Statistical reconstruction for low-dose CT

Jeffrey A. Fessler

EECS Department University of Michigan



FDA

Apr. 26, 2011

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

Full disclosure

- Research support from GE Healthcare
- Research support to GE Global Research
- Work supported in part by NIH grant R01-HL-098686
- Research support from Intel

Credits

Current students / post-docs

- Jang Hwan Cho
- Wonseok Huh
- Donghwan Kim
- Yong Long
- Madison McGaffin
- Sathish Ramani
- Stephen Schmitt
- Meng Wu

GE collaborators

- Jiang Hsieh
- Jean-Baptiste Thibault
- Bruno De Man

CT collaborators

- Mitch Goodsitt, UM
- Ella Kazerooni, UM
- Neal Clinthorne, UM
- Paul Kinahan, UW

Former PhD students (who did/do CT)

- Se Young Chun, Harvard / Brigham
- Hugo Shi, Enthought
- Joonki Noh, Emory
- Somesh Srivastava, JHU
- Rongping Zeng, FDA
- Yingying, Zhang-O'Connor, RGM Advisors
- Matthew Jacobson, Xoran
- Sangtae Ahn, GE
- Idris Elbakri, CancerCare / Univ. of Manitoba
- Saowapak Sotthivirat, NSTDA Thailand
- Web Stayman, JHU
- Feng Yu, Univ. Bristol
- Mehmet Yavuz, Qualcomm
- Hakan Erdoğan, Sabanci University

Former MS students

- Kevin Brown, Philips
- ...

A picture is worth 1000 words

(and perhaps several 1000 seconds of computation?)



Thin-slice FBP

Seconds

ASIR

A bit longer

Statistical

Much longer

Why statistical methods for CT?

- Accurate physical models
 - X-ray spectrum, beam-hardening, scatter, ...
 reduced artifacts? quantitative CT?
 - X-ray detector spatial response, focal spot size, ... improved spatial resolution?
 - detector spectral response (e.g., photon-counting detectors)
- Nonstandard geometries
 - transaxial truncation (big patients)
 - long-object problem in helical CT
 - o irregular sampling in "next-generation" geometries
 - o coarse angular sampling in image-guidance applications
 - limited angular range (tomosynthesis)
 - o "missing" data, e.g., bad pixels in flat-panel systems
- Appropriate statistical models
 - weighting reduces influence of photon-starved rays (FBP treats all rays equally)
 - o reducing image noise or dose

and more...

- Object constraints
 - nonnegativity
 - object support
 - piecewise smoothness
 - object sparsity (e.g., angiography)
 - sparsity in some basis
 - o motion models
 - dynamic models
 - 0 ...

Disadvantages?

- Computation time (super computer)
- Must reconstruct entire FOV
- Model complexity
- Software complexity
- Algorithm nonlinearities
 - Difficult to analyze resolution/noise properties (cf. FBP)
 - Tuning parameters
 - Challenging to characterize performance

"Iterative" vs "Statistical"

- Traditional successive substitutions iterations
 - o e.g., Joseph and Spital (JCAT, 1978) bone correction
 - usually only one or two "iterations"
 - not statistical
- Algebraic reconstruction methods
 - o Given sinogram data y and system model A, reconstruct object x by "solving" y = Ax
 - o ART, SIRT, SART, ...
 - iterative, but typically not statistical
 - Iterative filtered back-projection (FBP):

$$\mathbf{x}^{(n+1)} = \mathbf{x}^{(n)} + \mathbf{\alpha}$$
 FBP($\mathbf{y} - \mathbf{A}\mathbf{x}^{(n)}$)
step data forward
size project

- Statistical reconstruction methods
 - Image domain
 - Sinogram domain
 - Fully statistical (both)
 - Hybrid methods (e.g., AIR, SPIE 7961-18, Bruder et al.)

"Statistical" methods: Image domain

Denoising methods

$$\begin{array}{c} \mathsf{noisy} \\ \mathbf{y} \end{array} \to \begin{array}{c} \mathsf{FBP} \to \\ \mathsf{reconstruction} \\ \tilde{\mathbf{x}} \end{array} \to \begin{array}{c} \mathsf{iterative} \\ \mathsf{denoiser} \end{array} \to \begin{array}{c} \mathsf{final} \\ \mathsf{image} \\ \hat{\mathbf{x}} \end{array}$$

- Relatively fast, even if iterative
- Remarkable advances in denoising methods in last decade

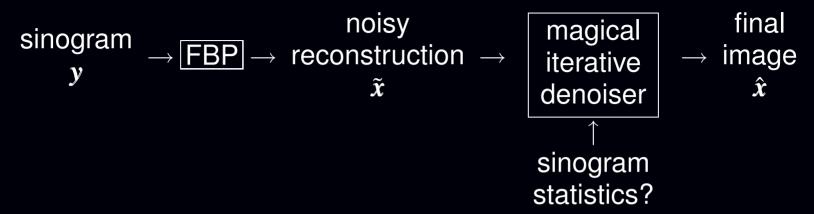




Zhu & Milanfar, T-IP, Dec. 2010, using "steering kernel regression" (SKR) method Challenges:

- Typically assume white noise
- Streaks in low-dose FBP appear like edges (highly correlated noise)

Image denoising methods "guided by data statistics"



- Image-domain methods are fast (thus practical)
- o ASIR? IRIS? ...
- The technical details are often a mystery...

