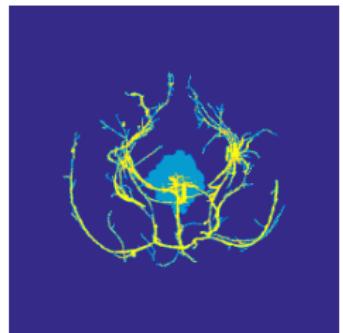
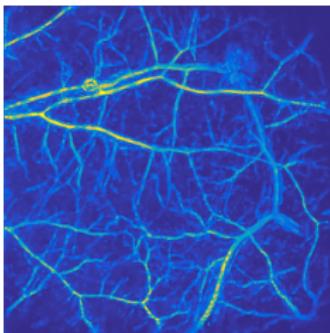
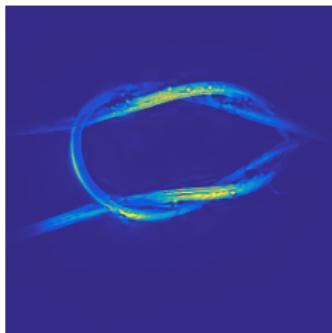


Accelerated High-Resolution Photoacoustic Tomography via Compressed Sensing



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joint with:

Simon Arridge, Paul Beard, Marta Betcke,
Ben Cox, Nam Huynh & Edward Zhang



CMIC

Centre for Medical Image Computing

SIAM-IS, Albuquerque,
May 24, 2016.



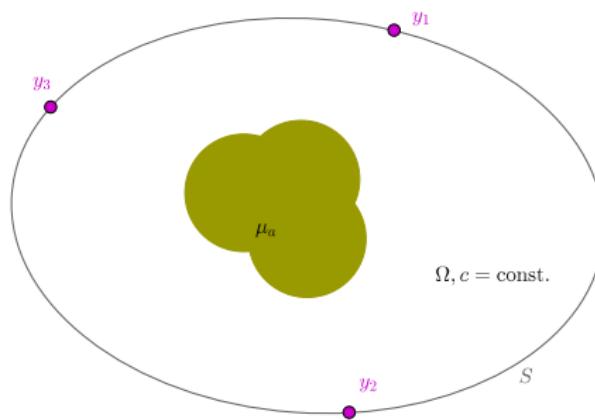
UCL CENTRE FOR
**INVERSE
PROBLEMS**

Optical Part

chromophore concentration: c_k

optical absorption coefficient: $\mu_a(c)$

Acoustic Part



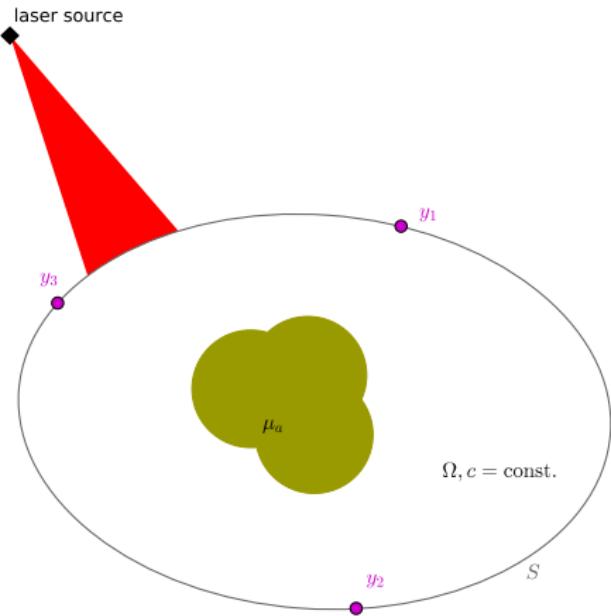
Optical Part

chromophore concentration: c_k

optical absorption coefficient: $\mu_a(c)$

pulsed laser excitation: $\Phi(\mu_a)$

Acoustic Part



Optical Part

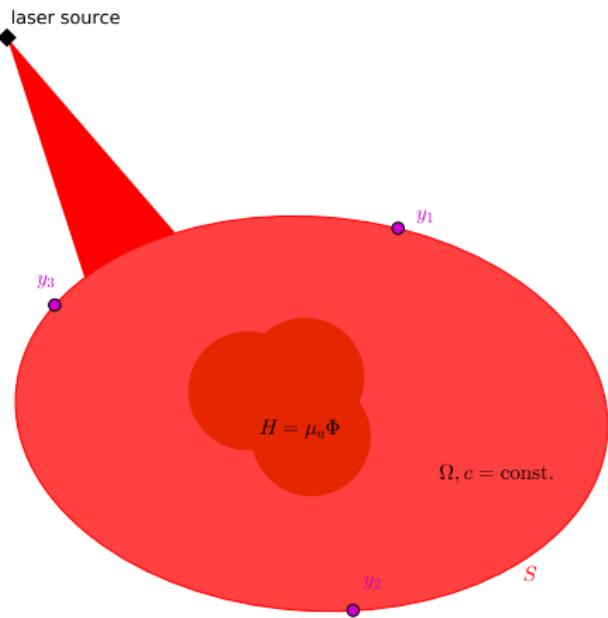
chromophore concentration: c_k

optical absorption coefficient: $\mu_a(c)$

pulsed laser excitation: $\Phi(\mu_a)$

thermalization: $H = \mu_a \Phi(\mu_a)$

Acoustic Part



Optical Part

chromophore concentration: c_k

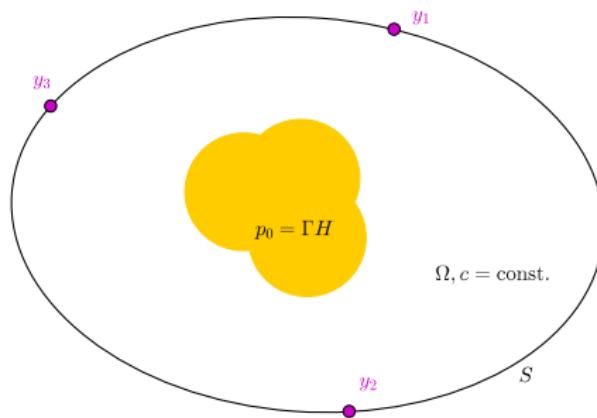
optical absorption coefficient: $\mu_a(c)$

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Acoustic Part

local pressure increase: $p_0 = \Gamma H$



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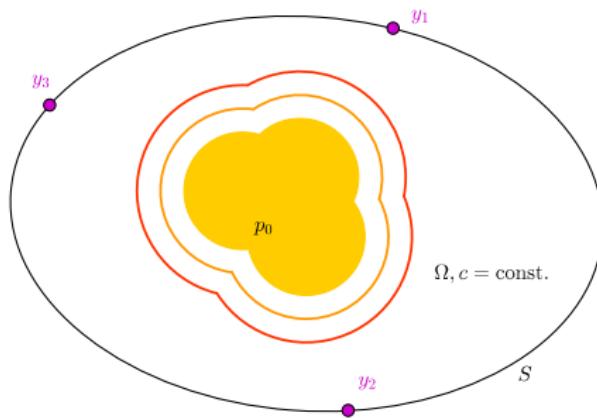
Acoustic Part

local pressure increase: $p_0 = \Gamma H$

elastic wave propagation:

$$\Delta p - \frac{1}{c^2} \frac{\partial^2 p}{\partial^2 t} = 0$$

$$p|_{t=0} = p_0, \quad \frac{\partial p}{\partial t}|_{t=0} = 0$$



Optical Part

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Acoustic Part

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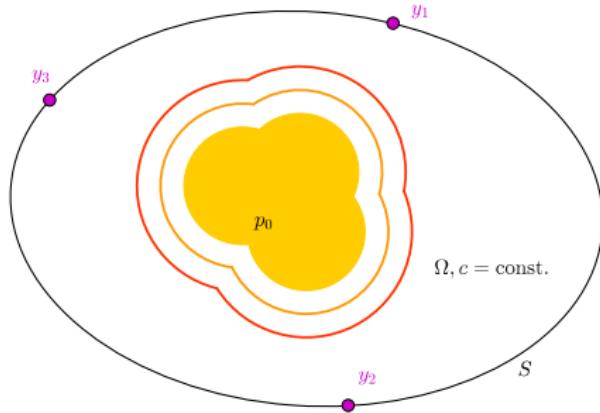
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measurement of pressure time courses:

$$f_i(t) = p(y_i, t)$$



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$$\Delta p - \frac{1}{c^2} \frac{\partial^2 p}{\partial^2 t} = 0$$

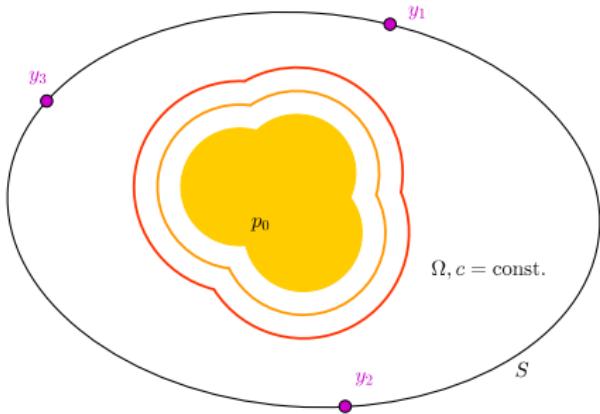
$$p|_{t=0} = p_0, \quad \frac{\partial p}{\partial t}|_{t=0} = 0$$

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Photoacoustic effect

- ▶ coupling of optical and acoustic modalities.
- ▶ "hybrid imaging"
- ▶ high optical contrast can be read by high-resolution ultrasound.



