# Deep Learning in CT Image Formation

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DEUTSCHES KREBSFORSCHUNGSZENTRUM **ER HELMHOLTZ-GEMEINSCHAFT** 

# **Fully Connected Neural Network**

- Each layer fully connects to previous layer
- Difficult to train (many parameters in W and b)
- Spatial relations not necessarily preserved



 $y(x) = f(W \cdot x + b)$  with  $f(x) = (f(x_1), f(x_2), ...)$  point-wise scalar, e.g.  $f(x) = x \vee 0 = \text{ReLU}$ 

# **Activation Functions**

Function	Equation	Plot	Function	Equation	Plot
Identity	f(x) = x		ReLU	$f(x) = \begin{cases} 0 & \text{for } x < 0\\ x & \text{for } x \ge 0 \end{cases}$	
Sigmoid	$f(x) = \frac{1}{1 + e^{-x}}$		Leaky ReLU	$f(x) = \begin{cases} \alpha x & \text{for } x < 0\\ x & \text{for } x \ge 0 \end{cases}$	
Hard sigmoid	$f(x) = \begin{cases} 0 & \text{for } x < -\alpha \\ \frac{\alpha + x}{2\alpha} & \text{for } -\alpha \le x < \\ 1 & \text{for } x \ge \alpha \end{cases}$		ELU	$f(x) = \begin{cases} \alpha(e^x - 1) & \text{for } x < 0\\ x & \text{for } x \ge 0 \end{cases}$	
Tanh	$f(x) = \frac{2}{1 + e^{-2x}} - 1$		Inverse square root LU	$f(x) = \begin{cases} \frac{x}{\sqrt{1+\alpha x^2}} & \text{for } x < 0\\ x & \text{for } x \ge 0 \end{cases}$	
Softsign	$f(x) = \frac{x}{1+ x }$			•••	••••
Softplus	$f(x) = \log(1 + \exp x)$				



### **Gradient Descent**

- Walk along the direction of the negative gradient
- Steepest descent
- Learning rate  $\eta$

$$\boldsymbol{w}^{\mathrm{new}} = \boldsymbol{w}^{\mathrm{old}} - \eta \, \boldsymbol{\nabla}_{\boldsymbol{w}} \, L(\boldsymbol{x}_n, \boldsymbol{y}_n, \boldsymbol{w})$$

- Easy to understand, but not optimal
- Methods in use
  - Batch gradient descent
  - Sochastic gradient descent
  - Mini-batch gradient descent
  - Conjugate gradient descent
  - Quasi Newton methods
  - Momentum methods

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# **Convolutional Neural Network (CNN)**

- Replace dense W in  $y(x) = f(W \cdot x + b)$  by a sparse matrix W with sparsity being of convolutional type.
- CNNs consist (mainly) of convolutional layers.
- Convolutional layers are not fully connected.
- Convolutional layers are connected by small, say 3x3, convolution kernels whose entries need to be found by training.
- CNNs preserve spatial relations to some extent.









<sup>1</sup>O. Ronneberger, P. Fischer, and T. Brox. U-net: Convolutional networks for biomedical image segmentation. Proc. MICCAI:234-241, 2015.



# What is an Autoencoder?

- In and output domain are the same, here x.
- Bottleneck z enforces the encoder and decoder to do a good job.

$$x - \mathbf{E} - z - \mathbf{D} - x' = D(z) = D(E(x))$$

#### • Examples:

- Principal component analysis (linear autoencoder), lossless
- PCA with dimensionality reduction (nonlinear due to clipping), lossy
- Image compression and decoding, e.g. jpeg, lossy
- Latent space typically not interpretable.



### What is a Variational Autoencoder?

- Make latent space regular.
- Allow to sample in latent space from a given distribution, here: normal distribution.

$$x - \mathbf{E} - (\mu, \sigma) \quad z \sim \mathcal{N}(\mu, \sigma) - \mathbf{D} \quad -x' = D(z) = \\ = D(\mathcal{N}(E(x)))$$

- The VAE is a generative model.
- It allows to generate new data by sampling new values from the normal distribution.



### **Loss Function**

 The neural network parameters (weights and biases) w are chosen by minimizing a loss function (cost function)

$$oldsymbol{w} = rg\min_{oldsymbol{w}} \sum_{n=1}^N L(oldsymbol{x}_n, oldsymbol{y}_n, oldsymbol{w})$$

- with  $x_n$  being the training data input,  $y(x_n, w)$  being the network output, and  $y_n$  being the so-called labels, i.e. the training target, and N being the number of training samples.
- An example for such a loss function is the MSE loss

$$L(\boldsymbol{x}_n, \boldsymbol{y}_n, \boldsymbol{w}) = \left( \boldsymbol{y}(\boldsymbol{x}_n, \boldsymbol{w}) - \boldsymbol{y}_n) \right)^2$$



# Generative Adversarial Network<sup>1</sup> (GAN)

 Useful, if no direct ground truth (GT) is available, the training data are unpaired, unsupervised learning