Challenges:

- FBP often does not use all data efficiently (e.g., Parker weighting)
- Low-dose CT statistics most naturally expressed in sinogram domain

"Statistical" methods: Sinogram domain

Sinogram restoration methods

- o Adaptive: J. Hsieh, Med. Phys., 1998; Kachelrieß, Med. Phys., 2001, ...
- o Iterative: P. La Riviere, IEEE T-MI, 2000, 2005, 2006, 2008
- Relatively fast even if iterative

Challenges:

- Limited denoising without resolution loss
- o Difficult to "preserve edges" in sinograms



FBP, 10 mA

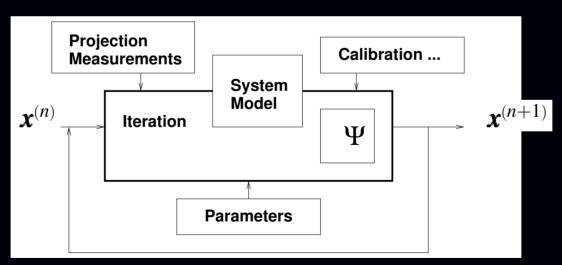


FBP from denoised sinogram

(True? Fully? Slow?) Statistical reconstruction

- Object model
- Physics/system model
- Statistical model
- Cost function (log-likelihood + regularization)
- Iterative algorithm for minimization

"Find the image \hat{x} that best fits the sinogram data y according to the physics model, the statistical model and prior information about the object"



- Repeatedly revisiting the sinogram data can use statistics fully
- Repeatedly updating the image can exploit object properties
- : greatest potential dose reduction, but repetition is expensive...

History: Statistical reconstruction for PET

- Iterative method for emission tomography (Kuhl, 1963)
- Weighted least squares for 3D SPECT (Goitein, NIM, 1972)
- Richardson/Lucy iteration for image restoration (1972, 1974)
- Poisson likelihood (emission) (Rockmore and Macovski, TNS, 1976)
- Expectation-maximization (EM) algorithm (Shepp and Vardi, TMI, 1982)
- Regularized (aka Bayesian) Poisson emission reconstruction (Geman and McClure, ASA, 1985)
- Ordered-subsets EM (OSEM) algorithm (Hudson and Larkin, TMI, 1994)
- Commercial release of OSEM for PET scanners circa 1997

Today, most commercial PET systems include unregularized OSEM.

15 years between key EM paper (1982) and commercial adoption (1997) (25 years if you count the R/L paper in 1972 which is the same as EM)

Key factors in PET

- OS algorithm accelerated convergence by order of magnitude
- Computers got faster (but problem size grew too)
- Key clinical validation papers?
- Key numerical observer studies?
- Nuclear medicine physicians grew accustomed to appearance of images reconstructed using statistical methods

History: Statistical reconstruction for CT*

• Iterative method for X-ray CT (Hounsfield, 1968)

ART for tomography (Gordon, Bender, Herman, JTB, 1970)

• ...

- Roughness regularized LS for tomography (Kashyap & Mittal, 1975)
- Poisson likelihood (transmission) (Rockmore and Macovski, TNS, 1977)
- EM algorithm for Poisson transmission (Lange and Carson, JCAT, 1984)
- Iterative coordinate descent (ICD) (Sauer and Bouman, T-SP, 1993)
- Ordered-subsets algorithms

(Manglos *et al.*, PMB 1995) (Kamphuis & Beekman, T-MI, 1998) (Erdoğan & Fessler, PMB, 1999)

• ...

Commercial introduction for CT scanners

circa 2010

RSNA 2010



Zhou Yu, Jean-Baptiste Thibault, Charles Bouman, Jiang Hsieh, Ken Sauer

Five Choices for Statistical Reconstruction

- 1. Object model
- 2. System physical model
- 3. Measurement statistical model
- 4. Cost function: data-mismatch and regularization
- 5. Algorithm / initialization

No perfect choices - one can critique all approaches!

Historically these choices are often left implicit in publications, but being explicit facilitates reproducibility

Choice 1. Object Parameterization

Finite measurements: $\{y_i\}_{i=1}^M$. Continuous object: $f(\vec{r}) = \mu(\vec{r})$.

"All models are wrong but some models are useful."

Linear *series expansion* approach. Represent $f(\vec{r})$ by $\mathbf{x} = (x_1, \dots, x_N)$ where

$$f(\vec{r}) pprox \tilde{f}(\vec{r}) = \sum_{j=1}^{N} x_j b_j(\vec{r}) \leftarrow$$
 "basis functions"

Reconstruction problem becomes "discrete-discrete:" estimate x from y

Numerous basis functions in literature. Two primary contenders:

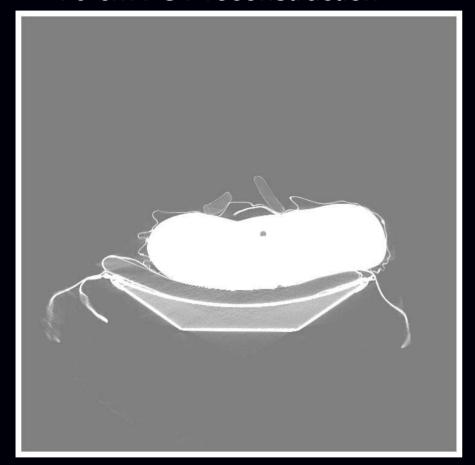
- voxels
- blobs (Kaiser-Bessel functions)
 - + Blobs are approximately band-limited (reduced aliasing?)
 - Blobs have larger footprints, increasing computation.

