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Introduction

Image reconstruction

Adaptive regularization

Deep-learning approaches

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#### Medical imaging overview









Most obvious place for machine learning is in post-processing (image analysis). Numerous special issues and surveys in medical imaging journals, *e.g.*, [1–9].





Machine learning for scan design

Choose best k-space phase encoding locations based on training images Hot topic in MRI recently [10–15].

Precursor by Yue Cao and David Levin, MRM Sep. 1993 [16-18].





Machine learning in medical image reconstruction

June 2018 special issue of IEEE Trans. on Medical Imaging [19].

Surveys: [20-27]

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.

#### Medical imaging overview

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A holy grail for machine learning in medical imaging?

- CT sinogram to vessel diameter [28, 29]
- k-space to ???





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### Generations of medical image reconstruction methods

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- 1. 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 based on imaging physics ("model based")
  - 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, ...

#### Two important milestones for clinical CT



• Deep-learning image reconstruction

FDA approved 2019 [31, 32]

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- interpretable (?) optimization formulations
- local prior information only (patch size)
- perhaps slower computation due to optimization iterations
- Train neural network (aka deep learning)
  - less interpretable
  - possibly more global prior information
  - slow training, but perhaps faster computation after trained

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#### Patch-based regularization and TV

Anisotropic discrete TV regularizer:  $R(\mathbf{x}) = \|\mathbf{T}\mathbf{x}\|_1$ where  $\mathbf{T}$  is finite-differences  $\equiv$  patches of size 2 × 1.

Larger patches provide more context for distinguishing signal from noise.

cf. CNN approaches

Patch-based regularizers:

- synthesis models
- analysis methods







## X-ray CT with learned sparsifying transforms



#### Data

- Population adaptive methods
- Patient adaptive methods
- Spatial structure
  - Patch-based models
  - Convolutional models
- Regularizer formulation
  - Synthesis (dictionary) approach
  - Analysis (sparsifying transform) approach

#### Patch-wise transform sparsity model

Assumption: if x is a plausible image, then each patch transform  $TP_m x$  is sparse.

- $P_m x$  extracts the *m*th of *M* patches from x
- **T** is a (often square) sparsifying transform matrix.





What  $\boldsymbol{T}$ ?

# Sparsifying transform learning (population adaptive)



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

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

- Encourage aggregate sparsity, not patch-wise sparsity (cf K-SVD [33])
- Non-convex due to unitary constraint and  $\|\cdot\|_0$
- Efficient alternating minimization algorithm [34]
  - z update : simple hard thresholding
  - **T** update : orthogonal Procrustes problem (SVD)
  - Subsequence convergence guarantees [34]

### Example of learned sparsifying transform





Parts of learned sparsifier  $T_*$ 



(2D slices in x-y, x-z, y-z, from 3D image volume)  $8 \times 8 \times 8$  patches  $\implies \mathbf{T}_*$  is  $8^3 \times 8^3 = 512 \times 512$ 

top 8  $\times$  8 slice of 256 of the 512 rows of  $\textit{\textbf{T}}_{*}\uparrow_{_{14/43}}$ 

### Regularizer based on learned sparsifying transform

Regularized inverse problem [35]:

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

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

 $T_*$  adapted to population training data

Alternating minimization optimizer:

- $\triangleright$   $z_m$  update : simple hard thresholding
- x update : quadratic problem (many options) Linearized augmented Lagrangian method (LALM) [36]



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



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

IEEE T-MI, June 2018 [35].



