

# RTG $\pi^3$ Compact Course A3

## Deep learning for Digital Pathology and Inverse Problems

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# Outline

- 1 Introduction
- 2 Data-driven approaches
- 3 Deep Image Prior
- 4 Application in Computed Tomography



## Section 1

### Introduction

# Inverse Problems

Consider an operator  $A : X \rightarrow Y$  between Hilbert spaces  $X$  and  $Y$ .  
Given measured noisy data

$$y^\delta = Ax^\dagger + \tau, \quad (1)$$

where  $\tau$ , with  $\|\tau\| \leq \delta$ , describes the noise in the measurement

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## Example: Computed Tomography

Radon transform

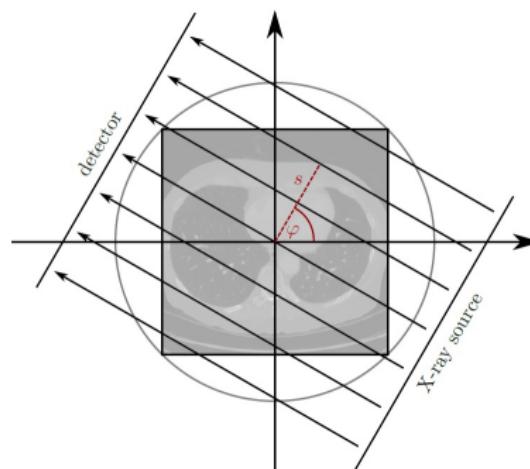


Figure: Parallel beam geometry

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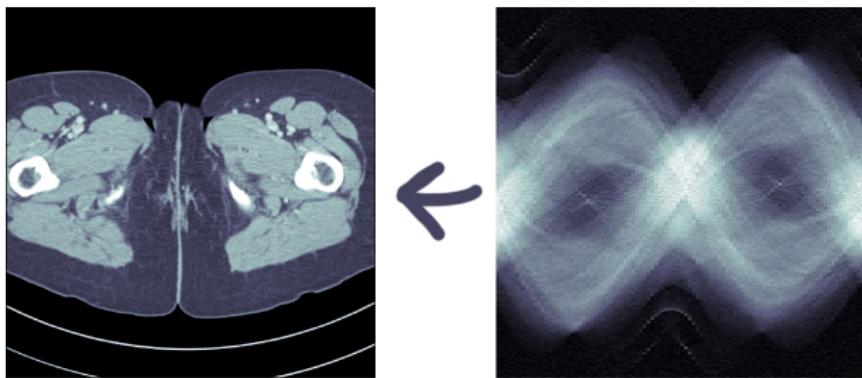


Figure: Human phantom and corresponding sinogram

# Classical approaches

- TSVD
- Tikhonov
- Landweber
- **Variational regularization:**

$$T_\alpha(y^\delta) = \arg \min \frac{1}{2} \|Ax - y^\delta\|^2 + \alpha R(x) \quad (2)$$

Examples of hand-crafted regularizers:  $\|x\|^2$ ,  $\|x\|_1$ ,  $\|\nabla x\|_1$

How to choose  $R$  and the regularization parameters, e.g.  $\alpha$ ?

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# Data-driven approaches<sup>1</sup>

Assume training data is given:  $\{x_i^\dagger, y_i^\delta\}_{i=1}^N$

**Data-driven parameter choice:**

$$\hat{\alpha} = \arg \min_{\alpha \in \mathbb{R}_+} \sum_{i=1}^N \ell(T_\alpha(y_i^\delta), x_i^\dagger) \quad (3)$$

Data-driven regularized inverse  $T_\Theta : Y \rightarrow X$

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## Section 2

### **Data-driven approaches**

## Recent approaches

- Learned methods

$$T_{\Theta} : Y \rightarrow X \quad (4)$$

- Learned regularizers

$$T_{\Theta}(y^{\delta}) = \arg \min_{x \in X} \ell(Ax, y^{\delta}) + R_{\Theta}(x) \quad (5)$$

- Generative Networks

$$T_{\Theta}(y^{\delta}) = \arg \min_{z \in Z} \ell(A\varphi_{\Theta}(z), y^{\delta}) + R(z) \quad (6)$$

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- Generative Networks

$$T_{\Theta}(y^{\delta}) = \arg \min_{z \in Z} \ell(A\varphi_{\Theta}(z), y^{\delta}) + R(z) \quad (9)$$

## Learned methods

- <sup>2</sup> Fully learned
- <sup>3</sup> Learned post-processing:  $T_\Theta = \mathcal{F}_\Theta \circ A^\dagger$
- <sup>4</sup> Learned iterative schemes

Training: Takes quite some time (even weeks)

Evaluation: Takes milliseconds

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## Learned methods: Fully learned

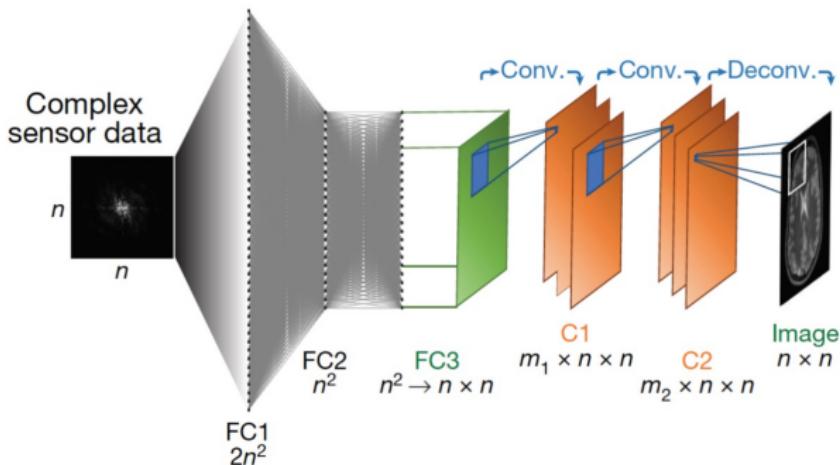


Figure: The AUTOMAP<sup>5</sup> architecture

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## Learned methods: Post-processing

Given data pairs  $\{(y_i^\delta, x_i^\dagger)\}$  and a pseudo inverse  $A^\dagger$ :

- Train a network  $\mathcal{F}_\Theta : X \rightarrow X$  by minimizing

$$\min_{\theta} \frac{1}{N} \sum_{i=1}^N \|\mathcal{F}_\Theta(A^\dagger y_i^\delta) - x_i^\dagger\|^2 \quad (10)$$

**Remark:** Is similar to denoising ( $A^\dagger y_i^\delta$  noisy version of  $x^\dagger$ )

**Architecture:** Autoencoder-like

**Result:**  $T_\Theta(y^\delta) = \mathcal{F}_\Theta(A^\dagger y^\delta)$

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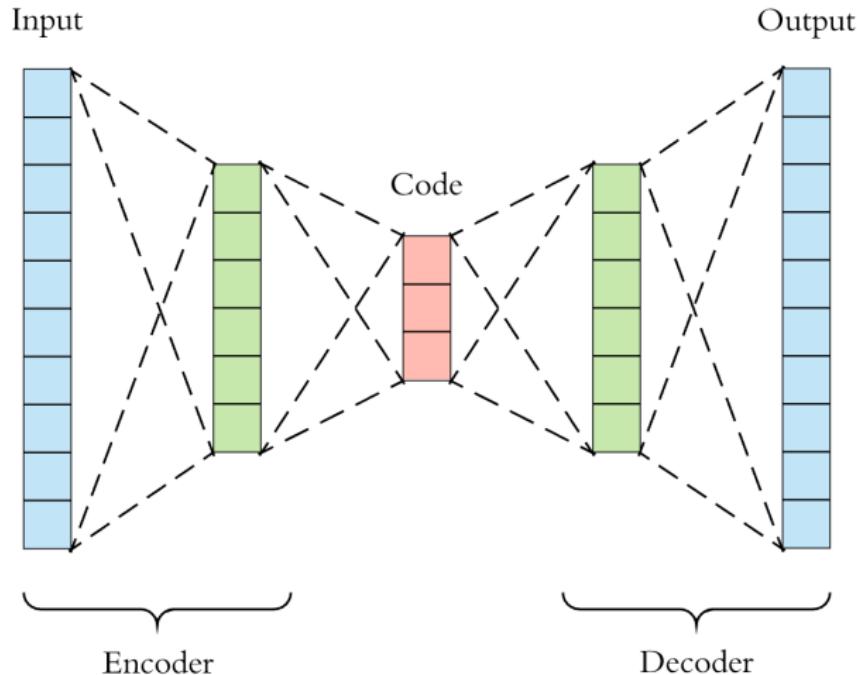
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## Autoencoder



# U-Net

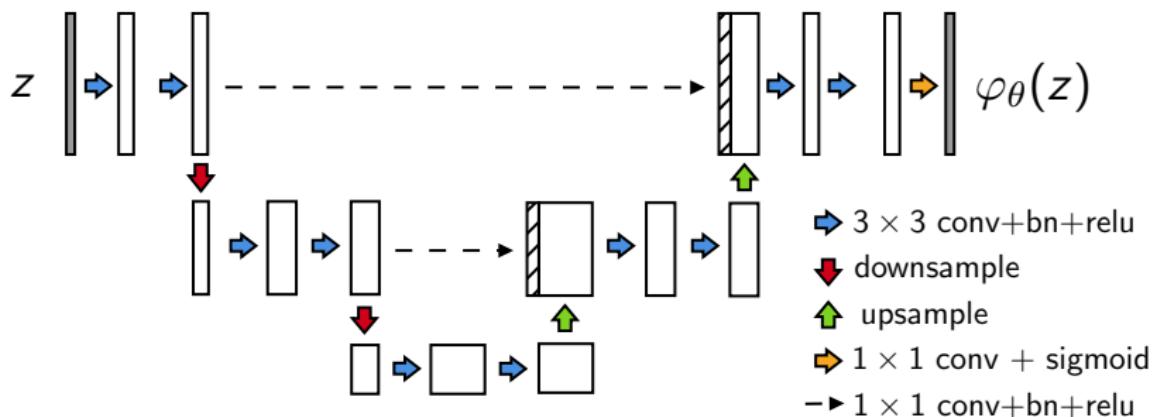
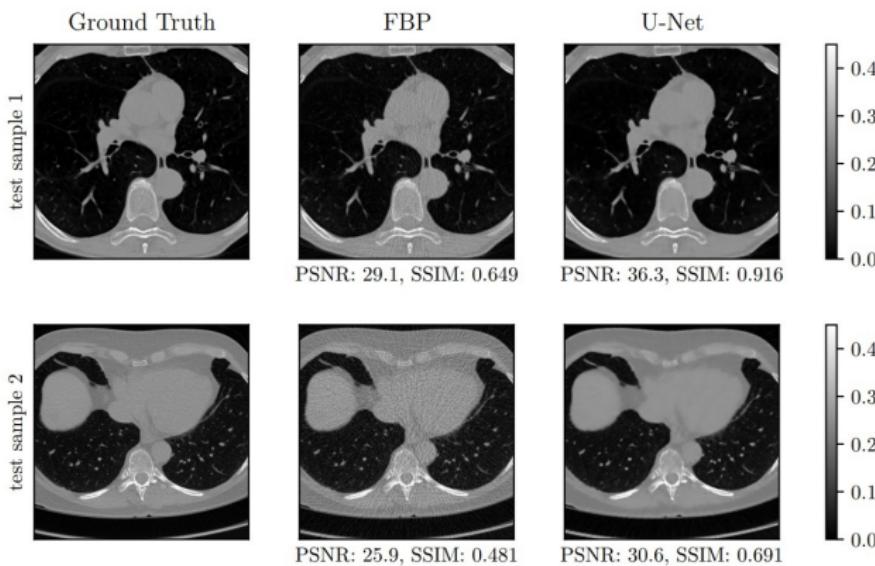


Figure: U-Net architecture

## Post-processing for CT<sup>6</sup>



<sup>6</sup> Johannes Leuschner, Maximilian Schmidt, Daniel Otero Baguer, and Peter Maass. *The LoDoPaB-CT Dataset: A Benchmark Dataset for Low-Dose CT Reconstruction Methods*. 2019. arXiv: 1910.01113 [eess.IV].



## The LoDoPaB-CT Dataset and DIV $\alpha\ell$

### The LoDoPaB-CT Dataset

A Benchmark Dataset for Low-Dose CT Reconstruction Methods

Johannes Leuschner <sup>\*1</sup>, Maximilian Schmidt <sup>†1</sup>, Daniel Otero Baguer <sup>‡1</sup>, and  
Peter Maaß <sup>§1</sup>

<sup>1</sup>University of Bremen, Center for Industrial Mathematics

October 2019

#### Abstract

Deep Learning approaches for solving Inverse Problems in imaging have become very effective and are demonstrated to be quite competitive in the field. Comparing these approaches is a challenging task since they highly rely on the data and the setup that is used for training. We

## Learned methods: Learned iterative schemes

Inspired from iterative optimization methods

$$\hat{x} = \arg \min \frac{1}{2} \|Ax - y^\delta\| + \alpha R(x) \quad (11)$$

Proximal gradient algorithm:

$$x^{k+1} = \underset{R, \alpha, \lambda}{\text{Prox}}(x^k - \lambda A^*(Ax^k - y^\delta)) \quad (12)$$

More general:

$$x^{k+1} = \varphi_\Theta(x^k, A^*(Ax^k - y^\delta)) \quad (13)$$

## Learned methods: Learned iterative schemes

Take a small fixed number of iterations  $k = 1, 2, \dots, L$ , e.g.  
 $L = 10$

Then let  $T_\Theta : Y \rightarrow X$  be

$$T_\Theta(y^\delta) = x^L \quad (14)$$

with

$$x^0 \leftarrow \text{initialized random} \quad (15)$$

$$x^{k+1} = \varphi_\Theta(x^k, A^*(Ax^k - y^\delta)) \quad (16)$$

Optimize  $\Theta$ :

$$\hat{\Theta} = \arg \min_{\theta} \frac{1}{N} \sum_{i=1}^N \| T_\Theta(y_i^\delta) - x_i^\dagger \|^2 \quad (17)$$

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## Recent approaches

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$$T_{\Theta} : Y \rightarrow X \quad (18)$$

- Learned regularizers

$$T_{\Theta}(y^{\delta}) = \arg \min_{x \in X} \ell(Ax, y^{\delta}) + R_{\Theta}(x) \quad (19)$$

- Generative Networks

$$T_{\Theta}(y^{\delta}) = \arg \min_{z \in Z} \ell(A\varphi_{\Theta}(z), y^{\delta}) + R(z) \quad (20)$$



## Learned regularizers

- <sup>7</sup> NETT (Network Tikhonov)
- <sup>8</sup> Adversarial regularizer

Training: Takes quite some time (even weeks)

Evaluation: Takes minutes

Data: Unsupervised data

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<sup>7</sup>Housen Li, Johannes Schwab, Stephan Antholzer, and Markus Haltmeier. "NETT: Solving Inverse Problems with Deep Neural Networks". In: *arXiv preprint arXiv:1803.00092* (Feb. 2018).

