# Computational medical imaging: from model-based approaches to machine learning

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#### Computational medical imaging

#### Model-based approaches

Introduction Compressed acoustic microscopy Results Other (successful) examples

#### Data-driven approaches

Introduction Limitation of model-based methods U-net

#### Conclusions

### What is medical imaging?

 Visualization of body parts, tissues or organs, for use in clinical diagonsis, treatment and disease monitoring



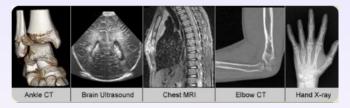
Anatomical vs Functional



### Medical imaging modalities

Nuclear medicine (SPECT, PET)

Radiology techniques (X-ray radiography, CT, MRI, Ultrasound)

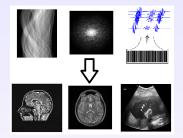


Scanners

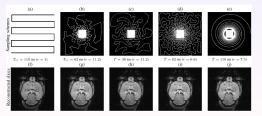


### **Computational medical imaging**

Data inversion



Incomplete data, non-traditional sensing, etc.

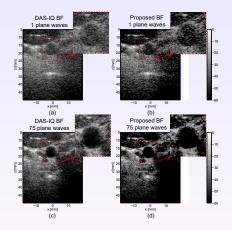


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#### Incomplete data in ultrasound

### Computational methods to compensate for the lack of data



<sup>.</sup> Winner of plane-wave imaging challenge in medical ultrasound, IEEE IUS, 2016.

<sup>.</sup> T. Szasz, A. Basarab, D. Kouamé, Beamforming through regularized inverse problems in ultrasound medical imaging, IEEE TUFFC, 2016.

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#### **Inverse problems**

$$\mathbf{y} = T(\mathbf{x}) + \mathbf{n}$$

- **b**  $\mathbf{y} \in \mathbb{C}^{M}$  is the observed data (image)
- ▶  $\mathbf{x} \in \mathbb{C}^{N}$  is the image of interest (not observed)
- ▶  $\mathbf{n} \in \mathbb{C}^M$  is the noise

### T is the observation (forward) operator

- known : estimate x from y
- unknown : estimate x and T from y

### Inverse problems in computational medical imaging are usually ill-posed

- T is not invertible, an infinity of solutions may exist
- A small perturbation on the data may cause an important variation on the estimate (e.g. Fourier measurements)

Solution

$$\hat{\mathbf{x}} = \arg\min_{\mathbf{x}} \|\mathbf{y} - T(\mathbf{x})\|_2^2 + \lambda r(\mathbf{x})$$

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#### **Basics on acoustic microscopy**

### Single-element transducer

- Very high frequency (> 50 MHz) : 250 MHz or 500 MHz
- Spherically-focused (F-number < 1.3)</p>

#### Raster scan

- RF data, reflected from each scanned position, yield 3D data volume
- Acoustic parameters estimated at each position form 2D maps

### Challenges

- Scanning time of 5 minutes for 1 mm by 1 mm sample
- Tissue properties may change during scanning
- Decrease the acquisition time and data amount, reduce equipment costs



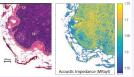


#### **Data acquisition**

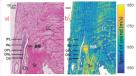
#### Raster scan

- For each (x<sub>1</sub>, x<sub>2</sub>) position, RF data composed of two main reflections is digitized and saved, yielding to a 3D data volume (x<sub>1</sub>, x<sub>2</sub>, t)
- Speed of sound, acoustic impedance and attenuation are estimated at each scan location (x<sub>1</sub>, x<sub>2</sub>) to form quantitative 2D parameter maps

Example of impedance map



### Example of speed of sound



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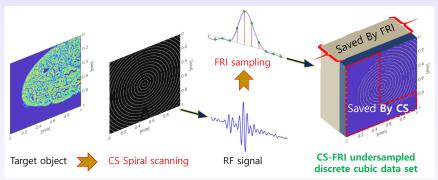
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Spiral pattern ( $\Phi$ ) scanning in **spatial domain**,  $y = \Phi x + n$ 

► Innovation rate ( $\rho = 1/T$ ) sampling in time domain,  $x_n = \left\langle \sum_{l=0}^{L-1} a_l h(t - t_l), \varphi\left(\frac{t}{T} - n\right) \right\rangle$ 

<sup>.</sup> J. Kim, J. Mamou, D. Kouamé, A. Achim, and A. Basarab, Spatio-temporal compressed quantitative acoustic microscopy, IEEE IUS, Oct 2019.

