Jeffrey A. Fessler

EECS Department, BME Department, Dept. of Radiology University of Michigan

http://web.eecs.umich.edu/~fessler

PETRIC Workshop / IEEE NSS MIC 2024-11-02

Acknowledgments: Jason Hu, Bowen Song, Xiaojian Xu, Liyue Shen

> arXiv 2406.02462 arXiv 2406.10211

### Dall-E 3's view of this talk





#### Introduction

Inverse problems

Generative models

Score matching / diffusion models

#### Patch-based models

Non-overlapping patch model Patch Diffusion Inverse Solver (PaDIS) CT reconstruction results Summary

Book

**Bibliography** 

► Applications: compressed sensing MRI, sparse-view CT, PET, inpainting, ...
All have *linear* forward models for data:

$$y = Ax + \varepsilon$$

y: sensor data (e.g., sinogram)

A: wide system matrix (known)

x: latent image (or image series in dynamic problems)

 $\varepsilon$ : noise with known distribution provides likelihood p(y|x)

► Maximum-likelihood estimation (physics-based fitting) is usually non-unique:

$$\hat{\mathbf{x}} = \arg\max_{\mathbf{x}} \log p(\mathbf{y}|\mathbf{x}) = \underbrace{\arg\min_{\mathbf{x}} \|\mathbf{A}\mathbf{x} - \mathbf{y}\|_{2}^{2}}_{\mathbf{x}}$$
(for gaussian noise)

▶ Minimum-norm least-squares solution is unique but usually impractical or useless:

$$\hat{\pmb{x}} = \pmb{A}^{+} \pmb{y} = \pmb{y}$$
 for inpainting problem

hand-crafted regularizers:

$$\hat{\boldsymbol{x}} = \arg\min_{\boldsymbol{x}} - \log p(\boldsymbol{y}|\boldsymbol{x}) + R(\boldsymbol{x}) = \arg\min_{\boldsymbol{x}} \frac{1}{2\sigma_{\varepsilon}^2} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_2^2 + R(\boldsymbol{x})$$

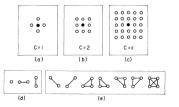
black-box data-driven supervised methods:

$$\mathbf{A}^+\mathbf{y} o \boxed{\mathsf{NN}} o \hat{\mathbf{x}}$$

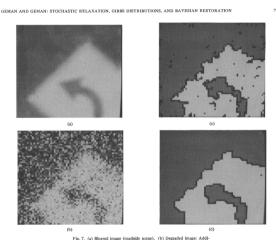
- unrolled deep learning methods (PNP, RED, MoDL, ...)
- ▶ Bayesian methods (e.g., MAP) based on a prior p(x), lately (?) relabeled as generative models

#### Markov random field models

(e.g.) Geman & Geman 1984 [1]



Mostly for inference?



tive noise. (c) Restoration including line process; 100 iterations. (d) Restoration including line process; 1000 iterations.

## Long history of generative models and inverse problems

MRF as generators?

[2] T-PAMI 1994

# An Empirical Study of the Simulation of Various Models Used for Images

A. J. Gray, J. W. Kay, and D. M. Titterington

Abstract— Markov random fields are typically used as priors in Bayesian image restoration methods to represent spatial information in the image. Commonly used Markov random fields are not in fact capable of representing the moderate-to-large scale clustering present in naturally occurring images and can also be time consuming to simulate,









### Gray, Kay, Titterington [2] T-PAMI 1994

... the local properties of spatial Markov models are undoubtedly plausible descriptors of the local associations typical of many images, which is the way in which the models are often used. Nevertheless. it would be reassuring if models used as priors did in fact provide a realistic representation of our prior assumptions and if their (empirical) properties were more widely known.

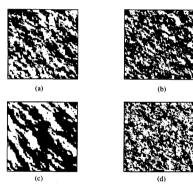


Fig. 4. Realizations of two-dimensional, one-parameter, autologistic Markov Mesh models: (a) binary, second-order model with  $\beta = \log 5$ ; (b) three-color second-order model with  $\beta = \log 5$ ; (c) binary second-order model with  $\beta = \log 3$ .





Computer ("AI") generated stills from hypothetical movie: Chilean director Alejandro Jodorowsky's 1976 version of "Tron" using midjourney.com as reported in 2023-01-13 NY Times article "This film does not exist" by director Frank Pavich.

### Generative models are hot in the news

- ➤ 2020-11-21 NY Times "Designed to Deceive: Do These People Look Real to You?" Article about generated (aka fake) faces.
- 2022-10-21 NY Times "A Coming-Out Party for Generative A.I., Silicon Valley's New Craze" (about "Stable Diffusion" image generator) https://nyti.ms/3SjsNOk
- ➤ 2023-01-09 NY Times "A.I. Turns Its Artistry to Creating New Human Proteins" https://nyti.ms/3IzY66m







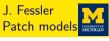
Théâtre d'Opéra Spatial Image created with Midjourney generative AI by Jason M. Allen.

Won 2022 Colorado State Fair fine art competition.

Wikimedia: "This file is in the public domain because it is the work of a computer algorithm or artificial intelligence and does not contain sufficient human authorship to support a copyright claim."



## Generative models are hot in imaging / inverse problems



Zhao, Ye, Bresler: Jan. 2023 IEEE SpMag survey paper [3]

- ► Generative adversarial network (GAN) models
- ► Variation auto-encoder (VAE) models [4]
- ► Normalizing flows [5, 6]
- Score-based diffusion models
  - o Zaccharie Ramzi et al., NeurIPS Workshop 2020 [7]
  - Yang Song & Liyue Shen et al., NeurIPS Workshop 2021, ICLR 2022 [8, 9]
  - o Ajil Jalal et al. ... Jon Tamir, NeurlPS 2021 [10]
  - o Hyungjin Chung & Jong Chul Ye, MIA, Aug. 2022 [11]
  - o Luo et al., MRM, 2023 [12]
  - o ...
- ► Kazerouni et al. [13] have github catalog, including >20 (!) survey papers
- ... (hopelessly incomplete lists)

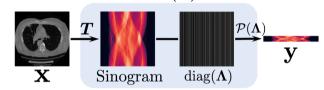
## Medical example: Low-dose sparse-view X-ray CT imaging

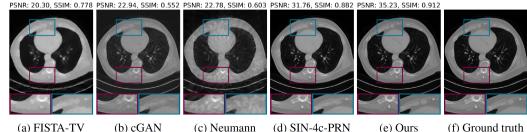
J. Fessler Patch models WHYPERTY

From Song & Shen et al., ICLR 2022 [9].

