Medical image reconstruction using adaptive signal models

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Declaration: No relevant financial interests or relationships to disclose

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#### Introduction

ML-based image reconstruction approaches

#### Adaptive regularization

Patch-based adaptive regularizers Convolutional adaptive regularizers Blind dictionary learning

### Other ML4MI topics

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Overview of medical imaging:



Most obvious place for machine learning is post-processing:



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Most obvious place for machine learning is post-processing:



(Many conference sessions; special issue of IEEE Trans. on Med. Imaging in May 2016 [1], ...)

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### Machine learning in medical image reconstruction



Another (initially less obvious?) place for machine learning (multiple conference sessions):



. . .

# Machine learning in medical image reconstruction

J. Fessler ML for IR

Another (initially less obvious?) place for machine learning (multiple conference sessions):

$$\begin{array}{ccc} \mathsf{raw} \ \mathsf{data} & & \mathsf{ML}\text{-}\mathsf{based} \\ \boldsymbol{y} & \rightarrow & \mathsf{image \ reconstruction} \end{array} \rightarrow & \hat{\boldsymbol{x}} \end{array}$$

Possibly easier (than diagnosis) due to lower bar:

- current reconstruction methods based on simplistic image models;
- human eyes are better at detection than at solving inverse problems.

# Machine learning in medical image reconstruction

J. Fessler ML for IR

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Possibly easier (than diagnosis) due to lower bar:

EMB NPSS

• current reconstruction methods based on simplistic image models;

• human eyes are better at detection than at solving inverse problems. June 2018 special issue of IEEE Trans. on Medical Imaging [2]:

IEEE TRANSACTIONS ON MEDICAL IMAGING, VOL. 37, NO. 6, JUNE 2018

1289

# Image Reconstruction Is a New Frontier of Machine Learning

Ge Wang<sup>®</sup>, *Fellow, IEEE*, Jong Chu Ye<sup>®</sup>, *Senior Member, IEEE*, Klaus Mueller<sup>®</sup>, *Senior Member, IEEE*, and Jeffrey A. Fessler<sup>®</sup>, *Fellow, IEEE* 

A more speculative opportunity for machine learning:

. . .



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A more speculative opportunity for machine learning:

$$\begin{array}{ccc} \mathsf{raw} \ \mathsf{data} & \mathsf{ML}\text{-}\mathsf{based} \\ \boldsymbol{y} & \stackrel{}{\rightarrow} & \overset{}{\mathsf{magic''}} & \stackrel{}{\rightarrow} & \mathsf{interpretation} \end{array}$$

- CT sinogram to vessel diameter [3]
- k-space to ???

See Wiro Niessen's keynote...

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### Machine learning in medical imaging: scan design

One more opportunity for ML in medical imaging:



. . .

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One more opportunity for ML in medical imaging:



Choose best k-space phase encoding locations based on training images:

- "Learning-based compressive MRI" [4, 5] (Volkan Cevher group, June 2018 IEEE T-MI) Single coil only so far; perhaps hard to generalize to parallel MRI?
- Yue Cao and David Levin, MRM Sep. 1993 "Feature recognizing MRI" [6-8]

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### Adaptive phase-encode selection





Sampling designed to optimize PSNR for basis pursuit (BP) reconstruction using shearlet transform, at 25% sampling rate.

Sampling design considers both the training data and the reconstruction method. No high spatial frequencies!?

(Images from Gözcü et al. [5].)





#### Introduction

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Forward problem (data acquisition):



SPECT, PET, X-ray CT, MRI, optical, ...

Inverse problem (image formation):



Image reconstruction topics: physics models, measurement statistical models, regularization / object priors, optimization...

# Generations of medical image reconstruction methods

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- 70's "Analytical" methods (integral equations) FBP for SPECT / PET / X-ray CT, IFFT for MRI, ...
- 2. 80's Algebraic methods (as in "linear algebra") Solve y = Ax
- 3. 90's Statistical methods
  - LS / ML methods
  - Bayesian methods (Markov random fields, ...)
  - regularized methods
- 4. 00's Compressed sensing methods (mathematical sparsity models)
- 5. 10's Adaptive / data-driven methods machine learning, deep learning, ...

# Accelerating MR imaging using adaptive regularization

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(a)  $4 \times$  under-sampled MR k-space (b) zero-filled reconstruction (c) "compressed sensing" reconstruction with TV regularization (d) adaptive regularization using dictionary learning

Ravishankar & Bresler, DLMRI, T-MI, May 2011,

[9, Fig. 10]

DL = dictionary learning

(not "deep learning")









### Ill-posed inverse problems





- **y** : measurements ε : noise
- **x** : unknown image
- A : system matrix (typically wide)

compressed sensing (*e.g.*, MRI)

k<sub>y</sub>

 $k_{x}$ 

deblurring (restoration)

- in-painting
- denoising (not ill posed)

(A Toeplitz)(A subset of rows of I)(A = I)

(**A** "random" rows of DFT)





If we have a prior p(x), then the MAP estimate is:

$$\hat{\boldsymbol{x}} = \operatorname*{arg\,max}_{\boldsymbol{x}} \operatorname{p}(\boldsymbol{x} \mid \boldsymbol{y}) = \operatorname*{arg\,max}_{\boldsymbol{x}} \log \operatorname{p}(\boldsymbol{y} \mid \boldsymbol{x}) + \log \operatorname{p}(\boldsymbol{x})$$
.