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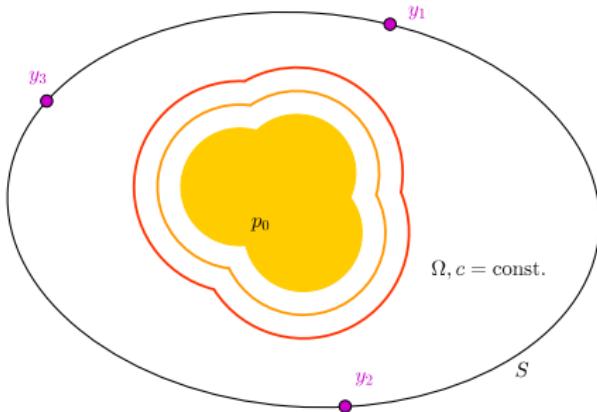
$$p|_{t=0} = p_0, \quad \frac{\partial p}{\partial t}|_{t=0} = 0$$

measurement of pressure time courses:

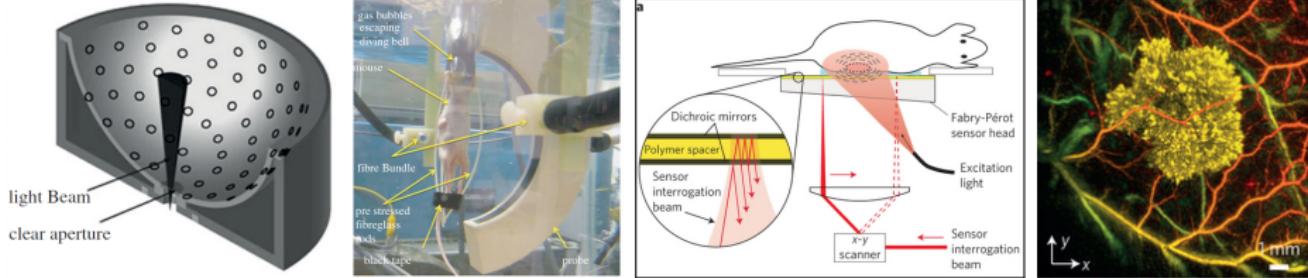
$$f_i(t) = p(y_i, t)$$

Photoacoustic effect

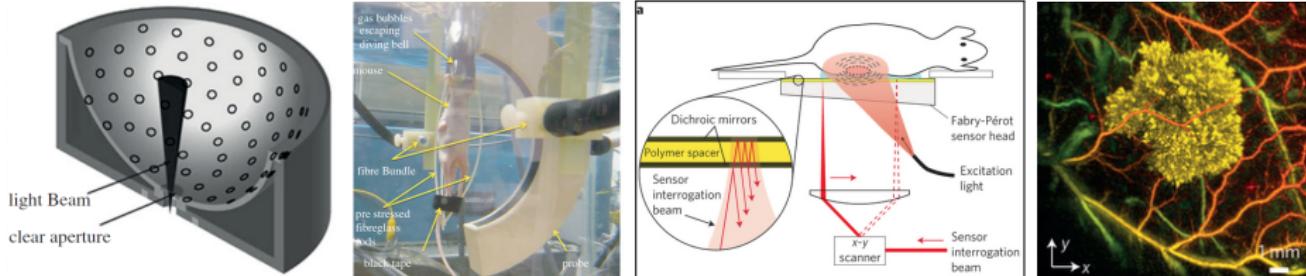
- ▶ coupling of optical and acoustic modalities.
- ▶ "hybrid imaging"
- ▶ high optical contrast can be read by high-resolution ultrasound.



Photoacoustic Sensing Systems

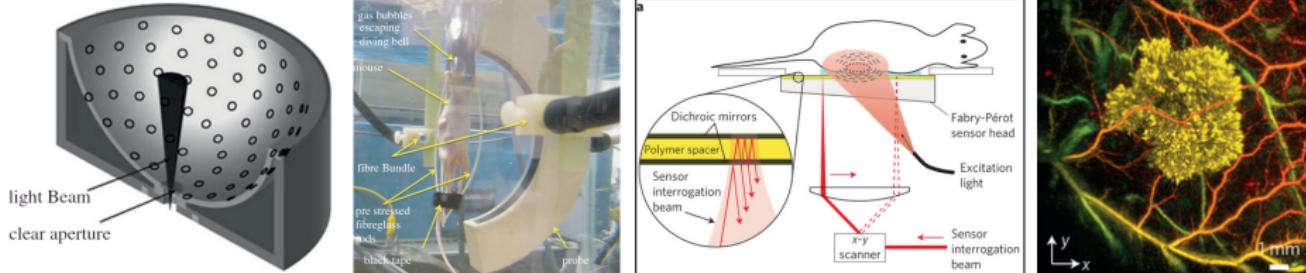


from: Beard, 2011, *Interface Focus*; Jathoul et al., 2015, *Nature Photonics*



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- ▶ High res 3D PA images require sampling acoustic waves with a frequency content in the **tens of MHz** over **cm scale** apertures.
- ▶ Nyquist criterion results in **tens of μm** scale sampling intervals
⇒ **several thousand detection points**.
- ▶ Sequential scanning currently takes **several minutes**.
- ▶ Crucial limitation for clinical, spectral and dynamical PAT (**4D PAT**).



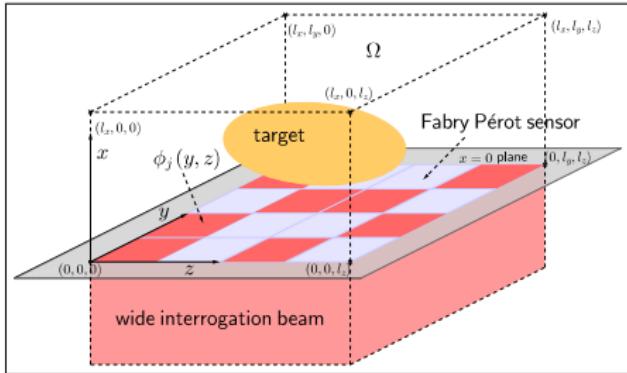
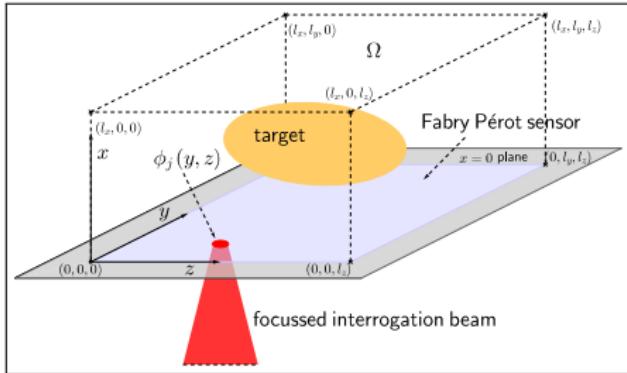
from: Beard, 2011, *Interface Focus*; Jathoul et al., 2015, *Nature Photonics*

Key observation and idea:

- ▶ Nyquist is too conservative (only band-limitlessness is assumed).
- ▶ Typical targets have additional structure, e.g., low spatial complexity (**sparsity**).
- ▶ Regularly sampled data is **highly redundant**.
- ▶ Non-redundant part could be sensed faster.
- ▶ Accelerated acquisition **without significant loss of image quality**.