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# Learned regularizers: NETT

## NETT (Network Tikhonov)

$$T_\Theta(y^\delta) = \arg \min_{x \in X} \frac{1}{2} \|Ax - y^\delta\|^2 + \psi(\varphi_\Theta(x)) \quad (21)$$

- $\varphi_\Theta : X \rightarrow Z$
- $\psi : Z \rightarrow [0, \infty]$  lower semi-continuous and coercive
- Regularizer  $R_\Theta = \psi(\varphi_\Theta(x))$  is non-convex

**Aim:** Construct a regularizer  $R_\Theta$  with small values for good  $x$  and large value for bad  $x$ .

# Learned regularizers: NETT

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## Learned regularizers: NETT

- $\varphi_\Theta(x)$  extracts artifacts from  $x$



- $\psi(\cdot) = \|\cdot\|^2$

## Learned regularizers: NETT

How to train  $\varphi_\Theta(x)$ ?

- Take training pairs  $\{(x_i^\dagger, 0)\}_{i=1}^N \cup \{(b_i, r_i)\}_{i=1}^N$
- $b_i = A^\dagger Ax_i$  (images with artifacts)
- $r_i = x_i - A^\dagger Ax_i$  (residual artifacts)
- Train network  $\varphi_\Theta$  by minimizing

$$\hat{\Theta} = \arg \min \frac{1}{N} \sum \|\varphi_\Theta(x_i^\dagger)\|^2 + \frac{1}{N} \sum \|\varphi_\Theta(b_i) - r_i\|^2 \quad (22)$$

## Learned regularizers: Adversarial

Given the training data  $\{(y_i^\delta, x_i^\dagger)\}$  the aim is to find a regularizer  $R$

$$R = \arg \min_{R \in \mathcal{R}} \sum_{i=1}^N \|\hat{x}_i - x_i^\dagger\| \quad (23)$$

s.t.

$$\hat{x}_i = \arg \min_{x \in X} \frac{1}{2} \|Ax - y_i^\delta\|^2 + R(x) \quad (24)$$

**Approach:** Train a neural network  $R_\Theta : X \rightarrow \mathbb{R}$  to discriminate between the distributions  $\mathbb{P}_n$  (good images) and  $\mathbb{P}_r$  (bad images)

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## Recent approaches

- Learned methods

$$T_{\Theta} : Y \rightarrow X \quad (25)$$

- Learned regularizers

$$T_{\Theta}(y^{\delta}) = \arg \min_{x \in X} \ell(Ax, y^{\delta}) + R_{\Theta}(x) \quad (26)$$

- Generative Networks

$$T_{\Theta}(y^{\delta}) = \arg \min_{z \in Z} \ell(A\varphi_{\Theta}(z), y^{\delta}) + R_{\Theta}(z) \quad (27)$$

## Generative networks

Consider a generative network  $\varphi_\Theta(z)$  previously trained

- $\Theta$  is fixed after the training phase
- We can obtain images by sampling  $z$

For solving inverse problems (e.g.<sup>9</sup>):

$$\hat{z} = \arg \min_z \frac{1}{2} \|A\varphi_\Theta(z) - y^\delta\|^2 + R(z) \quad (28)$$

$$\hat{x} = \varphi_\Theta(\hat{z}) \quad (29)$$

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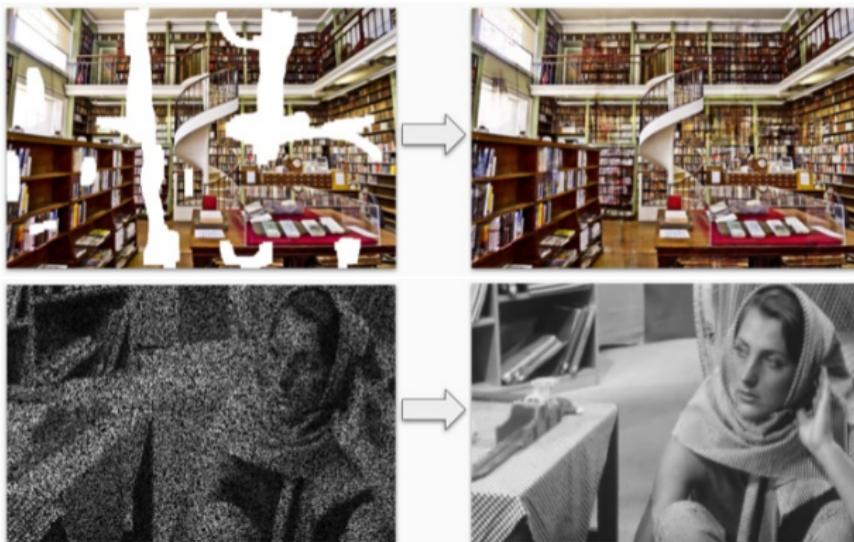
<sup>9</sup> Ashish Bora, Ajil Jalal, Eric Price, and Alexandros G. Dimakis. "Compressed Sensing using Generative Models". In: *Proceedings of the 34th International Conference on Machine Learning, ICML 2017, Sydney, NSW, Australia, 6-11 August 2017.* 2017, pp. 537–546.



## Section 3

# Deep Image Prior

## Examples <sup>10</sup>



<sup>10</sup>[https://dmitryulyanov.github.io/deep\\_image\\_prior](https://dmitryulyanov.github.io/deep_image_prior)

## Basic Idea<sup>11</sup>

Given measured noisy data

$$y^\delta = Ax^\dagger + \tau \quad (30)$$

- 1 Optimize a neural network  $\varphi_\Theta(z_0)$  with a fixed input  $z_0$

$$\hat{\Theta} = \arg \min_{\Theta} \frac{1}{2} \|A\varphi_\Theta(z_0) - y^\delta\|^2 \quad (31)$$

- 2 Set  $\hat{x} = \varphi_{\hat{\Theta}}(z_0)$  as the reconstruction

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<sup>11</sup>Dmitry Ulyanov, Andrea Vedaldi, and Victor S. Lempitsky. "Deep Image Prior". In: *CoRR* (2017). arXiv: 1711.10925.

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## Some insights

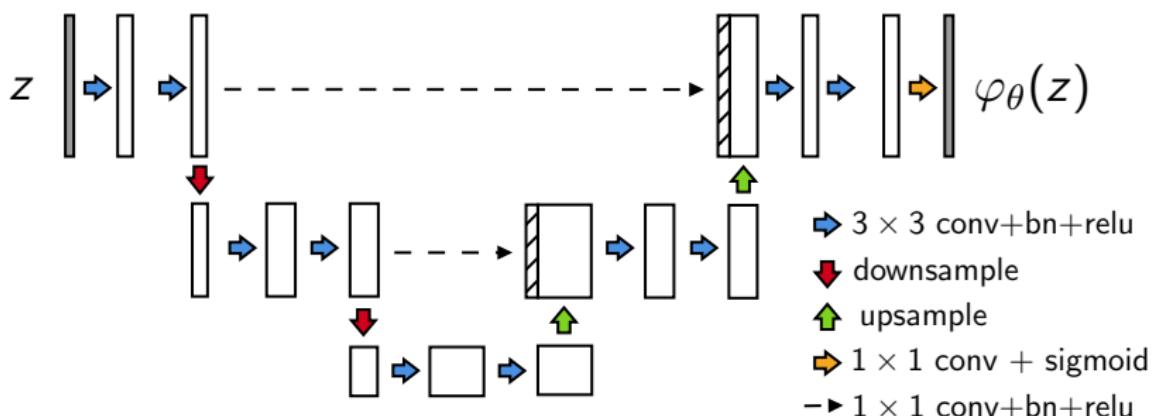
- The network  $\varphi_\Theta$  has a U-Net-like architecture
- It has enough expressive power to reproduce some noise
- Optimization method (ADAM<sup>12</sup>) with early stopping plays an important role
- Solving each instance requires training the network
- It takes a lot of time

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<sup>12</sup>Diederik P Kingma and Jimmy Ba. "Adam: A method for stochastic optimization". In: *arXiv preprint arXiv:1412.6980* (2014).

# Task dependent hyper-parameters

- U-Net-like architecture
  - Number of scales (e.g. 2, 3, 4, 5, 6, ...)
  - Filter size per scale (e.g. 3, 5, ...)
  - Number of filters per scale (e.g. 8, 16, 32, 64, 128, ...)
  - Number of filters per skip connection (e.g. 2, 4, ...)





## Section 4

# Application in Computed Tomography

## Radon transform

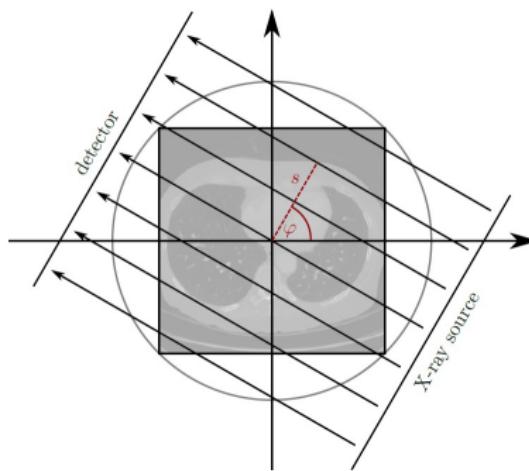


Figure: Parallel beam geometry

## Example

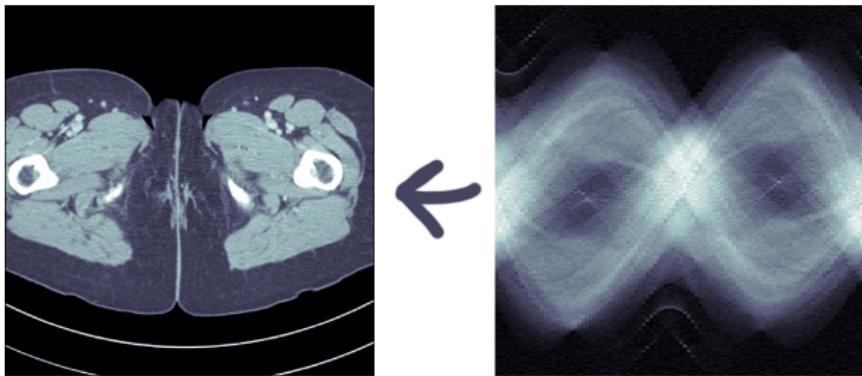


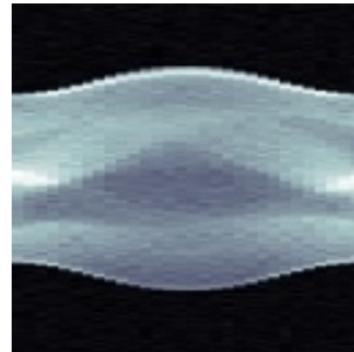
Figure: Human phantom and corresponding sinogram

## Example a): Shepp-Logan phantom

- Parallel beam geometry (30 angles, 183 detectors)
- 5% white noise
- Visualization window: [0.1, 0.4]



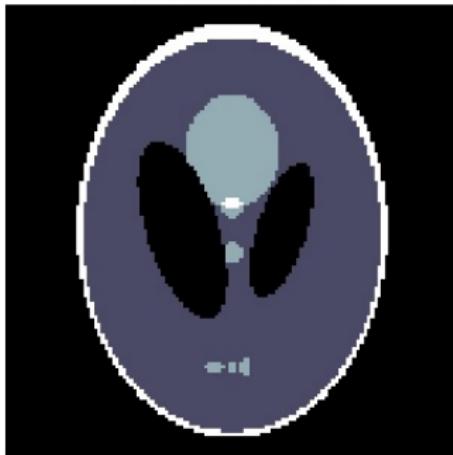
(a) Ground truth ( $128 \times 128$ )



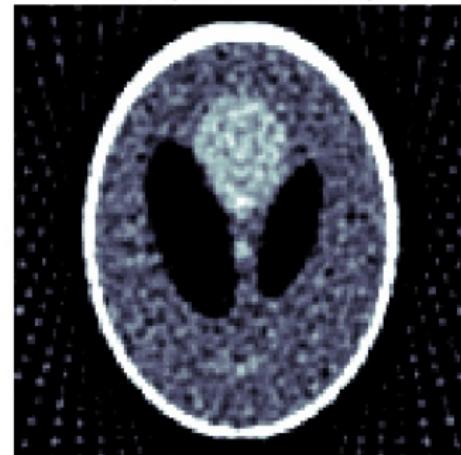
(b) Data ( $30 \times 183$ )

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Ground truth



FBP (PSNR: 19.75)



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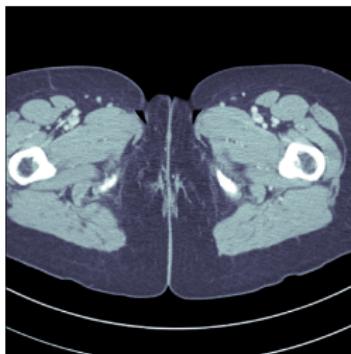


DIP (PSNR: 28.40)

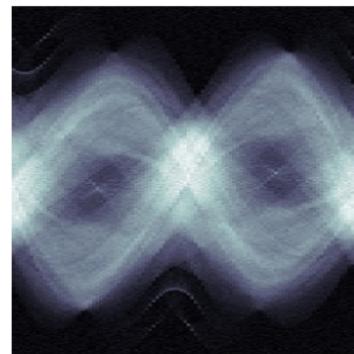


## Example b): Human phantom<sup>13</sup>

- Case i: Fan-beam geometry (100 angles, 1000 detectors)
- Case ii: Fan-beam geometry (1000 angles, 1000 detectors)
- 5% white noise



(a) Ground truth ( $512 \times 512$ )



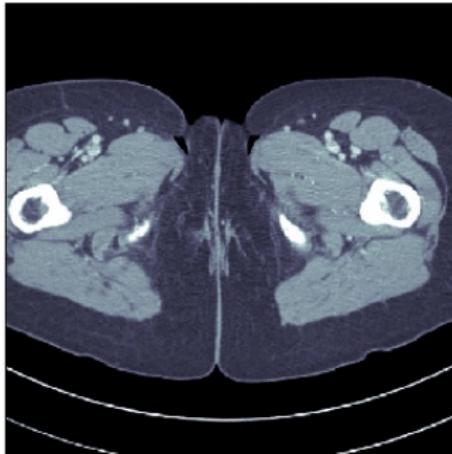
(b) Data ( $100 \times 1000$ )

<sup>13</sup> Jonas Adler and Ozan Öktem. "Learned primal-dual reconstruction". In: *IEEE transactions on medical imaging* 37.6 (2018), pp. 1322–1332.

