Medical imaging inverse problems using optimization and machine learning



J Fessler

Jeffrey A. Fessler William L. Root Professor of EECS

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

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

Work with Sai Ravishankar, II Yong Chun, Raj Nadakuditi, Yong Long, Xuehang Zheng, ...

CHEPS Seminar

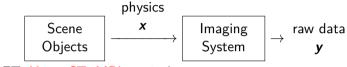
2019-10-08

Background



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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.

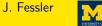
Generations of medical image reconstruction methods

- 1. 70's "Analytical" methods (integral equations) FBP for SPECT / PET / X-ray CT, IFFT for MRI, ...
- 80's Algebraic methods (as in "linear algebra") Solve y = Ax
- 3. 90's Statistical methods
 - LS / ML methods
 - regularized / Bayesian methods
- 4. 00's Compressed sensing methods (mathematical sparsity models)
- 5. 10's Adaptive / data-driven methods machine learning, deep learning, ...



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Improving X-ray CT image reconstruction



- A picture is worth 1000 words
- (and perhaps several 1000 seconds of computation?)



Thin-slice FBP Seconds

ASIR (denoise) A bit longer Statistical Much longer

Today's talk: less about computation, more about image quality Right image used edge-preserving regularization

Safety / health relevance: X-ray dose and diagnostic accuracy

History: Milestones in iterative image reconstruction

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Commercial availability of iterative methods for human scanners per FDA 510(k) dates:

► PET/SPECT

Unregularized OS-EM \approx 1997

► X-ray CT

Regularized MBIR [2011-11-09 for GE Veo] (Installed at UM in Jan. 2012)

► PET

Regularized EM variant (Q.Clear) 2014-03-21

MRI

Compressed sensing! (Sparsity-based regularization) [2017-01-27 for Siemens Cardiac Cine] [2017-04-20 for GE HyperSense]

► Ultrasound?

Accelerating MR imaging using adaptive regularization



(a) $4 \times$ under-sampled MR k-space

(b) zero-filled reconstruction

(c) "compressed sensing" reconstruction with TV regularization

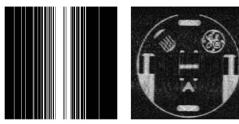
(d) adaptive dictionary learning regularization [1, Fig. 10]

Safety / health relevance:

 \circ scan time

 \circ motion

 \circ image quality









(b)

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Background

$\ensuremath{\mathsf{III}}\xspace$ problems and regularization

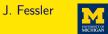
Classical "hand crafted" regularizers

Adaptive regularization

Patch-based adaptive regularizers Convolutional adaptive regularizers Blind dictionary learning

Summary

Ill-posed inverse problems

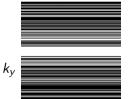






x : unknown image A : system matrix (typically wide)

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



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

 k_x

- deblurring (restoration)
- in-painting
- denoising (not ill posed)

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



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Why under-sample in MRI?

- Reduce scan time (?)
 - Patient comfort
 - Scan cost / throughput
 - Motion artifacts (Philips at ISMRM 2017)
- Improve spatial resolution (collect higher k-space lines)
- Improve scan diversity for quantitative MRI
- Improve temporal resolution trade-off in dynamic MRI

Why under-sample or reduce intensity in CT?

Reduce X-ray dose

(But under-sampling leads to ill-posed inverse problems...)



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)



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Priors for MAP estimation

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

$$\mathsf{p}(\boldsymbol{x}) \propto \mathrm{e}^{-\,\mathsf{R}(\boldsymbol{x})} \Longrightarrow -\log\mathsf{p}(\boldsymbol{x}) \equiv \mathsf{R}(\boldsymbol{x})$$

(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...

