is used as the initial input to the generative network G. x is the output of G.  $\hat{y}$  is the corresponding sinogram generated from x. On the other hand, x acts as the input to the denoiser D, and D(x) denotes the output of D. '×' denotes matrix multiplication. The final loss is composed

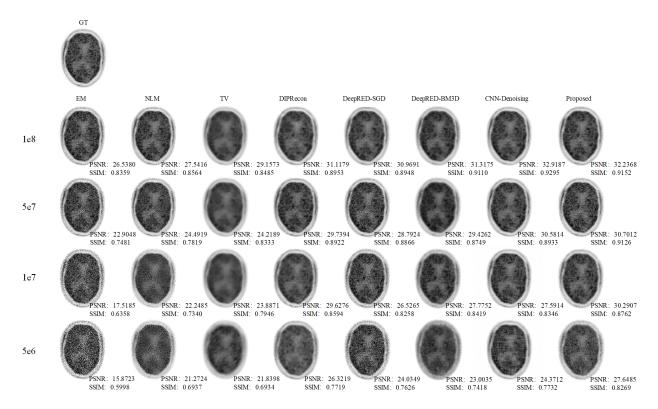


Fig. 6. Results with different methods with four count levels in the brain dataset study.

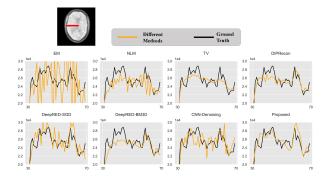


Fig. 7. Comparison of horizontal line profiles in brain PET images. The horizontal axis represents the pixel index, while the vertical axis represents the pixel intensity.

of  $x^T(x - D(x))$  and the Euclidean distance between y and  $\hat{y}$ . The factor  $\lambda$  controls the trade-off between two terms.

#### C. Network Architecture

Fig. 2 illustrates the architecture of the generative network G, which is derived from U-Net [19]. It contains four paired contracting and expansive blocks, and four skip connections are introduced between each pair. In the beginning, an image with a size of  $128 \times 128$  is entered to G, followed by repetitive blocks that are uniformly composed of a  $3 \times 3$  convolution layer, a batch normalization (BN) layer, a rectified linear unit (ReLU) layer and a  $2 \times 2$  max pooling layer with stride 2. Each step in the expansive phase consists of a bilinear interpolated result concatenated with the corresponding feature map from the contracting step. Eventually, a  $1 \times 1$  convolution

TABLE I PSNR and SSIM Values of Various Methods of Brain Data Under a Count Level of  $1 \times 10^8$ .

	Ax	ial	Sagi	ttal	Coronal		
Methods	PSNR	SSIM	PSNR	SSIM	PSNR	SSIM	
EM	26.5380	0.8359	23.6421	0.9087	30.3266	0.9332	
NLM	27.5416	0.8564	26.2239	0.9144	32.7966	0.9492	
TV	29.1573	0.8485	26.7681	0.9266	29.9412	0.9657	
DIPRecon	31.1179	0.8953	34.1441	0.9782	37.4770	0.9861	
DeepRED-SGD	30.9691	0.8948	33.1105	0.9715	32.5212	0.9531	
DeepRED-BM3D	31.3175	0.9110	32.7692	0.9675	34.3760	0.9753	
CNN-Denoising	32.9187	0.9295	33.7753	0.9739	37.0868	0.9847	
Proposed	32.2368	0.9152	35.2227	0.9819	38.2420	0.9876	

is executed to compress the 64-channel feature map to a 1-channel output. The differences between our generative network and U-Net are as follows: 1) bilinear interpolation is used instead of deconvolution upsampling to reduce the checkerboard artifact; and 2) the padding operation is used after each  $3 \times 3$  convolution to maintain the same size during propagation. Fig. 3 displays the architecture of our denoising network D, which employs DnCNN as the backbone. The only difference between our model and DnCNN is that we exclude residual connections

## IV. EXPERIMENTAL SETUP AND RESULTS

# A. Experimental Setup

The PET brain images used in this part were downloaded from The Cancer Imaging Archive (TCIA) [57]. Two patients' brain images were obtained using GE medical systems. One voxel size is  $1.92 \times 1.92 \times 3.27 mm^3$ , and the image size is