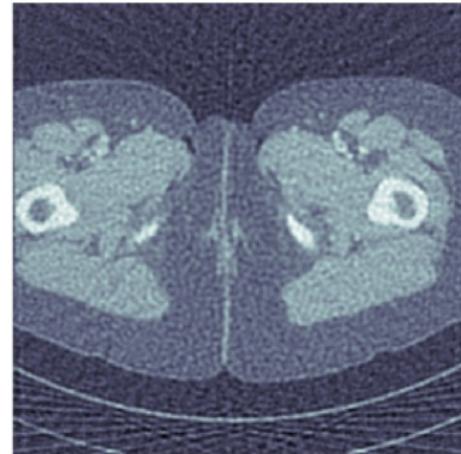
## Example b): Human phantom

Case i: 100 angles

Ground truth



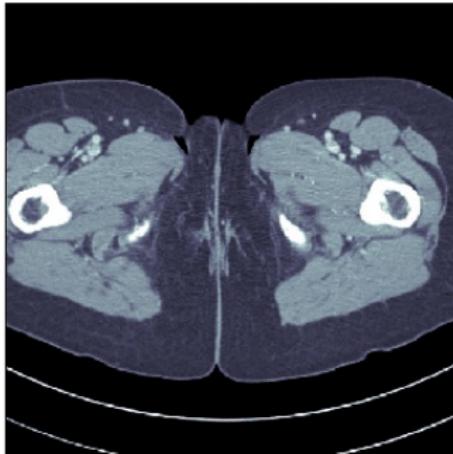
FBP (PSNR: 20.99)



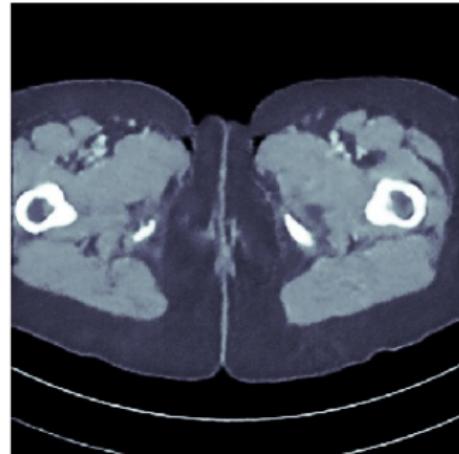
## Example b): Human phantom

Case i: 100 angles

Ground truth



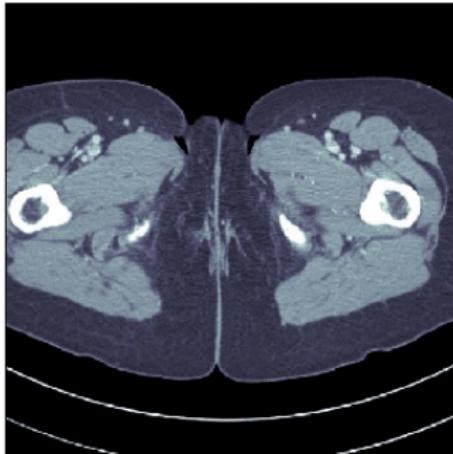
DIP (PSNR: 28.14)



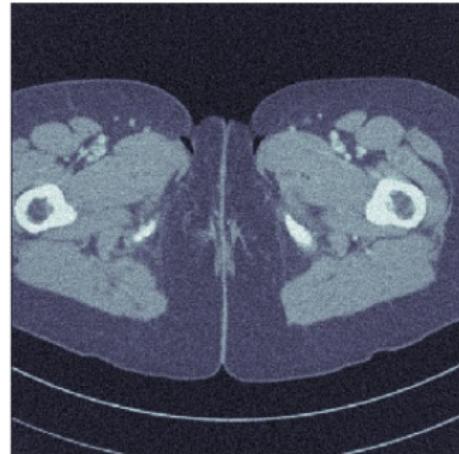
## Example b): Human phantom

Case ii: 1000 angles

Ground truth



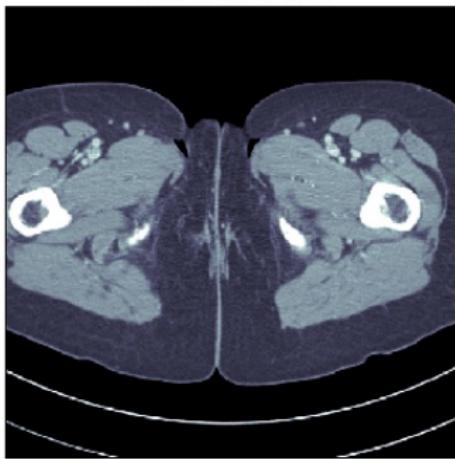
FBP (PSNR: 25.21)



## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 9.23)