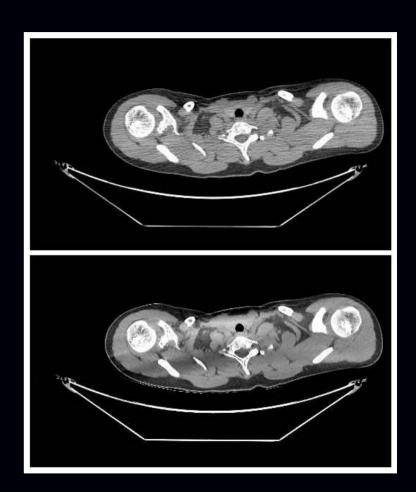
Open question: how small should the voxels be?

One practical compromise: wide FOV coarse-grid reconstruction followed by fine-grid refinement over ROI, e.g., Ziegler et al., Med. Phys., Apr. 2008

Global reconstruction: An inconvenient truth

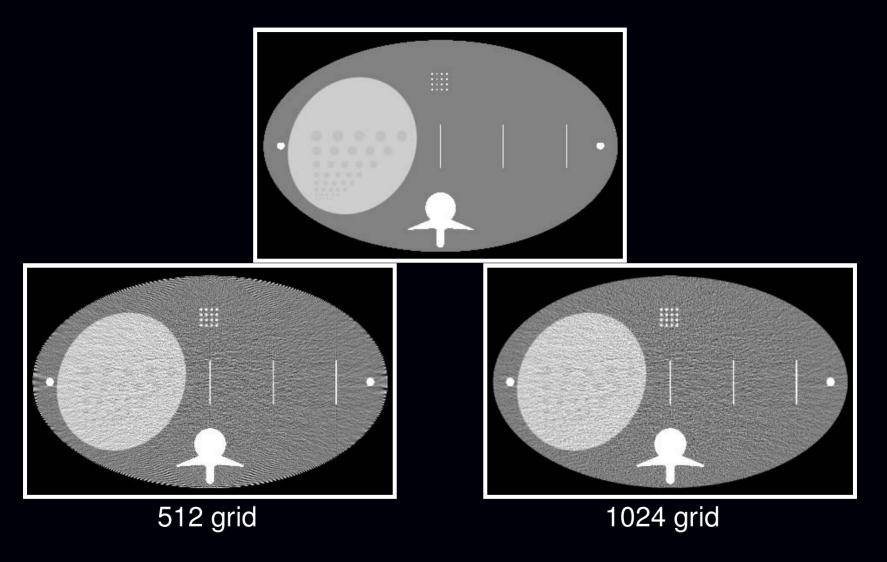
70-cm FOV reconstruction





Thibault et al., Fully3D, 2007

Voxel size matters?



Unregularized OS reconstructions. Zbijewski & Beekman, PMB, Jan. 2004

Choice 2. System model / Physics model

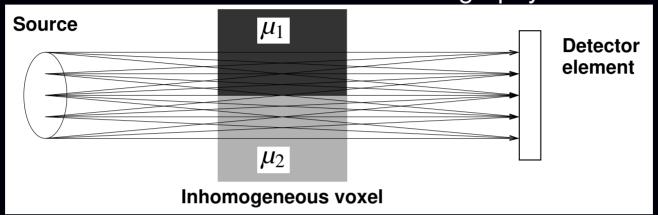
- scan geometry
- source intensity *I*₀
 - spatial variations (air scan)
 - intensity fluctuations
- resolution effects
 - finite detector size / detector spatial response
 - o finite X-ray spot size / anode angulation Inhomogeneous
 - detector afterglow
- spectral effects
 - X-ray source spectrum
 - bowtie filters
 - detector spectra response
- scatter
- ...

Challenges / trade-offs

- computation time versus
- accuracy/artifacts/resolution/contrast versus
- dose?

Exponential edge-gradient effect

Fundamental difference between emission tomography and CT:



Recorded intensity for *i*th ray:

(Joseph and Spital, PMB, May 1981)

$$I_{i} = \int_{\text{source}} \int_{\text{detector}} I_{0}(\vec{p}_{s}, \vec{p}_{d}) \exp\left(-\int_{\mathcal{L}(\vec{p}_{s}, \vec{p}_{d})} \mu(\vec{r}) \, d\ell\right) d\vec{p}_{d} \, d\vec{p}_{s}$$

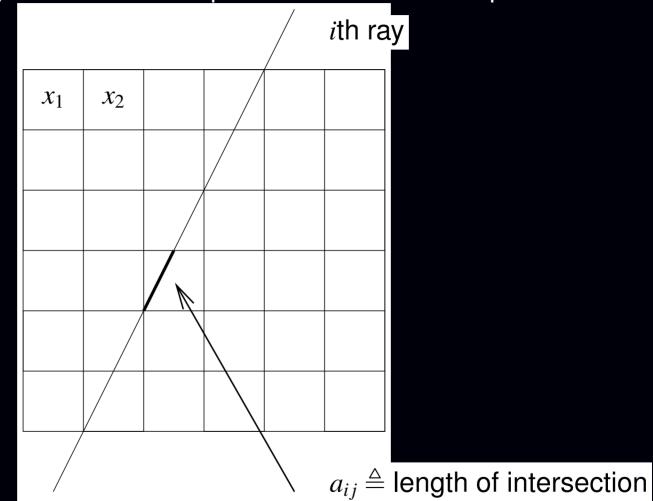
$$\neq I_{0} \exp\left(-\int_{\text{source}} \int_{\text{detector}} \int_{\mathcal{L}(\vec{p}_{s}, \vec{p}_{d})} \mu(\vec{r}) \, d\ell \, d\vec{p}_{d} \, d\vec{p}_{s}\right).$$