For gaussian measurement errors and a linear forward model:

$$-\log p(\boldsymbol{y} | \boldsymbol{x}) \equiv \frac{1}{2} \| \boldsymbol{y} - \boldsymbol{A} \boldsymbol{x} \|_{\boldsymbol{W}}^2$$

where  $\|\boldsymbol{y}\|_{\boldsymbol{W}}^2 = \boldsymbol{y}' \boldsymbol{W} \boldsymbol{y}$ 

and  $\boldsymbol{W}^{-1} = \text{Cov}\{\boldsymbol{y} \mid \boldsymbol{x}\}$  is known (**A** from physics, **W** from statistics)

### Priors for MAP estimation



▶ If all images **x** are "plausible" (have non-zero probability) then

$$p(\mathbf{x}) \propto e^{-R(\mathbf{x})} \Longrightarrow -\log p(\mathbf{x}) \equiv R(\mathbf{x})$$

(from fantasy / imagination / wishful thinking / data)

• MAP  $\equiv$  regularized weighted least-squares (WLS) estimation:

$$\hat{\boldsymbol{x}} = \arg \max_{\boldsymbol{x}} \log p(\boldsymbol{y} | \boldsymbol{x}) + \log p(\boldsymbol{x})$$
$$= \arg \min_{\boldsymbol{x}} \frac{1}{2} \|\boldsymbol{y} - \boldsymbol{A}\boldsymbol{x}\|_{\boldsymbol{W}}^2 + \mathsf{R}(\boldsymbol{x})$$

- A regularizer R(x), aka log prior, is essential for high-quality solutions to ill-conditioned / ill-posed inverse problems.
- ▶ Why ill-posed? Often high ambitions...

### Non-adaptive regularizers

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- Tikhonov regularization (IID gaussian prior)
- Markov random field (MRF) models
- Roughness penalty (cf MRF prior)
- Edge-preserving regularization
- Total-variation (TV) regularization
- Black-box denoiser like NLM, e.g., plug-and-play ADMM [10]
- Sparsity in ambient space
- Sparsifying transforms: wavelets, curvelets, ....
- Graphical models

All "hand crafted" from statistical / mathematical models ...

# Simpler methods for ML in image reconstruction

Many possible ways to use ML ideas in image reconstruction.

Basic "fast" methods:

- Enhance raw data (k-space, sinogram, ...)
- Enhance poorly reconstructed image
  - patch-based
  - image-based

Computation / quality trade-offs ?

. . .

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# Simpler methods for ML in image reconstruction

Many possible ways to use ML ideas in image reconstruction.

Basic "fast" methods:

- Enhance raw data (k-space, sinogram, ...)
- Enhance poorly reconstructed image
  - patch-based
  - image-based

Computation / quality trade-offs ?

Basic "slow" methods:

- Auto-tune regularization parameter(s)
- Provide an initial image for "conventional" iterative reconstruction

May not fully exploit the potential of ML

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ML-based "prior" image for iterative reconstruction [11]:

$$\hat{\boldsymbol{x}} = \operatorname*{arg\,min}_{\boldsymbol{x}} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_2^2 + \beta \|\boldsymbol{x} - \boldsymbol{x}_{\mathrm{prior}}\|_{
ho}^{
ho}$$

Fast for p = 2, but p = 1 more robust to errors in prior image Reminiscent of U. Wisconsin's PICCS methods, *e.g.*, [12]



ML-based "prior" image for iterative reconstruction [11]:

$$\hat{\boldsymbol{x}} = \operatorname*{arg\,min}_{\boldsymbol{x}} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{2}^{2} + \beta \|\boldsymbol{x} - \boldsymbol{x}_{\mathrm{prior}}\|_{p}^{p}$$

Fast for p = 2, but p = 1 more robust to errors in prior image Reminiscent of U. Wisconsin's PICCS methods, *e.g.*, [12]

▶ Unrolled loop (recurrent NN) with learned components [13–16]

### Nonlinear encoder methods for ML-based IR

- ML-based nonlinear encoder, *e.g.*, autoencoder or generative adversarial network (GAN) [17, 18]: nonlinear generalizations of subspace models
- learn G: maps low-dimensional latent parameter z into high-dimensional image x
- Synthesis form [19]:

$$\hat{oldsymbol{x}} = G(\hat{oldsymbol{z}}), \qquad \hat{oldsymbol{z}} = rgmin_{oldsymbol{z}} \|oldsymbol{A}G(oldsymbol{z}) - oldsymbol{y}\|_2^2$$

Challenges:  $\hat{x} \in \text{Range}(G)$ , non-convex minimization



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### Nonlinear encoder methods for ML-based IR

- ML-based nonlinear encoder, *e.g.*, autoencoder or generative adversarial network (GAN) [17, 18]: nonlinear generalizations of subspace models
- learn G: maps low-dimensional latent parameter z into high-dimensional image x
- Synthesis form [19]:

$$\hat{m{x}} = G(\hat{m{z}}), \qquad \hat{m{z}} = rgmin_{m{z}} \|m{A}G(m{z}) - m{y}\|_2^2$$

Challenges:  $\hat{x} \in \text{Range}(G)$ , non-convex minimization

Regularizer form:

$$\hat{\boldsymbol{x}} = \underset{\boldsymbol{x}}{\arg\min} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{2}^{2} + \beta R_{\text{encoder}}(\boldsymbol{x})$$
$$R_{\text{encoder}}(\boldsymbol{x}) = \underset{\boldsymbol{z}}{\min} \|\boldsymbol{x} - G(\boldsymbol{z})\|_{p}^{p}$$

Expensive non-convex double minimization, but more robust to encoder?