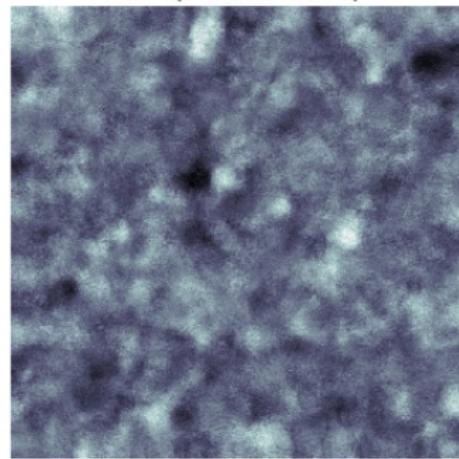


Figure: Iteration 0

## Example b): Human phantom

Case ii: 1000 angles

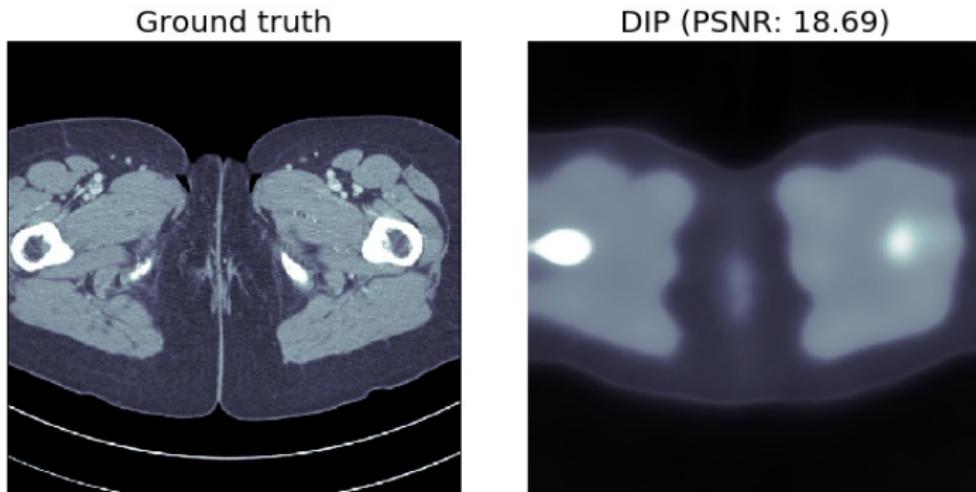


Figure: Iteration 100

## Example b): Human phantom

Case ii: 1000 angles

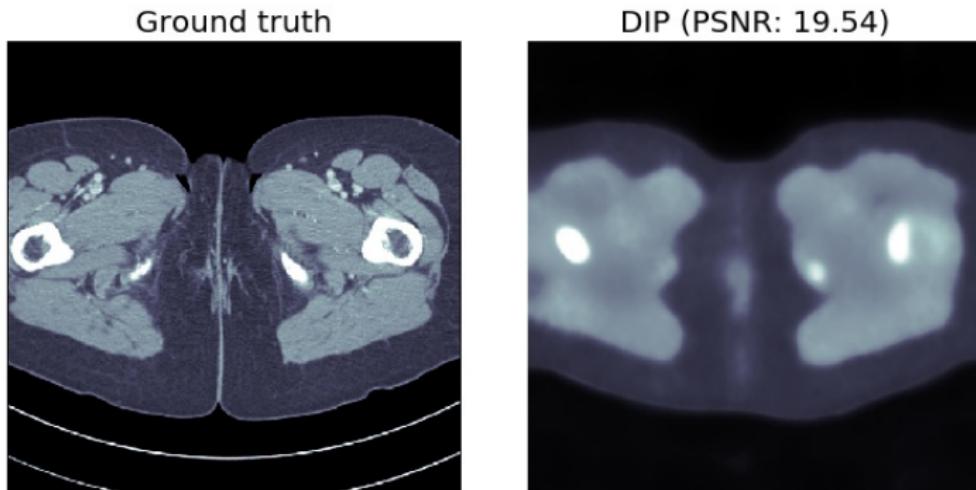


Figure: Iteration 200

## Example b): Human phantom

Case ii: 1000 angles

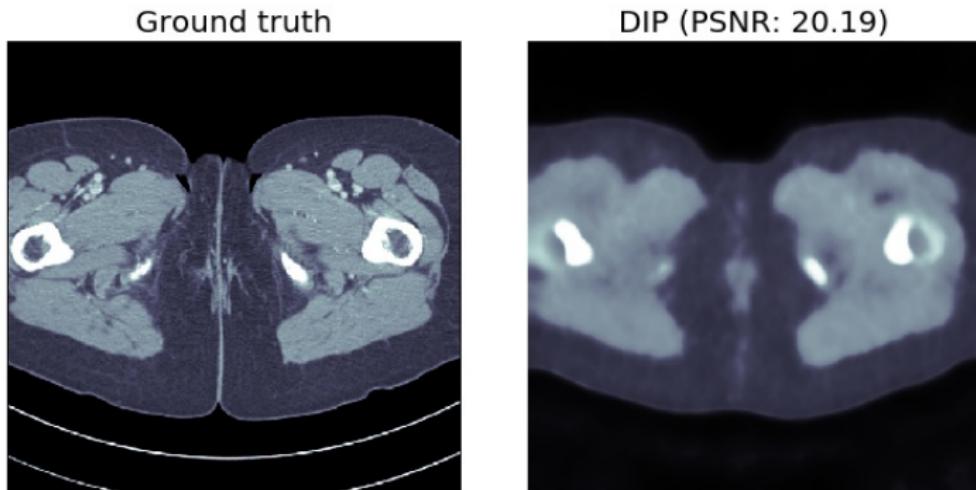


Figure: Iteration 300

## Example b): Human phantom

Case ii: 1000 angles

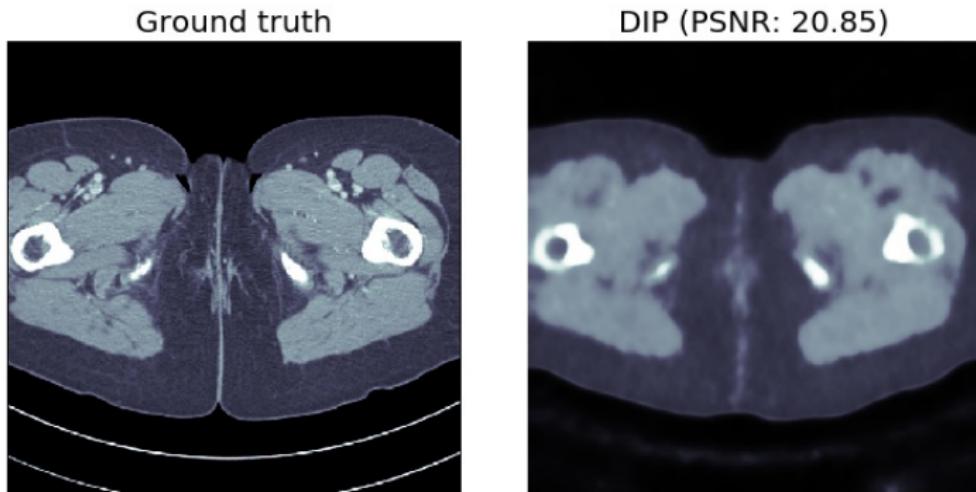


Figure: Iteration 400

## Example b): Human phantom

Case ii: 1000 angles

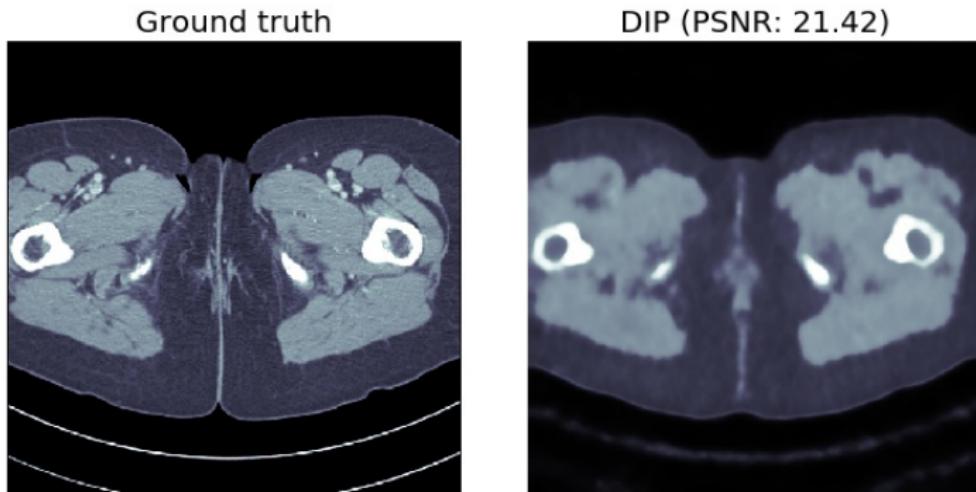


Figure: Iteration 500

## Example b): Human phantom

Case ii: 1000 angles

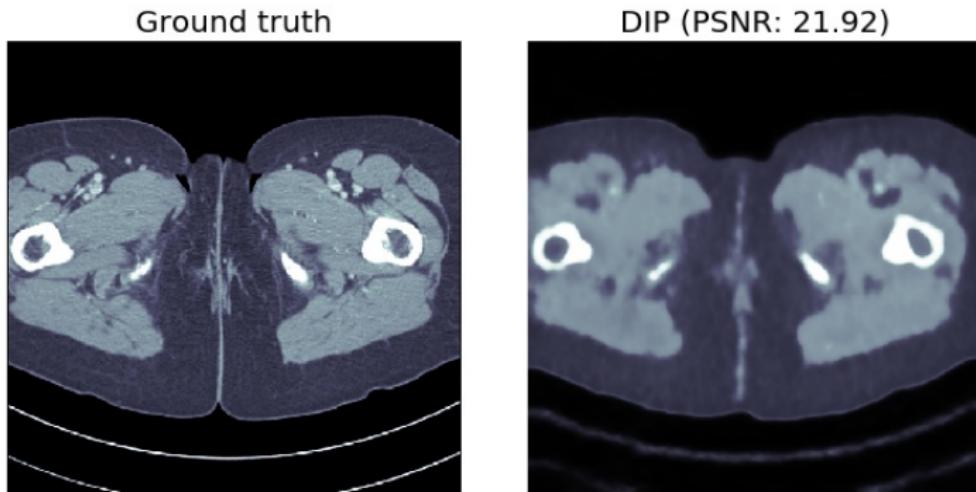


Figure: Iteration 600

## Example b): Human phantom

Case ii: 1000 angles

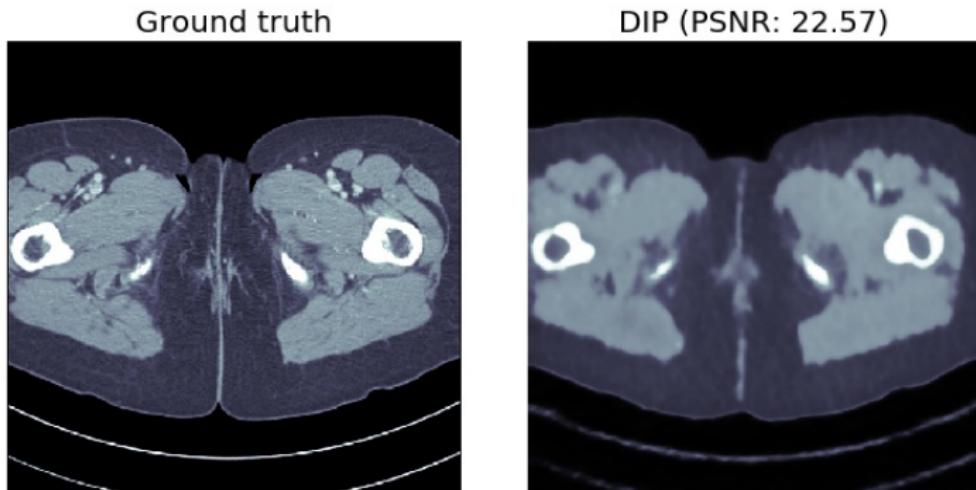


Figure: Iteration 700

## Example b): Human phantom

Case ii: 1000 angles

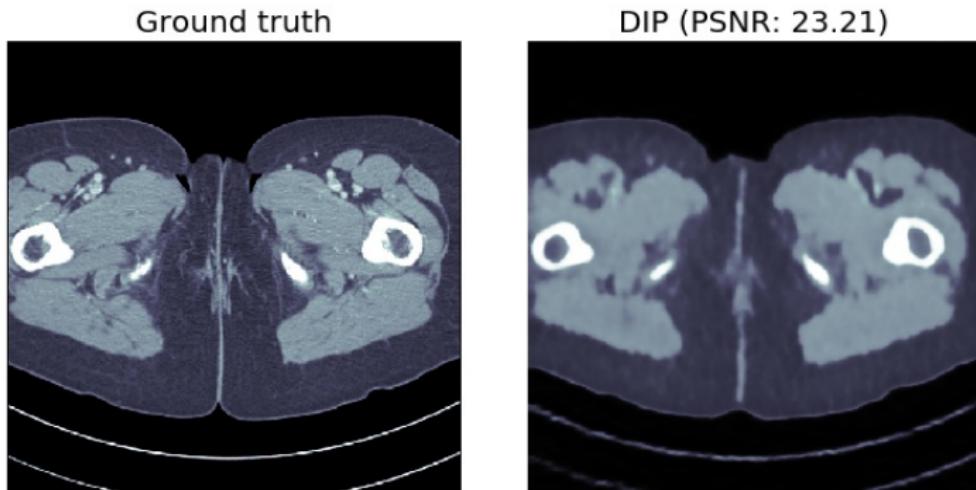


Figure: Iteration 800

## Example b): Human phantom

Case ii: 1000 angles

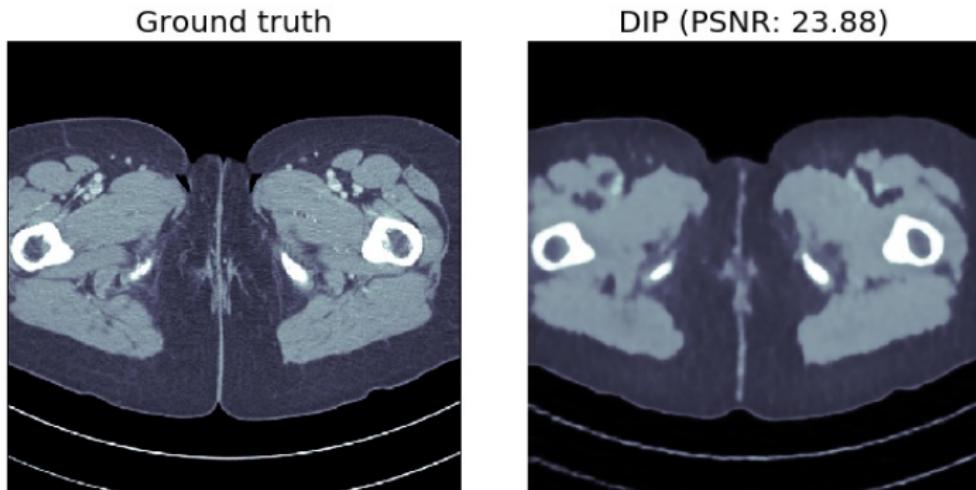


Figure: Iteration 900

## Example b): Human phantom

Case ii: 1000 angles

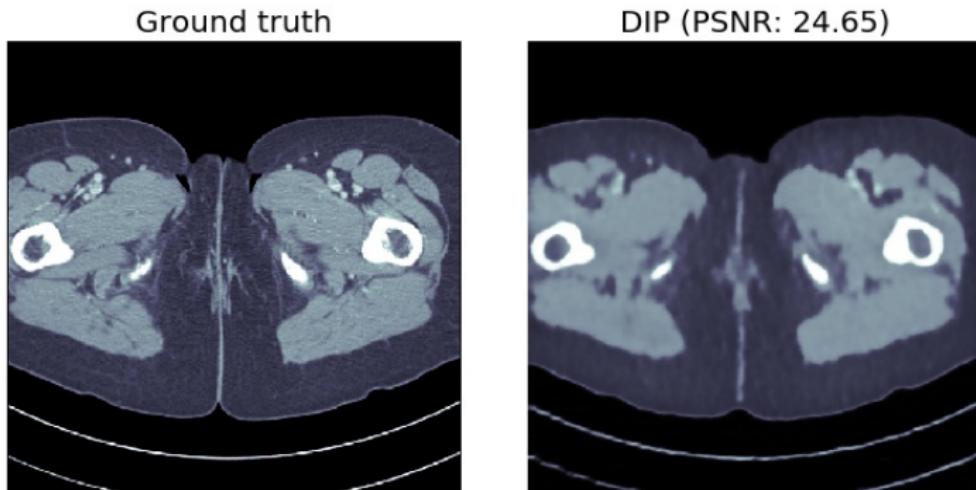


Figure: Iteration 1000

## Example b): Human phantom

Case ii: 1000 angles

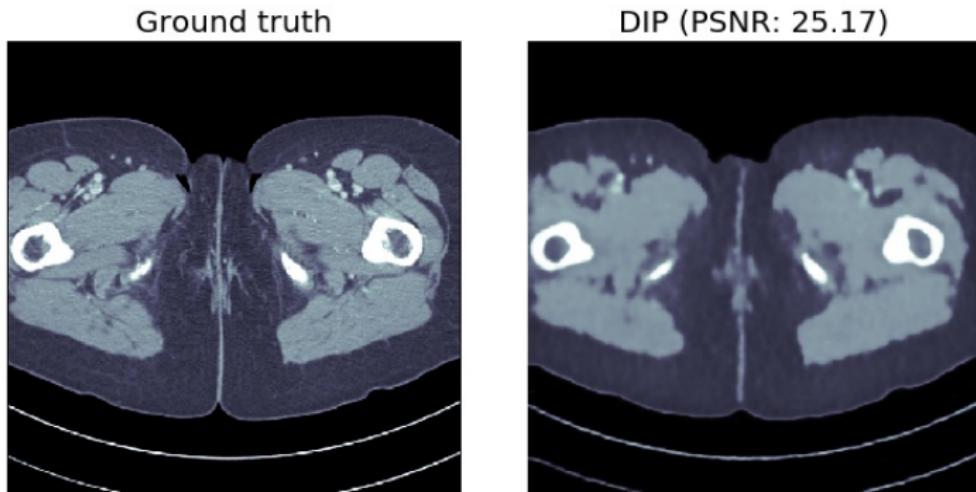


Figure: Iteration 1100

## Example b): Human phantom

Case ii: 1000 angles

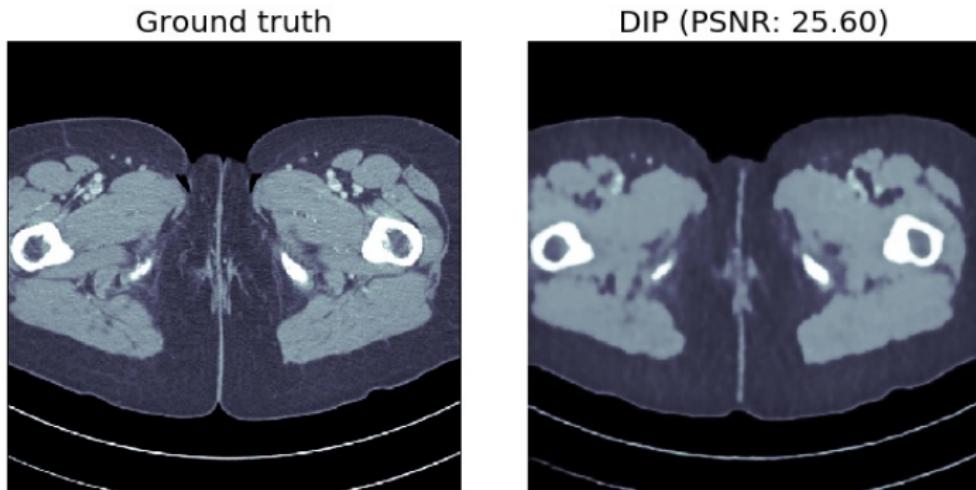


Figure: Iteration 1200

## Example b): Human phantom

Case ii: 1000 angles

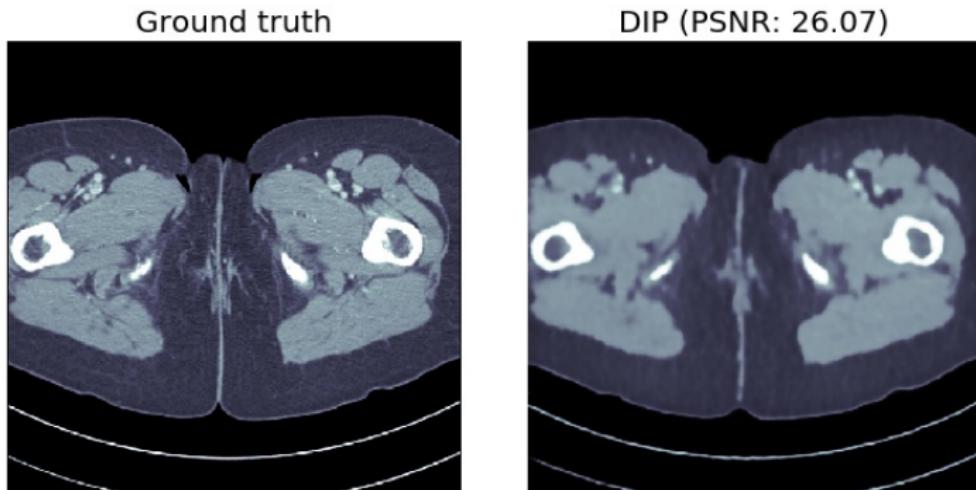


Figure: Iteration 1300

## Example b): Human phantom

Case ii: 1000 angles

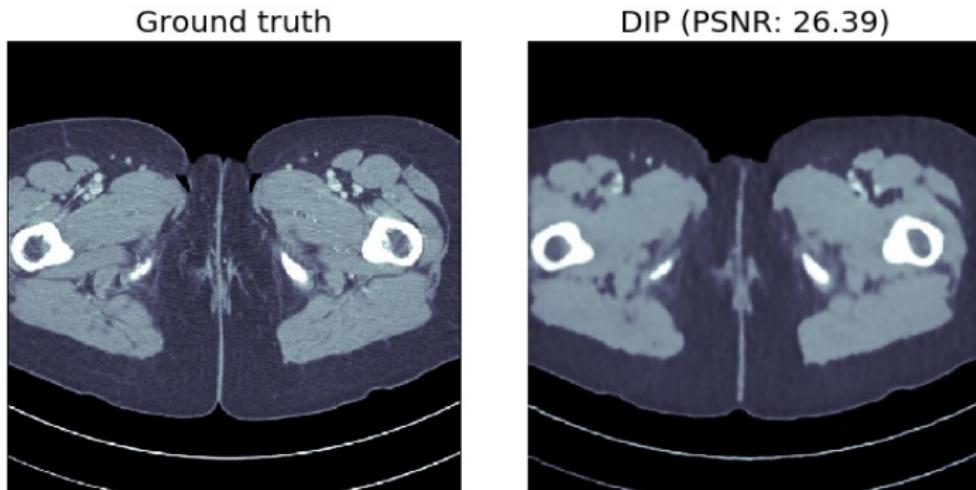


Figure: Iteration 1400

## Example b): Human phantom

Case ii: 1000 angles

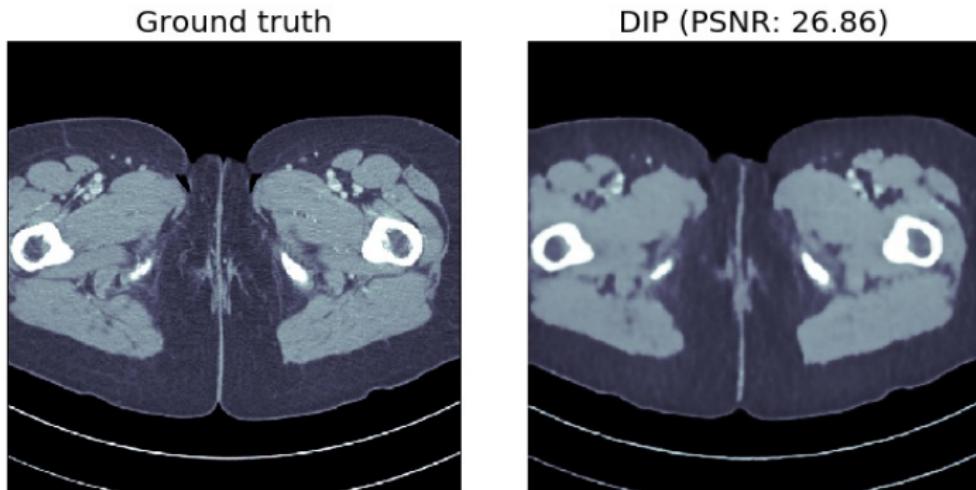


Figure: Iteration 1500

## Example b): Human phantom

Case ii: 1000 angles

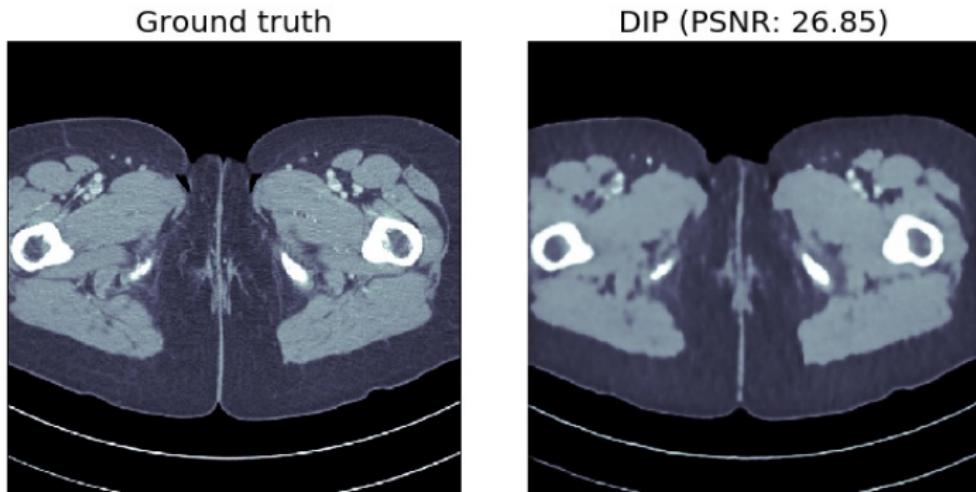
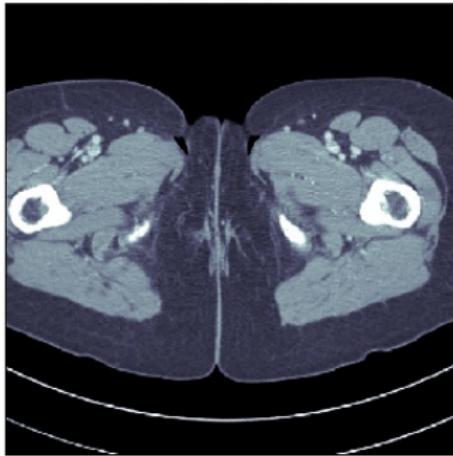