Usual "linear" approximation:

$$I_i \approx I_0 \exp\left(-\sum_{j=1}^N a_{ij}x_j\right), \qquad \underbrace{a_{ij} \triangleq \int_{\mathrm{source}} \int_{\det \mathrm{ector}} \int_{\mathcal{L}(\vec{p}_s, \vec{p}_d)} b_j(\vec{r}) \, \mathrm{d}\ell \, \mathrm{d}\vec{p}_d \, \mathrm{d}\vec{p}_s}_{\mathrm{elements}}$$
 elements of system matrix A

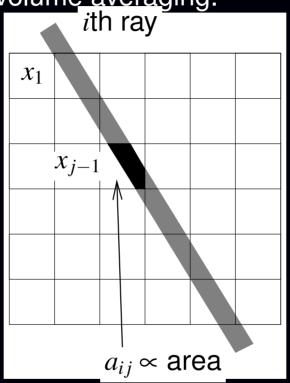
"Line Length" System Model

Assumes (implicitly?) that source is a point and detector is a point.



"Strip Area" System Model

Account for finite detector width. Ignores nonlinear partial-volume averaging.



Practical (?) implementations in 3D include

- Distance-driven method (De Man and Basu, PMB, Jun. 2004)
- Separable-footprint method (Long et al., T-MI, Nov. 2010)
- Further comparisons needed...

Lines versus strips

From (De Man and Basu, PMB, Jun. 2004)

MLTR of rabbit heart

Ray-driven (idealized point detector)



Distance-driven (models finite detector width)



Forward- / Back-projector "Pairs"

Typically iterative algorithms require two key steps.

• forward projection (image domain to projection domain):

$$\bar{\mathbf{y}} = \mathbf{A}\mathbf{x}, \qquad \bar{y}_i = \sum_{j=1}^N a_{ij}x_j = [\mathbf{A}\mathbf{x}]_i$$

backprojection (projection domain to image domain):

$$z = A'y,$$
 $z_j = \sum_{i=1}^{M} a_{ij}y_i$

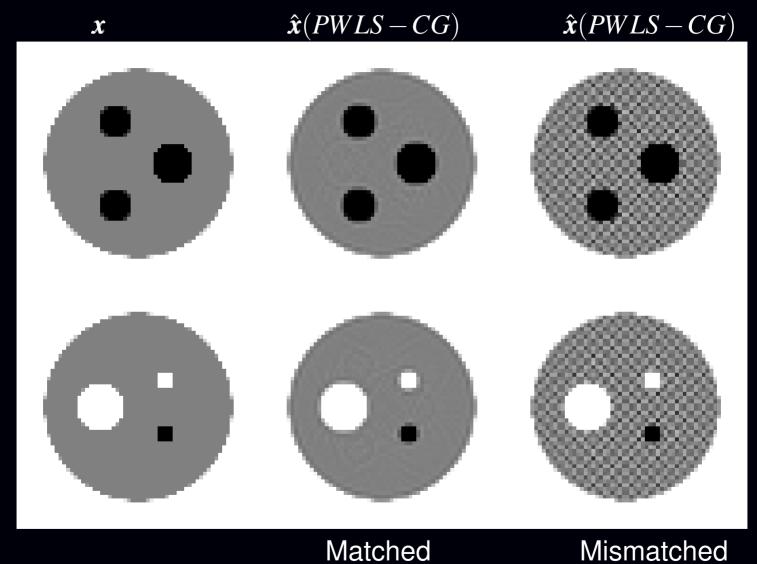
The term "forward/backprojection pair" often refers to some implicit choices for the object basis and the system model.

Sometimes A'y is implemented as By for some "backprojector" $B \neq A'$. Especially in SPECT and sometimes in PET.

Least-squares solutions (for example):

$$\hat{\boldsymbol{x}} = \underset{\boldsymbol{x}}{\operatorname{arg\,min}} \|\boldsymbol{y} - \boldsymbol{A}\boldsymbol{x}\|^2 = \left[\boldsymbol{A}'\boldsymbol{A}\right]^{-1}\boldsymbol{A}'\boldsymbol{y} \neq \left[\boldsymbol{B}\boldsymbol{A}\right]^{-1}\boldsymbol{B}\boldsymbol{y}$$

Mismatched Backprojector $B \neq A'$



cf. SPECT/PET reconstruction – usually unregularized

Projector/back-projector bottleneck

Challenges

- Projector/backprojector algorithm design
 - Approximations (e.g., transaxial/axial separability)
 - Symmetry
- Hardware / software implementation
 - o GPU, CUDA, OpenCL, FPGA, SIMD, pthread, OpenMP, MPI, ...
- Further "wholistic" approaches?
 e.g., Basu & De Man, "Branchless distance driven projection ...," SPIE 2006

• ..

Choice 3. Statistical Model

The physical model describes measurement mean, *e.g.*, for a monoenergetic X-ray source and ignoring scatter etc.:

$$\bar{I}_i([\boldsymbol{A}\boldsymbol{x}]_i) = I_0 e^{-\sum_{j=1}^N a_{ij}x_j}.$$

The raw noisy measurements $\{I_i\}$ are distributed around those means. Statistical reconstruction methods require a model for that distribution.

Challenges / Trade offs: using more accurate statistical models

- may lead to less noisy images
- may incur additional computation
- may involve higher algorithm complexity.