<sup>.</sup> J. Kim, J. Mamou, P. R. Hill, N. Canagarajah, D. Kouamé, A. Basarab, and A. Achim, Approximate message passing reconstruction of quantitative acoustic microscopy images, *IEEE Trans. on UFFC*, Mar 2018.

<sup>.</sup> J. Kim, J. Mamou, D. Kouamé, A. Achim, and A. Basarab, Reconstruction of quantitative acoustic microscopy images from rf signals sampled at innovation rate, *IEEE IUS*, Oct 2018.

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### Compressed sensing (CS)

- Sampling signals parsimoniously, acquiring only the relevant signal information, rather than sampling followed by compression
- Direct model

$$\boldsymbol{y} = \Phi \boldsymbol{x} + \boldsymbol{n}$$

- $\mathbf{y} \in \mathbb{R}^{M}$ : measurement vector
- $\Phi \in \mathbb{R}^{M \times N}$  : measurement matrix
- ▶  $\mathbf{x} \in \mathbb{R}^N$  : image to be reconstructed, M << N
- ▶  $\boldsymbol{n} \in \mathbb{R}^{M}$  : zero-mean additive white Gaussian noise

Reconstruction

$$\hat{\theta}_x = \min_{\theta_x} \|\theta_x\|_1$$
 subject to  $y = \Phi W^{-1} \theta_x$ 

• W is a sparsifying transform,  $\theta_x = W x$ 

<sup>.</sup> D.L. Donoho, Compressed sensing, IEEE Trans. on Information Theory, 2006.

<sup>.</sup> E. Candès, J. Romberg, T. Tao, Robust Uncertainty Principles : Exact Signal Reconstruction from Highly Incomplete Fourier Information, IEEE Trans. on Information Theory, 2006.

#### Sensing pattern

- Experimental constraints
  - Point-wise acquisition
  - RF data acquired continuously as the motor stages are moved
- Proposed solution
  - Spiral sensing originating in the center of the area to be sampled
  - The pace of the spreading is used to prescribe the measurement rate
- Example of spiral pattern for a measurement rate of 20%



<sup>.</sup> J-H. Kim, J. Mamou, P.R. Hill, N.Canagarajah, D. Kouamé, A. Basarab, A. Achim, Approximate Message Passing Reconstruction of Quantitative Acoustic Microscopy Images, *IEEE TUFFC*, 2018.

### **AMP** algorithm

- Iterative signal reconstruction algorithm
  - Turns the reconstruction problem into an iterative denoising approach

$$\boldsymbol{\theta}_{x}^{t+1} = \eta_{t} \Big( \Theta^{T} \boldsymbol{z}^{t} + \boldsymbol{\theta}_{x}^{t} \Big)$$
$$\boldsymbol{z}^{t} = \boldsymbol{y} - \Theta \boldsymbol{\theta}_{x}^{t} + \frac{1}{\delta} \boldsymbol{z}^{t-1} \big\langle \eta_{t-1}^{'} (\Theta^{T} \boldsymbol{z}^{t-1} + \boldsymbol{\theta}_{x}^{t-1} \big\rangle$$

- t : iterative index
- $\bullet \Theta = \Phi W^{-1}$
- $\eta_t(\cdot), \eta'_t(\cdot)$ : denoising function and its first derivative
- $\delta$  : measurement rate M/N
- ►  $z^t \in \mathbb{R}^M$  : current residual (error)

<sup>.</sup> D. L. Donoho, A. Maleki, A. Montanari, Message passing algorithms for compressed sensing, Proc. Nat. Academy Sci., 2009.