Figure: Iteration 1600

## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 27.35)

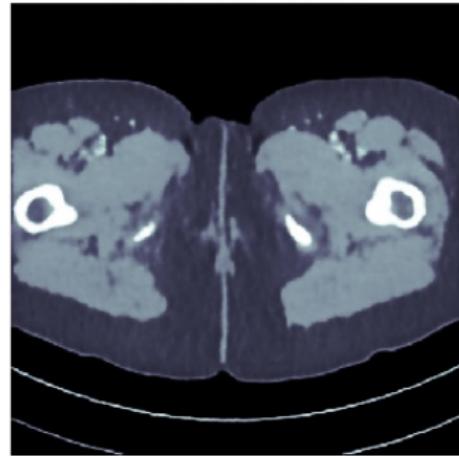
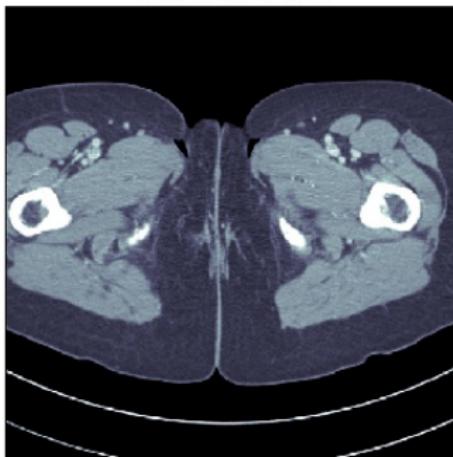


Figure: Iteration 1700

## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 27.49)

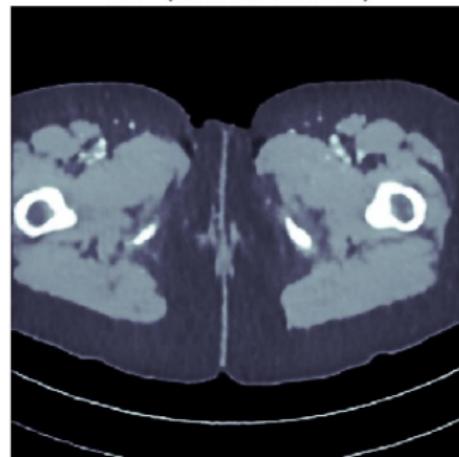


Figure: Iteration 1800

## Example b): Human phantom

Case ii: 1000 angles

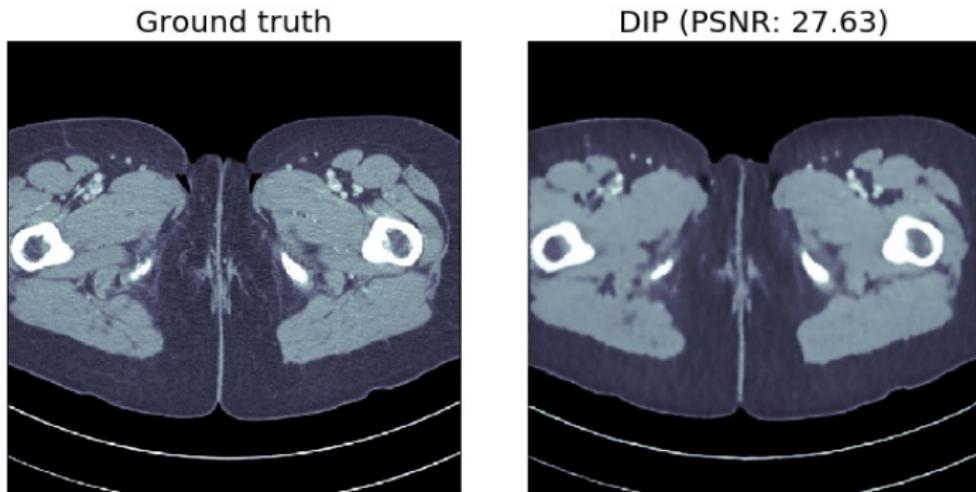


Figure: Iteration 1900

## Example b): Human phantom

Case ii: 1000 angles

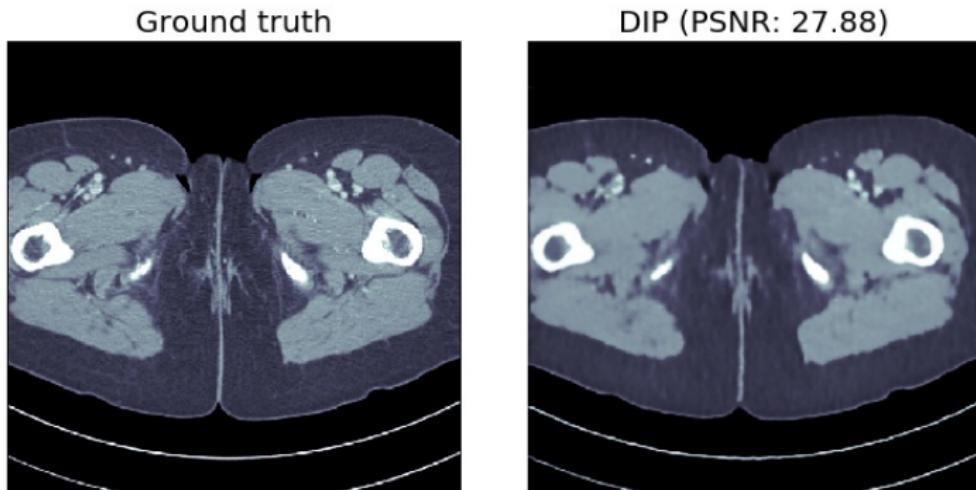


Figure: Iteration 2000

## Example b): Human phantom

Case ii: 1000 angles

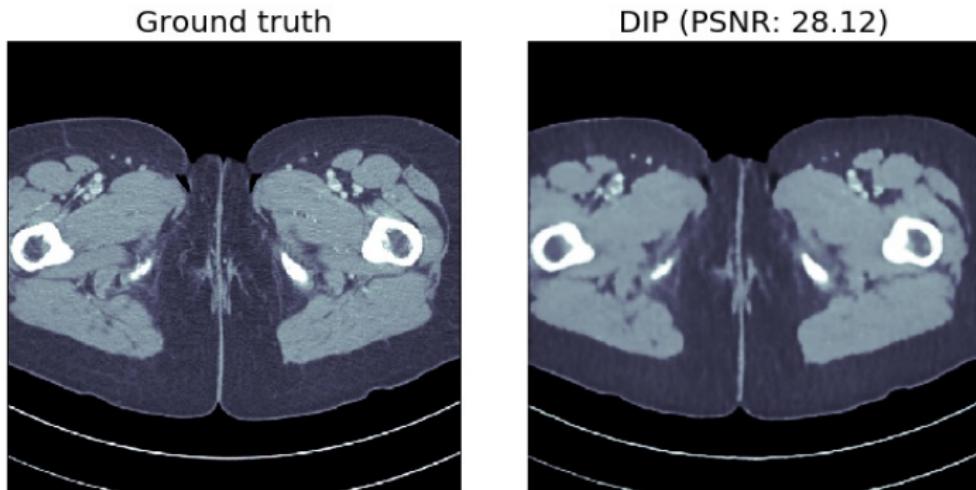


Figure: Iteration 2100

## Example b): Human phantom

Case ii: 1000 angles

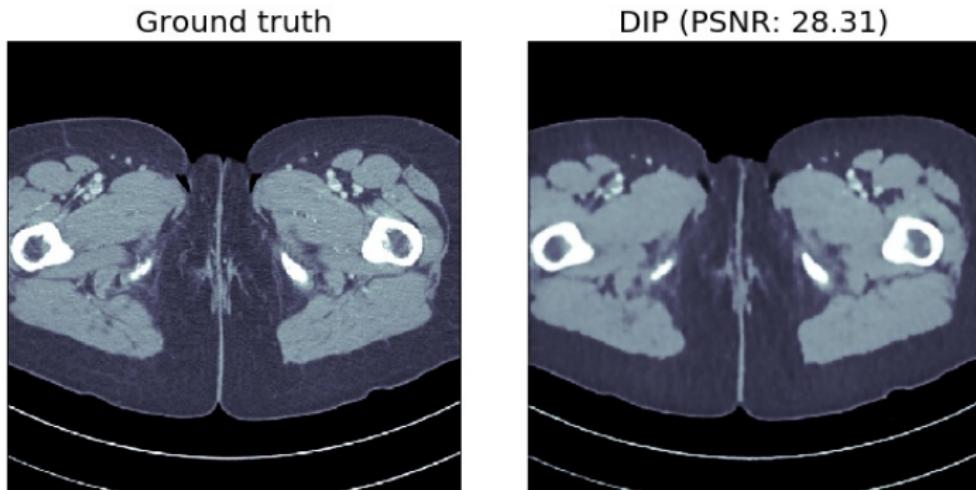


Figure: Iteration 2200

## Example b): Human phantom

Case ii: 1000 angles

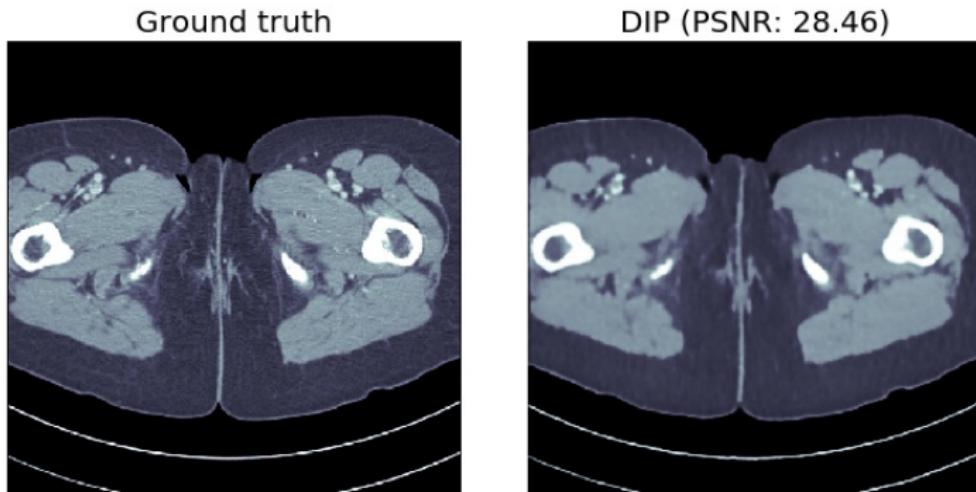
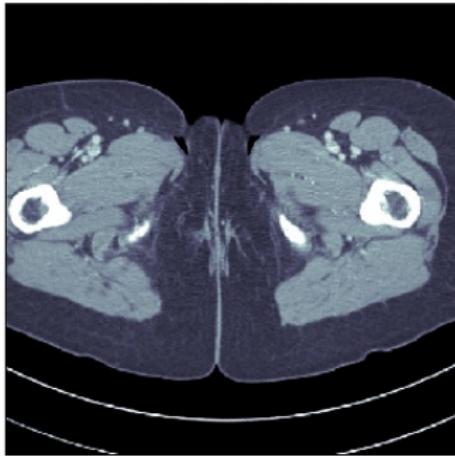


Figure: Iteration 2300

## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 28.68)