CT measurement statistics are very complicated, more so at low doses

- incident photon flux variations (Poisson)
- X-ray photon absorption/scattering (Bernoulli)
- energy-dependent light production in scintillator (?)
- shot noise in photodiodes (Poisson?)
- electronic noise in readout electronics (Gaussian?)
 Whiting, SPIE 4682, 2002; Lasio et al., PMB, Apr. 2007
- Inaccessibility of raw sinogram data

To log() or not to log() – That is the question

Models for "raw" data I_i (before logarithm)

- compound Poisson (complicated) Whiting, SPIE 4682, 2002; Elbakri & Fessler, SPIE 5032, 2003; Lasio et al., PMB, Apr. 2007
- Poisson + Gaussian (photon variability and electronic readout noise):

$$I_i \sim \mathsf{Poisson}\{ar{I}_i\} + \mathsf{N}(0, \mathbf{\sigma}^2)$$

Snyder et al., JOSAA, May 1993 & Feb. 1995

Shifted Poisson approximation (matches first two moments):

$$\tilde{I}_i \triangleq \left[I_i + \sigma^2\right]_+ \sim \mathsf{Poisson}\left\{\bar{I}_i + \sigma^2\right\}$$

Yavuz & Fessler, MIA, Dec. 1998

Ordinary Poisson (ignore electronic noise):

$$I_i \sim \mathsf{Poisson}\{ar{I}_i\}$$

Rockmore and Macovski, TNS, Jun. 1977; Lange and Carson, JCAT, Apr. 1984

Photon-counting detectors would simplify statistical modeling

All are somewhat complicated by the nonlinearity of the physics: $\bar{I}_i = e^{-[Ax]_i}$

After taking the log()

Taking the log leads to a linear model (ignoring beam hardening):

$$y_i \triangleq -\log\left(\frac{I_i}{I_0}\right) \approx [\mathbf{A}\mathbf{x}]_i + \mathbf{\varepsilon}_i$$

Drawbacks:

- Undefined if $I_i \leq 0$ (e.g., due to electronic noise)
- It is biased (by Jensen's inequality): $E[y_i] \ge -\log(\bar{I}_i/I_0) = [Ax]_i$
- Exact distribution of noise ε_i intractable

Practical approach: assume Gaussian noise model: $\varepsilon_i \sim N(0, \sigma_i^2)$

Options for modeling noise variance $\sigma_i^2 = Var\{\epsilon_i\}$

- consider both Poisson and Gaussian noise effects: $\sigma_i^2 = \frac{I_i + \sigma^2}{I_i^2}$ Thibault *et al.*, SPIE 6065, 2006
- consider just Poisson effect: $\sigma_i^2 = \frac{1}{I_i}$ (Sauer & Bouman, T-SP, Feb. 1993)
- pretend it is white noise: $\sigma_i^2 = \sigma_0^2$
- ignore noise altogether and "solve" y = Ax

Whether using pre-log data is better than post-log data is an open question.

Choice 4. Cost Functions

Components:

- Data-mismatch term
- Regularization term (and regularization parameter β)
- Constraints (e.g., nonnegativity)

Reconstruct image \hat{x} by minimizing a cost function:

$$\hat{x} \triangleq \underset{x \geq 0}{\operatorname{arg min}} \Psi(x)$$

$$\Psi(x) = \operatorname{DataMismatch}(y, Ax) + \beta \operatorname{Regularizer}(x)$$

Forcing too much "data fit" alone would give noisy images.

Equivalent to a Bayesian MAP (maximum a posteriori) estimator.

Distinguishes "statistical methods" from "algebraic methods" for "y = Ax."

Choice 4.1: Data-Mismatch Term

Standard choice is the negative log-likelihood of statistical model:

DataMismatch =
$$-L(\mathbf{x}; \mathbf{y}) = -\log p(\mathbf{y}|\mathbf{x}) = \sum_{i=1}^{M} -\log p(y_i|\mathbf{x})$$
.

For pre-log data I with shifted Poisson model:

$$-L(\boldsymbol{x};\boldsymbol{I}) = \sum_{i=1}^{M} (\bar{I}_i + \sigma^2) - [I_i + \sigma^2]_+ \log(\bar{I}_i + \sigma^2), \qquad \bar{I}_i = I_0 e^{-[\boldsymbol{A}\boldsymbol{x}]_i}$$

This can be non-convex if $\sigma^2 > 0$; it is convex if we ignore electronic noise $\sigma^2 = 0$. Trade-off ...

For post-log data y with Gaussian model:

$$-L(\mathbf{x};\mathbf{y}) = \sum_{i=1}^{M} w_i \frac{1}{2} (y_i - [\mathbf{A}\mathbf{x}]_i)^2 = \frac{1}{2} (\mathbf{y} - \mathbf{A}\mathbf{x})' \mathbf{W} (\mathbf{y} - \mathbf{A}\mathbf{x}), \qquad w_i = 1/\sigma_i^2$$

This is a kind of (data-based) weighted least squares (WLS). It is always convex in x. Quadratic functions are "easy" to minimize.

• ...

Choice 4.2: Regularization

How to control noise due to ill-conditioning?