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### Nonlinear encoder illustration



From jupyter notebook for [20] (13 layer CNN with  $\approx$  300K learned parameters) at

 ${\tt https://github.com/skolouri/swae/blob/master/MNIST_SlicedWassersteinAutoEncoder_Circle.ipynblocks$ 

 $\mapsto$  $m{x} = m{G}(m{z}) \in \mathbb{R}^{28 imes 28}$  $z \in \mathbb{R}^2$ 1.0 100 203 404

100

200

300

# Generative Adversarial Networks (GAN) example



### From Google's [21]:



Much more realistic than linear interpolation (averaging). "setting a new milestone in visual quality" [21].

# Generative Adversarial Networks (GAN) example



### From Google's [21]:



Non-physical output!





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- Population adaptive methods (*e.g.*, X-ray CT)
- Patient adaptive methods (e.g., dynamic MRI?)
- Spatial structure
  - Patch-based models
  - Convolutional models
- Regularizer formulation
  - Synthesis (dictionary) approach
  - Analysis (sparsifying transforms) approach

Many options...

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### Patch-wise transform sparsity model



Assumption: if  $\boldsymbol{x}$  is a plausible image, then each  $\Omega \boldsymbol{P}_m \boldsymbol{x}$  is sparse.

- $P_m x$  extracts the *m*th of *M* patches from x
- $\blacktriangleright \ \Omega$  is a square sparsifying transform matrix


# Sparsifying transform learning (population adaptive)



Given training images  $x_1, \ldots, x_L$  from a representative population, find transform  $\Omega_*$  that best sparsifies their patches:

$$\boldsymbol{\Omega}_{*} = \operatorname*{arg\,min}_{\boldsymbol{\Omega} \text{ unitary}} \min_{\left\{\boldsymbol{z}_{l,m}\right\}} \sum_{l=1}^{L} \sum_{m=1}^{M} \left\|\boldsymbol{\Omega}\boldsymbol{P}_{m}\boldsymbol{x}_{l} - \boldsymbol{z}_{l,m}\right\|_{2}^{2} + \alpha \left\|\boldsymbol{z}_{l,m}\right\|_{0}$$

- Encourage aggregate sparsity, not patch-wise sparsity (cf K-SVD [22])
- Non-convex due to unitary constraint and  $\|\cdot\|_0$
- Efficient alternating minimization algorithm [23]
  - z update is simply hard thresholding
  - $\Omega$  update is an orthogonal Procrustes problem (SVD)
  - Subsequence convergence guarantees [23]

# Example of learned sparsifying transform



#### 3D X-ray training data



Parts of learned sparsifier  $\Omega_*$ 



(2D slices in x-y, x-z, y-z, from 3D image volume)  $8 \times 8 \times 8$  patches  $\implies \Omega_*$  is  $8^3 \times 8^3 = 512 \times 512$ top 8  $\times$  8 slice of 256 of the 512 rows of  $\Omega_{*}\uparrow_{_{29/71}}$ 

### Regularizer based on learned sparsifying transform

Regularized inverse problem [24]:

$$\hat{\boldsymbol{x}} = \operatorname*{arg\,min}_{\boldsymbol{x}} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{\boldsymbol{W}}^2 + \beta \operatorname{\mathsf{R}}(\boldsymbol{x})$$

$$\mathsf{R}(\mathbf{x}) = \min_{\{\mathbf{z}_m\}} \sum_{m=1}^M \|\mathbf{\Omega}_* \mathbf{P}_m \mathbf{x} - \mathbf{z}_m\|_2^2 + \alpha \|\mathbf{z}_m\|_0.$$

 $\Omega_{\ast}$  adapted to population training data

Alternating minimization optimizer:

- *z<sub>m</sub>* update is simple hard thresholding
- x update is a quadratic problem: many options
   Linearized augmented Lagrangian method (LALM) [25]



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### Example: low-dose 3D X-ray CT simulation



X. Zheng, S. Ravishankar, Y. Long, JF:

IEEE T-MI, June 2018 [24]



### 3D X-ray CT simulation Error maps

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- Physics / statistics provides dramatic improvement
- Data adaptive regularization further reduces RMSE



Given training images  $x_1, \ldots, x_L$  from a representative population, find a set of transforms  $\{\hat{\Omega}_k\}_{k=1}^{K}$  that best sparsify image patches:

$$\begin{cases} \hat{\boldsymbol{\Omega}}_{k} \end{cases} = \underset{\{\boldsymbol{\Omega}_{k} \text{ unitary}\}}{\arg\min} \underset{\{k_{l,m} \in \{1,...,K\}\}}{\min} \underset{\{\boldsymbol{z}_{l,m}\}}{\min} \\ \sum_{l=1}^{L} \sum_{m=1}^{M} \left\| \boldsymbol{\Omega}_{k_{l,m}} \boldsymbol{P}_{m} \boldsymbol{x}_{l} - \boldsymbol{z}_{l,m} \right\|_{2}^{2} + \alpha \left\| \boldsymbol{z}_{l,m} \right\|_{0}$$

- Joint unsupervised clustering / sparsification
- Further nonconvexity due to clustering
- Efficient alternating minimization algorithm [26]

# Example: 3D X-ray CT learned set of transforms





### Example: 3D X-ray CT ULTRA for chest scan





Zheng et al., IEEE T-MI, June 2018 [24]



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Drawback of basic patch-based methods:  $512 \times 512 \times 512$  3D X-ray CT image volume  $8 \times 8 \times 8$  patches  $\implies 512^3 \cdot 8^3 \cdot 4 = 256$  Gbyte of patch data for stride=1

## Convolutional sparsity model

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Assumption: There is a set of filters  $\{\boldsymbol{h}_k\}_{k=1}^K$  such that the images  $\{\boldsymbol{h}_k * \boldsymbol{x}\}$  are sparse for a plausible image  $\boldsymbol{x}$ .