- Tikhonov regularization (IID gaussian prior)
- Roughness penalty (Basic MRF prior)
- Sparsity in ambient space
- Edge-preserving regularization
- ► Total-variation (TV) regularization
- Black-box denoiser like NLM



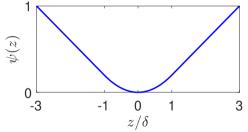
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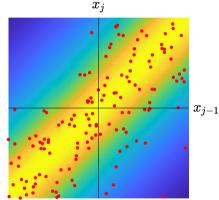
Edge-preserving regularization

Neighboring pixels tend to have similar values except near edges:

$$\mathsf{R}(\boldsymbol{x}) = \beta \sum_{j} \psi(x_j - x_{j-1})$$

Potential function ψ :





- Equivalent to improper prior (agnostic to DC value)
- Accounts for spatial correlations, but only very locally
- Used clinically now for low-dose X-ray CT image reconstruction



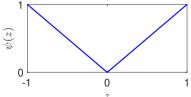
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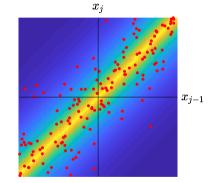
Total-variation (TV) regularization

Neighboring pixels tend to have similar values except near edges ("gradient sparsity"):

$$\begin{aligned} \mathsf{R}(\boldsymbol{x}) &= \beta \operatorname{TV}(\boldsymbol{x}) = \beta \left\| \boldsymbol{\Delta} \boldsymbol{x} \right\|_{1} \\ &= \beta \sum_{j} |x_{j} - x_{j-1}| \end{aligned}$$

Potential function ψ :





- Equivalent to improper prior (agnostic to DC value)
- Accounts for correlations, but only very locally
- Well-suited to piece-wise constant Shepp-Logan phantom!
- Used in many academic publications...



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- Transforms: wavelets, curvelets, ...
- Markov random field models
- Graphical models
- • •

All "hand crafted" ...



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Background

III-posed problems and regularization Classical "hand crafted" regularizers

Adaptive regularization

Patch-based adaptive regularizers Convolutional adaptive regularizers Blind dictionary learning

Summary

🕨 Data

- 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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X-ray CT with learned sparsifying transforms

Data

- Population adaptive methods
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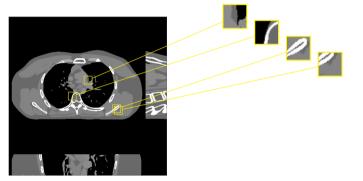


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Patch-wise transform sparsity model

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





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Sparsifying transform learning (population adaptive)

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

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

- Encourage aggregate sparsity, not patch-wise sparsity (cf K-SVD [2])
- Non-convex due to unitary constraint and $\|\cdot\|_0$
- Efficient alternating minimization algorithm [3]
 - z update is simply hard thresholding
 - Ω update is an orthogonal Procrustes problem (SVD)
 - Subsequence convergence guarantees [3]



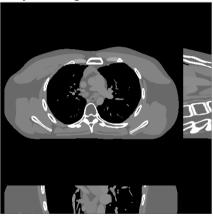
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Example of learned sparsifying transform

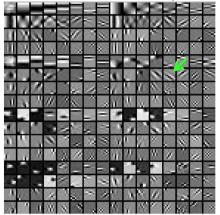




3D X-ray training data







(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 $\Omega_{*}\uparrow_{_{22/50}}$

Regularizer based on learned sparsifying transform

Regularized inverse problem [4]:

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

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

 Ω_{\ast} adapted to population training data

Alternating minimization optimizer:

- z_m update is simple hard thresholding
- x update is a quadratic problem: many options Linearized augmented Lagrangian method (LALM) [5]



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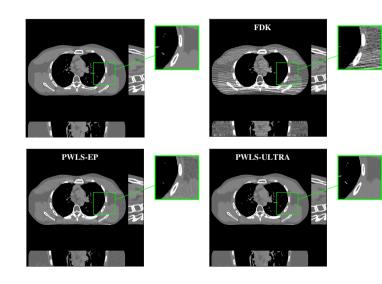
Example: low-dose 3D X-ray CT simulation



X. Zheng, S. Ravishankar,

Y. Long, JF:

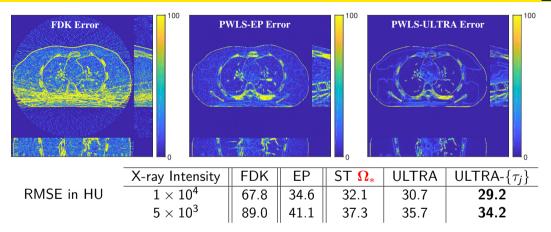
IEEE T-MI, June 2018 [4]