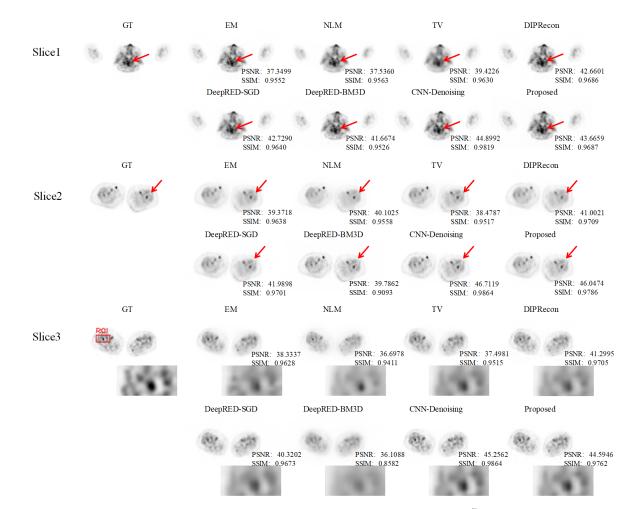
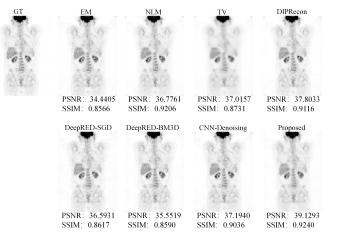


Fig. 8. Three axial slices of reconstructed whole-body PET images using different methods at a  $1 \times 10^7$  count level.

TABLE II PSNR and SSIM Values of Various Methods of Brain Data Under Different Count Levels.

Methods	$1 \times 10^8$		$5 \times 10^7$		$1 \times 10^7$		$5 \times 10^6$	
	PSNR	SSIM	PSNR	SSIM	PSNR	SSIM	PSNR	SSIM
EM	26.5380	0.8359	22.9048	0.7481	17.5185	0.6358	15.8723	0.5998
NLM	27.5416	0.8564	24.4919	0.7819	22.2485	0.7340	21.2724	0.6937
TV	29.1573	0.8485	24.2189	0.8333	23.8871	0.7946	21.8398	0.6934
DIPRecon	31.1179	0.8953	29.7394	0.8922	29.6276	0.8594	26.3219	0.7719
DeepRED-SGD	30.9691	0.8948	28.7924	0.8866	26.5265	0.8258	24.0349	0.7626
DeepRED-BM3D	31.3175	0.9110	29.4262	0.8749	27.7752	0.8419	23.0035	0.7418
CNN-Denoising	32.9187	0.9295	30.5814	0.8933	27.5914	0.8346	24.3712	0.7732
Proposed	32.2368	0.9152	30.7012	0.9126	30.2907	0.8762	27.6485	0.8269



 $128 \times 128 \times 47$ , the other voxel size is  $4.06 \times 4.06 \times 5.00 mm^3$ , and the image size is  $128 \times 128 \times 145$ . Each *FMISO* PET imaging session consisted of acquiring a low-dose CT transmission scan used for attenuation correction and a PET emission scan at approximately two hours after injection of  $^{18}FMISO$ . The system matrix *P* was modeled using Siddon's refined method to calculate the ray path integral [58]. The noise-free sinogram was generated by forward projecting the PET image. Each sinogram has a matrix size of 180 (radial bins)  $\times$  180 (azimuthal angles). Then, uniform events were added to the noise-free data as background, which accounts

Fig. 9. Reconstructed axial and coronal whole-body PET images from another patient using different methods at a  $1\times10^7$  count level.

for 10% of total true coincidences. Poisson noise was added by setting the total number of photons. In the brain dataset, four count levels were tested, including  $1 \times 10^8$ ,  $5 \times 10^7$ ,  $1 \times 10^7$  and  $5 \times 10^6$ . To further evaluate the performance of our proposed method, a whole-body dataset also downloaded from TCIA was tested. The whole-body data of ten patients