Established as **compressed sensing**, successful in similar modalities.

Novel Fabry-Pérot-Based Sensing Systems



$$f_j(t) = \int p(x = 0, y, z, t) \phi_j(y, z) \, dy \, dz$$

- ▶ Single-point sub-sampling (structured or random).
- ▶ Patterned interrogation similar to "single-pixel" Rice camera (via micromirror array).
- ▶ Multi-beam scanning + sub-sampling.

Applicable to other sequential scanning schemes, see **Huynh et al., 2014, 2015, 2016** for technical details.

Novel Fabry-Pérot-Based Sensing Systems

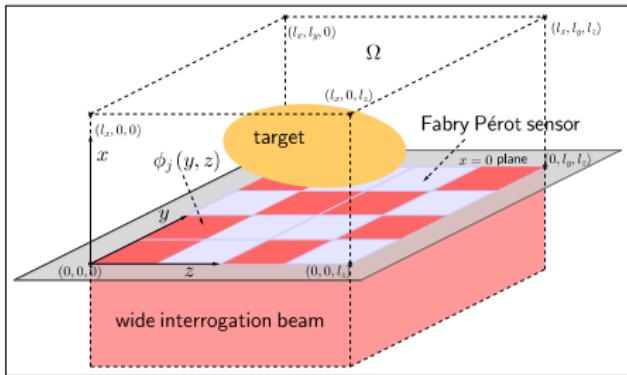
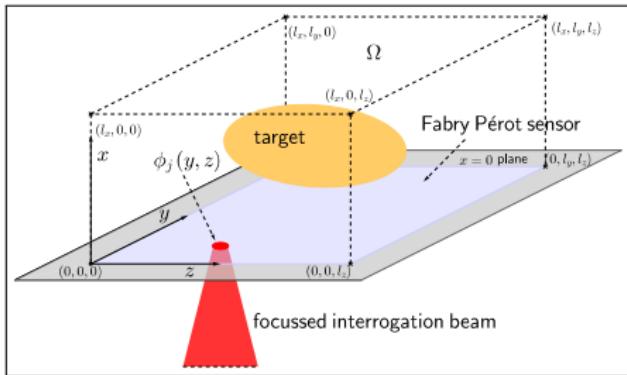


Image model: $f_i^c = C_i f_i = C_i (A p_i + \varepsilon_i)$ for each frame i .

Image reconstruction:

- ▶ $f_i^c \rightarrow f_i$, $f_i \rightarrow p_i$ by standard method, frame-by-frame.
- ▶ $f_i^c \rightarrow p_i$: standard or new method, frame-by-frame.
- ▶ $F^c \rightarrow F$, $f_i \rightarrow p_i$ by standard method, frame-by-frame.
- ▶ $F^c \rightarrow P$: Full spatio-temporal method.

Novel Fabry-Pérot-Based Sensing Systems

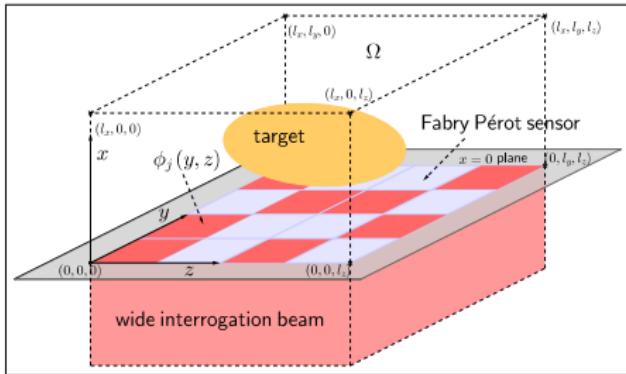
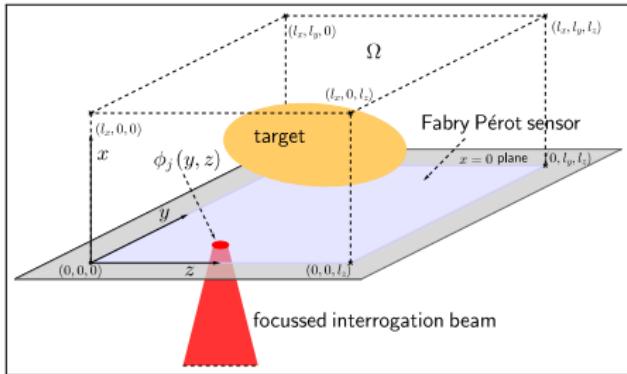


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- ▶ $F^c \rightarrow F$, $f_i \rightarrow p_i$ by standard method, frame-by-frame.
- ▶ $F^c \rightarrow P$: Full spatio-temporal method.

Analytic methods, e.g. eigenfunction expansion and closed-form filtered-backprojection, are too restrictive for us.

Time Reversal (TR):

- ▶ "Least restrictive PAT reconstruction"
- ▶ Sending the recorded waves "back" into volume.
- ▶ Requires a numerical model for acoustic wave propagation.

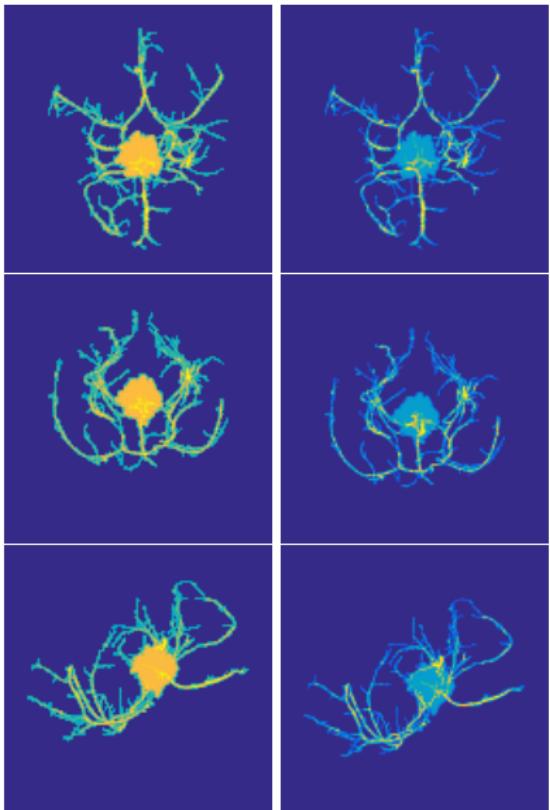
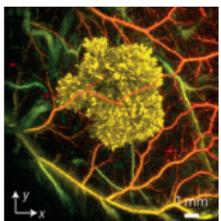
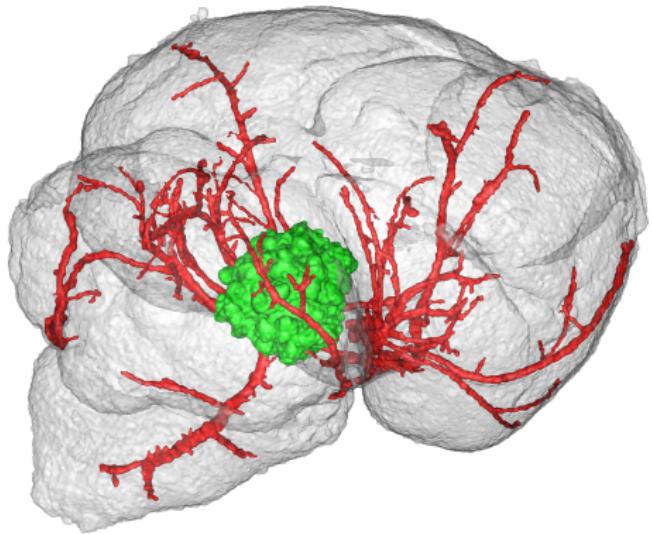
k-Wave[♣] implements a *k*-space pseudospectral method to solve the underlying system of first order conservation laws:

- ▶ Compute spatial derivatives in Fourier space: 3D FFTs.
- ▶ Modify finite temporal differences by *k*-space operator and use staggered grids for accuracy and robustness.
- ▶ Perfectly matched layer to simulate free-space propagation.
- ▶ Parallel/GPU computing leads to massive speed-ups.