Trained with 47K 2D images; 23 projection views ( $\approx$  17-fold dose reduction)

$$\boldsymbol{A} = \mathcal{P}(\boldsymbol{\Lambda})\boldsymbol{T}$$





(b) cGAN

(c) Neumann

(d) SIN-4c-PRN

(f) Ground truth

Bayesian inference methods use the posterior:

$$p(x|y) = \underbrace{p(y|x)}_{\text{physics}} \underbrace{p(x)}_{\text{prior}} / p(y)$$

- ▶ Here the prior p(x) is for quantifying (prior) probability, not necessarily for generation.
- $\triangleright$  A model for the posterior p(x|y) opens many doors:
  - Maximizing p(x|y) is maximum a posteriori (MAP) estimation
  - ▶ The conditional mean  $E[x|y] = \int x p(x|y) dx$  is the MMSE estimator
  - $\triangleright$  Sampling from the posterior p(x|y) facilitates uncertainty quantification in inference
- All these methods require the prior p(x), i.e., a prior model  $p(x;\theta)$ .

Bayesian inference methods use the posterior:

$$p(x|y) = \underbrace{p(y|x)}_{\text{physics}} \underbrace{p(x)}_{\text{prior}} / p(y)$$

- Here the prior p(x) is for quantifying (prior) probability, not necessarily for generation.
- $\triangleright$  A model for the posterior p(x|y) opens many doors:
  - Maximizing p(x|y) is maximum a posteriori (MAP) estimation
  - ▶ The conditional mean  $E[x|y] = \int x p(x|y) dx$  is the MMSE estimator
  - $\triangleright$  Sampling from the posterior p(x|y) facilitates uncertainty quantification in inference
- $\blacktriangleright$  All these methods require the prior p(x), i.e., a prior model  $p(x;\theta)$ .
- Or do they?

Sampling from a *prior*  $p(x; \theta)$  just needs its score function  $\nabla_x \log p(x; \theta)$ , using Langevin dynamics, aka stochastic gradient ascent of log-prior:

$$\mathbf{x}_t = \mathbf{x}_{t-1} + \alpha_t \underbrace{\nabla \log p(\mathbf{x}_{t-1}; \boldsymbol{\theta})}_{\text{score function}} + \beta_t \mathcal{N}(\mathbf{0}, \boldsymbol{I}), \quad t = 1, \dots, T.$$

- $\circ$  Draws samples from  $p(x; \theta)$  for suitable choices of  $\{\alpha_t\}$ ,  $\{\beta_t\}$ , and (large) T [14].
- $\circ$  If  $\alpha_t = 0$  and  $\beta_t = \beta$ , then akin to (isotropic) diffusion or Brownian motion

- ► Typical distribution models:  $p(x; \theta) = \frac{1}{Z(\theta)} e^{-U(x; \theta)}$ . Goal: learn  $\theta$  from training data  $x_1, \dots, x_T$
- ▶ For IID samples  $\{x_t\}$ , one could try to learn  $\theta$  by ML estimation:

$$\begin{split} \hat{\theta} &= \arg\max_{\theta} \mathsf{p}(\mathbf{x}_1, \dots, \mathbf{x}_T; \theta) = \arg\max_{\theta} \sum_{t=1}^{T} \mathsf{log}(\mathsf{p}(\mathbf{x}_t; \theta)) \\ &= \arg\max_{\theta} \left( -TZ(\theta) + \sum_{t=1}^{T} -U(\mathbf{x}_t; \theta) \right). \end{split}$$

Typically intractable due to the partition function  $Z(\theta)$ .

- ► Typical distribution models:  $p(x; \theta) = \frac{1}{Z(\theta)} e^{-U(x; \theta)}$ . Goal: learn  $\theta$  from training data  $x_1, \dots, x_T$
- ▶ For IID samples  $\{x_t\}$ , one could try to learn  $\theta$  by ML estimation:

$$\hat{m{ heta}} = rg \max_{m{ heta}} p(m{x}_1, \dots, m{x}_T; m{ heta}) = rg \max_{m{ heta}} \sum_{t=1}^T \log(p(m{x}_t; m{ heta}))$$

$$= rg \max_{m{ heta}} \left( -T m{Z}(m{ heta}) + \sum_{t=1}^T -U(m{x}_t; m{ heta}) \right).$$

Typically intractable due to the partition function  $Z(\theta)$ .

▶ In contrast, the score function is easier to handle:

$$s(x; \theta) \triangleq \nabla_x \log p(x; \theta) = \nabla_x (-\log Z(\theta) - U(x; \theta)) = -\nabla_x U(x; \theta).$$

- ▶ Given training data  $x_1, ..., x_T$ , learn score function  $s(x; \theta) \stackrel{?}{=} \nabla_x \log p(x; \theta)$

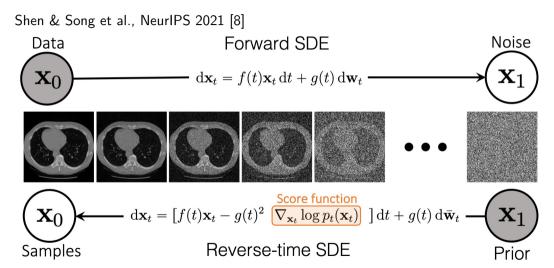
## Score matching

- ▶ Given training data  $x_1, ..., x_T$ , learn score function  $s(x; \theta) \stackrel{?}{=} \nabla_x \log p(x; \theta)$
- Explicit score matching (ESM) (Hyvärinen, 2005 [15])
- ► Implicit score matching (ISM)
- ▶ Denoising score matching (DSM) (Vincent, 2011 [16])
- ▶ Noise-conditional score matching (NCSM) (Song, 2019 [17, eqn. (5)]):

$$\ell(\boldsymbol{\theta}; \sigma) \triangleq \frac{1}{2} \, \mathsf{E}_{\mathsf{q}_0(\boldsymbol{x})} \bigg[ \mathsf{E}_{\boldsymbol{g}_{\sigma}(\boldsymbol{z})} \bigg[ \bigg\| \boldsymbol{s}(\boldsymbol{x} + \boldsymbol{z}; \boldsymbol{\theta}, \sigma) + \frac{\boldsymbol{z}}{\sigma^2} \bigg\|_2^2 \bigg] \bigg], \quad \mathcal{L}(\boldsymbol{\theta}; \{\sigma_I\}) = \frac{1}{L} \sum_{I=1}^L \sigma_I^2 \, \ell(\boldsymbol{\theta}; \sigma_I),$$

where  $s(x; \theta, \sigma)$  denotes a noise-conditional score network (NCSN).