<sup>1</sup>I. Goodfellow et al. Generative Adversarial Nets, arXiv 2014



# Generative Adversarial Network (GAN)

Typical loss function and minimax game:

 $\min_{G} \max_{D} L(D,G) := \mathcal{E}_x \ln \left(1 - D(G(x))\right) + \mathcal{E}_y \ln D(y)$ 

- Conditional GAN<sup>1</sup>
  - Conditinal GANs sample the generator input x not from a uniform distribution but from a conditional distribution, e.g. noisy CT images.
  - Need some measure to ensure similarity to input distribution (e.g. pixelwise loss added to the minimax loss function)

### Cycle GAN<sup>2</sup>

- Two GANs (X  $\rightarrow$  Y and Y  $\rightarrow$  X)
- Demand cyclic consistency, i.e.  $x = G_X(G_Y(x))$  and  $y = G_Y(G_X(x))$







### **MAR Example**

 Deep CNN-driven patch-based combination of the advantages of several MAR methods trained on simulated artifacts



- followed by segmentation into tissue classes
- followed by forward projection of the CNN prior and replacement of metal areas of the original sinogram
- followed by reconstruction

Yanbo Zhang and Hengyong Yu. Convolutional Neural Network Based Metal Artifact Reduction in X-Ray Computed Tomography. TMI 37(6):1370-1381, June 2018.









### **Metal artifacts are**



#### + increased susceptibility to sampling artifacts and motion.



# MAR without Machine Learning is a Good Alternative: Frequency Split Normalized MAR<sup>1,2</sup>

Uncorrected

#### **FSLIMAR**

FSNMAR



Patient with bilateral hip prosthesis, Somatom Definition Flash, (C=40/W=500).



FSMAR: Scheme



<sup>1</sup>E. Meyer, M. Kachelrieß. Normalized metal artifact reduction (NMAR) in computed tomography. Med. Phys. 37(10):5482-5493, Oct. 2010 <sup>2</sup>E. Meyer, M. Kachelrieß. Frequency split metal artifact reduction (FSMAR) in CT. Med. Phys. 39(4):1904-1916, April 2012.

# **Summary on Deep MAR**

- Most common uses for networks:
  - Improve image quality in image domain after MAR
  - Use network for the sinogram inpainting
  - Produce a prior image, e.g. for NMAR

#### Additional observations:

- Training data are often produced by segmenting an artifact-free CT image, adding metal and applying a polychromatic forward projection to different types of tissue separately.
- As of today, it seems hard to outperform NMAR, or hard to give convincing clinical examples.





- Architecture based on state-of-the-art networks for image classification (ResNet).
- 32 conv layers with skip connections
- About 2 million tunable parameters in total
- Input is arbitrarily-size stack of images, with a fixed number of adjacent slices in the channel/feature dimension.









#### Low dose images (1/4 of full dose)







#### **Denoised low dose**







#### Full dose







#### **Denoised full dose**



# **Canon's AiCE**

- Advanced intelligent Clear-IQ Engine (AiCE)
- Trained to restore low-dose CT data to match the properties of FIRST, the model-based IR of Canon.
- FIRST is applied to high-dose CT images to obtain a high fidelity training target



Information taken from https://global.medical.canon/products/computed-tomography/aice\_dlr

U = 100 kV CTDI = 0.6 mGy DLP = 24.7 mGy·cm D<sub>eff</sub> = 0.35 mSv





AIDR3De FC52 (image-based iterative)



AiCE Lung (deep learning)

Courtesy of Radboudumc, the Netherlands

# **GE's True Fidelity**

Based on a deep CNN

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- Trained to restore low-dose CT data to match the properties of Veo, the model-based IR of GE.
- No information can be obtained in how the training is conducted for the product implementation.

#### 2.5D DEEP LEARNING FOR CT IMAGE RECONSTRUCTION USING A MULTI-GPU IMPLEMENTATION

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#### ABSTRACT

While Model Based Iterative Reconstruction (MBIR) of CT scans has been shown to have better image quality than Filtered Back Projection (FBP), its use has been limited by its high computational cost. More recently, deep convolutional neural networks (CNN) have shown great promise in both denoising and reconstruction applications. In this research, we propose a fast reconstruction algorithm, which we call Deep

streaking artifacts caused by sparse projection views in CT images [8]. More recently, Ye, et al. [9] developed method for incorporating CNN denoisers into MBIR reconstruction as advanced prior models using the Plug-and-Play framework [10], [11].