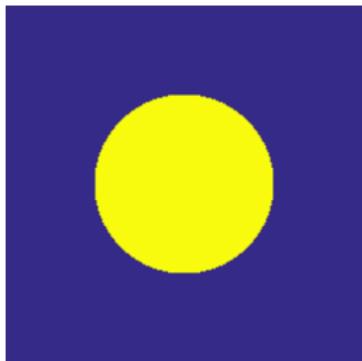


[♣]B. Treeby and B. Cox, 2010. k-Wave: MATLAB toolbox for the simulation and reconstruction of photoacoustic wave fields, *Journal of Biomedical Optics*.

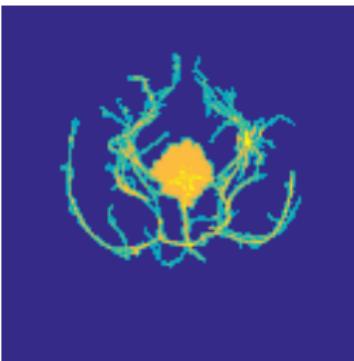
A Realistic Numerical Phantom



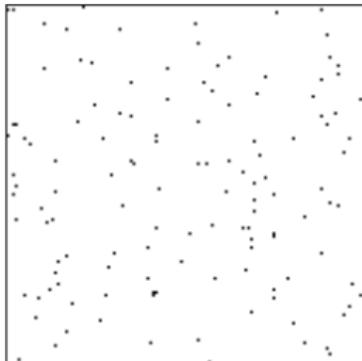
Time Reversal for Sub-Sampled Data



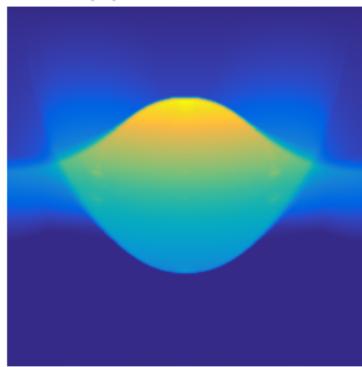
(a) IC, $n = 256^3$



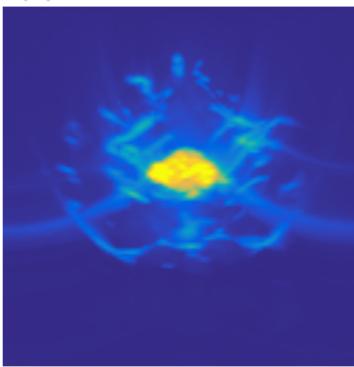
(b) high con., IC, $n = 128^3$



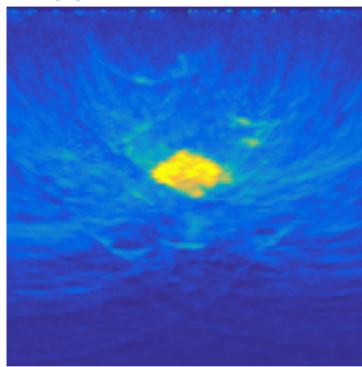
(c) sub-sampling, $128 \times$



(d) TR 1

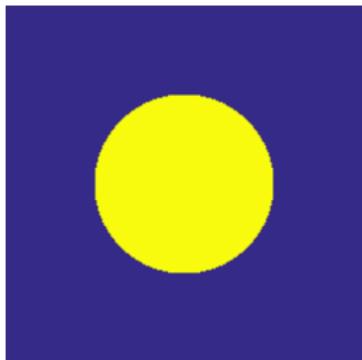
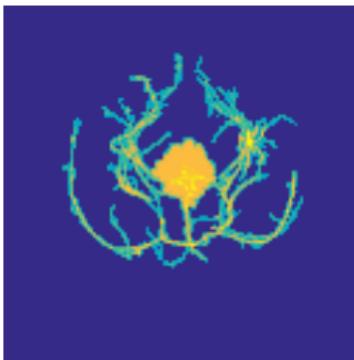
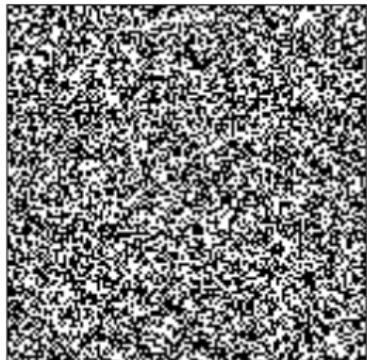


(e) TR 2

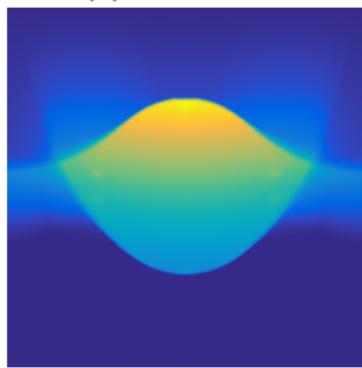


(f) TR 2, sub-sampled

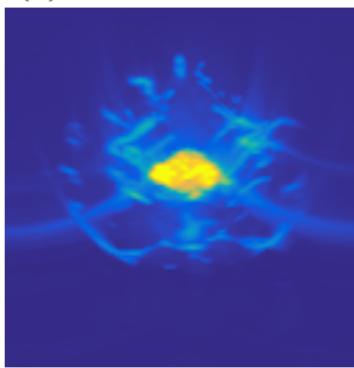
sensor on top; inverse crime data sampled at Nyquist; max intensity proj., side view

(a) IC, $n = 256^3$ (b) high con., IC, $n = 128^3$ 

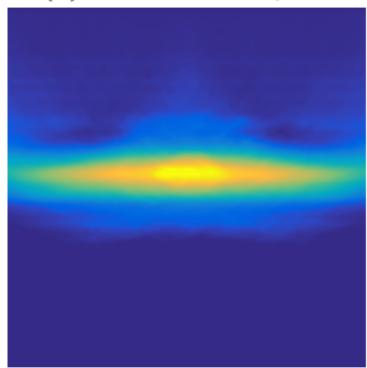
(c) sub-sampling, 1/128



(d) TR 1



(e) TR 2



(f) TR 2, sub-sampled

sensor on top; inverse crime data sampled at Nyquist; max intensity proj., side view

Solving variational regularization problems

$$\hat{p} = \operatorname{argmin}_{p \geq 0} \left\{ \frac{1}{2} \| C A p - f^c \|_2^2 + \lambda \mathcal{J}(p) \right\}$$

iteratively by first-order methods requires implementation of A and A^* .

k-Wave yields a discrete representation A_κ . For A^* , one can

- 1) adjoint k-Wave iteration to obtain $(A_\kappa)^*$ (algebraic adjoint):
 - ✓ high numerical accuracy.
 - ! tedious derivation, specific for k-Wave, limited insights.

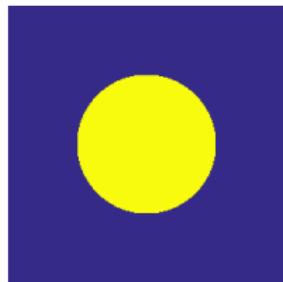
Huang, Wang, Nie, Wang, Anastasio, 2013. *IEEE Trans Med Imaging*

- 2) derive analytical adjoint and discretize it, e.g., $(A^*)_\kappa$.
 - ✓ good numerical accuracy.
 - ✓ simple proof, theoretical insights, generalizes to various numerical schemes.

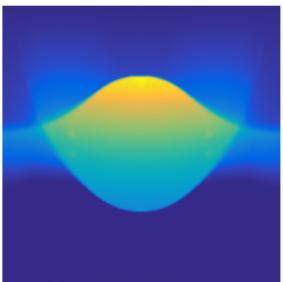
Arridge, Betcke, Cox, L, Treeby, 2015. *On the Adjoint Operator in Photoacoustic Tomography*, (*submitted, arXiv:1602.02027*).