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### Wavelet-based Cauchy AMP

### Assumptions

- Wavelet coefficients provide a sparse representation for natural images
- They can be accurately modelled using heavy tailed distributions
- Design the denoising function  $\eta$ 
  - Based on the hypothesis of Cauchy distributed wavelet coefficients of QAM images

$$P(w) = rac{\gamma}{w^2 + \gamma^2}$$

- w is the wavelet coefficient
- $\gamma$  is the dispersion parameter

<sup>.</sup> A. Achim and E. E. Kuruoglu, Image denoising using bivariate α-stable distributions in the complex wavelet domain, *IEEE Signal Processing Letters*, 2005.

#### **Cauchy-AMP** algorithm

### MAP estimator

 Observed coefficient v contaminated with additive Gaussian noise (n = v - w)

$$\hat{w} = \arg\max_{w} Pw|v(w|v)$$

Using Bayes' theorem and taking the logarithm form

$$\hat{w}(v) = \arg \max_{w} \left[ -\frac{(v-w)^2}{2\sigma_n^2} + \log\left(\frac{\gamma}{w^2+\gamma^2}\right) \right]$$

Cancel the first derivative w.r.t. w

$$\hat{w}^3 - v\hat{w}^2 + (\gamma^2 + 2\sigma_n^2)\hat{w} - \gamma^2 v = 0$$

Esimate of clean wavelet coefficient w

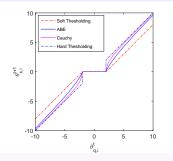
$$\hat{w} = \eta(v) = rac{v}{3} + s + t$$
  
 $\hat{w}^{'} = \eta^{'}(v) = 1/3 + s^{'} + t^{'}$ 

s and t depend on v and σ<sup>2</sup><sub>n</sub>

### Cauchy-AMP algorithm

The comparison of behaviour among four different denoisers

- Cauchy-based
- Soft thresholding (ST)
- Amplitude scale invariant Bayes estimator (ABE)



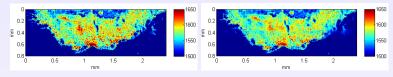
<sup>.</sup> D.L. Donoho, De-noising by soft-thresholding, IEEE Trans. on Information Theory, 1995.

<sup>.</sup> M. A. Figueiredo and R. D. Nowak, Wavelet-based image estimation : an empirical Bayes approach using Jeffreys' noninformative prior, IEEE Trans. on Image Processing, 2001.

- Compressed acoustic microscopy

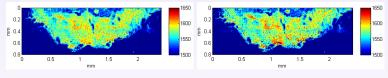
#### **Results at 30% measurement ratio**

### Human lymph node



(a) Original

(b) AMP-Cauchy



(c) AMP-ST

(d) AMP-ABE

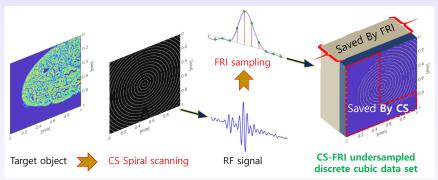
### Quantitative results

Method	SSIM	PSNR
AMP-Cauchy	0.714	40.42
AMP-ST	0.683	39.98
AMP-ABE	0.708	40.10

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<sup>.</sup> J. Kim, J. Mamou, D. Kouamé, A. Achim, and A. Basarab, Reconstruction of quantitative acoustic microscopy images from rf signals sampled at innovation rate, *IEEE IUS*, Oct 2018.

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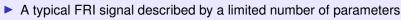
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### Finite rate of innovation (FRI)



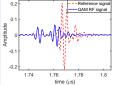
$$x(t) = \sum_{l=0}^{L-1} a_l h(t-t_l)$$

Unknown :

- $t_l$  : time delays
- a<sub>l</sub> : amplitude decays

Known :

h(t) : pulse shape



FRI sampling by kernel  $\varphi(t)$  and innovation rate ( $\rho = 1/T$ )

$$x_n = \left\langle \sum_{l=0}^{L-1} a_l h(t-t_l), \varphi(t/T-n) \right\rangle$$

- $\rho$  : total number of the unknown parameters  $(t_l, a_l)$
- FRI recovery techniques for estimating t<sub>l</sub> and a<sub>l</sub>
  - Annihilating filter, matrix pencil method, etc.