In this paper, we propose a fast reconstruction algorithm, which we call Deep Learning MBIR (DL-MBIR), for approximately achieving the improved quality of MBIR using a deep residual neural network. The DL-MBIR method is trained to



#### FBP

**ASIR V 50%** 

**True Fidelity** 

**Courtesy of GE Healthcare** 



Solomon et al. Noise and spatial resolution properties of a commercially available deep learning-based CT reconstruction algorithm. Med. Phys. 47(9):3961-3971, Sept. 2020



# **Philips' Precise Image**

 Noise-injected data serve as low dose examples while their original reconstructions are the labels. A CNN learns how to denoise the low dose images.





iDose<sup>4</sup> 1.4 mSv

iDose<sup>4</sup> 1.5 mSv

Taken from https://www.philips.com/c-dam/b2bhc/master/resource-catalog/landing/precise-suite/incisive\_precise\_image.pdf

Precise Image 0.75 mSv

iDose<sup>4</sup> 5.1 mSv

Precise Image 2.6 mSv











Precise Image 0.7 mSv

iDose<sup>4</sup> 5.4 mSv

Precise Image 2.6 mSv

### **True and Fake DECT**

Existing true DECT approaches (for more than one decade):

Existing fake DECT approaches (as of May 2022):

[1] J. Ma, Y. Liao, Y. Wang, S. Li, J. He, D. Zeng, Z. Bian, "Pseudo dual energy CT imaging using deep learning-based framework: basic material estimation", *SPIE Medical Imaging 2018*.

[2] W. Zhao, T. Lv, P. Gao, L. Shen, X. Dai, K. Cheng, M. Jia, Y. Chen, L. Xing, "A deep learning approach for dual-energy CT imaging using a single-energy CT data", *Fully3D 2019.* 

[3] D. Lee, H. Kim, B. Choi, H. J. Kim, "Development of a deep neural network for generating synthetic dual-energy chest x-ray images with single x-ray exposure", PMB 64(11), 2019.

[4] L. Yao, S. Li, D. Li, M. Zhu, Q. Gao, S. Zhang, Z. Bian, J. Huang, D. Zeng, J. Ma, "Leveraging deep generative model for direct energy-resolving CT imaging via existing energy-integrating CT images", *SPIE Medical Imaging 2020*.

[5] D. P. Clark, F. R. Schwartz, D. Marin, J. C. Ramirez-Giraldo, C. T. Badea, "Deep learning based spectral extrapolation for dual-source, dual-energy x-ray CT", Med. Phys. 47 (9): 4150–4163, 2020.

[6] C. K. Liu, C. C. Liu, C. H. Yang, H. M. Huang, "Generation of brain dual-energy CT from single-energy CT using deep learning", Journal of Digital Imaging 34(1):149–161, 2021.

[7] T. Lyu, W. Zhao, Y. Zhu, Z. Wu, Y. Zhang, Y. Chen, L. Luo, S. Li, L. Xing, "Estimating dual-energy CT imaging from single-energy CT data with material decomposition convolutional neural network", Medical Image Analysis 70:1–10, 2021.

[8] F. R. Schwartz, D. P. Clark, Y. Ding, J. C. Ramirez-Giraldo, C. T. Badea, D. Marin, "Evaluating renal lesions using deeplearning based extension of dual-energy FoV in dual-source CT—A retrospective pilot study", European Journal of Radiology 139:109734, 2021.

[9] Y. Li, X. Tie, K. Li, J. W. Garrett, G.-H. Chen, "Deep-En-Chroma: mining the spectral fingerprints in single-kV CT acquisitions using energy integration detectors", *SPIE Medical Imaging 2022*.







# **Algorithm for Partial DECT**



# **Deep Scatter Estimation**



???

In real time?





### **Motivation**

- X-ray scatter is a major cause of image quality degradation in CT and CBCT.
- Appropriate scatter correction is crucial to maintain the diagnostic value of the CT examination.



### Monte Carlo Scatter Estimation

- Simulation of photon trajectories according to physical interaction probabilities.
- 1 to 10 hours per tomographic data set Simulating a large number ries well approximat
  - suplete scatter distribution



# **Deep Scatter Estimation**

**Network architecture & scatter estimation framework** 



J. Maier, M. Kachelrieß et al. Deep scatter estimation (DSE). SPIE 2017 and Journal of Nondestructive Evaluation 37:57, July 2018. J. Maier, M. Kachelrieß et al. Robustness of DSE. Med. Phys. 46(1):238-249, January 2019.
### **Reconstructions of Measured Data**



#### *C* = 0 HU, *W* = 1000 HU

J. Maier, M. Kachelrieß et al. Deep scatter estimation (DSE). SPIE 2017 and Journal of Nondestructive Evaluation 37:57, July 2018. J. Maier, M. Kachelrieß et al. Robustness of DSE. Med. Phys. 46(1):238-249, January 2019.



## **Scatter in Dual Source CT (DSCT)**



forward and cross-scatter correction in dual source CT. Med. Phys. 48:4824-4842, July 2021.



finite size focal spot

pre patient collimation

dkfz.



J. Erath, T. Vöth, J. Maier, E. Fournié, M. Petersilka, K. Stierstorfer, and M. Kachelrieß. Deep learning-based forward and cross-scatter correction in dual source CT. Med. Phys. 48:4824–4842, July 2021.