Comparison for Conventional Data

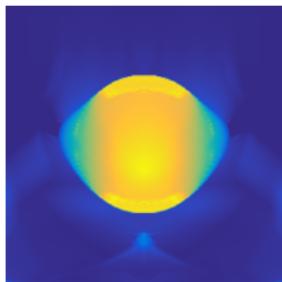
$$\hat{p} = \underset{p \geq 0}{\operatorname{argmin}} \left\{ \frac{1}{2} \|Ap - f\|_2^2 + \lambda \mathcal{J}(p) \right\}$$



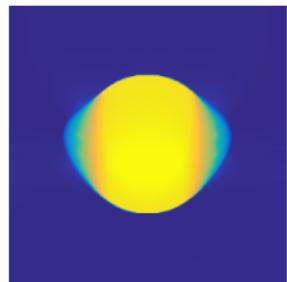
(a) $n = 256^3$



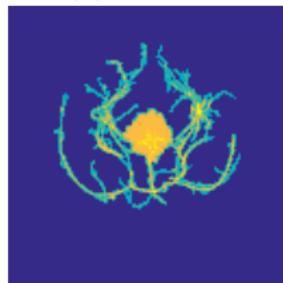
(b) TR



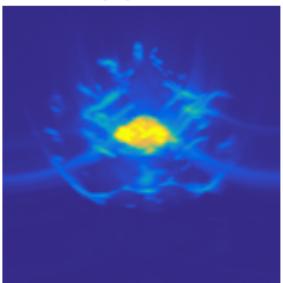
(c) LS+



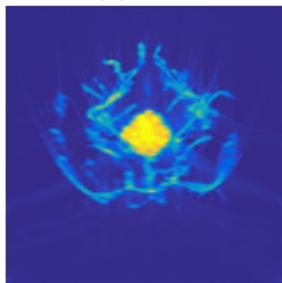
(d) TV+



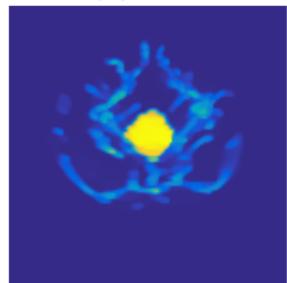
(e) $n = 128^3$



(f) TR



(g) LS+

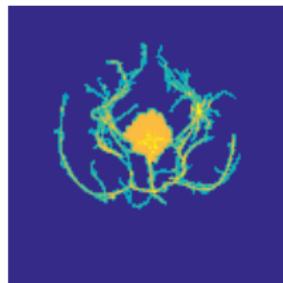


(h) TV+

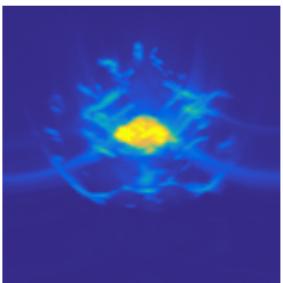
sensor on top; inverse crime data sampled at Nyquist; max intensity proj., side view

Sub Sampled Data, Best Case Scenario

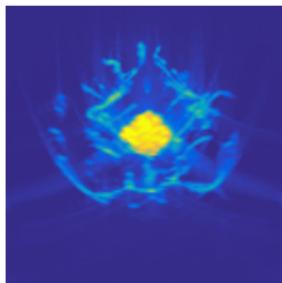
$$\hat{p} = \underset{p \geq 0}{\operatorname{argmin}} \left\{ \frac{1}{2} \| C A p - f^c \|_2^2 + \lambda \mathcal{J}(p) \right\}$$



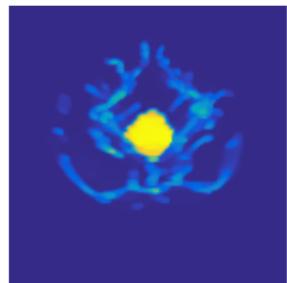
(a) $n = 128^3$



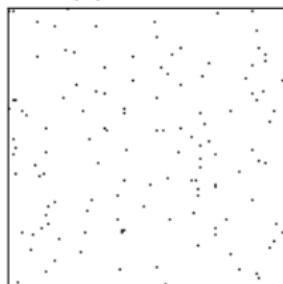
(b) TR



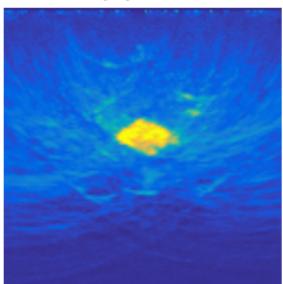
(c) L2+



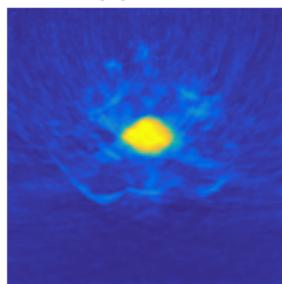
(d) TV+



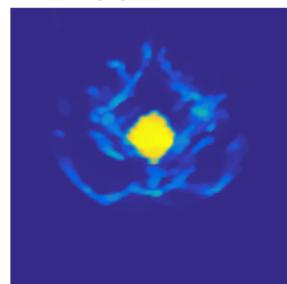
(e) SubSam, 128x



(f) TR



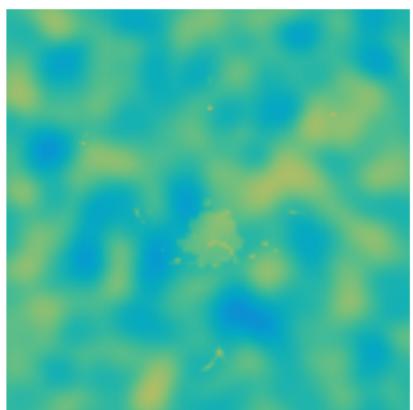
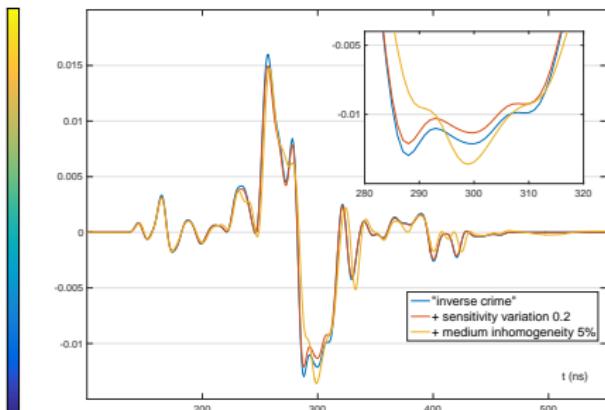
(g) L2+



(h) TV+

sensor on top; inverse crime data sampled at Nyquist; max intensity proj., side view

- ! Data created by the **same forward model** used for reconstruction.
- ! Conventional data was sampled at **Nyquist rates in space and time**.

(a) $c_0 + \tilde{c}$ 

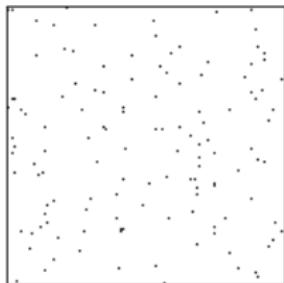
(b)

(c) pressure-time courses

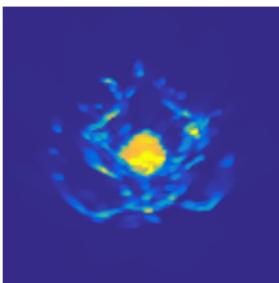
To obtain more realistic results:

- ▶ Generate data with perturbed, heterogeneous acoustic model.
- ▶ Model inhomogenous sensitivity and noise level of sensor channels.
- ▶ Conventional, "full" data is acquired below spatial Nyquist rate.

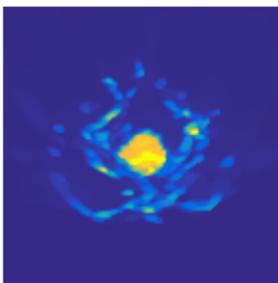
Conventional data acquired on 2×2 too coarse grid.



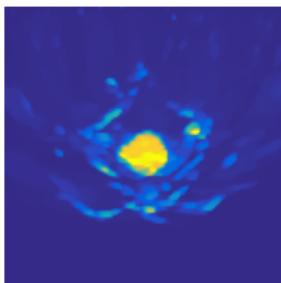
(d) single point



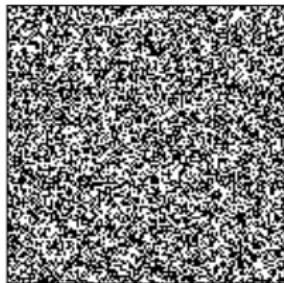
(e) TV+Br, 1x



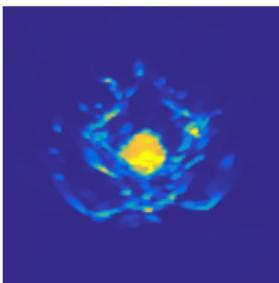
(f) TV+Br, 8x



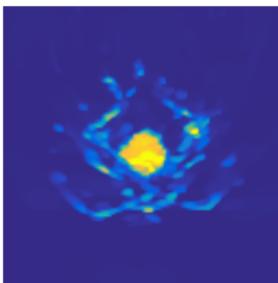
(g) TV+Br, 32x



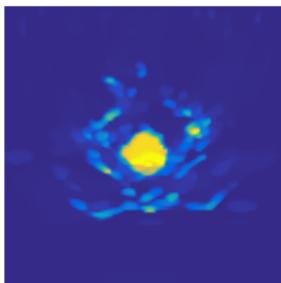
(h) patterned inter.