<sup>.</sup> M. Vetterli, P. Marziliano, and T. Blu, Sampling signals with finite rate of innovation, IEEE Trans. on Signal Proc., Jun 2002.

### Acoustic parameter estimation by AR estimator

• Retrieve Fourier coefficients X[k] from x[n] and  $\Phi(\omega)$ 

$$x[n] = \langle x(t), \varphi(t - nT) \rangle = \sum_{k \in Z} X[k] e^{\frac{j2\pi knT}{\tau}} \Phi^* \Big[ \frac{2\pi k}{\tau} \Big]$$

•  $\Phi^*(\omega)$  : transpose of Fourier transform of  $\varphi(t)$ 

• The power series form acquired from N[k] = X[k]/H[k]

$$N[k] = \sum_{l=0}^{L-1} a_l \{ \exp[\Delta f(\beta_l + j2\pi t_l)] \}^k = \sum_{l=0}^{L-1} a_l \lambda_l^k$$

AR model construction

$$N_k = \sum_{i=1}^n s_i N_{k-i} + \epsilon_k$$

Calculation of acoustic parameters from identified λ<sub>l</sub> and a<sub>l</sub>

<sup>.</sup> J-H. Kim, J. Mamou, D. Kouamé, A. Achim, A. Basarab, Autoregressive model-based reconstruction of quantitative acoustic maps from RF signals sampled at innovation rate, submitted to *IEEE TCI*, 2019.

<sup>.</sup> D. Rohrbach and J. Mamou, Autoregressive signal processing applied to high-frequency acoustic microscopy of soft tissues, IEEE Trans. on UFFC, 2018.

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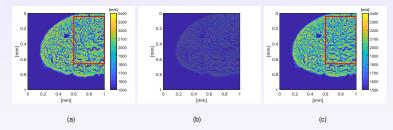
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#### Illustration of the images at each reconstruction stage

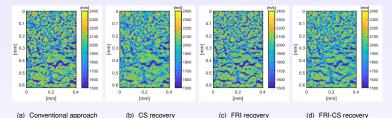
Speed of sound (SOS) maps of chicken tendon acquired with a 250 MHz transducer



- (a) : Conventional sampling at  $1\mu m$  by  $1\mu m$
- (b) : The spiral masking area recovered by FRI framework
- (c) : FRI-CS recovery

### Comparison of reconstructed images

### Chicken tendon 2D SOS maps



### Quantitative results

Methods	PSNR(dB)	NRMSE	SSIM	compressed ratio (%)	Acquisition times
CS (b)	26.31	0.0483	0.7166	40	5%
FRI (c)	24.19	0.0617	0.6741	6.5	100%
FRI-CS (d)	23.38	0.0678	0.5511	2.6	5%

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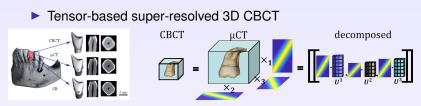
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#### **Super-resolution**



MR-ultrasound image fusion



Ultrasound







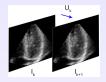
<sup>.</sup> J. Hatvani, A. Basarab, J.-Y. Tourneret, M. Gyongy, D. Kouamé, A Tensor Factorization Method for 3D Super-Resolution with Application to Dental CT, IEEE TMI, 2019.

<sup>.</sup> O. El Mansouri, A. Basarab, F. Vidal, D. Kouamé, J.-Y. Tourneret, Fusion of Magnetic Resonance and Ultrasound Images : a Preliminar Study on Simulated Data, IEEE ISBI, 2019.

#### **Cardiac motion estimation**

Let  $I_k \in \mathbb{R}^N$ ,  $I_{k+1} \in \mathbb{R}^N$  be two consecutive images

► The motion field to be estimated is  $\boldsymbol{U}_{\boldsymbol{k}} = (\boldsymbol{u}_{\boldsymbol{k}}^{T}, \boldsymbol{v}_{\boldsymbol{k}}^{T})^{T} \in \mathbb{R}^{2N}$ 



Minimization of an energy function

 $\min_{\boldsymbol{\alpha},\boldsymbol{U}} \{ E_{\text{data}}(\boldsymbol{U},\boldsymbol{I}) + \lambda_s E_{\text{spatial}}(\boldsymbol{U}) + \lambda_d E_{\text{sparse}}(\boldsymbol{U},\boldsymbol{D},\boldsymbol{\alpha}) \}$ 

<sup>.</sup> N. Ouzir, A. Basarab, O. Lairez, and J.-Y. Tourneret, Robust Optical Flow Estimation in Cardiac Ultrasound images Using a Sparse Representation, *IEEE TMI*, 2018.