### **Cross-DSE**

# **Ground Truth** Uncorrected xDSE (2D, xSSE) **Measurement-based** MAE = 42.6 HU MAE = 4.9 HU MAE = 10.6 HU

#### xDSE (2D, xSSE) maps primary + forward scatter + cross-scatter + cross-scatter approximation $\rightarrow$ cross-scatter

Images C = 40 HU, W = 300 HU, difference images C = 0 HU, W = 300 HU

J. Erath, T. Vöth, J. Maier, E. Fournié, M. Petersilka, K. Stierstorfer, and M. Kachelrieß. Deep learning-based forward and cross-scatter correction in dual source CT. Med. Phys. 48:4824–4842, July 2021.



### **Conclusions on DSE**

- DSE needs about 3 ms per CT and 10 ms per CBCT projection (as of 2020).
- DSE is a fast and accurate alternative to MC simulations.
- DSE outperforms kernel-based approaches in terms of accuracy and speed.
- Facts:
  - DSE can estimate scatter from a single (!) x-ray image.
  - DSE can accurately estimate scatter from a primary+scatter image.
  - DSE generalizes to all anatomical regions.
  - DSE works for geometries and beam qualities differing from training.
  - DSE may outperform MC even though DSE is trained with MC.
- DSE is not restricted to reproducing MC scatter estimates.
- DSE can rather be trained with any other scatter estimate, including those based on measurements.

J. Maier, M. Kachelrieß et al. Deep scatter estimation (DSE). SPIE 2017 and Journal of Nondestructive Evaluation 37:57, July 2018. J. Maier, M. Kachelrieß et al. Robustness of DSE. Med. Phys. 46(1):238-249, January 2019.



### **Scatter of Coarse ASG**



This paper received the "Highest Impact Paper Award" for the highest impact score at the 7th International Conference on Image Formation in X-Ray Com-puted Tomography in June 2022



**Scattered** 

photons

### **Scatter Artifacts of Coarse ASG**



Coarse ASG can lead to scatter-induced moiré artifacts.

Reconstruction: C = 40 HU, W = 300 HU





Scatter distribution averaged over all detector rows





Scatter distribution averaged over all detector rows

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### **Training and Validation Data**

- Monte Carlo simulation with the geometry of the photon counting CT scanner NAEOTOM Alpha (Siemens Healthineers)
- 12 patients for training and 4 for validation
- 14 z-positions with 36 projections each simulated for each patient
- 8064 paired scatter and primary data pairs
- Simulation of coarse ASG with macro pixel with detector dimension of 1376 x 144 pixels
- 6 different macro pixels locations
- Smooth only across same macro-pixel locations



Training and validation patients with high variety and different clinical situations, important to consider scatter-to-primary ratio

Example of validation data set:



 M(0,0)
 M(1,0)

 M(0,1)
 M(1,1)

 M(0,2)
 M(1,2)



### **DSE for coarse ASG**



This paper received the "Highest Impact Paper Award" for the highest impact score at the 7th International Conference on Image Formation in X-Ray Com-puted Tomography in June 2022



### **Results in Reconstructed Images**





Simulated Reconstruction C = 0 HU, W = 400 HU, Difference to GT C = 0 HU, W = 50 HU



## **Results in Reconstructed Images**



Simulated Reconstruction C = 0 HU, W = 400 HU, Difference to GT C = 0 HU, W = 50 HU



### Conclusions

- Coarse anti-scatter grid can lead to moiré artifacts due to scattered radiation.
- DSE reduces the mean absolute error (MAE) from about 9 HU to under 1 HU.
- The moiré pattern's amplitude can be reduced from 30 HU to less than 5 HU.



### **uDSE – Basis Principle**





### **Datasets**

- Training and testing data were generated using CT simulations based on 65 clinical CT reconstructions.
- Based on the corresponding voxel volumes, CBCT scans (120 kV, shifted detector, RFD = 1100 mm, RF = 700 mm, 360 views 360°) were simulated at five different z-positions within the abdomen region.
- Generation of one scatter corrupted dataset (30 patients) that was used as input to the generator network, one scatter-free dataset (30 patients) that was used to provide ideal reference for the critic network, and a scatter-corrupted dataset (remaining patients) for testing.



## Training

Training of conventional DSE as reference using the following loss function:

$$L_{\rm DSE}(\theta) = \sum_{n}^{B} \left| \frac{\text{DSE}_{\theta}(I_n) - S_n}{S_n} \right|$$

• Training of uDSE using a WGAN setup:

$$L_{\text{critic}}(\theta_c) = \sum_{n}^{B} C_{\theta_c}(G_{\theta_g}(I_n)) - C_{\theta_c}(f_{\text{real, n}})$$
$$L_{\text{gen}}(\theta_g) = -\sum_{n}^{B} C_{\theta_c}(G_{\theta_g}(I_n)),$$









### **Results** CT Reconstructions





### **Conclusions on uDSE**

- This study demonstrates the feasibility of learning CT scatter estimation in absence of labeled data.
- uDSE is able to remove most of the present scatter artifacts and yields similar CT value accuracy (mean error of 27.9~HU vs. 24.7~HU) as a state-of-the-art supervised scatter estimation approach
- In general uDSE is not restricted to CBCT but can be trained with any tomographic input and any scatter-free reference as long as both distributions are sufficiently equal after scatter correction.
- Thus, uDSE has the potential to extend the concept of neural network-based scatter estimation and correction to scenarios where labels are not available or cannot be generated with sufficient accuracy.