Noise-control methods in clinical use in PET reconstruction today:

- Stop an unregularized algorithm before convergence
- Over-iterate an unregularized algorithm then post-filter

Other possible "simple" solutions:

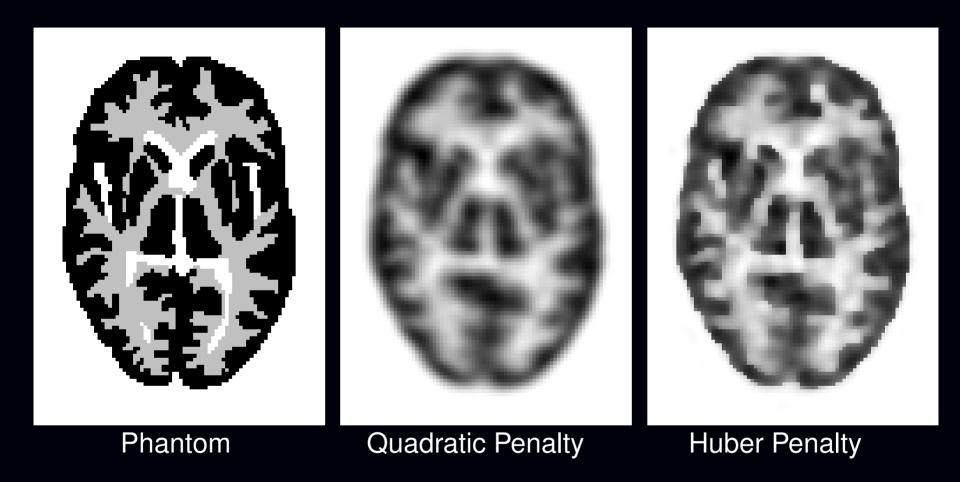
- Modify the raw data (pre-filter / denoise)
- Filter between iterations
- ...

Appeal:

- simple / familiar
- filter parameters have intuitive units (*e.g.*, FWHM), unlike a regularization parameter β
- Changing a post-filter does not require re-iterating, unlike changing a regularization parameter β

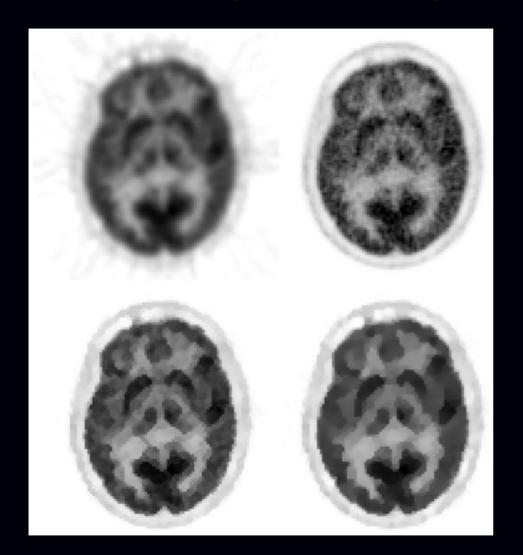
Dozens of papers on regularized methods for PET, but little clinical impact. (USC MAP method is available in mouse scanners.)

Edge-Preserving Reconstruction: PET Example



Quantification vs qualitative vs tasks...

More "Edge Preserving" PET Regularization



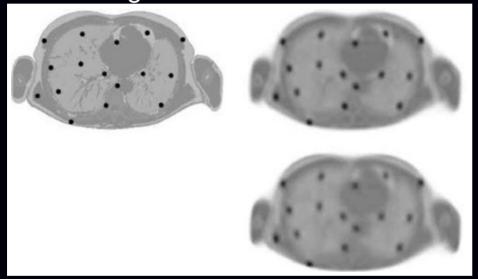
FBP	ML-EM
Median-root	Huber
prior	regularizer

Chlewicki et al., PMB, Oct. 2004; "Noise reduction and convergence of Bayesian algorithms with blobs based on the Huber function and median root prior"

Regularization in PET

Nuyts *et al.*, T-MI, Jan. 2009: MAP method outperformed post-filtered ML for lesion detection in simulation

Noiseless images:



Phantom	ML-EM filtered
	Regularized

Regularization options

Options for regularizer R(x) in increasing complexity:

- quadratic roughness
- convex, non-quadratic roughness
- non-convex roughness
- total variation
- convex sparsity
- non-convex sparsity

Challenges

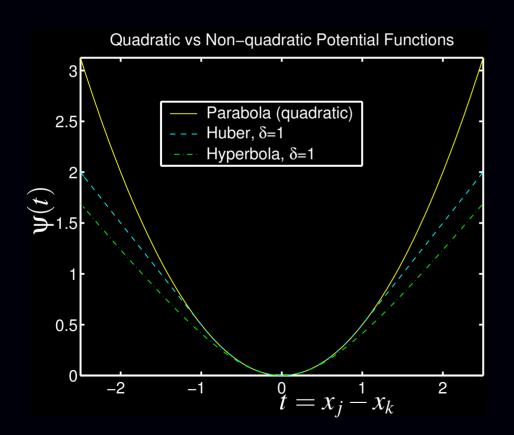
- Reducing noise without degrading spatial resolution
- Balancing regularization strength between and within slices
- Parameter selection
- Computational complexity (voxels have 26 neighbors in 3D)
- Preserving "familiar" noise texture
- Optimizing clinical task performance

Many open questions...