- ▶  $d(x; \theta) \triangleq x + \sigma^2 s(x; \theta, \sigma)$ : equivalent image denoiser by Tweedie's formula [18]
- ► Recommended choice [19]:  $\mathbf{s}(\mathbf{x}; \boldsymbol{\theta}, \sigma) \triangleq \tilde{\mathbf{s}}(\mathbf{x}; \boldsymbol{\theta}) / \sigma$ , where  $\tilde{\mathbf{s}}$  is unitless

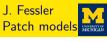


## Score-based diffusion models: trade-offs

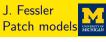
J. Fessler
Patch models

- ► No adversarial training needed
- ► High quality sample generation (if enough training data)

## Score-based diffusion models: trade-offs

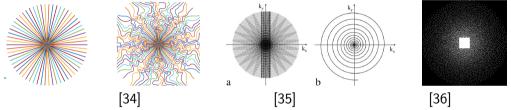


- ► No adversarial training needed
- ► High quality sample generation (if enough training data)
- Expensive sample generation (vs GAN models)
  - Distillation methods [20]
  - o Consistency models [21]
  - o Geometric decomposition [22]
  - o Multi-scale [23, 24] and pyramidal [25] and coarse-to-fine [26] models
  - Faster ODE solvers [27]
  - o Warm starts [28]
  - Latent diffusion models: use VAE and diffuse in latent space [29–31].
     Used in Stable Diffusion by start-up Stability AI
  - o 3D image reconstruction using 2D models [32, 33]
- ► Learning 3D (or 3D+T) whole-image generative models is challenging (training data, GPU memory, ...)



Jan. 2023 survey paper on generative models [3] does not mention "patch" once!?

### MRI k-space sampling:



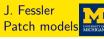
Patch-based models have long history in inverse problems, e.g.,

- patch GAN [37–39]
- patch dictionary models [40, 41]
- non-local means, BM3D
- Wasserstein patch prior [42, 43] ...

► Could patch-based generative models provide better robustness to distribution shifts, perhaps at the cost of reduced in-distribution performance?

Especially in applications with very limited training data? e.g., dynamic MRI

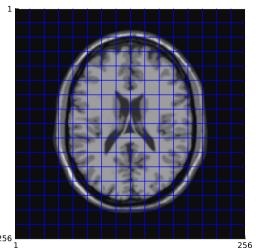
► Can we use the "latest" generative models, namely score-based models, for patches?



#### Warm up:

simple, but less effective, approach:

- Fixed patch size
- Fixed patch grid
- No position information (Fessler, Hu, Xu, BASP 2023 [46])



▶ Start with MRF formulation, aka *fields of experts* model [51–53] for image *x*:

$$p(\boldsymbol{x};\boldsymbol{\theta}) = \frac{1}{Z(\boldsymbol{\theta})} e^{-\sum_{c} V_{c}(\boldsymbol{x};\boldsymbol{\theta})} = \frac{1}{Z(\boldsymbol{\theta})} \prod_{c} e^{-V_{c}(\boldsymbol{x};\boldsymbol{\theta})}.$$

- $oldsymbol{ heta}$  : parameter vector that describes the prior
- $V_c$ : clique potential for the cth image patch
- $Z(\theta)$  : (intractable) partition function
- Assume (temporarily) statistical spatial stationarity (image shift invariance):

$$V_c(\mathbf{x}; \mathbf{\theta}) = V(\mathbf{G}_c \mathbf{x}; \mathbf{\theta})$$

- $oldsymbol{G}_c$ : wide binary matrix that grabs pixels of the cth patch from image  $oldsymbol{x}$
- $V(\mathbf{v}; \boldsymbol{\theta})$ : common patch clique function

► Resulting log-prior:

$$\log p(\mathbf{x}; \boldsymbol{\theta}) = -\log Z(\boldsymbol{\theta}) - \sum_{c} V(\mathbf{G}_{c}\mathbf{x}; \boldsymbol{\theta})$$

► Corresponding overall *image score function* arises from *patch score function*:

$$\mathbf{s}(\mathbf{x}; \boldsymbol{\theta}) \triangleq \nabla_{\mathbf{x}} \log p(\mathbf{x}; \boldsymbol{\theta}) = \sum_{c} \mathbf{G}_{c}' \mathbf{s}_{V}(\mathbf{G}_{c}\mathbf{x}; \boldsymbol{\theta}), \qquad \mathbf{s}_{V}(\mathbf{v}; \boldsymbol{\theta}) \triangleq -\nabla_{\mathbf{v}} V(\mathbf{v}; \boldsymbol{\theta}).$$

- ▶ All we must learn is the patch score function  $s_V(\mathbf{v}; \theta) : \mathbb{R}^n \to \mathbb{R}^n$ , e.g., a UNet.
- For non-overlapping patches:

$$\underbrace{\left\| \mathbf{s}(\mathbf{x} + \mathbf{z}; \boldsymbol{\theta}) + \mathbf{z}/\sigma^{2} \right\|_{2}^{2}}_{\text{image "denoise"}} = \left\| \sum_{c} \mathbf{G}_{c}' \mathbf{s}_{V}(\mathbf{G}_{c}(\mathbf{x} + \mathbf{z}); \boldsymbol{\theta}) + \mathbf{z}/\sigma^{2} \right\|_{2}^{2} \\
= \sum_{c} \underbrace{\left\| \mathbf{s}_{V}(\mathbf{x}_{c} + \mathbf{z}_{c}); \boldsymbol{\theta} \right) + \mathbf{z}_{c}/\sigma^{2} \right\|_{2}^{2}}_{\text{patch "denoise"}}, \quad \mathbf{z}_{c} \triangleq \mathbf{G}_{c} \mathbf{z}$$