## **Deep Dose Estimation**



??? In real time?





### **Estimation of Dose Distributions**

### Useful to study dose reduction techniques

- Tube current modulation
- Prefiltration and shaped filtration
- Tube voltage settings

- ...

### Useful to estimate patient dose

- Risk assessment requires segmentation of the organs (difficult)
- Often semiantropomorphic patient models take over
- The infamous k-factors that convert DLP into D<sub>eff</sub> are derived this way, e.g. k<sub>chest</sub> = 0.014 mSv/mGy/cm

- ...

- Could be useful for patient-specific CT scan protocol optimization
- However: Dose estimation does not work in real time!



### MC Dose Simulation for a 360° Scan



J. Maier, L. Klein, E. Eulig, S. Sawall, and M. Kachelrieß. Real-time estimation of patient-specific dose distributions for medical CT using the deep dose estimation. Med. Phys. 49(4):2259-2269, April 2022. Best Paper within Machine Learning at ECR 2019!



### **Deep Dose Estimation (DDE)**

 Combine fast and accurate CT dose estimation using a deep convolutional neural network

secon

Train the network to reightarrowgiven the



Depth concatenate

J. Maier, L. Klein, E. Eulig, S. Sawall, and M. Kachelrieß. Real-time estimation of patient-specific dose distributions for medical CT using the deep dose estimation. Med. Phys. 49(4):2259-2269, April 2022. Best Paper within Machine Learning at ECR 2019!



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### **Results** Thorax, tube A, 120 kV, no bowtie

#### **CT** image

#### First order dose

### MC ground truth





	МС	DDE
48 slices	1 h	0.25 s
whole body	20 h	5 s

MC uses 16 CPU kernels DDE uses one Nvidia Quadro P600 GPU

DDE training took 74 h for 300 epochs, 1440 samples, 48 slices per sample

#### **Relative error**



C = 0%W = 40%

J. Maier, L. Klein, E. Eulig, S. Sawall, and M. Kachelrieß. Real-time estimation of patient-specific dose distributions for medical CT using the deep dose estimation. Med. Phys. 49(4):2259-2269, April 2022. Best Paper within Machine Learning at ECR 2019!

### **Results** Pelvis, tube B, 120 kV, no bowtie

#### **CT** image

#### **First order dose**

			МС	DDE
		48 slices	1 h	0.25 s
Joe's Lie		whole body	20 h	5 s
		MC uses 16 DDE uses or GPU	CPU kernels ne Nvidia Quac	dro P600
		DDE training 1440 sample	took 74 h for s, 48 slices pe	300 epochs, er sample
	DDE			
MC ground truth	DDE	Rela	tive erro	r
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J. Maier, L. Klein, E. Eulig, S. Sawall, and M. Kachelrieß. Real-time estimation of patient-specific dose distributions for medical CT using the deep dose estimation. Med. Phys. 49(4):2259-2269, April 2022. Best Paper within Machine Learning at ECR 2019!

## DDE's Organ Dose and D<sub>eff</sub> MAPEs

Organ and ICRF	P weight	80 kV	100 kV	120 kV
Bone marrow	0.12	5.2%	6.7%	7.1%
Bone surface	0.01	5.7%	7.0%	7.2%
Brain	0.01	5.1%	4.9%	5.3%
Breast	0.12	1.0%	1.4%	2.1%
Colon	0.12	0.9%	1.7%	1.9%
Esophagus	0.04	1.3%	2.4%	2.3%
Gonads	0.08	3.2%	2.7%	2.2%
Liver	0.04	2.9%	1.1%	0.8%
Lung	0.12	1.7%	3.5%	4.0%
Remainder	0.12	0.9%	1.9%	2.3%
Salivary glands	0.01	4.9%	5.1%	5.3%
Skin	0.01	2.8%	3.3%	4.2%
Stomach	0.12	2.3%	1.1%	0.8%
Thyroid gland	0.04	3.1%	3.0%	2.3%
Urinary bladder	0.04	1.7%	1.7%	1.3%
Effec	tive dose	1.2%	2.5%	2.7%

Weighting factors and mean absolute percentage error of the DDE organ dose values with respect to the ground truth Monte Carlo organ dose values.



## **Conclusions on DDE**

- DDE provides accurate dose predictions
  - for circle scans
  - for sequence scans
  - for partial scans (less than 360°)
  - for limited angle scans (less than 180°)
  - for spiral scans
  - for different tube voltages
  - for scans with and without bowtie filtration
  - for scans with tube current modulation
- In practice it may therefore be not necessary to perform separate training runs for these cases.
- Thus, accurate real-time patient dose estimation may become feasible with DDE.