- For more plausible images,  $\{h_k * x\}$  is more sparse.
- \* denotes convolution
- Inherently shift invariant and no patches



# Sparsifying filter learning (population adaptive)

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Given training images  $\mathbf{x}_1, \ldots, \mathbf{x}_L$  from a representative population, find filters  $\{\hat{\mathbf{h}}_k\}_{k=1}^K$  that best sparsify them:

$$\left\{ \hat{\boldsymbol{h}}_{k} \right\} = \underset{\{\boldsymbol{h}_{k}\} \in \mathcal{H}}{\arg\min} \min_{\{\boldsymbol{z}_{l,k}\}} \sum_{l=1}^{L} \sum_{k=1}^{K} \|\boldsymbol{h}_{k} \ast \boldsymbol{x}_{l} - \boldsymbol{z}_{l,k}\|_{2}^{2} + \alpha \|\boldsymbol{z}_{l,k}\|_{0}$$

► To encourage filter diversity:

• 
$$\mathcal{H} = \{ \boldsymbol{H} : \boldsymbol{H}\boldsymbol{H}' = \boldsymbol{I} \}, \ \boldsymbol{H} = [\boldsymbol{h}_1 \ \dots \ \boldsymbol{h}_K]$$

- *cf.* tight-frame condition  $\sum_{k=1}^{K} \| \boldsymbol{h}_k * \boldsymbol{x} \|_2^2 \propto \| \boldsymbol{x} \|_2^2$
- Encourage aggregate sparsity, period
- ▶ Non-convex due to constraint  $\mathcal{H}$  and  $\|\cdot\|_0$
- Efficient alternating minimization algorithm [27]
  - z update is simply hard thresholding
  - Filter update uses diagonal majorizer, proximal map (SVD)
  - Subsequence convergence guarantees [27]

### Examples of learned sparsifying filters









### Regularizer based on learned sparsifying filters

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Regularized inverse problem [27]:

$$\hat{\boldsymbol{x}} = \underset{\boldsymbol{x} \succeq \boldsymbol{0}}{\arg\min} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{\boldsymbol{W}}^2 + \beta \operatorname{\mathsf{R}}(\boldsymbol{x})$$
$$\operatorname{\mathsf{R}}(\boldsymbol{x}) = \underset{\{\boldsymbol{z}_k\}}{\arg\min} \sum_{k=1}^{K} \left\| \hat{\boldsymbol{h}}_k * \boldsymbol{x} - \boldsymbol{z}_k \right\|_2^2 + \alpha \|\boldsymbol{z}_k\|_0.$$

 $\left\{ \hat{m{h}}_k 
ight\}$  adapted to population training data

Block proximal gradient with majorizer (BPG-M) optimizer:

- z<sub>k</sub> update is simple hard thresholding
- x update is a quadratic problem: diagonal majorizer

I. Y. Chun, JF, 2018, arXiv 1802.05584 [27]

### Example: sparse-view 2D X-ray CT simulation





EР





Adaptive CAOL

FBP



123 views (out of usual 984)  $\implies$  8× dose reduction

RMSE (in HU):			
FBP	82.8		
EP	40.8		
Adaptive filters	35.2		

- Physics / statistics provides dramatic improvement
- Data-adaptive regularization further reduces RMSE

## Extension to multiple layers (cf CNN) I



Convolutional sparsity model:  $h_k * x$  is sparse for  $k = 1, ..., K_1$ Learning 1 "layer" of filters:

$$\{\hat{\boldsymbol{h}}_{k}^{[1]}\} = \underset{\{\boldsymbol{h}_{k}^{[1]}\} \in \mathcal{H}}{\arg\min} \min_{\{\boldsymbol{z}_{l,k}^{[1]}\}} \sum_{l=1}^{L} \sum_{k=1}^{K_{1}} \left\|\boldsymbol{h}_{k}^{[1]} * \boldsymbol{x}_{l} - \boldsymbol{z}_{l,k}^{[1]}\right\|_{2}^{2} + \alpha \left\|\boldsymbol{z}_{l,k}^{[1]}\right\|_{0}^{2}$$

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Learning 2 layers of filters [27]:

$$\left( \{ \hat{\boldsymbol{h}}_{k}^{[1]} \}, \{ \hat{\boldsymbol{h}}_{k}^{[2]} \} \right) = \arg\min_{\{\boldsymbol{h}_{k}^{[1]} \}, \{ \boldsymbol{h}_{k}^{[2]} \} \in \mathcal{H}} \min_{\{\boldsymbol{z}_{l,k}^{[1]} \}, \{ \boldsymbol{z}_{l,k}^{[2]} \}} \sum_{\substack{l=1 \ k=1}}^{L} \sum_{k=1}^{K_{1}} \left\| \boldsymbol{h}_{k}^{[1]} * \boldsymbol{x}_{l} - \boldsymbol{z}_{l,k}^{[1]} \right\|_{2}^{2} + \alpha \left\| \boldsymbol{z}_{l,k}^{[1]} \right\|_{0}^{2} + \sum_{l=1}^{L} \sum_{k=1}^{K_{2}} \left\| \boldsymbol{h}_{k}^{[2]} * \left( \boldsymbol{P}_{k} \boldsymbol{z}_{l}^{[1]} \right) - \boldsymbol{z}_{l,k}^{[2]} \right\|_{2}^{2} + \alpha \left\| \boldsymbol{z}_{l,k}^{[2]} \right\|_{0}^{2}$$

Here  $P_k$  is a pooling operator for the output of first layer Block proximal gradient with majorizer (BPG-M) optimizer

I. Y. Chun, JF, 2018, arXiv 1802.05584  $\left[ 27 \right]$ 

Use multi-level learned filters as (interpretable?) regularizer for CT.