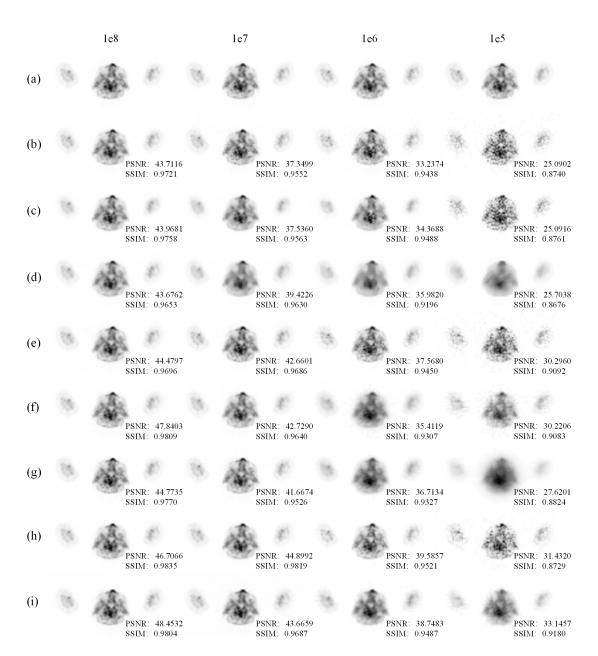


Fig. 10. Results with different methods with four count levels in the whole-body dataset study. (a) Ground Truth, (b) EM, (c) NLM, (d) TV, (e) DIPRecon, (f) DeepRED-BM3D, (g) DeepRED-SGD, (h) CNN-Denoising, (i) Proposed.

were acquired as approximately 300 slices with a size of  $128 \times 128$ . Before the examination, the patients underwent fasting for at least 6 hours, and the blood glucose of each patient was less than 11 mmol/L. Whole-body emission scans were acquired 60 minutes after the intravenous injection of 18F-FDG (4.44 MBq/kg, 0.12 mCi/kg), with patients in the supine position in the PET scanner. FDG doses and uptake times were 168.72-468.79 MBq (295.8±64.8 MBq) and 27-171 min (70.4±24.9 minutes), respectively. 18F-FDG with a radiochemical purity of 95% was provided. Patients were allowed to breathe normally during PET and CT acquisitions. Attenuation correction of PET images was performed using CT data with the hybrid segmentation method. We followed the same procedure as in a previous brain dataset study to generate low count sinograms. In the whole-body dataset,

count levels of  $1\times 10^8,\, 1\times 10^7,\, 1\times 10^6$  and  $1\times 10^5$  were tested.

To quantify the performance of different methods, the peak signal-to-noise ratio (PSNR) and similarity structure index (SSIM) were employed. The proposed model was implemented by the PyTorch Python library [59] on a PC (Intel Xeon E5-2620 CPU and a GeForce GTX 1080Ti GPU). The network was trained using the Adam [60] optimizer, and the learning rate was set to 0.0001. Six state-of-the-art methods were involved for comparison, including EM [5], TV [9], NLM [2], DIPRecon [37], DeepRED-BM3D [40] and CNN-Denoising [61]. EM is a commonly used reconstruction method deployed in most commercial scanners. TV is a classic regularization model usually employed in different inverse problems. NLM is an efficient image denoising method that

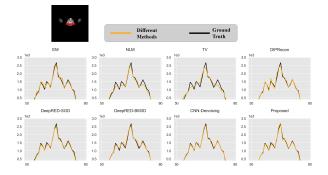


Fig. 11. Comparison of horizontal line profiles in whole-body PET images. The horizontal axis represents the pixel index, while the vertical axis represents the pixel intensity.

TABLE III PSNR and SSIM Values of Various Methods of Whole-body Data Under a Count Level of  $1 \times 10^7$ .