## Patient Risk-Minimizing Tube Current Modulation

### **1.** Coarse reconstruction from two scout views

 E.g. X. Ying, et al. X2CT-GAN: Reconstructing CT from biplanar xrays with generative adversarial networks. CVPR 2019.

### 2. Segmentation of radiation-sensitive organs

 E.g. S. Chen, M. Kachelrieß et al., Automatic multi-organ segmentation in dual-energy CT (DECT) with dedicated 3D fully convolutional DECT networks. Med. Phys. 2019.

# 3. Calculation of the effective dose per view using the deep dose estimation (DDE)

 J. Maier, E. Eulig, S. Dorn, S. Sawall and M. Kachelrieß. Real-time patient-specific CT dose estimation using a deep convolutional neural network. IEEE Medical Imaging Conference Record, M-03-178: 3 pages, Nov. 2018.

## 4. Determination of the tube current modulation curve that minimizes the radiation risk

L. Klein, C. Liu, J. Steidel, L. Enzmann, M. Knaup, S. Sawall, A. Maier, M. Lell, J. Maier, and M. Kachelrieß. Patient-specific radiation risk-based tube current modulation for diagnostic CT. Med. Phys. 49(7):4391-4403, July 2022.











View angle



### Patient 04 - Abdomen



#### C = 25 HU, W = 400 HU

<sup>1</sup>L. Klein, C. Liu, J. Steidel, L. Enzmann, M. Knaup, S. Sawall, A. Maier, M. Lell, J. Maier, and M. Kachelrieß. Patient-specific radiation risk-based tube current modulation for diagnostic CT. Med. Phys. 49(7):4391-4403, July 2022.



### **Conclusions on RiskTCM**

- **Risk-specific TCM minimizes the patient risk.** ightarrow
- With *D*<sub>eff</sub> as a risk model riskTCM can reduce risk by up to 30%, compared with the gold standard mAsTCM.
- Other risk models, in particular age-, weight- and sexspecific models, can be used with riskTCM It is up to the vendors to take action!
- Note:  $\bullet$ 
  - good for the patient
  - detector flux equalizing TCM = good for the detector

#### ECR 2022 – Best Research Presentation Abstract

within the topic Physics in Medical Imaging with the presentation:

Risk-minimising tube current modulation (riskTCM) for CT - potential dose reduction across different tube voltages (16765)

L. Klein1, C. Liu2, J. Steidel1, L. Enzmann1, S. Sawall1, J. Maier1, A. Maier2, M. Lell3, M. Kachelrieß1; 1Heidelberg/DE, 2Erlangen/DE, 3Nuremberg/DE



<sup>1</sup>L. Klein, C. Liu, J. Steidel, L. Enzmann, M. Knaup, S. Sawall, A. Maier, M. Lell, J. Maier, and M. Kachelrieß. Patient-specific radiation risk-based tube current modulation for diagnostic CT. Med. Phys. 49(7):4391-4403, July 2022.



## **Deep Cardiac Motion Compensation**





### Motivation



C = 0 HU, W = 1200 HU

Motion artifacts

### High noise levels

Table 3: Reason for  $\ensuremath{\mathsf{FFR}_{\mathsf{cr}}}$  Rejection in the ADVANCE Registry and Clinical Cohort

	$FFR_{_{CT}}$ Rejected*		
Reason for Rejection	ADVANCE Registry $(n = 80)$	Clinical Cohort ( <i>n</i> = 892)	
Inadequate image quality <sup>†</sup>			
Blooming	4 (5.0)	29 (3.0)	
Clipped structure	ч (Э.0)	39 (4.3)	
Motion artifacts	63 (78.0)	729 (81.4)	
Image noise	2 (2.5)	198 (22.1)	
Inappropriate submission			
Stent or previous coronary artery bypass graft	5 (6.2)	116 (13.0)	
present	a (a a)		
Cardiac hardware present	2 (2.5)	29 (3.2)	

The rejection rate was 892 of 10416 cases submitted

\* G. Pontone et al., "Determinants of Rejection Rate for Coronary CT Angiography Fractional Flow Reserve Analysis", *Radiology*, 292(3), 597–605 (2019)



\*

### Partial Angle-Based Motion Compensation (PAMoCo)



Animated rotation time = 100 × real rotation time

### dkfz.

## Partial Angle-Based Motion Compensation (PAMoCo)







### Partial Angle-Based Motion Compensation (PAMoCo)

ho Motion vector field  $\, {f s}_1({f r})$ 





Apply motion vector fields (MVFs) to partial angle reconstructions
## Partial Angle-Based Motion Compensation (PAMoCo)



#### **Prior work:**

[1] S. Kim et al., "Cardiac motion correction based on partial angle reconstructed images in x-ray CT", Med. Phys. 42 (5): 2560–2571 (2015).