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### Patch-wise dictionary sparsity model



Assumption: if  $\boldsymbol{x}$  is a plausible image, then each patch has

 $P_m x \approx D z_m$ 

for a sparse coefficient vector  $\boldsymbol{z}_m$ . (Synthesis approach.)

- $P_m x$  extracts the *m*th of *M* patches from x
- **D** is a (typically overcomplete) dictionary for patches



# MR reconstruction using adaptive dictionary regularizer

Dictionary-blind MR image reconstruction:

$$\hat{\boldsymbol{x}} = \arg\min_{\boldsymbol{x}} \frac{1}{2} \|\boldsymbol{y} - \boldsymbol{A}\boldsymbol{x}\|_{2}^{2} + \beta R(\boldsymbol{x})$$

$$R(\boldsymbol{x}) = \min_{\boldsymbol{D} \in \mathcal{D}} \min_{\boldsymbol{z}' \in \mathcal{C}} \sum_{m=1}^{M} \left( \|\boldsymbol{P}_{m}\boldsymbol{x} - \boldsymbol{D}\boldsymbol{z}_{m}\|_{2}^{2} + \lambda^{2} \|\boldsymbol{z}_{m}\|_{0} \right)$$

where  $P_m$  extracts *m*th of *M* image patches.

In words: of the many images...

Alternating (nested) minimization:

- Fixing  $\boldsymbol{x}$  and  $\boldsymbol{D}$ , update each row of  $\boldsymbol{Z} = [\boldsymbol{z}_1 \dots \boldsymbol{z}_M]$  sequentially via hard-thresholding.
- Fixing x and Z, update D using SOUP-DIL [28].
- Fixing **Z** and **D**, updating **x** is a quadratic problem.
  - Efficient FFT solution for single-coil Cartesian MRI.
  - Use CG for non-Cartesian and/or parallel MRI.
- Non-convex, but monotone decreasing and some convergence theory [28].



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ML for IR

### 2D CS MRI results I

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### 2D CS MRI results II





(SNR compared to fully sampled image.) Using  $\|\boldsymbol{z}_m\|_0$  leads to higher SNR than  $\|\boldsymbol{z}_m\|_1$ . Adaptive case is non-convex anyway...

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lm.	Samp.	Acc.	0-fill	Sparse MRI	PANO	DLMRI	SOUP- DILLI	SOUP- DILLO
а	Cart.	7×	27.9	28.6	31.1	31.1	30.8	31.1
b	Cart.	2.5×	27.7	31.6	41.3	40.2	38.5	42.3
с	Cart.	2.5×	24.9	29.9	34.8	36.7	36.6	37.3
с	Cart.	4×	25.9	28.8	32.3	32.1	32.2	32.3
d	Cart.	2.5×	29.5	32.1	36.9	38.1	36.7	38.4
е	Cart.	2.5×	28.1	31.7	40.0	38.0	37.9	41.5
f	2D rand.	5×	26.3	27.4	30.4	30.5	30.3	30.6
g	Cart.	2.5x	32.8	39.1	41.6	41.7	42.2	43.2
Ref.				[29]	[30]	[9]	[28]	[28]

# 2D CS MRI results IV

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Summary: 2D static MR reconstruction from under-sampled data with adaptive dictionary learning and convergent algorithm, faster than K-SVD approach of DLMRI.

# Summary of patch-based, data-driven adaptive regularizers

Use training data to learn:

- dictionary **D** (for patches)
- sparsifying transform(s)  $\Omega$  (for patches)

• or convolutional versions thereof [32, 33]

ML-based regularized optimization problem using M image patches:

$$\hat{\boldsymbol{x}} = \underset{\boldsymbol{x}}{\arg\min} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{2}^{2} + \beta R_{\mathrm{ML}}(\boldsymbol{x})$$

$$R_{\mathrm{ML-DL}}(\boldsymbol{x}) = \underset{\{\boldsymbol{z}_{m}\}}{\min} \sum_{m=1}^{M} \|\boldsymbol{P}_{m}\boldsymbol{x} - \boldsymbol{D}\boldsymbol{z}_{m}\|_{2}^{2} + \alpha \|\boldsymbol{z}_{m}\|_{0}$$

$$R_{\mathrm{ML-ST}}(\boldsymbol{x}) = \underset{\{\boldsymbol{z}_{m}\}}{\min} \sum_{m=1}^{M} \|\boldsymbol{\Omega}\boldsymbol{P}_{m}\boldsymbol{x} - \boldsymbol{z}_{m}\|_{2}^{2} + \alpha \|\boldsymbol{z}_{m}\|_{0}$$

Alternative: blind adaptive learned dictionary [9] or learned sparsifying transform [34]. Double minimization (so very "deep?") More interpretable than CNNs?