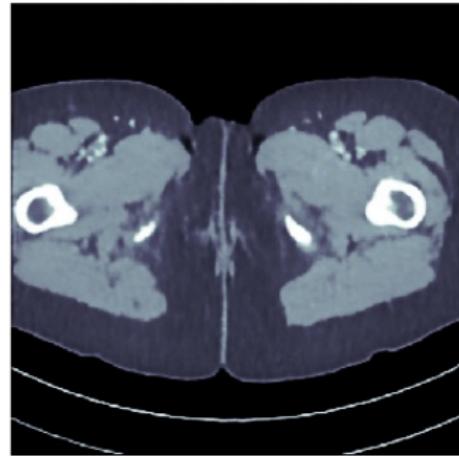


Figure: Iteration 2400

## Example b): Human phantom

Case ii: 1000 angles

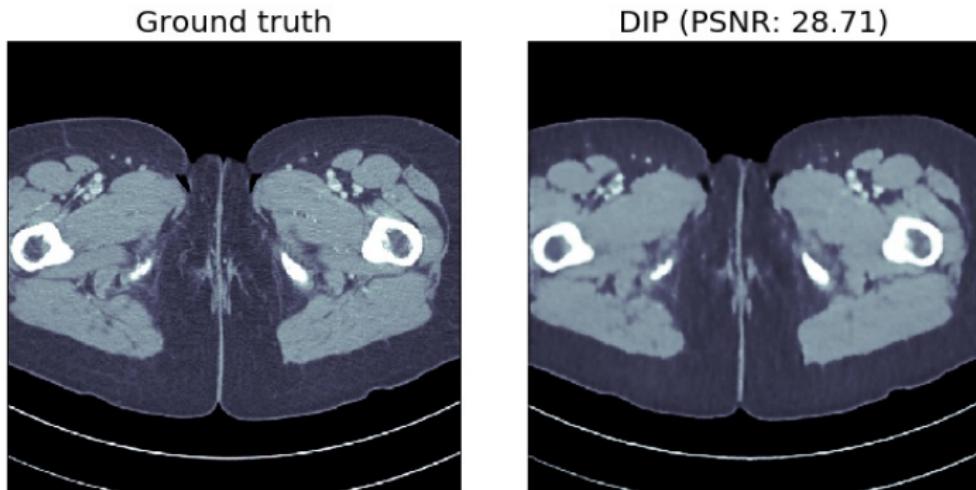


Figure: Iteration 2500

## Example b): Human phantom

Case ii: 1000 angles

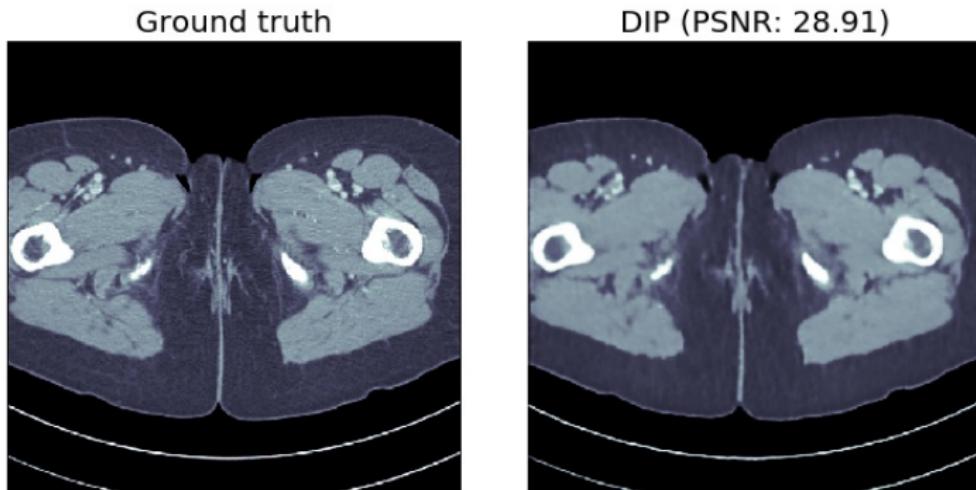


Figure: Iteration 2600

## Example b): Human phantom

Case ii: 1000 angles

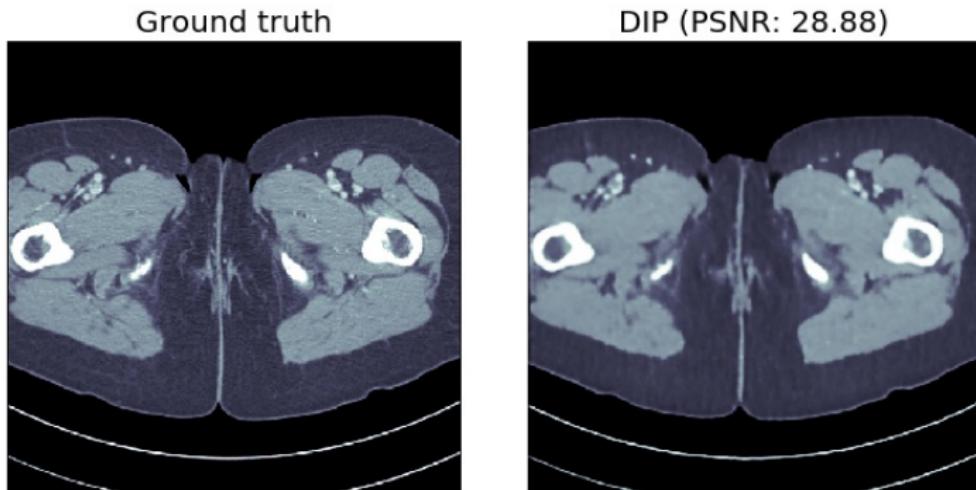


Figure: Iteration 2700

## Example b): Human phantom

Case ii: 1000 angles

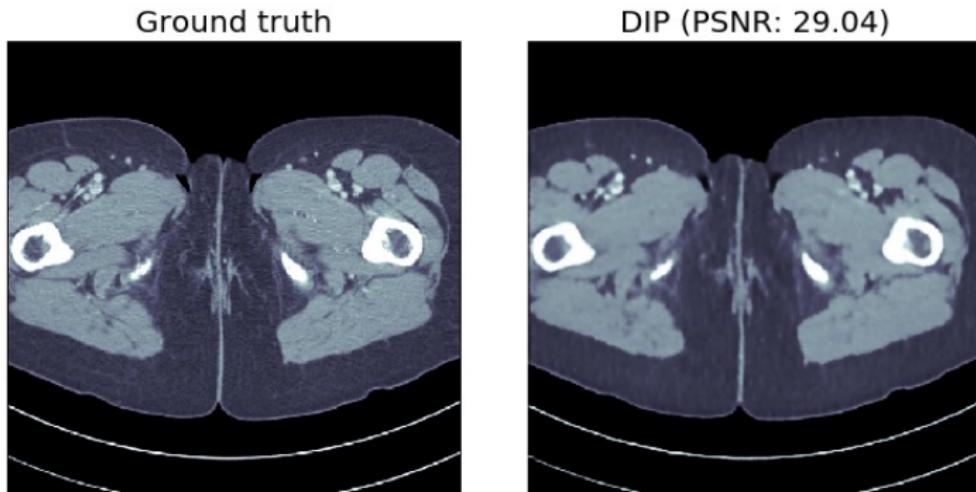


Figure: Iteration 2800

## Example b): Human phantom

Case ii: 1000 angles

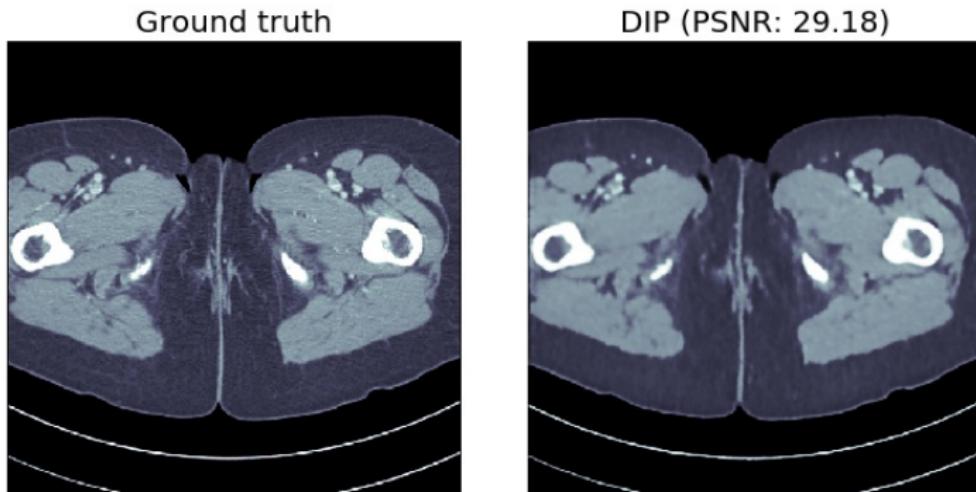
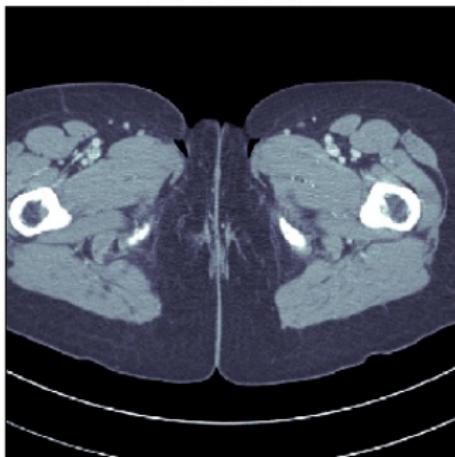


Figure: Iteration 2900

## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 29.34)