[2] J. Hahn et al., "Motion compensation in the region of the coronary arteries based on partial angle reconstructions from short-scan CT data", Med. Phys. 44 (11): 5795–5813 (2017).

[3] S. Kim et al., "Cardiac motion correction for helical CT scan with an ordinary pitch", IEEE TMI 37 (7): 1587–1596 (2018).

→ Limitation: Challenging / timeconsuming optimization



## **Deep Partial Angle-Based Motion Compensation (Deep PAMoCo)**

PARs centered Neural network to predict parameters of a motion model around coronary artery Fully  $\mathbf{x} = \mathbf{s}_{0,x}$ connected  $a = s_{0,y}$  $\mathbf{x} = s_{0,z}$  $\mathbf{s} = \mathbf{s}_{2,x}$  $\grave{\mathbf{x}} \equiv s_{2,u}$ 📙 3 × 3 × 3 Convolution, Batch norm, ReLU 🌔 2 × 2 × 2 Max pooling 🍃 Flatten 🛛 🗙 Dropout (25 %)

**Reinsertion of patch into** initial reconstruction



[1] M. Jaderberg et al., "Spatial transformer networks", NIPS 2015: 2017–2025 (2015).

## **Training Data Generation**

- Removal of coronary arteries from real CT reconstructions.
- Insertion of artificial coronary arteries with different shape, size, and contrast.
- Simulation of CT scans with coronary artery motion.





### Patient 1

### Original







#### C = 0 HU, W = 1400 HU

J. Maier, S. Lebedev, J. Erath, E. Eulig, S. Sawall, E. Fournié, K. Stierstorfer, M. Lell, and M. Kachelrieß. Deep learning-based coronary artery motion estimation and compensation for short-scan cardiac CT. Med. Phys. 48(7):3559-3571, July 2021.



## Patient 2

### Original







#### C = 0 HU, W = 1600 HU

J. Maier, S. Lebedev, J. Erath, E. Eulig, S. Sawall, E. Fournié, K. Stierstorfer, M. Lell, and M. Kachelrieß. Deep learning-based coronary artery motion estimation and compensation for short-scan cardiac CT. Med. Phys. 48(7):3559-3571, July 2021.



### Patient 3

#### Original







#### C = 0 HU, W = 1000 HU

J. Maier, S. Lebedev, J. Erath, E. Eulig, S. Sawall, E. Fournié, K. Stierstorfer, M. Lell, and M. Kachelrieß. Deep learning-based coronary artery motion estimation and compensation for short-scan cardiac CT. Med. Phys. 48(7):3559-3571, July 2021.



### Patient 4 (Iterative Recon) Measurements at a Siemens Somatom AS, patient 1



C = 0 HU, W = 1200 HU



## **Are the Methods Reliable?**

- Studies about explainability of Al in CT image formation are more than sparse.
- My thoughts:
  - Cosmetic corrections: Unclear if noise reduction, metal artifact reduction etc. is removing/adding lesions. The whole process is a black box.
  - Physical corrections: A clear physical meaning and rawdata fidelity appear more reliable. Examples:
    - » MAR or detruncation networks where the NN output is used only to forward project and inpaint/extrapolate the rawdata
    - Scatter correction that estimates a smooth physically realistic (trained with MC) scatter signal in intensity domain
    - » Motion correction networks that estimate motion vectors rather than manipulating the voxel values



Explainable AI for CT: Analyzing CT Image Denoising Networks by Reconstructing their Invariances

- Elias Eulig, Björn Ommer, and Marc Kachelrieß
- RSNA 2022







#### FBP(10 mAs)

IRLNet(10 mAs, T-Net)

IRLNet(10 mAs, A-Net)



## **Motivation**

### In general:

- Deep learning methods are employed for many problems in medical image formation, including image-based noise reduction.
- However, they lack interpretability due to black-box nature of DNNs. Recent advancement in generative modelling signal false confidence.

### Aim of this work:

- Lay fundamentals for post-hoc interpretability and robustness analysis of denoising DNNs.
- Use two simple denoising networks *f* as initial examples:
  - Chen's simple 3-layer CNN trained with  $\mathcal{L}_2$  loss<sup>1</sup>
  - Yang's Wasserstein GAN with additional perceptual loss<sup>2</sup>
- See what they have learned to represent and what to ignore: For a given output x´there are many inputs x that produce the same output x´ = f(x).
- Employ low dose CT image and projection dataset for all studies.<sup>3</sup>



<sup>2</sup>Q. Yang et al., "Low-Dose CT Image Denoising Using a Generative Adversarial Network [...]", in IEEE TMI, vol. 37, no. 6, 2018.



<sup>&</sup>lt;sup>1</sup>H. Chen et al., "Low-dose CT denoising with convolutional neural network", ISBI 2017, 2017.

<sup>&</sup>lt;sup>3</sup>C. McCollough et al., "Data from low dose CT image and projection data [data set]," The Cancer Imaging Archive, 2020.

## Recap 1: What is an Autoencoder (AE)?

- In and output domain are the same, here x.
- Bottleneck z enforces the encoder and decoder to do a good job.