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Unrolled loop method with 20 layers trained with  $1.3\cdot10^6$  MR image 8  $\times$  8 patches <code>Ravishankar</code> et al., ISBI 2018 [15]



#### Tested with 5 different MR images:











### Training an unrolled loop II

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	M
	UNIVERSITY OF
I	MICHIGAN

Undersampling	Image	Zero-filled	Sparse MRI	UTMRI	Unrolled
3.3×	1	25.6	26.7	28.3	28.2
	2	25.2	26.6	27.9	27.8
	3	26.0	27.3	29.3	28.9
	4	25.4	26.7	28.2	28.1
	5	27.2	28.9	30.6	30.3
Avg. PSNR change	-	-	1.36	2.98	2.78
5×	1	24.7	25.9	27.6	27.5
	2	24.2	25.5	27.2	27.0
	3	24.9	26.3	28.5	28.0
	4	24.4	25.7	27.6	27.4
	5	26.2	27.9	29.8	29.5
Avg. PSNR change	-	-	1.38	3.26	3.0
Approx recon time	-	-	100s	240s	50s

Results:

Sparse MRI [35] total variation (TV) and wavelets UTMRI [26] (union of learned sparsifying transforms): adaptive, not "deep"



Quantitative MRI:images  $\rightarrow$  estimation $\rightarrow$  parameters (T1, T2, ...)

- Traditional nonlinear estimation methods:
  - nonlinear least squares
  - dictionary matching (quantized maximum likelihood via variable projection)
- Machine-learning methods
  - deep neural network regression [36–39] Requires long training times
  - parameter estimation via kernel regression (PERK) Gopal Nataraj et al., ISBI 2017, IEEE T-MI 2018 [40, 41]

#### Parameter estimation via kernel regression (PERK) example J. Fessler ML for IR



### PERK applied to myelin water imaging



6 parameters (T1 slow/fast, T2 slow/fast,  $M_0$ , fast fraction) Estimated from 3 optimized dual-echo steady state (DESS) scans [42]



PERK training: 33.8s, testing 0.99s / slice





MESE scan took 32m (16m  $\times$ 2) DESS scan took 3m15s Take away: "traditional" machine learning is still useful...



#### Introduction

ML-based image reconstruction approaches

#### Adaptive regularization

Patch-based adaptive regularizers Convolutional adaptive regularizers Blind dictionary learning

#### Other ML4MI topics

#### Summary

Bibliography



- Machine learning has great potential for medical imaging
- Much excitement but many challenges
- Image reconstruction seems especially suitable for ML ideas
- Data-driven, adaptive regularizers beneficial for low-dose CT and under-sampled MRI
- More comparisons between model-based methods with adaptive regularizers and CNN-based methods needed
- Machine learning tools like kernel regression remain useful


- Overviews: [43, 45, 46]
- ► Generative models: [20, 47]:
- Deep learning myths [48]

▶ ...

- ▶ NN complexity analysis / function approximation [49–51] [52]
- Application to MR fingerprinting [36, 39]
- ▶ MR reconstruction / enhancement using CNN [16, 53–60]
- Dynamic MR reconstruction using CNN [61]

#### Resources



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



# **Bibliography I**



- H. Greenspan, B. van Ginneken, and R. M. Summers. "Guest editorial deep learning in medical imaging: overview and future promise of an exciting new technique." In: IEEE Trans. Med. Imag. 35.5 (May 2016), 1153–9.
- [2] G. Wang, J. C. Ye, K. Mueller, and J. A. Fessler. "Image reconstruction is a new frontier of machine learning." In: IEEE Trans. Med. Imag. 37.6 (June 2018), 1289–96.
- [3] E. Haneda, B. Claus, P. FitzGerald, G. Wang, and B. De Man. "CT sinogram analysis using deep learning." In: Proc. 5th Intl. Mtg. on Image Formation in X-ray CT. 2018, 419–22.
- [4] L. Baldassarre, Y-H. Li, J. Scarlett, B. Gozcu, I. Bogunovic, and V. Cevher. "Learning-based compressive subsampling." In: IEEE J. Sel. Top. Sig. Proc. 10.4 (June 2016), 809–22.
- [5] B. Gozcu, R. K. Mahabadi, Y-H. Li, E. Ilicak, T. Cukur, J. Scarlett, and V. Cevher. "Learning-based compressive MRI." In: IEEE Trans. Med. Imag. 37.6 (June 2018), 1394–406.
- [6] Y. Cao and D. N. Levin. "Feature-recognizing MRI." In: Mag. Res. Med. 30.3 (Sept. 1993), 305–17.
- [7] Y. Cao, D. N. Levin, and L. Yao. "Locally focused MRI." In: Mag. Res. Med. 34.6 (Dec. 1995), 858-67.
- [8] Y. Cao and D. N. Levin. "Using an image database to constrain the acquisition and reconstruction of MR images of the human head." In: IEEE Trans. Med. Imag. 14.2 (June 1995), 350–61.
- 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.
- [10] S. H. Chan, X. Wang, and O. A. Elgendy. "Plug-and-play ADMM for image restoration: fixed-point convergence and applications." In: IEEE Trans. Computational Imaging 3.1 (Mar. 2017), 84–98.
- [11] G. Yang, S. Yu, H. Dong, G. Slabaugh, P. L. Dragotti, X. Ye, F. Liu, S. Arridge, J. Keegan, Y. Guo, and D. Firmin. "DAGAN: Deep de-aliasing generative adversarial networks for fast compressed sensing MRI reconstruction." In: IEEE Trans. Med. Imag. 37.6 (June 2018), 1310–21.