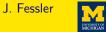
3D X-ray CT simulation Error maps

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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:

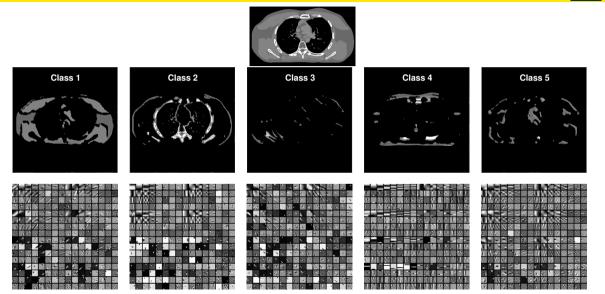
$$\begin{cases} \hat{\boldsymbol{\Omega}}_{k} \end{cases} = \underset{\{\boldsymbol{\Omega}_{k} \text{ unitary}\}}{\arg\min} \underset{\{k_{l,m} \in \{1,...,\mathcal{K}\}\}}{\min} \underset{\{\boldsymbol{z}_{l,m}\}}{\min} \\ \sum_{l=1}^{L} \sum_{m=1}^{M} \left\| \boldsymbol{\Omega}_{k_{l,m}} \boldsymbol{P}_{m}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

- Joint unsupervised clustering / sparsification
- Further nonconvexity due to clustering
- Efficient alternating minimization algorithm [6]

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

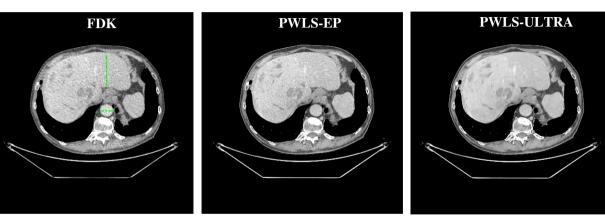






Example: 3D X-ray CT ULTRA for chest scan





Zheng et al., IEEE T-MI, June 2018 [4]







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Summary

X-ray CT with learned convolutional filters



Data

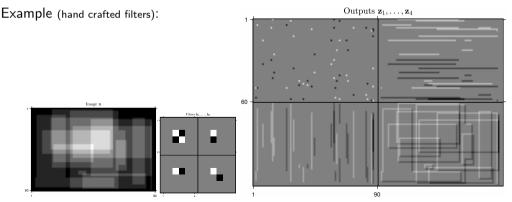
- Population adaptive methods
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Drawback of basic patch-based methods: $512 \times 512 \times 512$ 3D X-ray CT image volume $8 \times 8 \times 8$ patches $\implies 512^3 \cdot 8^3 \cdot 4 = 256$ Gbyte of patch data for stride=1

Convolutional sparsity model

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





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Sparsifying filter learning (population adaptive)

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Given training images x_1, \ldots, x_L from a representative population, find filters $\{\hat{h}_k\}_{k=1}^K$ that best sparsify them:

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

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 [7]
 - z update is simply hard thresholding
 - Filter update uses diagonal majorizer, proximal map (SVD)
 - Subsequence convergence guarantees [7]

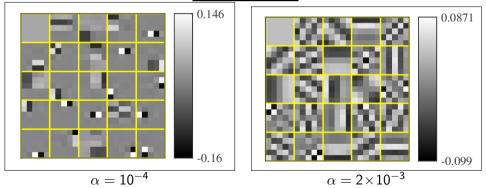
Examples of learned sparsifying filters











Regularizer based on learned sparsifying filters

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Regularized inverse problem [7]:

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

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

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

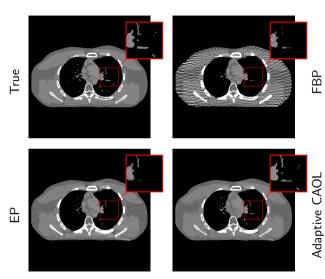
- \triangleright z_k update is simple hard thresholding
- x update is a quadratic problem: diagonal majorizer