### • Examples:

- Principal component analysis (linear autoencoder), lossless
- PCA with dimensionality reduction (nonlinear due to clipping), lossy
- Image compression and decoding, e.g. jpeg, lossy
- Latent space typically not interpretable.



## Recap 2: What is a Variational AE (VAE)?

- Make latent space regular.
- Allow to sample in latent space from a given distribution, here: normal distribution.

$$x - \mathbf{E} - (\mu, \sigma) \quad \bar{z} \sim \mathcal{N}(\mu, \sigma) - \mathbf{D} \quad -\bar{x} = D(z) = \\ = D(\mathcal{N}(E(x)))$$

- The VAE is a generative model.
- It allows to generate new data by sampling new values from the normal distribution.



### Method Recovering Invariances

- Our work is based on Rombach et al.<sup>1</sup>
- Given a function or network  $f(x) = \Psi(\Phi(x))$  we analyze its internal latent representations  $z = \Phi(x)$ .
- Train a VAE to learn a complete data representation  $\bar{z} = E(x)$  of low dose images.
- Disentangle information captured in z and invariances v by learning a mapping  $v = t(\overline{z}|z), \ \mathcal{L}(v) = \mathcal{N}(0, 1)$
- $t(\cdot|z)$  is realized by a conditional invertible neural network (cINN).
- Generate new images varying only by their invariances

$$\bar{x} = D(t^{-1}(v|z)) \qquad v \sim \mathcal{N}(0,1)$$

 $f(x) = \Psi(\Phi(x))$ **Conv k**3 f32 Conv k9 f6 conv k9 f6 inv k3 f1  $\mathcal{Z}$  $v = t(\bar{z}|z)$ ED



Alternative: Use VAE in high dose domain, i.e. VAE<sub>v</sub>, to visualize the invariances.



### Method Recovering Invariances

- 1. Our work is based on Rombach et al.<sup>1</sup>
- 2. Train denoising methods Chen et al. & Yang et al.
- 3. Train VAE to learn a complete data representation of the low dose images *x*.
- 4. For each denoising method and layer in the network we wish to evaluate, train a cINN to recover the invariances.
- 5. For a given test image, sample 250 invariances v, apply the inverse mapping  $t^{-1}$  and apply the pretrained decoder D.

### $t^{-1}$ maps $\mathcal{N}(0,1)$ onto $p(\bar{z}|z)$ .





Building block of INN: Invertible block,  $\xi_{12}$  and  $\delta_{12}$  are CNNs or NNs

 $x_1 \exp(\xi_2(\hat{x}_2)) + \delta_2(\hat{x}_2) = \hat{x}_1$  $x_2 \exp(\xi_1(x_2)) + \delta_1(x_1) = \hat{x}_2$ 

<sup>1</sup>Rombach et al. "Making sense of CNNs: Interpreting deep representations and their invariances with INNs", ECCV 2020.







Alternative: Use VAE in high dose domain, i.e. VAE<sub>v</sub>, to visualize the invariances.

### **Results** Denoising (Yang et al.) $f = \Psi \circ \Phi$





### **Results** Denoising (Yang et al.) $f = \Psi \circ \Phi$



Arrows point at selected differences between prediction and ground truth.



## Results

5 6 7 8

2 3 4

Sampling Invariances in Yang et al.'s Net



Same samples of v used for the rows corresponding to wiretapping after layers 1, 4 and 7.







### Results

Sampling Invariances in Target Domain in Chen et al.'s Net



$$x' = f(x) = f(\mathsf{R}^{-1}\mathcal{P}\,\mathsf{R}\,\bar{y}) \quad \forall \ \bar{y}$$

Wiretapping after last layer.



## **Conclusions & Outlook**

#### Conclusions

- Designed a method to highlight invariances of a given network.
- Algorithm agnostic, not restricted to denosing.
- Architecture agnostic, not restricted to CT.
- Both denoising methods are invariant to some anatomical features to some extent.

#### Outlook

- Improve interpretability by
  - improving the embedding of the VAEs,
  - mapping sampled invariance images to semantically meaningful space (disentangled representations of e.g. tumors).
- One could use the undesired invariances to finetune the denoising methods.





## **Conclusions on Deep CT**

- Machine learning plays and will play a significant role in CT image formation.
- High potential for
  - Artifact correction
  - Noise and dose reduction
  - Real-time dose assessment (also for RT)
- Care has to be taken
  - Underdetermined acquisition, e.g. sparse view or limited angle CT, require the net to make up information!
  - Nice looking images do not necessarily represent the ground truth.
  - Data consistency layers and variational networks with rawdata access may ensure that the information that is made up is consistent with the measured data.

- ...

# Thank You!

Job opportunities through DKFZ's international PhD or Postdoctoral Fellowship programs (marc.kachelriess@dkfz.de). Parts of the reconstruction software were provided by RayConStruct<sup>®</sup> GmbH, Nürnberg, Germany.