(i) TV+Br, 1x

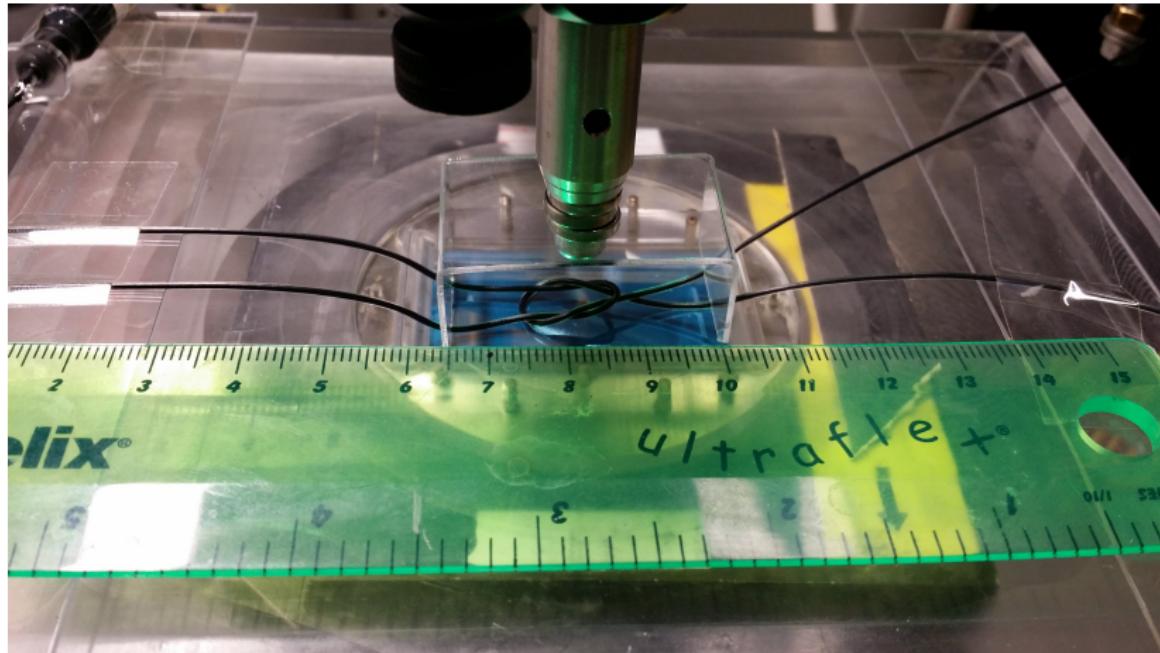


(j) TV+Br, 8x



(k) TV+Br, 32x

sensor on top; max intensity proj., side view

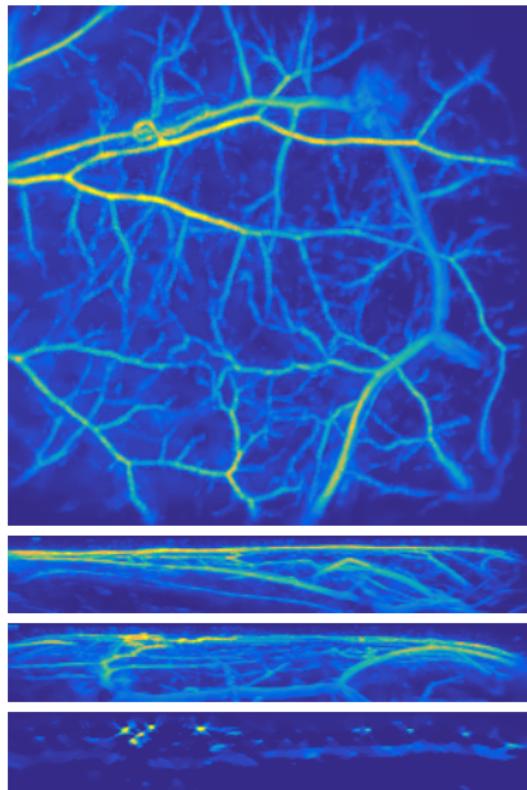


- ▶ Two polythene tubes filled with 10/100% ink.
- ▶ Stop-motion-style data acquisition of pulling one tube end.
- ▶ 45 frames (15min for conventional scanning per frame).
- ▶ Conventional data reconstructions to validate sub-sampling.

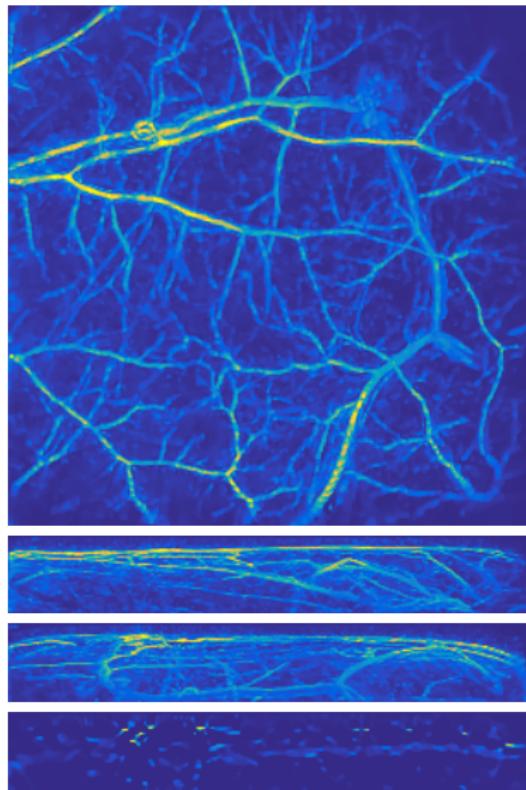
TR & TV denoising

TV+

In Vivo Measurements: Conventional Data



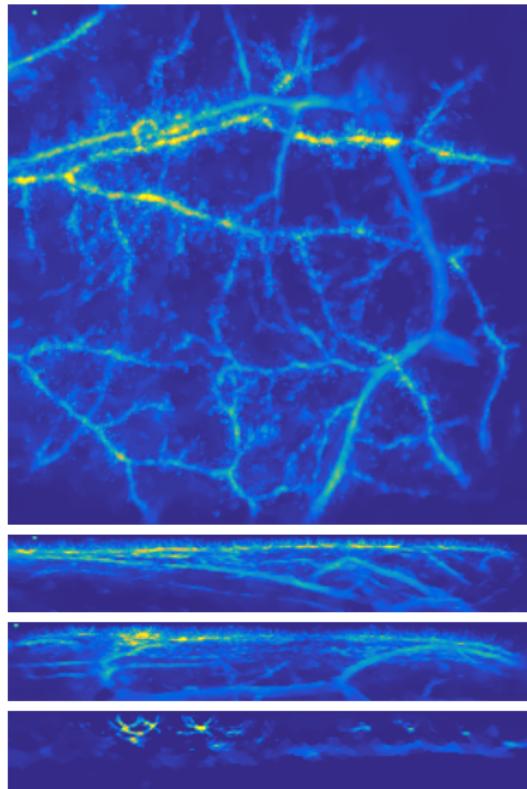
TR & TV denoising



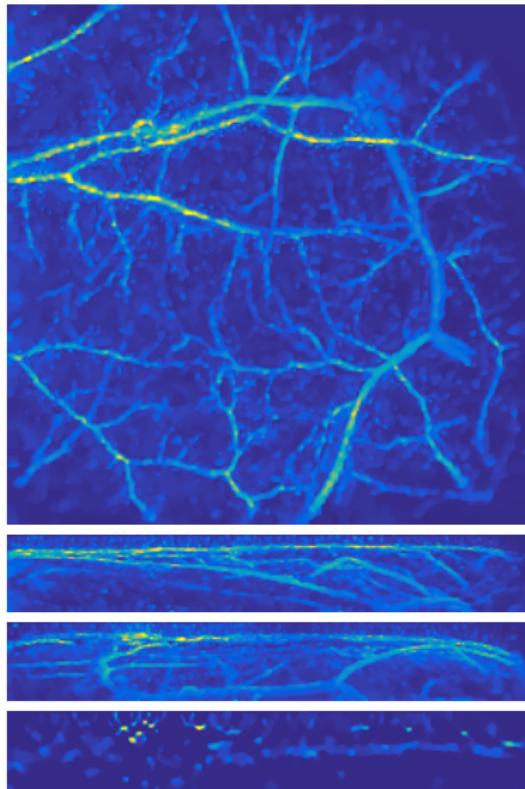
Bregman TV+

Thanks to Olumide Ogunlade for the excellent data!