I. Y. Chun, JF, 2018, arXiv 1802.05584 [7]

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



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35 / 50



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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

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Extension to multiple layers (cf CNN) I

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

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

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Learning 2 layers of filters [7]:

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

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 $\left[7\right]$

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





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MR with adapted dictionary



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Patch-wise dictionary sparsity model

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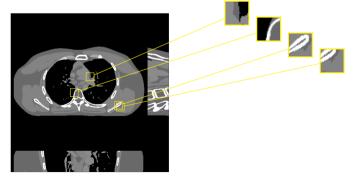
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Assumption: if \boldsymbol{x} is a plausible image, then each patch has

 $P_m x \approx D z_m$

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

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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 \boldsymbol{P}_m extracts *m*th of *M* image patches.

In words: of the many images...

Alternating (nested) minimization:

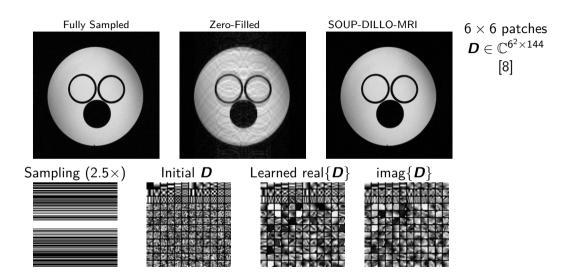
- Fixing x and D, update each row of $Z = [z_1 \dots z_M]$ sequentially via hard-thresholding.
- Fixing *x* and *Z*, update *D* using SOUP-DIL [8].
- Fixing Z and D, updating x is a quadratic problem.
 - Efficient FFT solution for single-coil Cartesian MRI.
 - Use CG for non-Cartesian and/or parallel MRI.

Non-convex, but monotone decreasing and some convergence theory [8].

2D CS MRI results I

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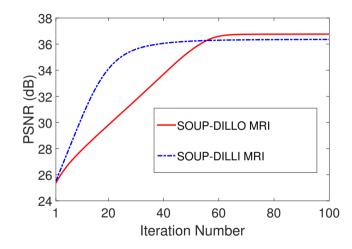




2D CS MRI results II

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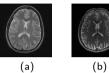


(SNR compared to fully sampled image.) Using $\|\boldsymbol{z}_m\|_0$ leads to higher SNR than $\|\boldsymbol{z}_m\|_1$. Adaptive case is non-convex anyway...

2D CS MRI results III

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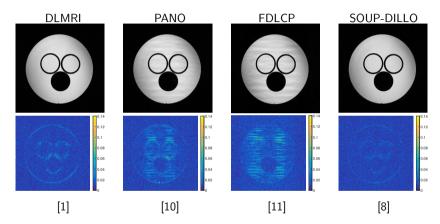
(g)

| 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. | | | | [9] | [10] | [1] | [8] | [8] |

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.





Data-driven / adaptive regularization

- Beneficial for low-dose CT and under-sampled MRI reconstruction
- Dictionary atom structure (e.g., low rank) further helpful for dynamic MRI
- Block proximal methods provide reasonably computational efficiency
- Convergence theory (unlike KSVD)

Future work:

- Synthesis (*e.g.*, dictionary) vs analysis (*e.g.*, transform learning) formulations Begs for some principled model comparison...
- Online methods for reduced memory, better adaptation [12-15]
- Adaptive methods versus "deep" methods?
- Prospective use



June 2018 special issue of IEEE Trans. on Medical Imaging [16]:



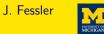
IEEE TRANSACTIONS ON MEDICAL IMAGING, VOL. 37, NO. 6, JUNE 2018

1289

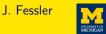
Image Reconstruction Is a New Frontier of Machine Learning

Ge Wang[®], *Fellow, IEEE*, Jong Chu Ye[®], *Senior Member, IEEE*, Klaus Mueller[®], *Senior Member, IEEE*, and Jeffrey A. Fessler[®], *Fellow, IEEE*

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