# **Bibliography II**



- [12] G-H. Chen, J. Tang, and S. Leng. "Prior image constrained compressed sensing (PICCS): A method to accurately reconstruct dynamic CT images from highly undersampled projection data sets." In: Med. Phys. 35.2 (Feb. 2008), 660–3.
- [13] K. Gregor and Y. LeCun. "Learning fast approximations of sparse coding." In: Proc. Intl. Conf. Mach. Learn. 2010.
- [14] Y. Chen and T. Pock. "Trainable nonlinear reaction diffusion: A flexible framework for fast and effective image restoration." In: IEEE Trans. Patt. Anal. Mach. Int. 39.6 (June 2017), 1256–72.
- [15] S. Ravishankar, A. Lahiri, C. Blocker, and J. A. Fessler. "Deep dictionary-transform learning for image reconstruction." In: Proc. IEEE Intl. Symp. Biomed. Imag. 2018, 1208–12.
- [16] K. Hammernik, T. Klatzer, E. Kobler, M. P. Recht, D. K. Sodickson, T. Pock, and F. Knoll. "Learning a variational network for reconstruction of accelerated MRI data." In: Mag. Res. Med. 79.6 (June 2018), 3055–71.
- [17] I. J. Goodfellow, J. Pouget-Abadie, M. Mirza, B. Xu, D. Warde-Farley, S. Ozair, A. Courville, and Y. Bengio. Generative adversarial networks. 2014.
- [18] X. Chen, Y. Duan, R. Houthooft, J. Schulman, I. Sutskever, and P. Abbeel. "InfoGAN: interpretable representation learning by information maximizing generative adversarial nets." In: Neural Info. Proc. Sys. 2016, 2172–80.
- [19] A. Bora, A. Jalal, E. Price, and A. G. Dimakis. "Compressed sensing using generative models." In: Proc. Intl. Conf. Mach. Learn. Vol. 70. 2017, 537–46.
- [20] S. Kolouri, P. E. Pope, C. E. Martin, and G. K. Rohde. Sliced-Wasserstein autoencoder: an embarrassingly simple generative model. 2018.
- [21] D. Berthelot, T. Schumm, and L. Metz. BEGAN: boundary equilibrium generative adversarial networks. 2017.
- [22] 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.
- [23] S. Ravishankar and Y. Bresler. "I<sub>0</sub> sparsifying transform learning with efficient optimal updates and convergence guarantees." In: IEEE Trans. Sig. Proc. 63.9 (May 2015), 2389–404.

# **Bibliography III**



- [24] X. Zheng, S. Ravishankar, Y. Long, and J. A. Fessler. "PWLS-ULTRA: An efficient clustering and learning-based approach for low-dose 3D CT image reconstruction." In: IEEE Trans. Med. Imag. 37.6 (June 2018), 1498–510.
- [25] H. Nien and J. A. Fessler. "Relaxed linearized algorithms for faster X-ray CT image reconstruction." In: IEEE Trans. Med. Imag. 35.4 (Apr. 2016), 1090–8.
- [26] S. Ravishankar and Y. Bresler. "Data-driven learning of a union of sparsifying transforms model for blind compressed sensing." In: IEEE Trans. Computational Imaging 2.3 (Sept. 2016), 294–309.
- [27] I. Y. Chun and J. A. Fessler. "Convolutional analysis operator learning: acceleration, convergence, application, and neural networks." In: IEEE Trans. Im. Proc. (2018). Submitted.
- [28] S. Ravishankar, R. R. Nadakuditi, and J. A. Fessler. "Efficient sum of outer products dictionary learning (SOUP-DIL) and its application to inverse problems." In: IEEE Trans. Computational Imaging 3.4 (Dec. 2017), 694–709.
- [29] M. Lustig and J. M. Pauly. "SPIRiT: Iterative self-consistent parallel imaging reconstruction from arbitrary k-space." In: Mag. Res. Med. 64.2 (Aug. 2010), 457–71.
- [30] X. Qu, Y. Hou, F. Lam, D. Guo, J. Zhong, and Z. Chen. "Magnetic resonance image reconstruction from undersampled measurements using a patch-based nonlocal operator." In: *Med. Im. Anal.* 18.6 (Aug. 2014), 843–56.
- [31] Z. Zhan, J-F. Cai, D. Guo, Y. Liu, Z. Chen, and X. Qu. "Fast multiclass dictionaries learning with geometrical directions in MRI reconstruction." In: IEEE Trans. Biomed. Engin. 63.9 (Sept. 2016), 1850–61.
- [32] I. Y. Chun and J. A. Fessler. "Convolutional dictionary learning: acceleration and convergence." In: IEEE Trans. Im. Proc. 27.4 (Apr. 2018), 1697–712.
- [33] I. Y. Chun and J. A. Fessler. Convolutional analysis operator learning: acceleration, convergence, application, and neural networks. 2018.
- [34] S. Ravishankar and Y. Bresler. "Efficient blind compressed sensing using sparsifying transforms with convergence guarantees and application to MRI." In: SIAM J. Imaging Sci. 8.4 (2015), 2519–57.