In Vivo Measurements: 4x



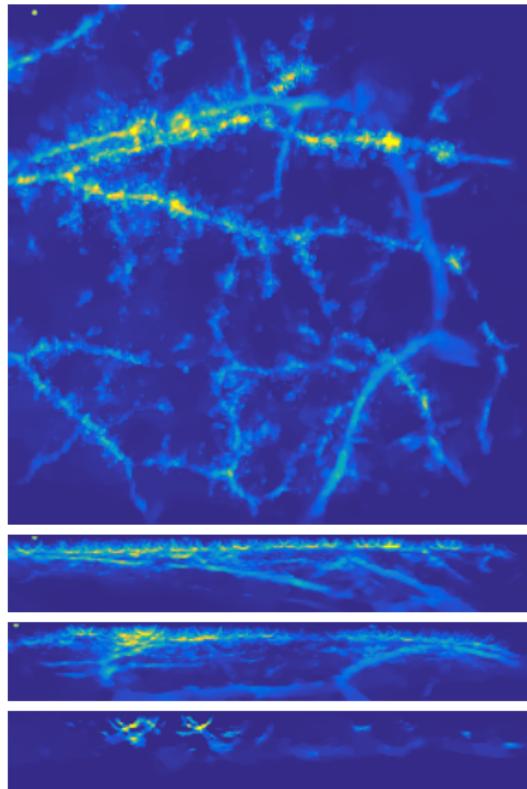
TR & TV denoising



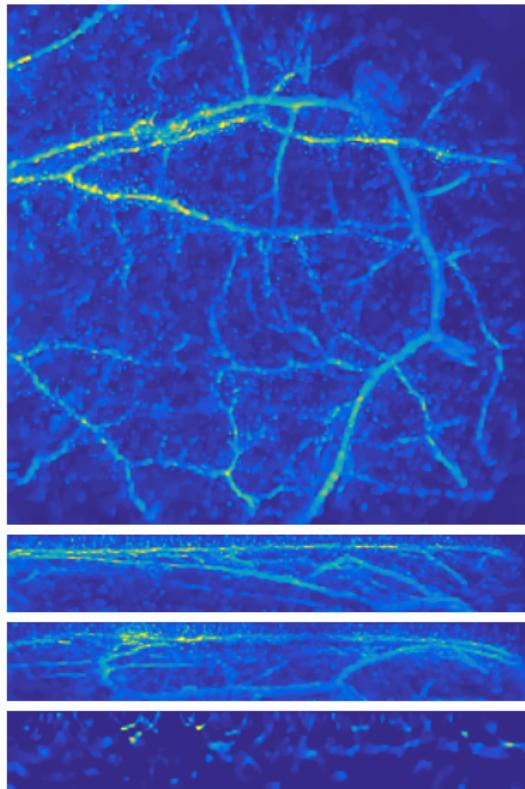
Bregman TV+

Thanks to Olumide Ogunlade for the excellent data!

In Vivo Measurements: 8x



TR & TV denoising



Bregman TV+

Thanks to Olumide Ogunlade for the excellent data!

Continuous data acquisition

⇒ tradeoff between spatial and temporal resolution.

Different dynamic models:

- ▶ Structured Low-Rank (functional imaging with static anatomies/QPAT).
- ▶ Tracer uptake/wash-in models.
- ▶ Perfusion models.
- ▶ Needle guidance
- ▶ Optical flow constraints for joint image reconstruction and motion estimation.

$$P = W \cdot V, \quad P \in \mathbb{R}^{N \times K}, \quad W \in \mathbb{R}^{N \times R}, \quad V \in \mathbb{R}^{R \times K}, \quad R \leq \min(N, K)$$

Example, $N = 10\,000$, $K = 25$, $R = 1$:

Can we acquire multi-spectral data as fast as one conventional scan?

- ▶ spatial sub-sampling by factor $K = 25$.
- ▶ 4 instead of 100 scanning locations per wave length.
- ▶ geometric information scattered over data set.

$$\hat{p}_i = \underset{p \geq 0}{\operatorname{argmin}} \left\{ \|C_i A p - f_i^c\|_2^2 \right\} \quad \forall i = 1, \dots, K$$

Neither geometry nor spectrum can be recovered!

$$\hat{P} = \underset{P \geq 0}{\operatorname{argmin}} \left\{ \frac{1}{2} \| CAP - F^c \|_{fro}^2 + \lambda |P|_* \right\}$$

λ such that $\text{rank}(P) = 1$ + Bregman iter to restore contrast.

Better, but...

$$P^{k+1} = \Pi \left(P^k - \nu \nabla \frac{1}{2} \| CAP^k - F^c \|_2^2 \right) = \Pi \left(P^k - \nu A^T C^T (CAP^k - F^c) \right)$$

- ✓ Π projection onto convex set, e.g., \mathbb{R}_+^N .
- ✓ Π proximal mapping for convex functional, e.g., nuclear norm, TV.
- ! Π projection onto **non-convex** set, e.g., **non-negative matrix factorization**.

Recovers both geometry and spectrum!

Aim: Recover (relative) chromophore concentrations, e.g., blood oxygen saturation (sO_2).

Study: Recover known concentrations in tube phantom. PA reconstruction only first step in procedure.

...but data is messy & computations are heavy, so no results yet :/

Joint ongoing struggle with Martina Bargeman Fonseca, Robert Ellwood, Emma Malone, Lu An, Ben Cox, Simon Arridge and Paul Beard.

Challenges of fast, high resolution 3D PA sensing:

- ▶ Nyquist requires several thousand detection points.
- ▶ Sequential schemes are **slow**.
- ▶ Crucial limitation for clinical, spectral and dynamical PAT.

Acceleration through sub-sampling:

- ▶ Exploit **low spatio-temporal complexity** to beat Nyquist.
- ▶ Acceleration by sub-sampling the incident wave field to **maximize non-redundancy** of data.
- ▶ Requires development of **novel scanners**.
- ▶ Demonstrated for Fabry-Pérot interferometer.

Results:

- ▶ Standard reconstruction methods fail on sub-sampled data.
- ▶ Adjoint PAT operator allows to use variational/iterative approaches.
- ▶ Sparse variational regularization/iterative non-convex projections give promising results for sub-sampled data.
- ▶ Demonstrated on simulated, experimental phantom and in-vivo data.

Challenges:

- ▶ Realizing this potential with experimental data requires
 - ▶ Model refinement/calibration.
 - ▶ Pre-processing to align data and model.
 - ▶ More suitable spatio-temporal constraints.
- ▶ High computational complexity.

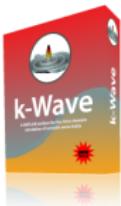


Arridge, Beard, Betcke, Cox, Huynh, L, Ogunlade, Zhang, 2016.

Accelerated High-Resolution Photoacoustic Tomography via Compressed Sensing, submitted, arXiv:1605.00133.



Arridge, Betcke, Cox, L, Treeby, 2015. *On the Adjoint Operator in Photoacoustic Tomography, submitted, arXiv:1602.02027.*



We gratefully acknowledge the support of NVIDIA Corporation with the donation of the Tesla K40 GPU used for this research.

Thank you for your attention!



Arridge, Beard, Betcke, Cox, Huynh, L, Ogunlade, Zhang, 2016.

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Variational approaches,

$$\hat{p} = \operatorname{argmin}_p \left\{ \frac{1}{2} \| C A p - f^c \|_2^2 + \lambda \mathcal{J}(p) \right\},$$

suffer from **systematic bias** (e.g., contrast loss for TV):

- ! Problem for **quantitative use**.
- ✓ Iterative enhancement through **Bregman iterations**:

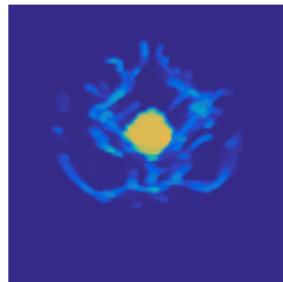
$$\begin{aligned} p^{k+1} &= \operatorname{argmin}_p \left\{ \frac{1}{2} \| C A p - (f^c + b^k) \|_2^2 + \lambda \mathcal{J}(p) \right\} \\ b^{k+1} &= b^k + (f^c - C A p^{k+1}) \end{aligned}$$

Potential for sub-sampling demonstrated in several other applications.

