Model-based image reconstruction for low-dose X-ray CT

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- Kevin Brown, Philips
- Meng Wu, Stanford
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Statistical image reconstruction: a CT revolution

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



Thin-slice FBP

Seconds

A bit longer

ASIR

Statistical Much longer

(Same sinogram, so all at same dose)

Why statistical/iterative methods for CT?

- Accurate physics models
 - X-ray spectrum, beam-hardening, scatter, ...
 - \implies reduced artifacts? quantitative CT?
 - X-ray detector spatial response, focal spot size, ...
 - \implies improved spatial resolution?
 - \circ detector spectral response (*e.g.*, photon-counting detectors) \implies improved contrast?

Nonstandard geometries

- transaxial truncation (wide patients)
- long-object problem in helical CT
- irregular sampling in "next-generation" geometries
- coarse angular sampling in image-guidance applications
- limited angular range (tomosynthesis)
- o "missing" data, e.g., bad pixels in flat-panel systems
- Appropriate models of (data dependent) measurement statistics

 weighting reduces influence of photon-starved rays (*cf.* FBP)
 reducing image noise or X-ray dose

and more...

- Object constraints / priors
 - \circ nonnegativity
 - object support
 - \circ piecewise smoothness
 - object sparsity (*e.g.*, angiography)
 - sparsity in some basis
 - \circ motion models
 - \circ dynamic models

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Henry Gray, Anatomy of the Human Body, 1918, Fig. 413.

These constraints may help reduce image artifacts or noise or dose.

Disadvantages?

- Computation time (super computer)
- Must reconstruct entire FOV
- Complexity of models and software
- Algorithm nonlinearities
 - Difficult to analyze resolution/noise properties (*cf.* FBP)
 - Tuning parameters
 - Challenging to characterize performance / assess image quality

"Iterative" vs "Statistical"

- Traditional *successive substitutions* iterations
 - e.g., Joseph and Spital (JCAT, 1978) bone correction
 - usually only one or two "iterations"
 - not statistical
- Algebraic reconstruction methods
 - \circ Given sinogram data y and system model A, reconstruct object x by

"solving"
$$y = Ax$$

- ART, SIRT, SART, ...
- \circ iterative, but typically not statistical \Longrightarrow limited (if any) dose reduction
- Iterative filtered back-projection (FBP):



- Statistical reconstruction methods now come in several flavors:
 - Image domain
 - Sinogram domain
 - Fully statistical (both)
 - Hybrid methods (*e.g.*, AIR, SPIE 7961-18, 2011, Bruder *et al.*) Examine these in more detail next to consider dose.

"Statistical" methods: Image domain

Denoising methods

$$egin{array}{c} {
m sinogram} y &
ightarrow {
m FBP}
ightarrow {
m reconstruction}
ightarrow {
m iterative} {
m denoiser}
ightarrow {
m final} \ {
m denoiser}
ightarrow {
m image} {
m transform {
m sinogram} {
m sinogra$$

- 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)

Example (denoising by wavelet soft thresholding with spin cycling)



• Image denoising methods "guided by data statistics"



- Image-domain methods are fast (thus very practical)
 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

Several studies of potential dose reduction with image-domain methods:

- Hara et al., AJR 193(3) 2009, 32-65 % dose reduction
- Silva et al., AJR 194(1) 2010, 50-65 % dose reduction
- Leipsic et al., AJR 195(3) 2010, 27 % dose reduction

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"Statistical" methods: Sinogram domain

Sinogram restoration methods

noisy	adaptive	cleaned	final
sinogram \rightarrow	or iterative	\rightarrow sinogram \rightarrow FBP -	\rightarrow image
У	denoiser	ŷ	$\hat{\pmb{x}}$

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

Challenges:

- \circ Limited denoising without resolution loss \implies modest dose reduction?
- Difficult to "preserve edges" in sinograms





FBP, 10 mA FBP from denoised sinogram Wang *et al.*, T-MI, Oct. 2006, using PWLS-GS on sinogram

"Fully" (?) statistical image 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 measurement statistics fully
- Repeatedly updating the image can exploit object properties
- .: greatest potential dose reduction, but repetition is expensive...

Sub-mSv example

3D helical X-ray CT scan of abdomen/pelvis: 100 kVp, 25-38 mA, 0.4 second rotation, 0.625 mm slice, 0.6 mSv.