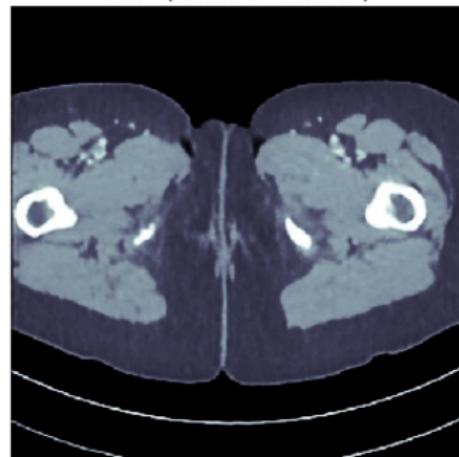


Figure: Iteration 3000

## Example b): Human phantom

Case ii: 1000 angles

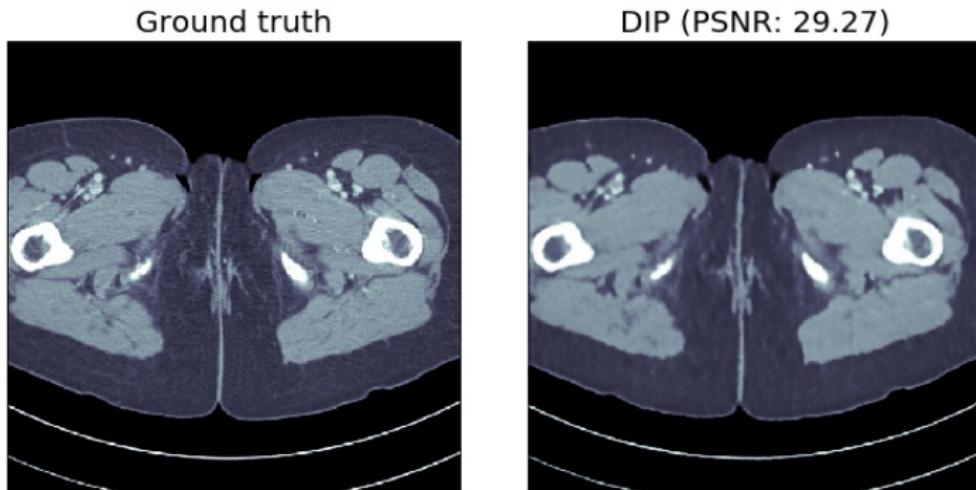


Figure: Iteration 3100

## Example b): Human phantom

Case ii: 1000 angles

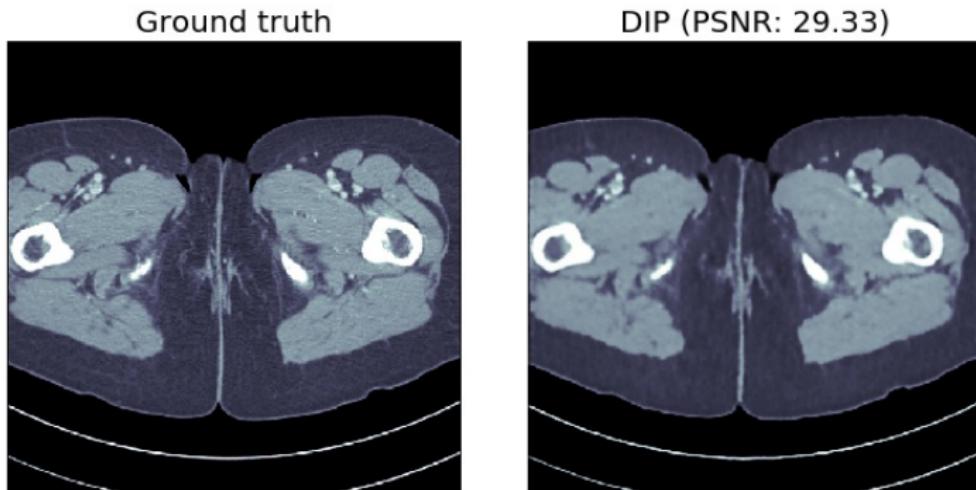


Figure: Iteration 3200

## Example b): Human phantom

Case ii: 1000 angles

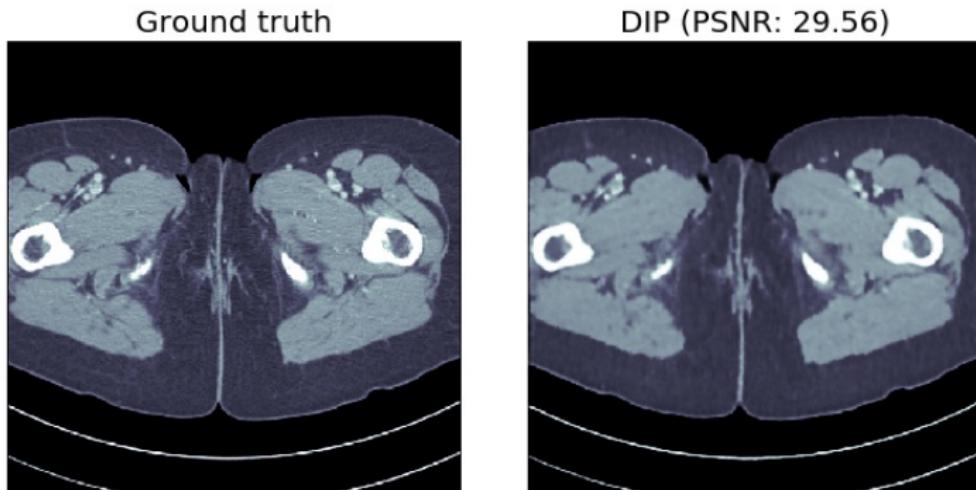


Figure: Iteration 3300

## Example b): Human phantom

Case ii: 1000 angles

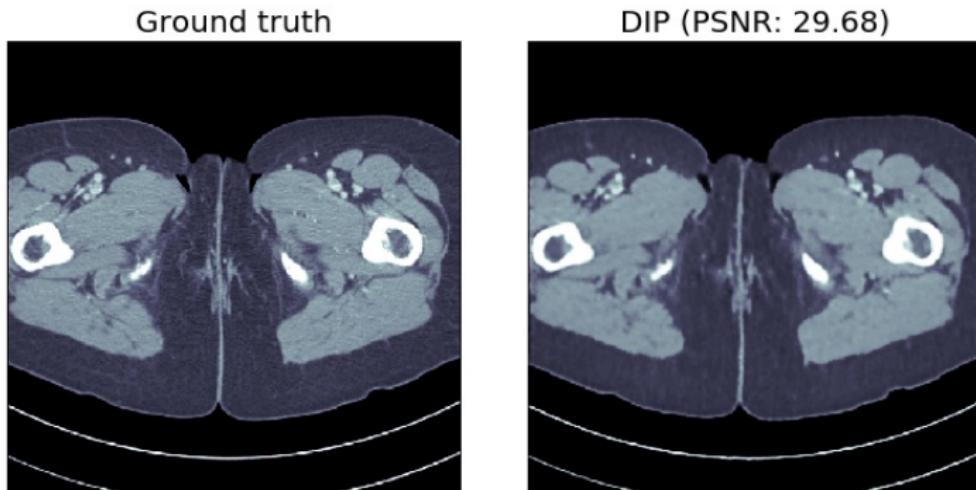


Figure: Iteration 3400

## Example b): Human phantom

Case ii: 1000 angles

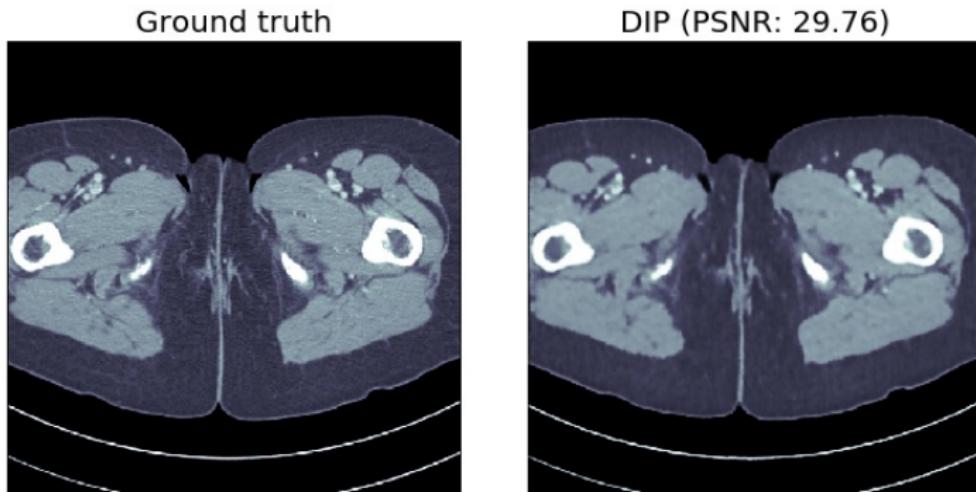
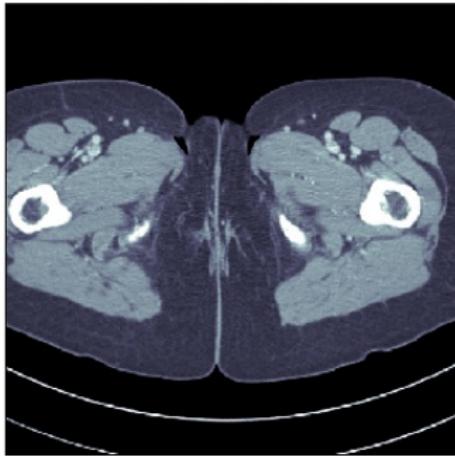


Figure: Iteration 3500

## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 29.80)