▶ For training image patches  $\{v_1, ..., v_T\}$ , apply denoising score matching (DSM) of Vincent, 2011 [16], typically for a range of noise variances  $\sigma^2$  [14]:

$$\hat{\boldsymbol{\theta}} = \arg\min_{\boldsymbol{\theta}} \frac{1}{T} \sum\nolimits_{t=1}^{T} \mathsf{E}_{\sigma \sim p(\sigma)} \bigg[ \sigma^2 \, \mathsf{E}_{\mathbf{z} \sim \mathcal{N}(0, \sigma^2 \mathbf{I}_n)} \bigg[ \frac{1}{2} \, \bigg\| \mathbf{s}_V(\mathbf{v}_t + \mathbf{z}; \boldsymbol{\theta}, \sigma) + \frac{\mathbf{z}}{\sigma^2} \bigg\|_2^2 \bigg] \bigg] \, .$$

- Final patch score model is  $\mathbf{s}_V(\mathbf{v}; \hat{\boldsymbol{\theta}}, \sigma_{\min})$ .

# Patch-based score learning (simple)

▶ For training image patches  $\{v_1, ..., v_T\}$ , apply denoising score matching (DSM) of Vincent, 2011 [16], typically for a range of noise variances  $\sigma^2$  [14]:

$$\hat{\boldsymbol{\theta}} = \arg\min_{\boldsymbol{\theta}} \frac{1}{T} \sum\nolimits_{t=1}^{T} \mathsf{E}_{\sigma \sim p(\sigma)} \bigg[ \sigma^2 \, \mathsf{E}_{\mathbf{z} \sim \mathcal{N}(0, \sigma^2 \mathbf{I}_n)} \bigg[ \frac{1}{2} \, \bigg\| \mathbf{s}_V(\mathbf{v}_t + \mathbf{z}; \boldsymbol{\theta}, \sigma) + \frac{\mathbf{z}}{\sigma^2} \bigg\|_2^2 \bigg] \bigg] \, .$$

- Final patch score model is  $\mathbf{s}_V(\mathbf{v}; \hat{\boldsymbol{\theta}}, \sigma_{\min})$ .
- Network input is just image patches, never the entire image ⇒ scales to large 2D images, 3D, 4D, etc.

▶ For training image patches  $\{v_1, ..., v_T\}$ , apply denoising score matching (DSM) of Vincent, 2011 [16], typically for a range of noise variances  $\sigma^2$  [14]:

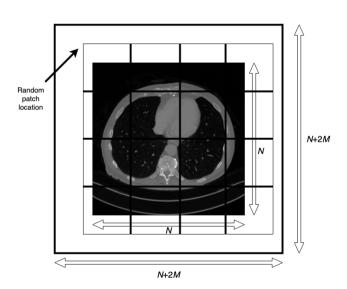
$$\hat{\boldsymbol{\theta}} = \arg\min_{\boldsymbol{\theta}} \frac{1}{T} \sum\nolimits_{t=1}^{T} \mathsf{E}_{\sigma \sim p(\sigma)} \bigg[ \sigma^2 \, \mathsf{E}_{\mathbf{z} \sim \mathcal{N}(\mathbf{0}, \sigma^2 \mathbf{I}_n)} \bigg[ \frac{1}{2} \, \bigg\| \mathbf{s}_V(\mathbf{v}_t + \mathbf{z}; \boldsymbol{\theta}, \sigma) + \frac{\mathbf{z}}{\sigma^2} \bigg\|_2^2 \bigg] \bigg] \, .$$

- Final patch score model is  $\mathbf{s}_V(\mathbf{v}; \hat{\boldsymbol{\theta}}, \sigma_{\min})$ .
- Network input is just image patches, never the entire image ⇒ scales to large 2D images, 3D, 4D, etc.
- Drawbacks:
  - Visible patch boundaries
  - o Fixed patch size slows learning
  - o Suboptimal stationarity assumption (cf. vertebrae)

- ▶ zero-pad image x
- use multiple grid locations

### Inspirations:

- Wavelet "cycle spinning"[47, 54–57]
- o Wang, NeurIPS 2023 [58]

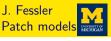


- $\triangleright$   $N_1 \times N_2$ : original image size
- $ightharpoonup P_1 \times P_2$ : patch size
- $ightharpoonup K_i \triangleq 1 + |N_i/P_i|, i = 1, 2: \# non-overlapping patches for original image$
- $ightharpoonup (N_1 + 2M_1) \times (N_2 + 2M_2)$ : padded image size;  $M_i \triangleq K_i P_i N_i$
- ► Product probability model:

$$p(\mathbf{x}) \triangleq \frac{1}{Z} \underbrace{\prod_{m=1}^{M_1 M_2}}_{\text{grid}} \underbrace{\left(\underbrace{p_{m,B}(\mathbf{x}_{m,B})}_{\text{border}} \underbrace{\prod_{k=1}^{K_1 K_2} p_{m,k}(\mathbf{x}_{m,k})}_{\text{patches}}\right)}_{\text{patches}} = \frac{1}{Z} \underbrace{\prod_{m=1}^{M_1 M_2} \prod_{k=1}^{K_1 K_2} \underbrace{e^{-V(\mathbf{x}_{m,k}; \mathbf{m}, \mathbf{k})}}_{\text{position}}}_{\text{encoding}}$$

- $\circ x_{m,B}$ : border pixels for mth shift (all zero)
- o  $\mathbf{x}_{m,k}$ : kth patch for mth shift

# Probability model with padding & grids & positions

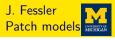


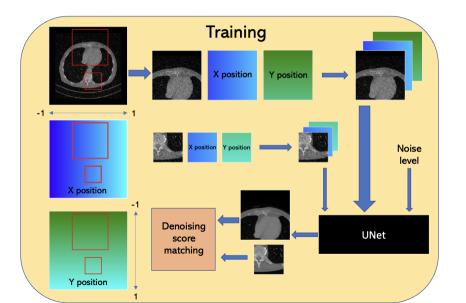
- $\triangleright$   $N_1 \times N_2$ : original image size
- $ightharpoonup P_1 \times P_2$ : patch size
- $ightharpoonup K_i \triangleq 1 + |N_i/P_i|, i = 1,2: \# non-overlapping patches for original image$
- $(N_1 + 2M_1) \times (N_2 + 2M_2)$ : padded image size;  $M_i \triangleq K_i P_i N_i$
- ▶ Product probability model:

$$p(\mathbf{x}) \triangleq \frac{1}{Z} \underbrace{\prod_{m=1}^{M_1 M_2}}_{\text{grid}} \underbrace{\left(\underbrace{p_{m,B}(\mathbf{x}_{m,B})}_{\text{border}} \underbrace{\prod_{k=1}^{K_1 K_2} p_{m,k}(\mathbf{x}_{m,k})}_{\text{patches}}\right)}_{\text{patches}} = \frac{1}{Z} \underbrace{\prod_{m=1}^{M_1 M_2} \prod_{k=1}^{K_1 K_2} \underbrace{e^{-V(\mathbf{x}_{m,k}; \mathbf{m}, \mathbf{k})}}_{\text{position}}}_{\text{encoding}}$$