<sup>.</sup> N. Ouzir, A. Basarab, H. Liebgott, B. Harbaoui, and J.-Y. Tourneret, Motion estimation in echocardiography using sparse representation and dictionary learning, IEEE TIP, 2018.

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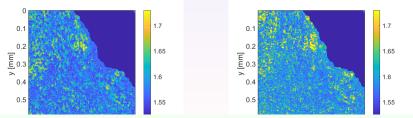
### Spatial resolution in acoustic microscopy

### Dependent on the central frequency

- Increasing the frequency comes with
  - Increased costs associated with the transducer and the necessary electronics
  - Experimental difficulties (e.g., sensitivity to nm scale vibrations and temperature)

### Example of impedance images on a section of cancerous human lymph node

Thin section of soft tissue (12 µm) affixed to a microscopy slide



### 250 MHz

500 MHz

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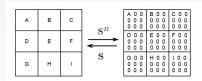
#### **Forward model**

 $\mathbf{y} = \mathbf{S}\mathbf{H}\mathbf{x} + \mathbf{n}$ 

- ▶  $\mathbf{y} \in \mathbb{R}^{N_l \times 1}$  : is the measured image,  $N_l = m_l \times n_l$
- ▶  $\mathbf{x} \in \mathbb{R}^{N_h \times 1}$  : super-resolved image to be estimated,  $N_h = d^2 N_l$
- ▶  $\mathbf{n} \in \mathbb{R}^{N_l \times 1}$ : Gaussian noise

### Degradation operators

- ▶  $\mathbf{H} \in \mathbb{R}^{N_h \times N_h}$ : 2D circulant convolution matrix (PSF of the transducer)
- ▶  $\mathbf{S} \in \mathbb{R}^{N_l \times N_h}$  : subsampling operator



### SR optimization problem

 Super-resolved slice x<sub>t</sub> estimation (time subscript is omitted in the following)

$$\min_{\mathbf{x}} \frac{1}{2} \|\mathbf{y} - \mathbf{SHx}\|_2^2 + \tau \phi(\mathbf{Ax})$$

Total variation regularization

$$\phi(\mathsf{A}\mathbf{x}) = \sqrt{\|\mathsf{D}_{\mathrm{h}}\mathbf{x}\|^2 + \|\mathsf{D}_{\mathrm{v}}\mathbf{x}\|^2}$$

$$\mathbf{A} = [\mathbf{D}_{\mathrm{h}}, \mathbf{D}_{\mathrm{v}}]^{\mathcal{T}} \in \mathbb{R}^{2N_{\mathrm{h}} imes N_{\mathrm{h}}}$$

Constrained optimization

$$\min_{\mathbf{x},\mathbf{u}} \quad \frac{1}{2} \|\mathbf{y} - \mathbf{SHx}\|_2^2 + \tau \phi(\mathbf{u})$$
 subject to  $\mathbf{Ax} = \mathbf{u}$ 

Associated augmented Lagrangian function

$$\mathcal{L}(\mathbf{x}, \mathbf{u}, \boldsymbol{\lambda}) = rac{1}{2} \|\mathbf{y} - \mathbf{SHx}\|_2^2 + au \phi(\mathbf{u}) + rac{\mu}{2} \|\mathbf{Ax} - \mathbf{u} + \boldsymbol{\lambda}\|_2^2$$