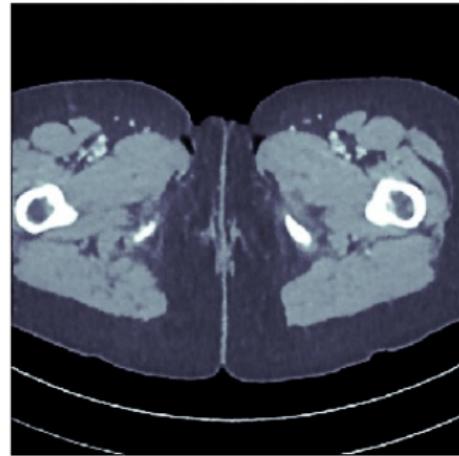


Figure: Iteration 3600

## Example b): Human phantom

Case ii: 1000 angles

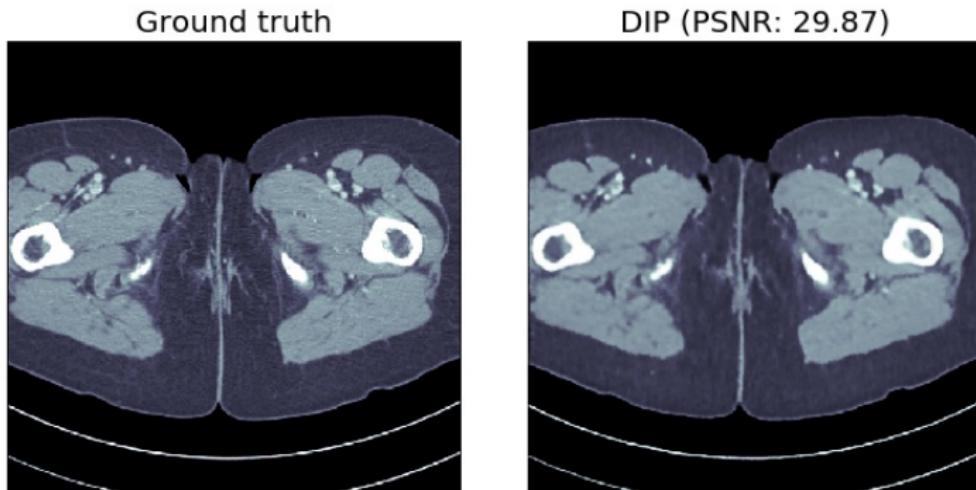


Figure: Iteration 3700

## Example b): Human phantom

Case ii: 1000 angles

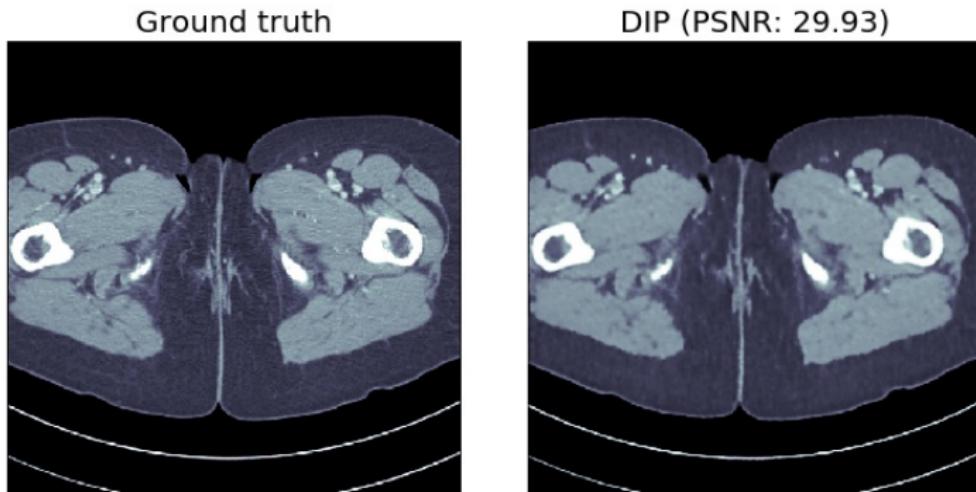


Figure: Iteration 3800

## Example b): Human phantom

Case ii: 1000 angles

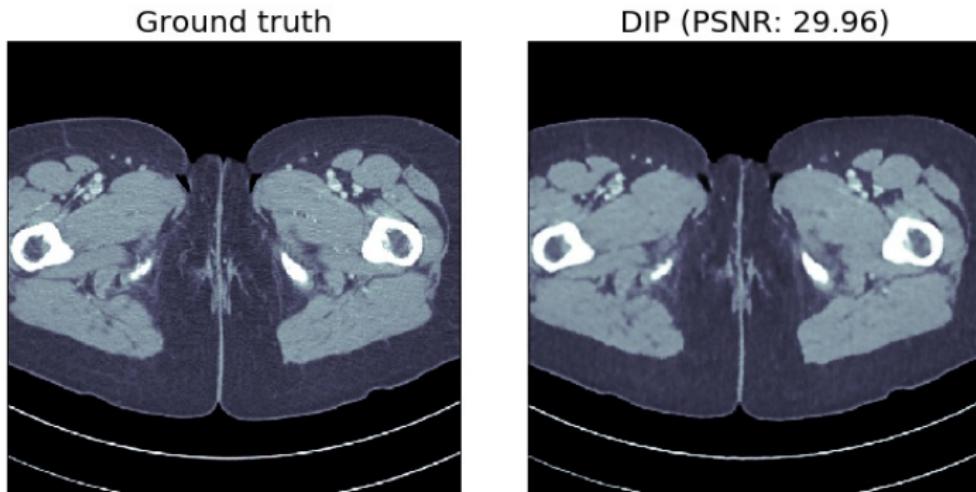


Figure: Iteration 3900

## Example b): Human phantom

Case ii: 1000 angles

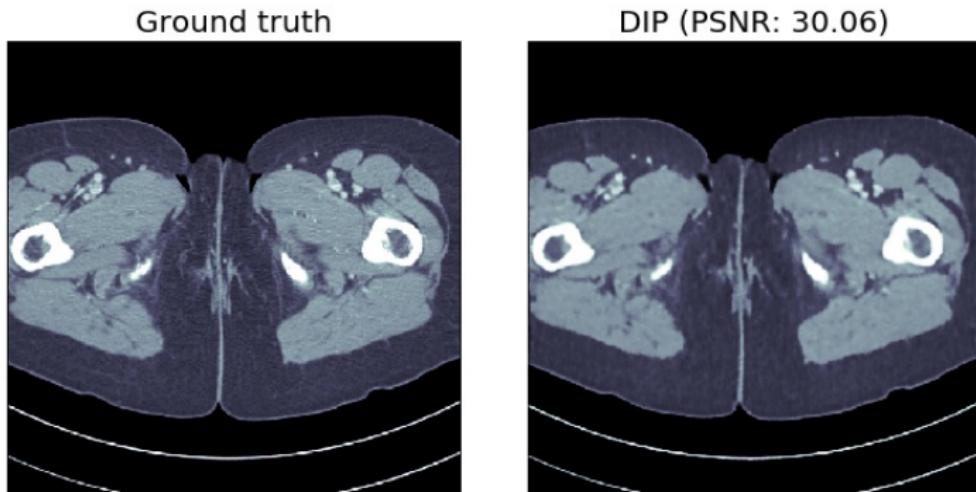


Figure: Iteration 4000

## Example b): Human phantom

Case ii: 1000 angles

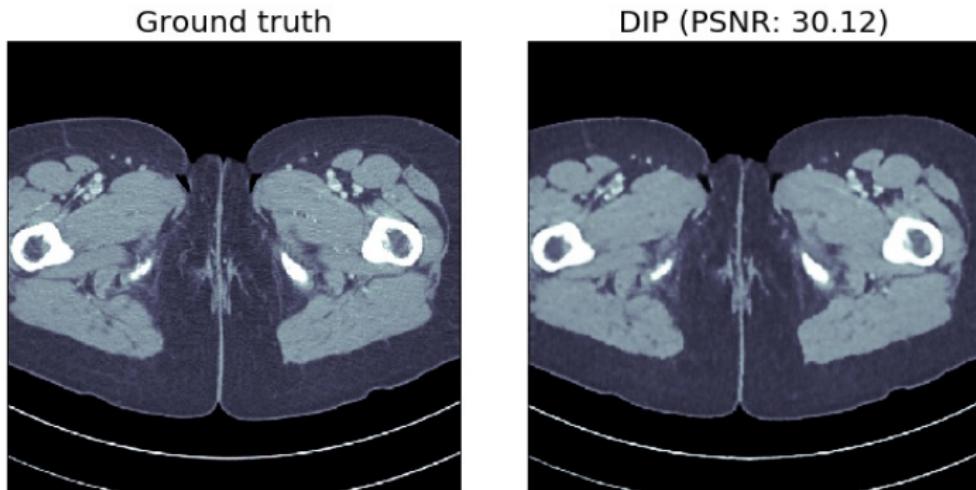


Figure: Iteration 4100

## Example b): Human phantom

Case ii: 1000 angles

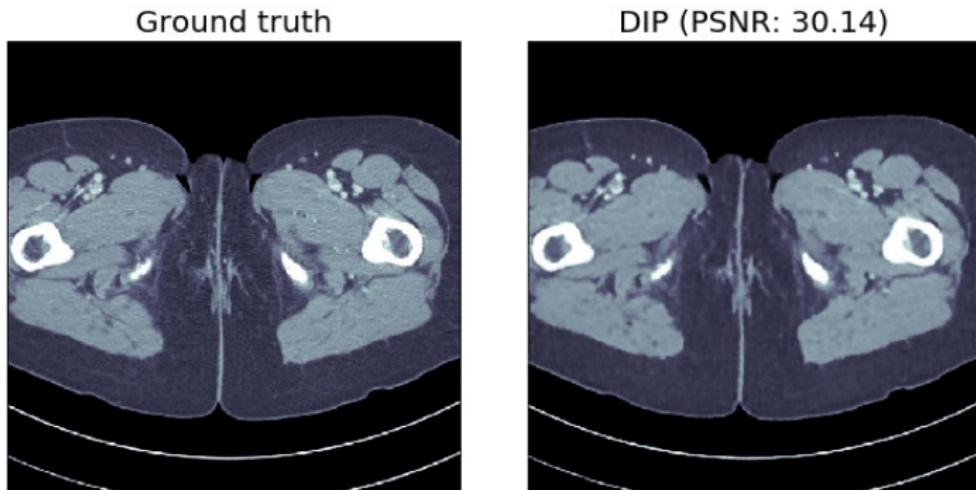


Figure: Iteration 4200

## Example b): Human phantom

Case ii: 1000 angles

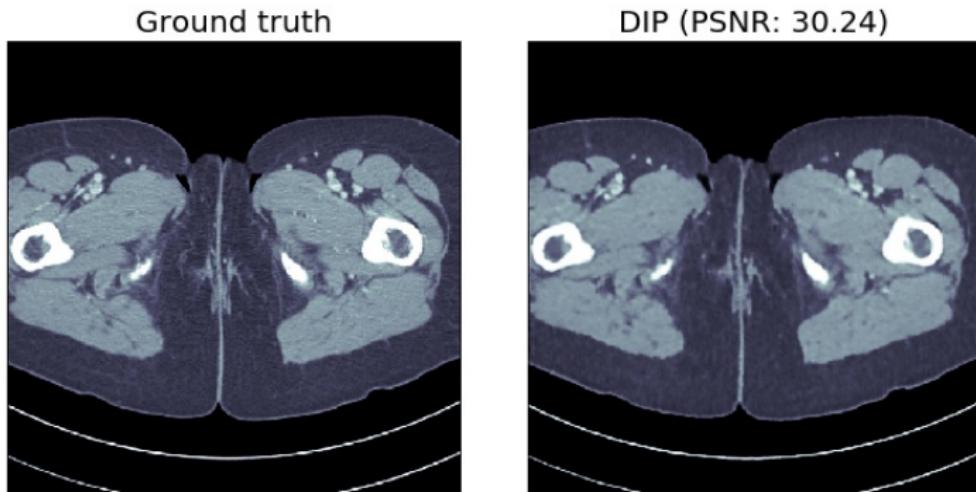


Figure: Iteration 4300

## Example b): Human phantom

Case ii: 1000 angles

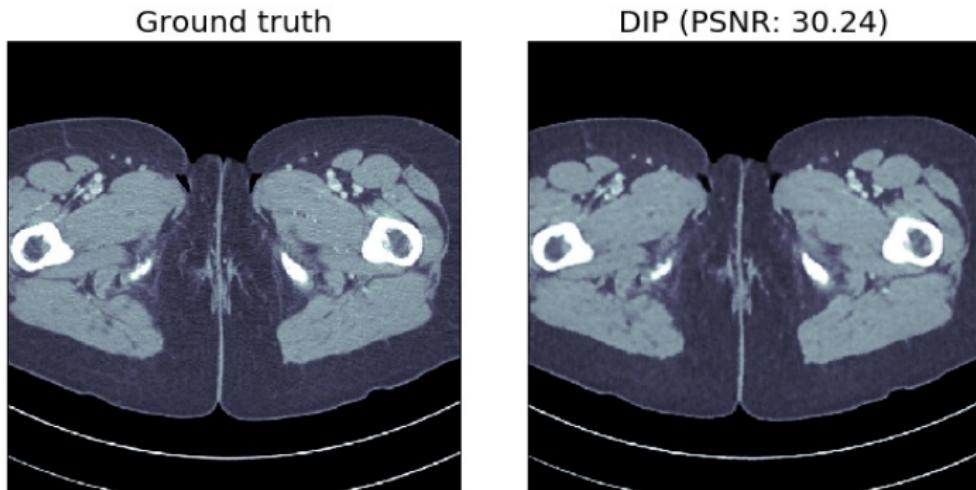


Figure: Iteration 4400

## Example b): Human phantom

Case ii: 1000 angles

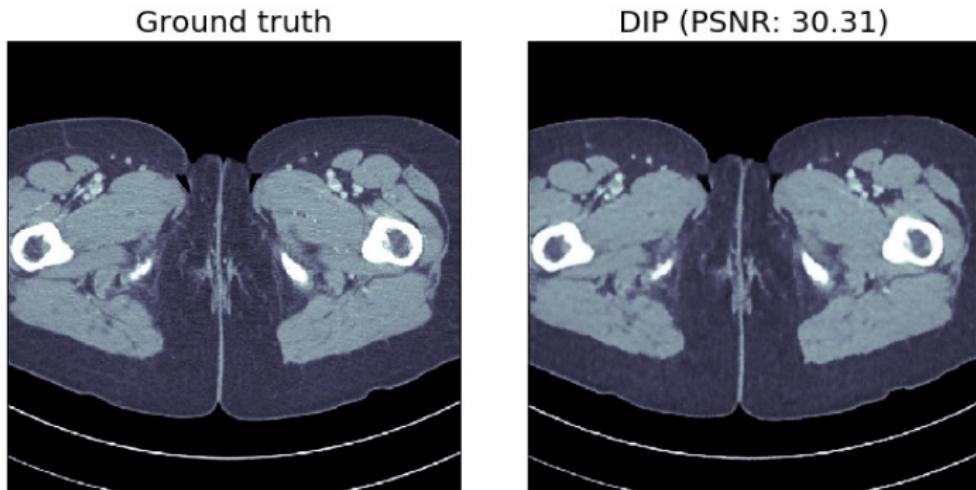


Figure: Iteration 4500

## Example b): Human phantom

Case ii: 1000 angles

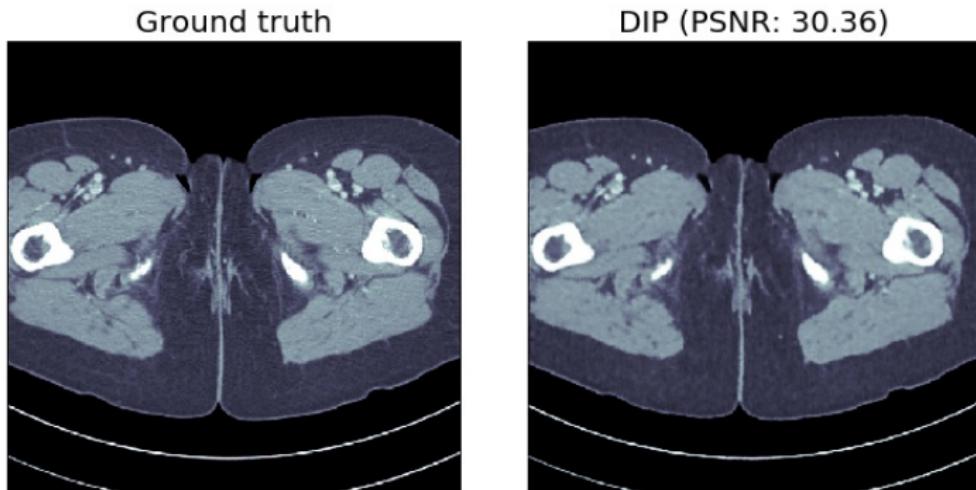


Figure: Iteration 4600

## Example b): Human phantom

Case ii: 1000 angles

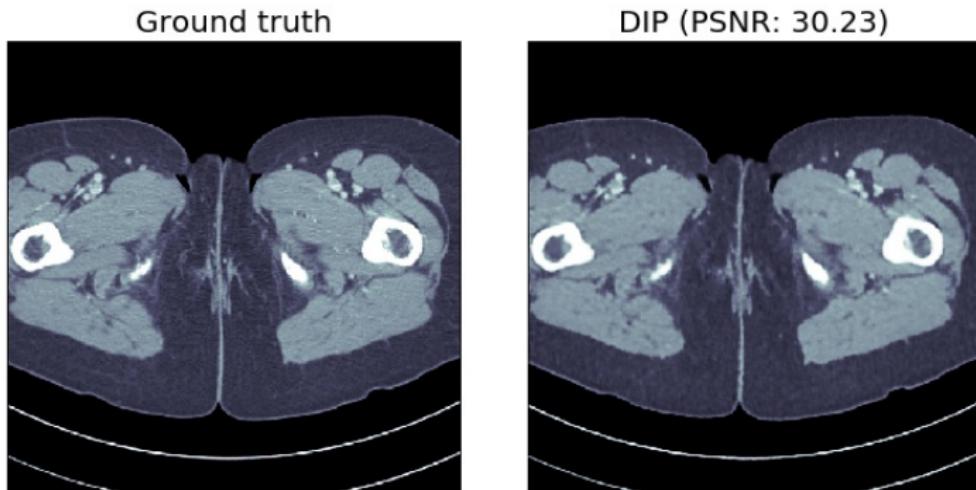


Figure: Iteration 4700

## Example b): Human phantom

Case ii: 1000 angles

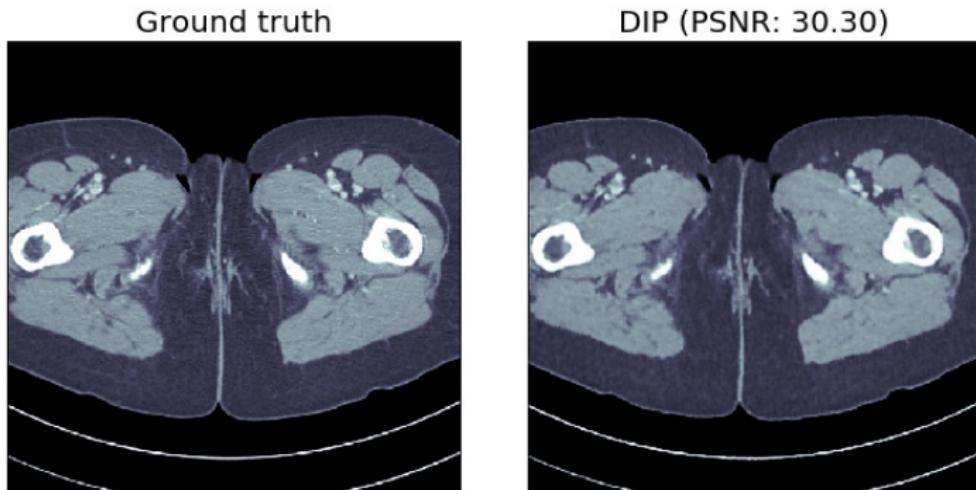


Figure: Iteration 4800

## Example b): Human phantom

Case ii: 1000 angles

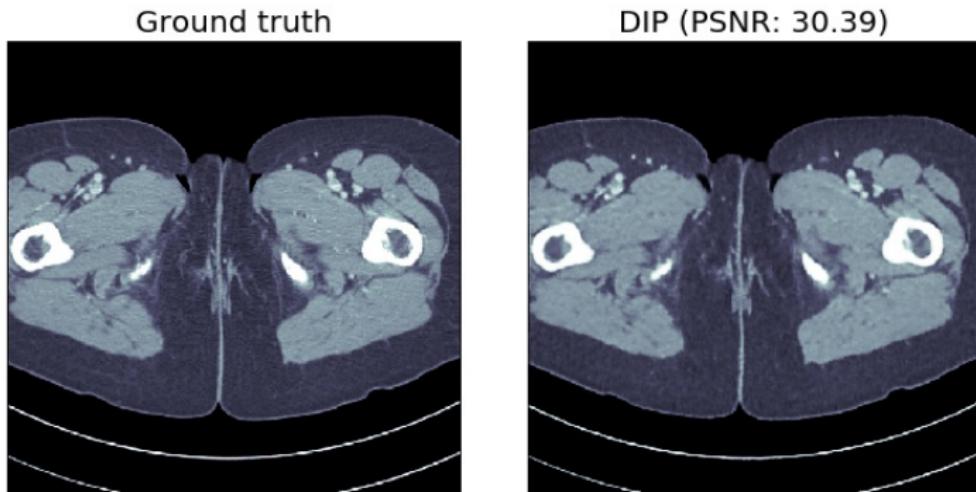


Figure: Iteration 4900

## Example b): Human phantom

Case ii: 1000 angles

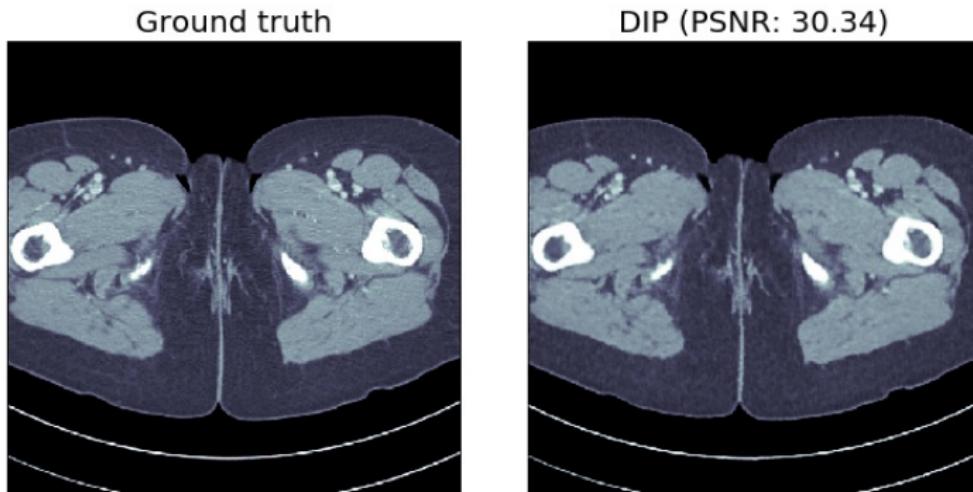
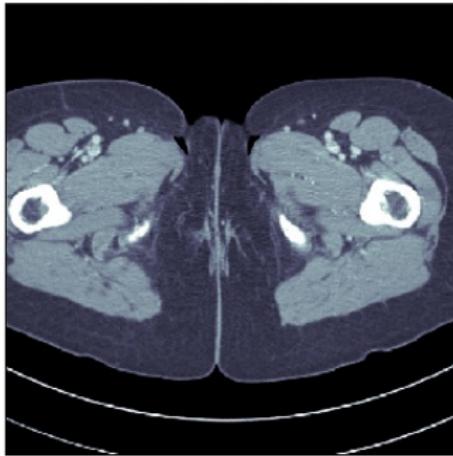


Figure: Iteration 5000

## Example b): Human phantom

Case ii: 1000 angles

Ground truth



DIP (PSNR: 31.69)

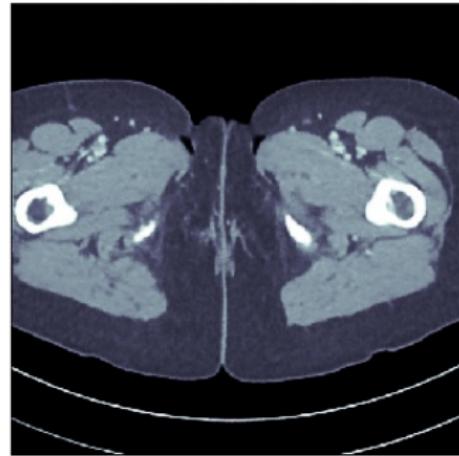


Figure: Final result

## Example b): Human phantom

Case ii: 1000 angles (Running time  $\approx 7$  min)

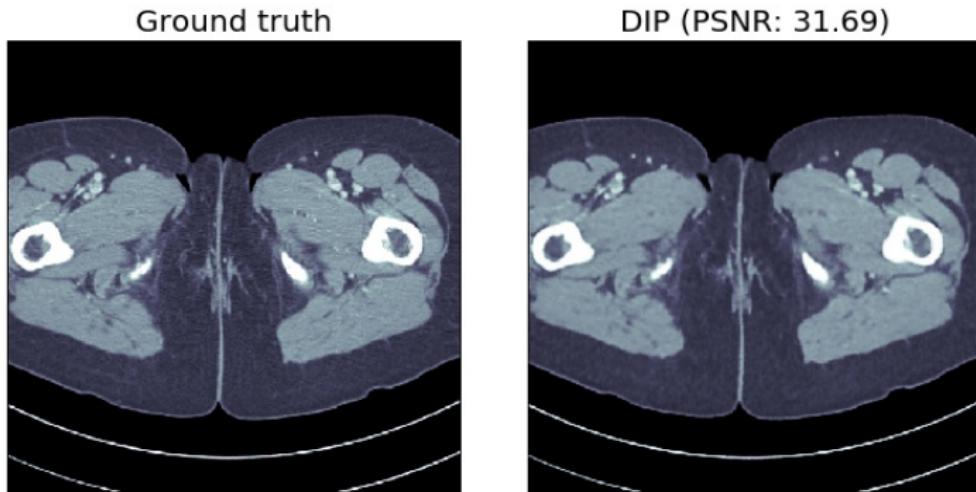


Figure: Final result

## Implementation

### Libraries:

- DIP source code<sup>14</sup>
- Operator Discretization Library (ODL)<sup>15</sup>

### Training parameters:

- Iterations: 5000
- Learning rate:  $10^{-3}$
- Regularization noise:  $10^{-2}$

### Architecture:

- Number of scales: 5
- Filter size per scale: 3
- Number of filters per scale: 128
- Number of filters per skip connection: 4

### Hardware:

- Nvidia GeForce GTX 1080

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<sup>14</sup><https://github.com/DmitryUlyanov/deep-image-prior>

<sup>15</sup><https://github.com/odlgroup/odl>



Thanks!