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

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



$$\left\{\hat{\boldsymbol{T}}_{k}\right\} = \underset{\{\boldsymbol{T}_{k} \text{ unitary}\}}{\arg\min} \min_{\{\boldsymbol{z}_{l,m}\}} \sum_{l=1}^{L} \sum_{m=1}^{M} \left( \min_{k \in \{1,\dots,K\}} \|\boldsymbol{T}_{k}\boldsymbol{P}_{m}\boldsymbol{x}_{l} - \boldsymbol{z}_{l,m}\|_{2}^{2} + \alpha \|\boldsymbol{z}_{l,m}\|_{0} \right)$$

- Joint unsupervised clustering / sparsification
- Further nonconvexity due to clustering
- Efficient alternating minimization algorithm [37]

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## 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 [35] (Special issue on machine learning for image reconstruction) Matlab code: http://web.eecs.umich.edu/~fessler/irt/reproduce/ https://github.com/xuehangzheng/PWLS-ULTRA-for-Low-Dose-3D-CT-Image-Reconstruction





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Overview:

- ▶ image-domain learning [38–40]...
- k-space or data-domain learning e.g., [41], [42], [43]
- transform learning (direct from k-space to image) e.g., AUTOMAP [44], [45–47]
- hybrid-domain learning (unrolled loop, *e.g.*, variational network) alternate between denoising/dealiasing and reconstruction from k-space *e.g.*, [42, 48–52] ...

#### DL for IR: image-domain learning





Figure courtesy of Jong Chul Ye, KAIST University.

- + simple and fast
- $-\,$  aliasing is spatially widespread, requires deep network



#### Investigating Robustness to Unseen Pathologies in Model-Free Deep Multicoil Reconstruction

Gopal Nataraj<sup>1</sup> and Ricardo Otazo<sup>1,2</sup>

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#### Introduction

Speed is often claimed as a key advantage of deep learning (DL) for undersampled parallel MRI reconstruction [1]. However, the only DL approach that to our knowledge has studied generalizability to pathologies unseen in training [2] requires repeated application of the MR acquisition model and its adjoint, just as in iterative methods. In contrast, model-free DL reconstruction has the potential to be much faster. Prior model-free DL work [3] proposes to learn a mapping directly from k-snace but with



[53] ISMRM 2020 Workshop on Data Sampling & Image Reconstruction

#### Dangers of image-domain learning II





Figure 3: Reconstructions in a case of anaplastic astrocytoma, a rare malignant brain tumor. SPARSE-SENSE and DL reconstructions are from the same 4x-accelerated retrospectively undersampled acquisition. DL achieves lower whole-volume MAE than SPARSE-SENSE, but fails to properly reconstruct regions near the tumor.

Use NN output as a "prior" for iterative reconstruction [38, 54]:

$$\hat{\boldsymbol{x}}_{\beta} = \operatorname*{arg\,min}_{\boldsymbol{x}} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{2}^{2} + \beta \|\boldsymbol{x} - \boldsymbol{x}_{\mathsf{NN}}\|_{2}^{2} = (\boldsymbol{A}'\boldsymbol{A} + \beta\boldsymbol{I})^{-1}(\boldsymbol{A}'\boldsymbol{y} + \beta\boldsymbol{x}_{\mathsf{NN}})$$

For single-coil Cartesian case:

• no iterations are needed (solve with FFTs)

- ${\sf lim}_{\beta\to 0}\, \hat{\textbf{\textit{x}}}_\beta$  replaces missing k-space data with FFT of  $\textbf{\textit{x}}_{NN}$
- Iterations needed for parallel MRI and/or non-Cartesian sampling (PCG)

Learn residual (aliasing artifacts), then subtract [55, 56]



## DL for IR: k-space / sinogram domain learning



Figure courtesy of Jong Chul Ye, KAIST University.

- + simple and fast ("nonlinear GRAPPA")
- perhaps harder to represent local image features?

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#### DL for IR: transform learning





Figure courtesy of Jong Chul Ye, KAIST University.

- + in principle, purely data driven; potential to avoid model mismatch
- high memory requirement for fully connected layers

#### DL for IR: hybrid domain learning





Figure courtesy of Jong Chul Ye, KAIST University.