Roughness Penalty Functions

$$R(\mathbf{x}) = \sum_{j=1}^{N} \frac{1}{2} \sum_{k \in \mathcal{N}_j} \mathbf{\psi}(x_j - x_k)$$

 $\mathcal{N}_j \triangleq neighborhood$ of jth pixel (e.g., left, right, up, down) ψ called the *potential function*



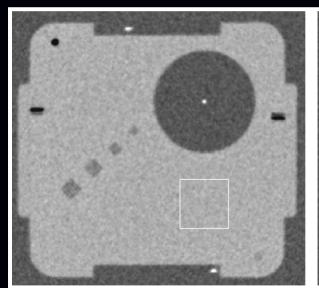
quadratic: $\psi(t) = t^2$ hyperbola: $\psi(t) = \sqrt{1 + (t/\delta)^2}$ (edge preservation)

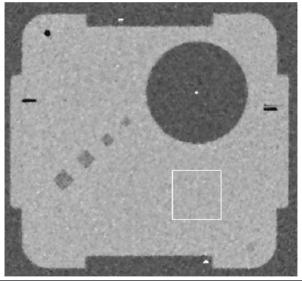
Regularization parameters: Dramatic effects

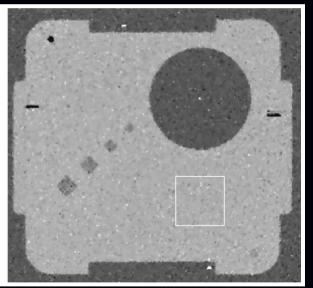
Thibault et al., Med. Phys., Nov. 2007

"q generalized gaussian" potential function with tuning parameters: β, δ, p, q :

$$\beta \Psi(t) = \beta \frac{\frac{1}{2} |t|^p}{1 + |t/\delta|^{p-q}}$$







p = q = 2 $p = 2, q = 1.2, \delta = 10 \text{ HU}$

$$p = q = 1.1$$

noise: 11.1 (#lp/cm): 4.2

10.9 7.2

10.8 8.2

Summary thus far

- 1. Object parameterization
- 2. System physical model
- 3. Measurement statistical model
- 4. Cost function: data-mismatch / regularization / constraints

Reconstruction Method ≜ **Models** + **Cost Function** + **Algorithm**

5. Minimization algorithms:

$$\hat{\mathbf{x}} = \operatorname*{arg\,min}_{\mathbf{x}} \Psi(\mathbf{x})$$

Choice 5: Minimization algorithms

Conjugate gradients

- Converges slowly for CT
- o Difficult to precondition due to weighting and regularization
- Difficult to enforce nonnegativity constraint
- Very easily parallelized

Ordered subsets

- Initially converges faster than CG if many subsets used
- o Does not converge without relaxation etc., but those slow it down
- \circ Computes regularizer gradient $\nabla R(x)$ for every subset expensive?
- Easily enforces nonnegativity constraint
- Easily parallelized

Coordinate descent

(Sauer and Bouman, T-SP, 1993)

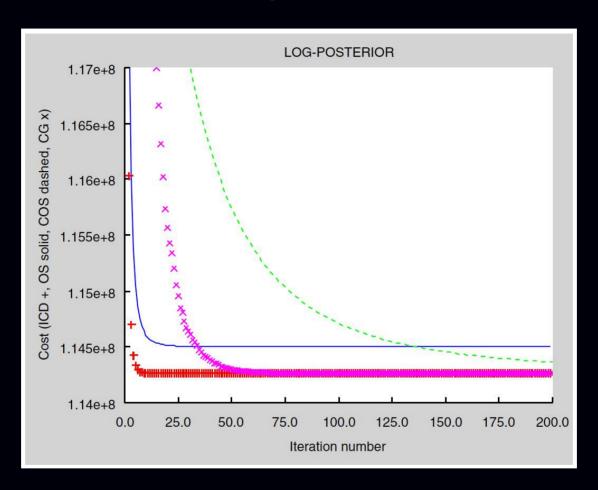
- o Converges high spatial frequencies rapidly, but low frequencies slowly
- Easily enforces nonnegativity constraint
- Challenging to parallelize

Block coordinate descent

(Benson et al., NSS/MIC, 2010)

- Spatial frequency convergence properties depend...
- Easily enforces nonnegativity constraint
- More opportunity to parallelize than CD

Convergence rates

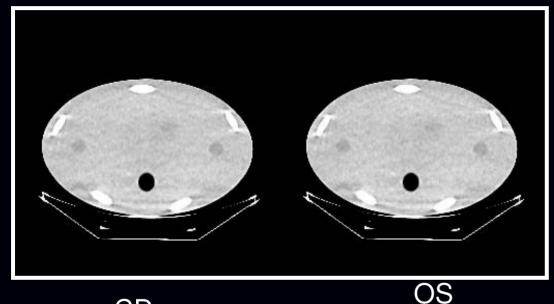


(De Man et al., NSS/MIC 2005)

In terms of iterations: CD < OS < CG < Convergent OS In terms of compute time? (it depends...)

Ordered subsets convergence

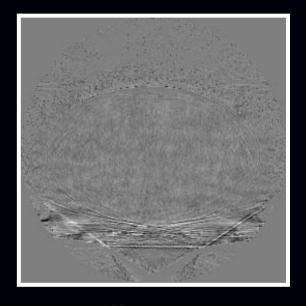
Theoretically OS does not converge, but it may get "close enough," even with regularization.



CD 200 iter

display: 930 HU \pm 58 HU

41 subsets 200 iter



 $\begin{array}{c} \text{difference} \\ 0 \pm 10 \text{HU} \end{array}$

(De Man et al., NSS/MIC 2005)

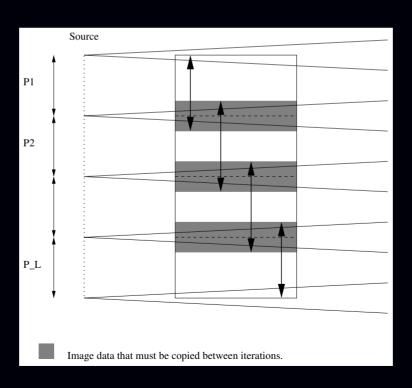
Ongoing saga...