- $\circ x_{m,B}$ : border pixels for mth shift (all zero)
- $\circ \mathbf{x}_{m,k}$ : kth patch for mth shift
- Learn position-dependent patch score function  $s(\mathbf{v}; \boldsymbol{\theta}, m, k) = -\nabla_{\mathbf{v}} V(\mathbf{v}; m, k)$

# Patch Diffusion Inverse Solver (PaDIS): Training



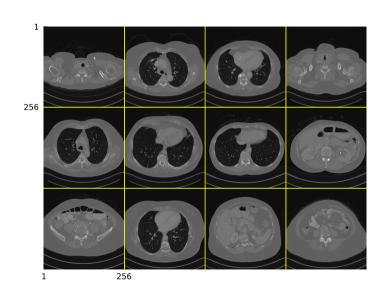


NeurIPS 2024 [60]

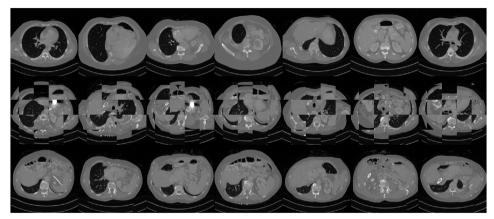
## Training images (CT)

AAPM 2016 CT challenge data [61]; 10 3D volumes, rescaled to 256<sup>3</sup>

Example slices:

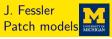


# Image generation (unconditional sampling from prior)



- Top: generation with a network trained on whole images (2D...)
- Middle: patch-only version of [58] (non-overlapping patches).
- $\circ$  Bottom: proposed PaDIS method.

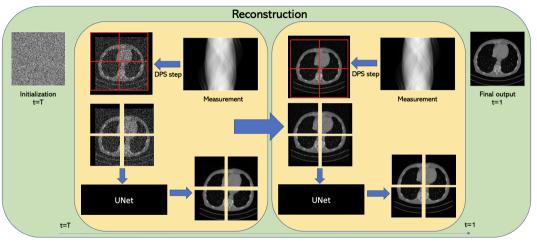
## Training time results



- 2 A40 GPUs using PyTorch and ADAM
- ▶ whole image model: 24 − 36 hours
- ightharpoonup patch-based model: pprox 12 hours

## Patch Diffusion Inverse Solver (PaDIS): Reconstruction

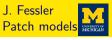




Diffusion posterior sampling (DPS) (Chung et al., ICLR 2023 [62]) with Langevin dynamics, modified to use patch score with random grid shifts.

**Input**:  $\mathbf{y}$ ,  $\mathbf{A}$ , T,  $\sigma_1 < \sigma_2 < \ldots < \sigma_T$ ,  $\epsilon > 0$ ,  $\{\zeta_t > 0\}$ ,  $P_1, P_2, M_1, M_2$ . trained noise-conditional, position-encoded patch denoiser  $d(\cdot; \theta_*, m, k, \sigma)$ Initialize random image  $\mathbf{x} \sim \mathcal{N}(\mathbf{0}, \sigma_{\tau}^2 \mathbf{I})$ for t = T : 1 do Randomly select grid integer  $m \in \{1, ..., M_1 M_2\}$ for  $k = 1 : (K_1K_2)$  do (parallelizable) Extract patch  $x_{m,k}$ Denoise patch:  $\mathbf{d}_{m,k} \triangleq \mathbf{d}(\mathbf{x}_{m,k}; \boldsymbol{\theta}_*, m, k, \sigma_t)$ end for Combine denoised patches to get denoised image d Compute image score function:  $\mathbf{s} = (\mathbf{d} - \mathbf{x})/\sigma_{+}^{2}$ Data term:  $\mathbf{x} := \mathbf{x} - \zeta_t \nabla_{\mathbf{x}} \| \mathbf{A} \ \mathbf{d}(\mathbf{x}) - \mathbf{y} \|_2^2$ Sample  $\mathbf{z} \sim \mathcal{N}(\mathbf{0}, \sigma_t^2 \mathbf{I})$ Step size  $\alpha_t \triangleq \epsilon \, \sigma_t^2$ Langevin update:  $\mathbf{x} := \mathbf{x} + \frac{\alpha_t}{2}\mathbf{s} + \sqrt{\alpha_t}\mathbf{z}$ end for

### CT Experiments



#### Default setup:

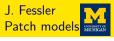
- 9 of 10 volumes for training  $\Longrightarrow$  2304 slices
- 25 slices of 10th volume for testing
- 512 element parallel-beam CT detector
- A from Operator Discretization Library (ODL)
- $56 \times 56$  patch size
- U-Net of Karras 2022 [59]
- Step size  $\zeta_t = \zeta/\|\boldsymbol{Ad}(\boldsymbol{x}_t) \boldsymbol{y}\|_2$
- 1000 neural function evaluations (NFEs) [59]