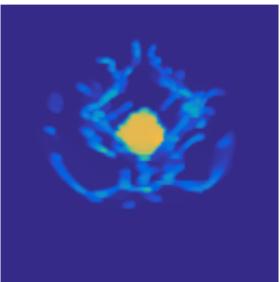


Osher, Burger, Goldfarb, Xu, Yin, 2006. *An iterative regularization method for total variation-based image restoration, Multiscale Modeling and Simulation, 4(2):460-489.*

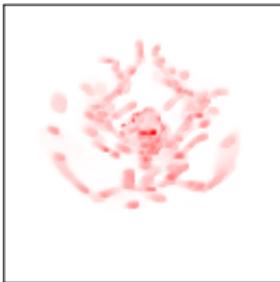
Contrast Enhancement by Bregman Iterations



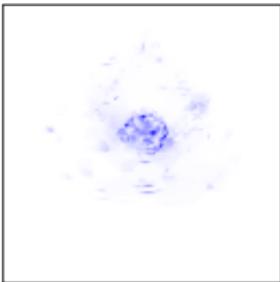
(a) $\text{TV}+$, cnv data



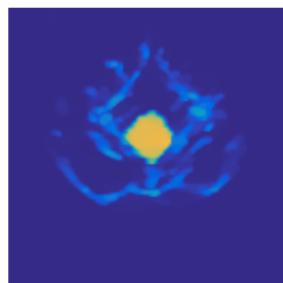
(b) $\text{TV}+\text{Br}$,
cnv
data



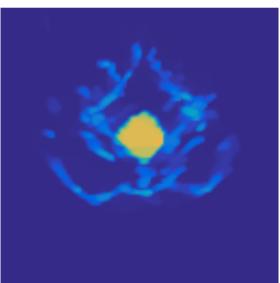
(c) $(p_{\text{TV}+\text{Br}} - p_{\text{TV}+})_+$,
cnv data



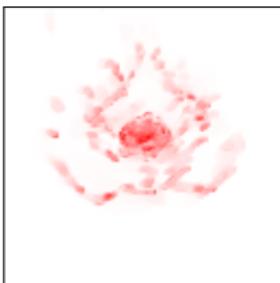
(d) $(p_{\text{TV}+\text{Br}} - p_{\text{TV}+})_-$,
cnv data



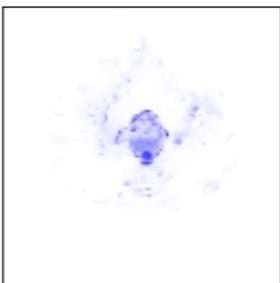
(e) $\text{TV}+$, rSP-128



(f) $\text{TV}+\text{Br}$, rSP-128



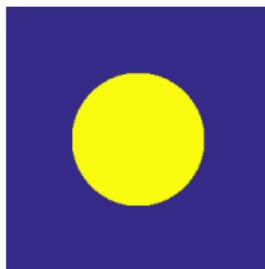
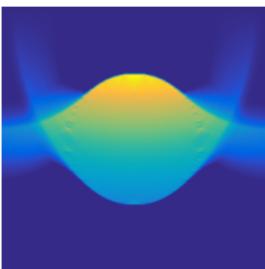
(g) $(p_{\text{TV}+\text{Br}} - p_{\text{TV}+})_+$,
rSP-128



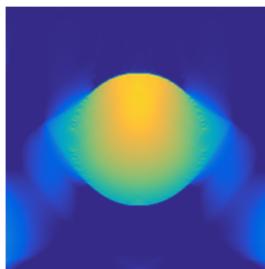
(h) $(p_{\text{TV}+\text{Br}} - p_{\text{TV}+})_-$,
rSP-128

sensor on top; inverse crime data sampled at Nyquist; max intensity proj., side view

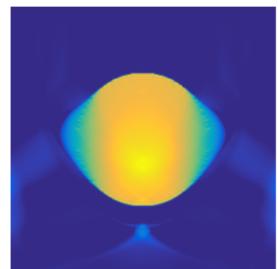
$$p^{k+1} = \Pi \left(p^k - \theta B \left(A p^k - f \right) \right)$$

(a) Ground truth p_0 

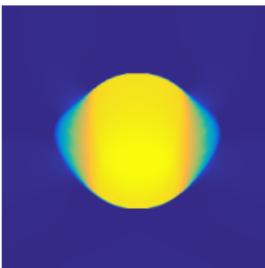
(b) TR



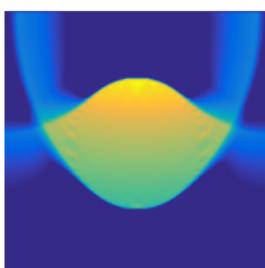
(c) iTR



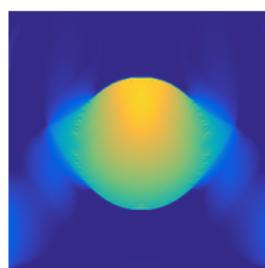
(d) iTR+



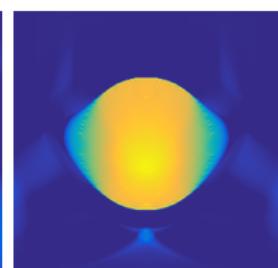
(e) TV+



(f) BP



(g) LS



(h) LS+

sensor on top; 2D slices at $y = 128$ through the 3D reconstructions.

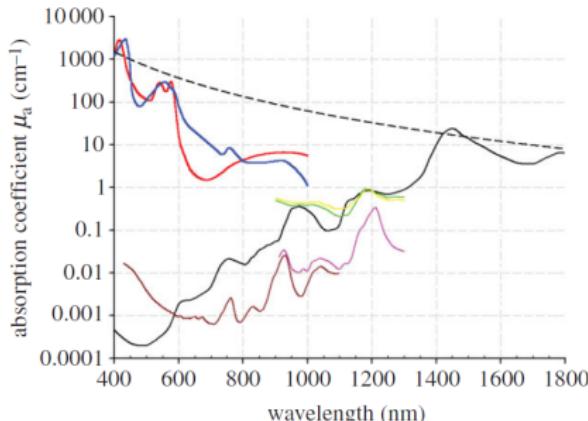


Figure 1. Absorption coefficient spectra of endogenous tissue chromophores. Oxyhaemoglobin (HbO_2), red line: (<http://omlc.ogi.edu/spectra/hemoglobin/summary.html>; 150 g l^{-1}), deoxyhaemoglobin (HHb), blue line: (<http://omlc.ogi.edu/spectra/hemoglobin/summary.html>; 150 g l^{-1}), water, black line [22] (80% by volume in tissue), lipid^(a), brown line [23] (20% by volume in tissue), lipid^(b), pink line [24], melanin, black dashed line (<http://omlc.ogi.edu/spectra/melanin/mua.html>; μ_a corresponds to that in skin). Collagen (green line) and elastin (yellow line) spectra from [24].

- ▶ High contrast between blood and water/lipid.
- ▶ Light-absorbing structures embedded in soft tissue.
- ▶ Gap between oxygenated and deoxygenated blood
~~ functional imaging.
- ▶ Different wavelengths allow quantitative spectroscopic examinations.
- ▶ Use of contrast agents for molecular imaging.

from: Paul Beard, 2011. *Biomedical photoacoustic imaging, Interface Focus*.

- ▶ Up to now, conventional data was sampled at **Nyquist rates in space and time** (numerical phantoms were band-limited in space).
- ▶ In experiments, conventional data is usually already sub-sampled in space but over-sampled in time.
- ▶ Reconstruction on a finer spatial grid to exploit high frequency content of time series.

Example:

- ▶ Scan a $20\text{mm} \times 20\text{mm}$ with $\delta_x = 150\mu\text{m}$ (133×133 locations).
- ▶ Measured with temporal resolution of $\delta_t = 12\text{ns} \approx 83\text{MHz}$.
- ▶ Low-pass filtered to 20MHz .
- ▶ Reconstructing a signal limited to 20MHz with a sound speed of 1540m s^{-1} would require $\delta_x = 38.5\mu\text{m}$ and $\delta_t = 25\text{ns}$.