- + physics-based use of k-space data & image-domain priors
- + interpretable connections to optimization approaches
- more computation to due to "iterations" (layers) and repeated Ax, A'r

#### Convolutional sparsity revisted



Cost function for convolutional sparsity regularization:

$$\arg\min_{\boldsymbol{x}} \frac{1}{2} \|\boldsymbol{A}\boldsymbol{x} - \boldsymbol{y}\|_{\boldsymbol{W}}^{2} + \beta \left(\min_{\boldsymbol{\zeta}} \sum_{k=1}^{K} \frac{1}{2} \|\boldsymbol{h}_{k} \ast \boldsymbol{x} - \boldsymbol{\zeta}_{k}\|_{2}^{2} + \alpha \|\boldsymbol{\zeta}_{k}\|_{1}\right)$$

Alternating minimization, aka block coordinate descent (BCD), updates:

Sparse code: 
$$\boldsymbol{\zeta}_{k}^{(n+1)} = \operatorname{soft} \{ \boldsymbol{h}_{k} * \boldsymbol{x}^{(n)}, \alpha \}$$

Image: 
$$\mathbf{x}^{(n+1)} = \arg\min_{\mathbf{x}} F(\mathbf{x}; \mathbf{y}, \mathbf{z}^{(n)})$$
  
 $F(\mathbf{x}; \mathbf{y}, \mathbf{z}^{(n)}) \triangleq \frac{1}{2} \|\mathbf{A}\mathbf{x} - \mathbf{y}\|_{\mathbf{W}}^{2} + \beta \left(\sum_{k=1}^{K} \frac{1}{2} \|\mathbf{h}_{k} * \mathbf{x} - \boldsymbol{\zeta}_{k}^{(n+1)}\|_{2}^{2} + \alpha \|\boldsymbol{\zeta}_{k}^{(n+1)}\|_{1}\right)$   
 $= \frac{1}{2} \|\mathbf{A}\mathbf{x} - \mathbf{y}\|_{\mathbf{W}}^{2} + \beta \frac{1}{2} \|\mathbf{x} - \mathbf{z}^{(n)}\|_{2}^{2} \quad (\text{quadratic but } \text{large} \Longrightarrow \text{majorize})$   
 $\mathbf{z}^{(n)} = \mathcal{R}(\mathbf{z}^{(n)}) = \sum_{k=1}^{K} \text{flip}(\mathbf{h}_{k}) * \text{soft}\{\mathbf{h}_{k} * \mathbf{x}^{(n)}\} \quad (\text{denoise} \Longrightarrow \text{learn})$ 





Unrolled loop network with momentum and guadratic majorizer [57, 58]:

Diagonal majorizer for CT:  $M = Diag\{A'WA1\} + \beta I \succeq A'WA + \beta I$ 

**Learn** image mapper ("refiner")  $\mathcal{R}$  from training data (supervised). cf CNN: filter  $\rightarrow$  threshold  $\rightarrow$  filter



- Image mapper  $\mathcal{R}$  is shallow
  - $\implies$  less risk of over-fitting / hallucination
- ▶ Momentum accelerates convergence ⇒ fewer "layers" (outer iterations)
- First unrolled loop approach to have convergence theory (under suitable assumptions on *R*)
- Image update uses original measurements y and imaging physics A

[57, 58] II Yong Chun, Zhengyu Huang, Hongki Lim, J A Fessler Momentum-Net: Fast and convergent iterative neural network for inverse problems http://arxiv.org/abs/1907.11818,

IEEE Tr. on PAMI, 2020 http://doi.org/10.1109/TPAMI.2020.3012955

#### Momentum-Net preliminary results

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Illustration of benefits of momentum:



### Momentum-Net preliminary image results





Sparse-view CT with 123/984 views,  $I_0 = 10^5$ , 800-1200 mod. HU display.





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- CT image reconstruction has evolved greatly in the 50+ years since Allan Cormack's seminal papers [59, 60]
  - physics
  - statistics
  - regularization and optimization
  - data adaptive methods inspired by machine learning
- 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
- More comparisons between model-based methods with adaptive regularizers and CNN-based methods needed

#### Resources



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



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