(SPIE, ISBI, Fully 3D, ...) 43

Optimization algorithms

Challenges:

- theoretical convergence (to establish gold standards)
- practical: near convergence in few iterations
- highly parallelizable
- efficient use of hardware: memory bandwidth, cache, ...
- predictable stopping rules
- partitioning of helical CT data across multiple compute nodes

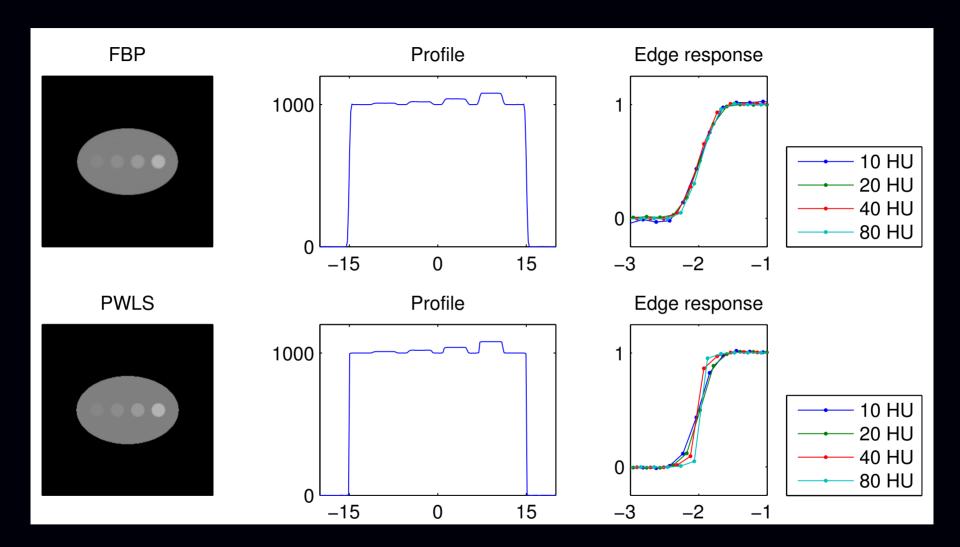


Example

(movie in pdf)

82-subset OS with two different (but similar) edge-preserving regularizers. One frame per every 10th iteration.

Resolution characterization: 2D CT



Challenge:

Shape of edge response depends on contrast for edge-preserving regularization.

Assessing image quality

Challenges:

- Resolution (PSF, edge response, MTF)
- Noise (predictions)
- Task-based performance measures
 Known-location versus unknown-location tasks

• ...

"How low can the dose go" - quite challenging to answer

Some open problems

- Modeling
 - Statistical modeling for very low-dose CT
 - Resolution effects
 - Spectral CT
 - Object motion
- Parameter selection / performance characterization
 - Performance prediction for nonquadratic regularization
 - Effect of nonquadratic regularization on detection tasks
 - Choice of regularization parameters for nonquadratic regularization
- Algorithms
 - o optimization algorithm design
 - software/hardware implementation
 - Moore's law alone will not suffice (dual energy, dual source, motion, dynamic, smaller voxels ...)
- Clinical evaluation
- ...

Current CT research in my group

Recent work

- Y. Long, J. A. Fessler, and J. M. Balter. 3D forward and back-projection for X-ray CT using separable footprints. *IEEE Trans. Med. Imag.*, 29(11):1839–50, Nov. 2010.
- Y. Lu, H-P. Chan, J. A. Fessler, L. Hadjiiski, J. Wei, M. Goodsitt, A. Schmitz, B. E. H. Claus, and F. W. Wheeler. Adaptive diffusion regularization for enhancement of microcalcifications in digital breast tomosynthesis (DBT) reconstruction. In *Proc. SPIE* 7961, 2011.
- Y. Lu, H-P. Chan, J. A. Fessler, L. Hadjiiski, J. Wei, M. Goodsitt, A. Schmitz, B. E. H. Claus, and F. W. Wheeler. Adaptive diffusion regularization for enhancement of microcalcifications in digital breast tomosynthesis (DBT) reconstruction. In *Proc. SPIE* 7961, 2011.
- W. Huh and J. A. Fessler. Iterative image reconstruction for dual-energy x-ray CT using regularized material sinogram estimates. ISBI 2011.
- D. Kim and J. A. Fessler. Accelerated ordered-subsets algorithm based on separable quadratic surrogates for regularized image reconstruction in X-ray CT. ISBI 2011.
- J-K. Kim, Z. Zhang, and J. A. Fessler. Hardware acceleration of iterative image reconstruction for X-ray computed tomography. ICASSP, 2011.

Forthcoming work

- J. A. Fessler and D. Kim. Axial block coordinate descent (ABCD) algorithm for X-ray CT image reconstruction. *Fully 3D*, 2011.
- S. Ramani and J. A. Fessler. Convergent iterative CT reconstruction with sparsity-based regularization. *Fully 3D*, 2011.
- M. Wu and J. A. Fessler. GPU acceleration of 3D forward and backward projection using separable footprints for X-ray CT image reconstruction. Fully 3D (workshop), 2011.

Work in progress

- Spectral CT from a single sinogram using bow-tie filter
- Motion-compensated cardiac CT reconstruction
- Noise predictions for iterative CT reconstruction
- Application to lung CT (NIH R01 with GE GRC)
 - lung nodule quantification
 - airway quantification
 - observer studies?