## **Bibliography IV**



- [35] M. Lustig, D. Donoho, and J. M. Pauly. "Sparse MRI: The application of compressed sensing for rapid MR imaging." In: Mag. Res. Med. 58.6 (Dec. 2007), 1182–95.
- [36] P. Virtue, S. X. Yu, and M. Lustig. "Better than real: Complex-valued neural nets for MRI fingerprinting." In: Proc. IEEE Intl. Conf. on Image Processing. 2017, 3953–7.
- [37] A. Lahiri, J. A. Fessler, and L. Hernandez-Garcia. "Optimized design of MRF scan parameters for ASL signal acquisition." In: ISMRM Workshop on MR Fingerprinting. 2017.
- [38] A. Lahiri, J. A. Fessler, and L. Hernandez-Garcia. "Optimized scan design for ASL fingerprinting and multiparametric estimation using neural network regression." In: Proc. Intl. Soc. Mag. Res. Med. 2018, p. 309.
- [39] O. Cohen, B. Zhu, and M. S. Rosen. "MR fingerprinting Deep RecOnstruction NEtwork (DRONE)." In: Mag. Res. Med. 80.3 (Sept. 2018), 885–94.
- [40] G. Nataraj, J-F. Nielsen, and J. A. Fessler. "Dictionary-free MRI parameter estimation via kernel ridge regression." In: Proc. IEEE Intl. Symp. Biomed. Imag. 2017, 5–9.
- [41] G. Nataraj, J-F. Nielsen, C. D. Scott, and J. A. Fessler. "Dictionary-free MRI PERK: Parameter estimation via regression with kernels." In: IEEE Trans. Med. Imag. 37.9 (Sept. 2018), 2103–14.
- [42] G. Nataraj, J-F. Nielsen, M. Gao, and J. A. Fessler. Fast, precise myelin water quantification using DESS MRI and kernel learning. Submitted. 2018.
- [43] G. Wang, M. Kalra, and C. G. Orton. "Machine learning will transform radiology significantly within the next five years." In: Med. Phys. 44.6 (June 2017), 2041–4.
- [44] A. Esteva, B. Kuprel, R. A. Novoa, J. Ko, S. M. Swetter, H. M. Blau, and S. Thrun. "Dermatologist-level classification of skin cancer with deep neural networks." In: Nature 542.7639 (2017), 115–8.
- [45] G. Wang. "A perspective on deep imaging." In: IEEE Access 4 (Nov. 2016), 8914–24.

## **Bibliography V**



- [46] M. T. McCann, K. H. Jin, and M. Unser. "Convolutional neural networks for inverse problems in imaging: A review." In: IEEE Sig. Proc. Mag. 34.6 (Nov. 2017), 85–95.
- [47] I. Deshpande, Z. Zhang, and A. Schwing. "Generative modeling using the sliced Wasserstein distance." In: Proc. IEEE Conf. on Comp. Vision and Pattern Recognition. 2018.
- [48] S. Rakhlin. MythBusters: A Deep Learning Edition. Slides dated Jan 18-19, 2018. 2018.
- [49] N. Golowich, A. Rakhlin, and O. Shamir. Size-independent sample complexity of neural networks. 2017.
- [50] T. Liang, T. Poggio, A. Rakhlin, and J. Stokes. Fisher-Rao metric, geometry, and complexity of neural networks. 2017.
- [51] M. Raghu, B. Poole, J. Kleinberg, S. Ganguli, and J. Sohl-Dickstein. "On the expressive power of deep neural networks." In: Proc. Intl. Conf. Mach. Learn. Vol. 70, 2017, 2847–54.
- [52] S. Liang and R. Srikant. "Why deep neural networks for function approximation?" In: Proc. Intl. Conf. on Learning Representations. 2017.
- [53] S. Ravishankar, I. Y. Chun, and J. A. Fessler. "Physics-driven deep training of dictionary-based algorithms for MR image reconstruction." In: Proc., IEEE Asilomar Conf. on Signals, Systems, and Comp. Invited. 2017, 1859–63.
- [54] M. Mardani, E. Gong, J. Y. Cheng, S. Vasanawala, G. Zaharchuk, M. Alley, N. Thakur, S. Han, W. Dally, J. M. Pauly, and L. Xing. Deep generative adversarial networks for compressed sensing automates MRI. 2017.
- [55] B. Zhu, J. Z. Liu, S. F. Cauley, B. R. Rosen, and M. S. Rosen. "Image reconstruction by domain-transform manifold learning." In: Nature 555 (Mar. 2018), 487–92.
- [56] Y. Han, J. Yoo, H. H. Kim, H. J. Shin, K. Sung, and J. C. Ye. "Deep learning with domain adaptation for accelerated projection-reconstruction MR." In: Mag. Res. Med. 80.3 (Sept. 2018), 1189–205.
- [57] K. H. Jin and M. Unser. "3D BPConvNet to reconstruct parallel MRI." In: Proc. IEEE Intl. Symp. Biomed. Imag. 2018, 361-4.



- [58] H. Jeelani, J. Martin, F. Vasquez, M. Salerno, and D. S. Weller. "Image quality affects deep learning reconstruction of MRI." In: Proc. IEEE Intl. Symp. Biomed. Imag. 2018, 357–60.
- [59] T. M. Quan, T. Nguyen-Duc, and W-K. Jeong. "Compressed sensing MRI reconstruction using a generative adversarial network with a cyclic loss." In: IEEE Trans. Med. Imag. 37.6 (June 2018), 1488–97.
- [60] T. Eo, Y. Jun, T. Kim, J. Jang, H-J. Lee, and D. Hwang. "KIKI-net: cross-domain convolutional neural networks for reconstructing undersampled magnetic resonance images." In: Mag. Res. Med. (2018).
- [61] J. Schlemper, J. Caballero, J. V. Hajnal, A. N. Price, and D. Rueckert. "A deep cascade of convolutional neural networks for dynamic MR image reconstruction." In: IEEE Trans. Med. Imag. 37.2 (Feb. 2018), 491–503.