## Quantitative results on three different inverse problems

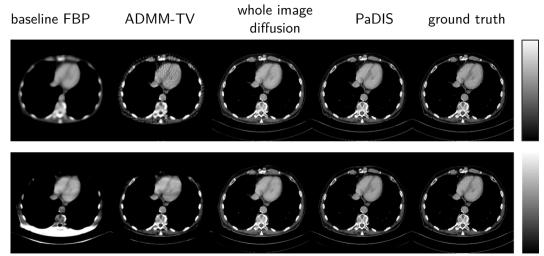
Method	CT, 20 Views		CT, 8 Views		Deblurring		Superresolution	
	PSNR↑	SSIM↑	PSNR↑	SSIM↑	PSNR↑	SSIM↑	PSNR↑	SSIM↑
Baseline	24.93	0.595	21.39	0.415	24.54	0.688	25.86	0.739
ADMM-TV	26.82	0.724	23.09	0.555	28.22	0.792	25.66	0.745
PnP-ADMM [63]	26.86	0.607	22.39	0.489	28.82	0.818	26.61	0.785
PnP-RED [64]	27.99	0.622	23.08	0.441	29.91	0.867	26.36	0.766
Whole image diffusion	32.84	0.835	25.74	0.706	30.19	0.853	29.17	0.827
Langevin dynamics [17]	33.03	0.846	27.03	0.689	30.60	0.867	26.83	0.744
Predictor-corrector [11]	32.35	0.820	23.65	0.546	28.42	0.724	26.97	0.685
VE-DDNM [65]	31.98	0.861	27.71	0.759	-	-	26.01	0.727
Patch Averaging [50]	33.35	0.850	28.43	0.765	29.41	0.847	27.67	0.802
Patch Stitching	32.87	0.837	26.71	0.710	29.69	0.849	27.50	0.780
PaDIS (Ours)	33.57	0.854	29.48	0.767	30.80	0.870	29.47	0.846

(Averages across all test images.)

# More inverse problem experiments

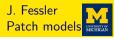


Method	CT, 60 Views		CT, Fan Beam		Heavy Deblurring	
	PSNR↑	SSIM↑	PSNR↑	SSIM↑	PSNR↑	SSIM↑
Baseline	25.89	0.746	20.07	0.521	21.14	0.569
ADMM-TV	30.93	0.833	25.78	0.719	26.03	0.724
Whole image diffusion	35.83	0.894	26.89	0.835	28.35	0.808
PaDIS (Ours)	39.28	0.941	29.91	0.932	28.91	0.818



Top: 60 view CT Bottom: fan-beam CT

 $\approx 400~\text{HU}$  window width



#### Patchsize

Ρ	PSNR↑	SSIM↑	
8	32.57	0.844	
16	32.57	0.829	
32	32.72	0.853	
56	33.57	0.854	
96	33.36	0.854	
256	32.84	0.835	

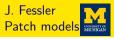
### Positional encoding

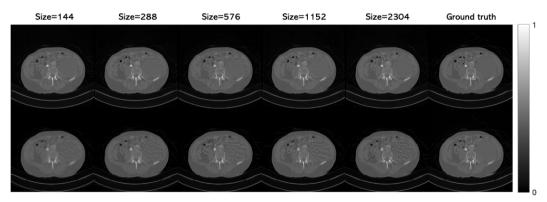
	PSNR↑	SSIM↑	
no position enc.	23.25	0.459	
no position $+$ init	24.51	0.518	
with position	33.57	0.854	

## Effect of training dataset size on CT reconstruction

Dataset	Pato	ches	Whole image		
size	56 >	< 56	$256 \times 256$		
	PSNR↑	SSIM↑	PSNR↑	SSIM↑	
144	32.28	0.841	29.12	0.804	
288	32.43	0.837	31.09	0.829	
576	33.03	0.846	31.81	0.835	
1152	33.01	0.849	31.36	0.834	
2304	33.57	0.854	32.84	0.835	

### 20 view CT reconstruction: training dataset sizes





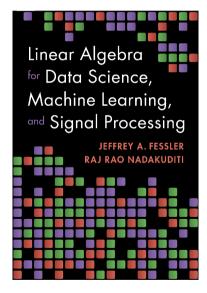
Top: PaDIS

Bottom: whole image diffusion model

- ► Training whole-image generative models for 3D or dynamic PET would be challenging
- Patch-based models should be feasible if suitable training data available
- ▶ Ideally use the best resolution / senstivity scanner, e.g., total-body PET
- Challenge: out-of distribution test data
- ▶ Patch-based diffusion models beat whole-image models for mismatched distribution inverse problems arXiv 2410.11730 [66]

- ▶ Generative models are promising for under-determined inverse problems
- ▶ Learning patch score models is feasible with denoising score matching
- ► For limited training data, patch-models can outperform whole-image models
- ▶ Integrate invariances: amplitude scale / rotation / flip / DC offset ...
- Explore trade-offs between generalizability and in-distribution performance
- Extend to 3D, 3D+Time, 3D+Multicontrast

Tutorial Julia code: https://github.com/JeffFessler/ScoreMatching.jl



- Online demos: https://github.com/JeffFessler/ book-la-demo
- Topics include: low-rank matrix approximation, robust PCA, photometric stereo, video foreground/background separation, spectral clustering, matrix completion, ...
- Available from Cambridge Univ. Press

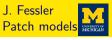


Talk and code available online at http://web.eecs.umich.edu/~fessler



- [1] S. Geman and D. Geman. "Stochastic relaxation, Gibbs distributions, and Bayesian restoration of images." In: IEEE Trans. Patt. Anal. Mach. Int. 6.6 (Nov. 1984), 721–41.
- [2] A. J. Gray, J. W. Kay, and D. M. Titterington. "An empirical study of the simulation of various models used for images." In: IEEE Trans. Patt. Anal. Mach. Int. 16.5 (May 1994), 507–12.
- [3] Z. Zhao, J. C. Ye, and Y. Bresler. "Generative models for inverse imaging problems: from mathematical foundations to physics-driven applications." In: IEEE Sig. Proc. Mag. 40.1 (Jan. 2023), 148–63.
- [4] E. D. Zhong, T. Bepler, B. Berger, and J. H. Davis. "CryoDRGN: reconstruction of heterogeneous cryo-EM structures using neural networks." In: Nature Meth. 18.2 (2021), 176–85.
- [5] D. Rezende and S. Mohamed. "Variational inference with normalizing flows." In: Proc. Intl. Conf. Mach. Learn. 2015, 1530-8.
- [6] F. Altekruger, A. Denker, P. Hagemann, J. Hertrich, P. Maass, and G. Steidl. "PatchNR: learning from very few images by patch normalizing flow regularization." In: *Inverse Prob.* 39.6 (May 2023), p. 064006.
- [7] Z. Ramzi, B. Remy, F. Lanusse, J-L. Starck, and P. Ciuciu. "Denoising score-matching for uncertainty quantification in inverse problems." In: NeurIPS 2020 Workshop on Deep Learning and Inverse Problems. 2020.
- [8] Y. Song, L. Shen, L. Xing, and S. Ermon. "Solving inverse problems in medical imaging with score-based generative models." In: NeurIPS Deep Inv. Work. 2021.
- [9] Y. Song, L. Shen, L. Xing, and S. Ermon. "Solving inverse problems in medical imaging with score-based generative models." In: Proc. Intl. Conf. on Learning Representations. 2022.
- [10] A. Jalal, M. Arvinte, G. Daras, E. Price, A. Dimakis, and J. Tamir. "Robust compressed sensing MR imaging with deep generative priors." In: NeurIPS Workshop Deep Inverse. 2021.
- [11] H. Chung and J. C. Ye. "Score-based diffusion models for accelerated MRI." In: Med. Im. Anal. 80 (Aug. 2022), p. 102479.