	Ax		Coronal		
Methods	PSNR	SSIM	PSNR	SSIM	
EM	38,3518	0.9606	34.4405	0.8566	
NLM	38.1121	0.9510	36.7761	0.9206	
TV	38.4664	0.9554	37.0157	0.8731	
DIPRecon	41.6539	0.9700	37.8033	0.9116	
DeepRED-SGD	41.6796	0.9671	36.5931	0.8617	
DeepRED-BM3D	39.1874	0.9067	35.5519	0.8590	
CNN-Denoising	45.6224	0.9849	37,1940	0.9036	
Proposed	44.7693	0.9745	39.1293	0.9240	

has been applied in many imaging modalities. DIPRecon is based on a Poisson-based unsupervised iterative reconstruction whose regularization term is formulated as DIP. DeepRED is an efficient iteration framework equipped with a selectable denoiser, and in our experiment, the well-known BM3D is employed as the denoiser dubbed DeepRED-BM3D. CNN denoising is an end-to-end denoising network based on supervised learning. Moreover, to validate the effectiveness of SGLD, we also compare the proposed model with SGD, named DeepRED-SGD. Particularly, when we trained CNN-Denoising, we randomly selected 180 images for the brain and 300 images for the whole-body.

#### B. Brain Data Results

Figs. 4 and 5 show the reconstructed brain PET images from different patients using various methods with  $1 \times 10^8$  count levels from axial, coronal and sagittal views, respectively. In

TABLE IV PSNR and SSIM of Various Methods of Whole-body Data Under Different Count Levels.

	$1 \times 10^8$		$1 \times 10^7$		$1 \times 10^{6}$		$1 \times 10^5$	
Methods	PSNR	SSIM	PSNR	SSIM	PSNR	SSIM	PSNR	SSIM
EM	43.7116	0.9721	37.3499	0.9552	33.2374	0.9438	25.0902	0.8740
NLM	43.9681	0.9758	37.5360	0.9563	34.3688	0.9488	25.0916	0.8761
TV	43.6762	0.9653	39.4226	0.9630	35.9820	0.9196	25.7038	0.8676
DIPRecon	44.4797	0.9696	42.6601	0.9686	37.5680	0.9450	30.2960	0.9092
DeepRED-SGD	47.8403	0.9809	42.7290	0.9640	35.4119	0.9307	30.2206	0.9083
DeepRED-BM3D	44.7735	0.9770	41.6674	0.9526	36.7134	0.9327	27.6201	0.8824
CNN-Denoising	46.7066	0.9835	44.8992	0.9819	39.5857	0.9521	31.4320	0.8729
Proposed	48.4532	0.9804	43.6659	0.9687	38.7483	0.9487	33.1475	0.9180

general, methods based on CNNs garner better results than conventional methods that blur the image or produce severe noise. Note that images reconstructed by DeepRED-SGD gain a negligible advantage over DeepRED-BM3D, probably because we chose a relatively better result for DeepRED-BM3D since it produces oscillatory curves, while we chose the result for DeepRED-SGD when it converged to a stable solution. It can also be observed that our proposed method outperforms all the other traditional and unsupervised methods in terms of detail recovery and noise suppression. The results with different count levels are shown in Fig. 6. Consistent with the case of the  $1 \times 10^8$  count level, CNN-based methods perform better than conventional methods, and our proposed method still achieves better results than other traditional and unsupervised methods. When the count level reaches  $1 \times 10^7$ , the performance of EM, DeepRED-SGD and DeepRED-BM3D drops significantly, while the other methods exhibit much less performance degradation. Our proposed method has visible improvement on DeepRED, which confirms the merit brought by SGLD. The quantitative results listed in TABLE I and TABLE II show coherent trends to the visual inspection that our method achieves the competitive scores in terms of both metrics to CNN-denoising better results than other methods. Pixel values along the red line in Fig. 7 of the ground truth versus the images reconstructed using different methods are plotted in Fig. 7. The proposed method yields a profile closer to the ground truth than the other methods. It can be observed that our method has a more robust performance against the decline of the count level. Although CNN-denoising also has close performance to our method, it belongs to supervised learning and requires large amounts of perfectly paired samples, which are almost practically impossible.