#### **ADMM-based algorithm**

Iterate  

$$\begin{aligned} \mathbf{x}^{k+1} &= \arg\min_{\mathbf{x}} \|\mathbf{y} - \mathbf{SH}\mathbf{x}\|_{2}^{2} + \mu \|\mathbf{A}\mathbf{x} - \mathbf{u}^{k} + \mathbf{d}^{k}\|_{2}^{2} \\ \mathbf{u}^{k+1} &= \arg\min_{\mathbf{u}} \tau \phi(\mathbf{u}) + \frac{\mu}{2} \|\mathbf{A}\mathbf{x}^{k+1} - \mathbf{u} + \mathbf{d}^{k}\|_{2}^{2} \\ \mathbf{d}^{k+1} &= \mathbf{d}^{k} + (\mathbf{A}\mathbf{x}^{k+1} - \mathbf{u}^{k+1}) \\ \text{until stopping criterion is satisfied.} \end{aligned}$$

Update **u** using the vector-soft-thresholding operator

$$\nu = [\mathbf{D}_{h}\mathbf{x}^{k+1} + \mathbf{d}_{h}^{k}, \mathbf{D}_{v}\mathbf{x}^{k+1} + \mathbf{d}_{v}^{k}]$$
$$\mathbf{u}^{k+1}[i] = \max\{\mathbf{0}, \|\nu[i]\|_{2} - \tau/\mu\} \frac{\nu[i]}{\|\nu[i]\|_{2}}$$

Update x using a closed-form solution exploiting the following property of S

$$\mathsf{FS}^{H}\mathsf{SF}^{H}=rac{1}{d}\mathsf{J}_{d}\cdot\mathsf{I}_{N_{l}}$$

<sup>.</sup> N. Zhao, Q. Wei, A. Basarab, N. Dobigeon, D. Kouamé, J.-Y. Tourneret, Fast Single Image Super-resolution using a New Analytical Solution for  $\ell_2$ - $\ell_2$  Problems, *IEEE TIP*, 2016.

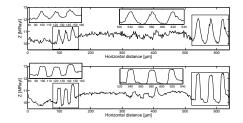
### **Resolution target results**

#### USAF 1951 resolution phantom

Super-resolution factor d = 2



### Horizontal profiles



#### Model-based super-resolution

#### Fails on ex vivo samples

Convolution with the PSF not sufficient to model the 250 MHz images

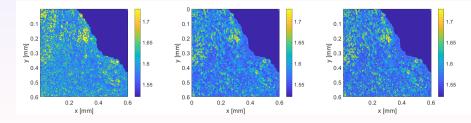
Example of impedance images on a section of cancerous human lymph node

Thin section of soft tissue (12 µm) affixed to a microscopy slide

500 MHz

250 MHz

## Enhanced 250 MHz



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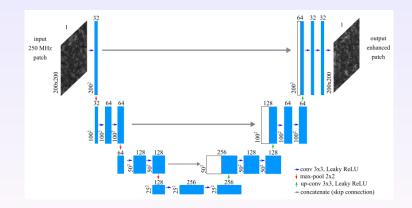
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#### **Data-driven super-resolution**

#### Fully convolution neural network (U-net) trained on 250 and 500 MHz images



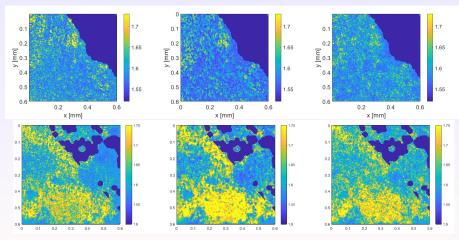
<sup>.</sup> J. Mamou, T. Pellegrini, D. Kouamé, A. Basarab, A convolutional neural network for 250-MHz quantitative acoustic-microscopy resolution enhancement, IEEE EMBC, 2019.

#### **Results**

500 MHz

250 MHz

### Enhanced 250 MHz



### Conclusions

### Computational imaging

- In most of medical applications data is not sufficient to form the image (noise, incomplete data)
- Computational methods are used to avoid the ill-posedness of the resulting inverse problem

#### Model-based approaches

- Models include knowledge about the physics : fidelity, tractability ?
- Regularization terms are required and usually use adaptive or non-adaptive transforms : appropriate choice ?

#### Data-driven approaches

- More flexibility, but usually require learning databases
- How to include knowledge about the physics?
- Forward model, regularization, both?

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