### Bibliography II

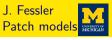


- [12] G. Luo, M. Blumenthal, M. Heide, and M. Uecker. "Bayesian MRI reconstruction with joint uncertainty estimation using diffusion models." In: Mag. Res. Med. 90.1 (July 2023), 295–311.
- [13] A. Kazerouni, E. K. Aghdam, M. Heidari, R. Azad, M. Fayyaz, I. Hacihaliloglu, and D. Merhof. "Diffusion models in medical imaging: A comprehensive survey." In: Med. Im. Anal. 88 (Aug. 2023), p. 102846.
- [14] Y. Song, J. Sohl-Dickstein, D. P. Kingma, A. Kumar, S. Ermon, and B. Poole. "Score-based generative modeling through stochastic differential equations." In: Proc. Intl. Conf. on Learning Representations. 2021.
- [15] A. Hyvärinen. "Estimation of non-normalized statistical models by score matching." In: J. Mach. Learning Res. 6.24 (2005), 695–709.
- [16] P. Vincent. "A connection between score matching and denoising autoencoders." In: Neural Comput. 23.7 (July 2011), 1661–74.
- [17] Y. Song and S. Ermon. "Generative modeling by estimating gradients of the data distribution." In: NeurIPS. 2019.
- [18] B. Efron. "Tweedie's formula and selection bias." In: J. Am. Stat. Assoc. 106.496 (2011), 1602-14.
- [19] Y. Song and S. Ermon. "Improved techniques for training score-based generative models." In: NeurIPS. Vol. 33. 2020, 12438–48.
- [20] T. Salimans and J. Ho. "Progressive distillation for fast sampling of diffusion models." In: Proc. Intl. Conf. on Learning Representations. 2022.
- [21] Y. Song, P. Dhariwal, M. Chen, and I. Sutskever. Consistency models. 2023.
- [22] H. Chung, S. Lee, and J. C. Ye. "Decomposed diffusion sampler for accelerating large-scale inverse problems." In: *Proc. Intl. Conf. on Learning Representations.* 2024.
- [23] F. Guth, S. Coste, V. D. Bortoli, and Stéphane Mallat. "Wavelet score-based generative modeling." In: NeurIPS. 2022.
- [24] Z. Kadkhodaie, F. Guth, Stephane Mallat, and E. P. Simoncelli. "Learning multi-scale local conditional probability models of images." In: Proc. Intl. Conf. on Learning Representations. 2023.

### Bibliography III

- [25] D. Ryu and J. C. Ye. Pyramidal denoising diffusion probabilistic models, 2022.
- [26] S. Lee, H. Chung, J. Kim, and J. C. Ye. "Progressive deblurring of diffusion models for coarse-to-fine image synthesis." In: NeurIPS Workshop SBM. 2022.
- [27] C. Lu, Y. Zhou, F. Bao, J. Chen, C. Li, and J. Zhu. "DPM-solver: A fast ODE solver for diffusion probabilistic model sampling in around 10 steps." In: NeurIPS. 2022.
- [28] H. Chung, B. Sim, and J. C. Ye. "Come-closer-diffuse-faster: accelerating conditional diffusion models for inverse problems through stochastic contraction." In: Proc. IEEE Conf. on Comp. Vision and Pattern Recognition. 2022, 12403–12.
- [29] A. Vahdat, K. Kreis, and J. Kautz. "Score-based generative modeling in latent space." In: NeurIPS. 2021.
- [30] R. Rombach, A. Blattmann, D. Lorenz, P. Esser, and Bjorn Ommer. "High-resolution image synthesis with latent diffusion models." In: Proc. IEEE Conf. on Comp. Vision and Pattern Recognition. 2022, 10674–85.
- [31] K. C. Tezcan, N. Karani, C. F. Baumgartner, and E. Konukoglu. "Sampling possible reconstructions of undersampled acquisitions in MR imaging with a deep learned prior." In: IEEE Trans. Med. Imag. 41.7 (July 2022), 1885–96.
- [32] H. Chung, D. Ryu, M. T. McCann, M. L. Klasky, and J. C. Ye. "Solving 3D inverse problems using pre-trained 2D diffusion models." In: Proc. IEEE Conf. on Comp. Vision and Pattern Recognition. 2023, 22542–51.
- [33] S. Lee, H. Chung, M. Park, J. Park, W-S. Ryu, and J. C. Ye. "Improving 3D imaging with pre-trained perpendicular 2D diffusion models." In: Proc. Intl. Conf. Comp. Vision. 2023.
- [34] G. Wang, T. Luo, J-F. Nielsen, D. C. Noll, and J. A. Fessler. "B-spline parameterized joint optimization of reconstruction and k-space trajectories (BJORK) for accelerated 2D MRI." In: IEEE Trans. Med. Imag. 41.9 (Sept. 2022), 2318–30.
- [35] W. Wu and K. L. Miller. "Image formation in diffusion MRI: A review of recent technical developments." In: J. Mag. Res. Im. 46.3 (Sept. 2017), 646–62.