## C. Whole-body Data Results

Fig. 8 shows three typical whole-body slices from different body parts using different methods with count level of  $1 \times$  $10^7$ . For the first and second cases, red arrows indicate the regions that can be used to distinguish the performances of different methods. For the third case, a region of interest (ROI) marked by a red box is chosen to better visualize the improvement achieved by our method. Additionally, methods based on CNNs recover more details than conventional methods. Compared to DeepRED-BM3D, images reconstructed by DeepRED-SGD are closer to the ground truth, which agrees with the widely accepted fact that CNN-based denoisers generally bring better performance than some popular denoisers, such as BM3D. Notably, consisitent with the previous experiments on Brain Data, our method performs similar to CNN-Denoising, which belongs to supervised learning. Fig. 9 shows a reconstructed whole-body image from a coronal view. Our method achieves the best visual effect, followed by DIPRecon, while DeepRED-SGD and DeepRED-BM3D generate much noise. Experiments on different count levels have been conducted, and the results are displayed in Fig. 10. As the count level decreases, the performance of each method drops significantly. When the count level reaches  $1 \times 10^5$ , DeepRED-BM3D can only reconstruct rough outlines without

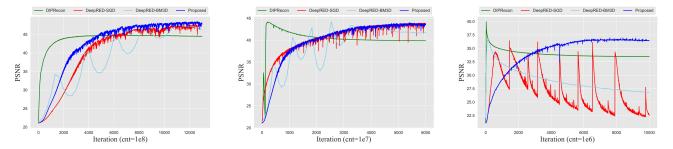


Fig. 12. PSNR curves with the increase in iterations for DIP-based methods. Left: the count level of  $1 \times 10^8$ . Middle: the count level of  $1 \times 10^7$ . Right: the count level of  $1 \times 10^6$ .

any details, and severe noise occurs in DIPRecon, while our method has a better trade-off between detail preservation and noise suppression than other methods. TABLE III and TABLE IV show the means of PSNR and SSIM values in the wholebody dataset using different methods. Our method has better scores than all the other methods except CNN denoising. In addition, we select a line profile across the image, as shown in Fig. 11, which further demonstrates that our proposed method obtains the best result.

# D. Convergence analysis

The PSNR curves with the increase in iterations for each method are shown in Fig. 12. Our method converges to a stable solution in all three count levels, but other methods produce oscillatory curves to different degrees. For a count level of  $1 \times 10^8$ , our proposed method achieves noticeable advantages over all the other methods. For a lower count level of  $1 \times 10^7$ , the scores of all the methods significantly decrease. DeepRED-SGD and our method yield similar performance and better performance than other methods. DIPRecon reaches the best result first but then declines, which indicates the occurrence of overfitting. When the count level is  $1 \times 10^6$ , DeepRED-SGD overfits while our method still converges, which shows the advantage brought by SGLD.

### V. DISCUSSION AND CONCLUSION

In this work, we propose a PET reconstruction method directly from a sinogram under the DeepRED framework from a Bayesian perspective. Two well-known networks, U-Net and DnCNN, are selected as backbones of the generative network and denoiser in our framework, respectively. Notably, more state-of-the-art network components, such as octave convolution, attention modules and even transformers, may further improve the reconstruction performance. Since the main purpose of this work is to validate the effectiveness of our proposed framework, exploring the impacts of different network architectures will be our future work. In addition, only 2D cases are considered in this work as a pilot study. Introducing extra data from the spatial or temporal dimension will probably further improve the results, but this strategy will inevitably boost the computational cost. Since our method cannot be implemented in a patchwise manner (patch-based image denoising), 3D or 4D PET reconstruction will need much more video memory, which may be impractical for regular users, and light network architecture will be preferable for this situation. Moreover, no anatomic prior is used in our method, which means our method is fully unsupervised. It was mentioned in [37] that taking MRI prior as complementary input could produce better results than random input. In future work, we will evaluate the performance of introducing anatomic images, such as CT or MRI images, as auxiliary priors. The selection of parameters is also an important topic for our method. In the current approach, the parameters are manually optimized to fit the specific dataset or count level. How to develop an adaptive parameter selection strategy will also be a future topic of ours.

In conclusion, a PET reconstruction model that combines the DeepRED framework with approximate Bayesian inference is proposed in this paper. Our model restores PET images from a single sinogram in a fully unsupervised mode. A learnable denoiser, which has a differentiable form, is trained to substitute the conventional denoisers used in DeepRED. To further alleviate overfitting, Gaussian noise is injected into the gradient updates, which derives an MCMC sampler from SGD. Brain data and whole-body data experiments show that our proposed method produces better visual effects in detail preservation and noise suppression. Quantitative results also show that our proposed method outperforms several state-ofthe-art methods in terms of PSNR and SSIM.

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