## Bibliography IV



- [36] S. Bhadra, W. Zhou, and M. A. Anastasio. "Medical image reconstruction with image-adaptive priors learned by use of generative adversarial networks." In: Proc. SPIE 11312 Medical Imaging: Phys. Med. Im. 2020, p. 113120V.
- [37] C. Li and M. Wand. "Precomputed real-time texture synthesis with Markovian generative adversarial networks." In: Proc. European Comp. Vision Conf. 2016, 702–16.
- [38] P. Isola, J-Y. Zhu, T. Zhou, and A. A. Efros. "Image-to-image translation with conditional adversarial networks." In: *Proc. IEEE Conf. on Comp. Vision and Pattern Recognition.* 2017, 5967–76.
- [39] A. Elnekave and Y. Weiss. "Generating natural images with direct patch distributions matching." In: Proc. European Comp. Vision Conf. Vol. 13677. 2022.
- [40] M. Aharon, M. Elad, and A. Bruckstein. "K-SVD: an algorithm for designing overcomplete dictionaries for sparse representation." In: IEEE Trans. Sig. Proc. 54.11 (Nov. 2006), 4311–22.
- [41] S. Ravishankar and Y. Bresler. "MR image reconstruction from highly undersampled k-space data by dictionary learning." In: IEEE Trans. Med. Imag. 30.5 (May 2011), 1028–41.
- [42] J. Hertrich, A. Houdard, and C. Redenbach. "Wasserstein patch prior for image superresolution." In: IEEE Trans. Computational Imaging 8 (2022), 693–704.
- [43] F. Altekruger and J. Hertrich. "WPPNets and WPPFlows: The power of Wasserstein patch priors for superresolution." In: SIAM J. Imaging Sci. 16.3 (2023), 1033–67.
- [44] G. Vaksman, M. Zibulevsky, and M. Elad. "Patch ordering as a regularization for inverse problems in image processing." In: SIAM J. Imaging Sci. 9.1 (2016), 287–319.
- [45] M. Piening, F. Altekrüger, J. Hertrich, P. Hagemann, A. Walther, and G. Steidl. Learning from small data sets: Patch-based regularizers in inverse problems for image reconstruction. 2023.
- [46] J. A. Fessler, J. Hu, and X. Xu. "Generalizability (or not?) of patch-based image models." In: BASP. Invited presentation. 2023.

- [47] U. S. Kamilov, E. Bostan, and M. Unser. "Variational justification of cycle spinning for wavelet-based solutions of inverse problems." In: IEEE Signal Proc. Letters 21.11 (Nov. 2014), 1326–30.
- [48] A. Saucedo, S. Lefkimmiatis, N. Rangwala, and K. Sung. "Improved computational efficiency of locally low rank MRI reconstruction using iterative random patch adjustments." In: IEEE Trans. Med. Imag. 36.6 (2017), 1209–20.
- [49] J. L. Rumberger, X. Yu, P. Hirsch, M. Dohmen, V. E. Guarino, A. Mokarian, L. Mais, J. Funke, and D. Kainmueller. "How shift equivariance impacts metric learning for instance segmentation." In: Proc. Intl. Conf. Comp. Vision. 2021, 7108–16.
- [50] O. Ozdenizci and R. Legenstein. "Restoring vision in adverse weather conditions with patch-based denoising diffusion models." In: IEEE Trans. Patt. Anal. Mach. Int. 45.8 (Jan. 2023), 10346–57.
- [51] G. E. Hinton. "Training products of experts by minimizing contrastive divergence." In: Neural Computation 14.8 (Aug. 2002), 1771–800.
- [52] S. Roth and M. J. Black. "Fields of experts." In: Intl. J. Comp. Vision 82.2 (Jan. 2009), 205–29.
- [53] D. P. Kingma and Y. LeCun. "Regularized estimation of image statistics by score matching." In: NeurIPS. 2010, 1126-34.
- [54] R. R. Coifman and D. L. Donoho. Translation-invariant denoising, 1995.
- [55] M. A. T. Figueiredo and R. D. Nowak. "An EM algorithm for wavelet-based image restoration." In: IEEE Trans. Im. Proc. 12.8 (Aug. 2003), 906–16.
- [56] U. Kamilov, E. Bostan, and M. Unser. "Wavelet shrinkage with consistent cycle spinning generalizes total variation denoising." In: IEEE Signal Proc. Letters 19.4 (Apr. 2012), 187–90.
- [57] F. Ong and M. Lustig. "Beyond low rank + sparse: multiscale low rank matrix decomposition." In: IEEE J. Sel. Top. Sig. Proc. 10.4 (June 2016), 672–87.
- [58] Z. Wang, Y. Jiang, H. Zheng, P. Wang, P. He, Z. Wang, W. Chen, and M. Zhou. "Patch diffusion: faster and more data-efficient training of diffusion models." In: NeurlPS. Vol. 36. 2023, 72137–54.

- [59] T. Karras, M. Aittala, T. Aila, and S. Laine. "Elucidating the design space of diffusion-based generative models." In: NeurIPS. 2022.
- [60] J. Hu, B. Song, X. Xu, L. Shen, and J. A. Fessler. "Learning image priors through patch-based diffusion models for solving inverse problems." In: NeurIPS. To appear, 2024.
- [61] C. H. McCollough, A. C. Bartley, R. E. Carter, B. Chen, T. A. Drees, P. Edwards, D. R. Holmes, A. E. Huang, F. Khan, S. Leng, K. L. McMillan, G. J. Michalak, K. M. Nunez, L. Yu, and J. G. Fletcher. "Low-dose CT for the detection and classification of metastatic liver lesions: Results of the 2016 Low Dose CT Grand Challenge." In: Med. Phys. 44.10 (Oct. 2017), e339–52.
- [62] H. Chung, J. Kim, M. T. Mccann, M. L. Klasky, and J. C. Ye. "Diffusion posterior sampling for general noisy inverse problems." In: Proc. Intl. Conf. on Learning Representations. 2023.
- [63] X. Xu, J. Liu, Y. Sun, B. Wohlberg, and U. S. Kamilov. "Boosting the performance of plug-and-play priors via denoiser scaling." In: Proc., IEEE Asilomar Conf. on Signals, Systems, and Comp. 2020, 1305–12.
- [64] Y. Hu, J. Liu, X. Xu, and U. S. Kamilov. "Monotonically convergent regularization by denoising." In: Proc. IEEE Intl. Conf. on Image Processing. 2022, 426–30.
- [65] Y. Wang, J. Yu, and J. Zhang. "Zero-shot image restoration using denoising diffusion null-space model." In: Proc. Intl. Conf. Mach. Learn. 2023.
- [66] J. Hu, B. Song, J. A. Fessler, and L. Shen. Patch-based diffusion models beat whole-image models for mismatched distribution inverse problems. 2024.