FBP

ASIR

Statistical

MBIR example: Routine chest CT

Helical chest CT study with dose = 0.09 mSv. Typical CXR effective dose is about 0.06 mSv. Source: Health Physics Society. http://www.hps.org/publicinformation/ate/q2372.html



FBP

MBIR

Veo (MBIR) images courtesy of Jiang Hsieh, GE Healthcare

History: Statistical reconstruction for PET

- Iterative method for emission tomography (Kuhl, 1963)
- FBP for PET
- Weighted least squares for 3D SPECT
- 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 (all?) 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)

(Chesler, 1971)

(Goitein, NIM, 1972)

Key factors in PET

ML-EM:

- 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





Llacer et al., 1993

FBP:

Whole-body PET example



FBP

ML-OSEM

Meikle et al., 1994

Key factor in PET: modeling measurement statistics

History: Statistical reconstruction for CT*

- Iterative method for X-ray CT
- 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)
- Ordered-subsets algorithms

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

(Sauer and Bouman, T-SP, 1993)

(Hounsfield, 1968)

circa 2010

Sep. 2011

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• ...
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- Commercial introduction of ICD for CT scanners
- FDA 510(k) clearance of Veo (Prof. William Shuman of UW quoted in GE's press release.)

(* numerous omissions, including many denoising methods) 18

RSNA 2010



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

https://engineering.purdue.edu/BME/AboutUs/News/HomepageFeatures/ResultsofPurdueResearchUnveiledatRSNA

Five Choices for Statistical Image 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.

Next: overview of each choice, focusing on how they may affect dose.

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

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

For a statistical approach to interior tomography, see Xu et al., IEEE T-MI, May 2011.

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
 - \circ finite detector size / detector spatial response
 - finite X-ray spot size / anode angulation
 - detector afterglow / gantry rotation
- spectral effects
 - X-ray source spectrum
 - \circ bowtie filters
 - detector spectra response
- scatter

• ...

Challenges / trade-offs

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

Detector size modeling matters

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):

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

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 and CT.

Least-squares solutions (for example):

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

Mismatched Backprojector $B \neq A'$

\hat{x} (PWLS-CG) \hat{x} (PWLS-CG)



Mismatched

Matched *cf.* SPECT/PET reconstruction – usually unregularized

X

Choice 3. Statistical Model

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

 $\overline{I_i} = I_0 e^{-\sum_{j=1}^N a_{ij} x_j}.$

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, particularly 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, 2007; Massoumzadeh *et al.*, PMB 2009
- 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):

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

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):

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

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 simpler linear model (ignoring beam hardening):

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

Drawbacks:

- Undefined if $I_i \leq 0$ (due to electronic noise); bad for very low-dose scans
- It is *biased* (by Jensen's inequality): $E[y_i] \ge -\log(\bar{I}_i/I_0) = [\mathbf{A}\mathbf{x}]_i$
- Exact distribution of log-domain noise ε_i is intractable.

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

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

- consider both Poisson and Gaussian noise effects: $\sigma_i^2 = \frac{\bar{I}_i + \sigma^2}{\bar{I}_i^2}$ (Thibault *et al.*, SPIE 6065, 2006)
- consider just Poisson effect: $\sigma_i^2 = rac{1}{\overline{I_i}}$
- 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, especially for very low-dose scans (*cf.* experience with PET rod scans).

(Sauer & Bouman, T-SP, Feb. 1993)

Choice 4. Cost Functions

Components:

- 4.1 *Data-mismatch* term
- 4.2 *Regularization* term (and regularization parameter β)
- 4.3 *Constraints* (*e.g.*, nonnegativity: minor effect on noise / dose)

Reconstruct image \hat{x} by finding minimizer of a cost function:

$$\hat{\boldsymbol{x}} \triangleq \underset{\substack{\boldsymbol{x} \geq \boldsymbol{0} \\ \text{Constraints}}}{\operatorname{constraints}} \Psi(\boldsymbol{x})$$

 $\Psi(\mathbf{x}) = \text{DataMismatch}(\mathbf{y}, \mathbf{A}\mathbf{x}) + \beta \text{Regularizer}(\mathbf{x})$

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."

Such optimization-based formulation appears to be key to low-dose CT. Theoretically optimizes bias-variance trade-off (Eldar, IEEE T-SP, Jul. 2004).

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:

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

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.

• ...

• Effect on dose is related to statistical model accuracy

Choice 4.2: Regularization

How to control noise due to ill-conditioning in tomography? Perhaps the most important factor influencing "how low (dose) can we go"

Noise-control methods (used in clinical 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

Regularization options

Options for regularizer $R(\mathbf{x})$ 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 immediate neighbors in 3D)
- Preserving "familiar" noise texture
- Optimizing clinical task performance

Many open questions...

Which regularization method can enable the lowest-dose scans?

Roughness Penalty Functions

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

N_j ≜ neighborhood of *j*th pixel (e.g., left, right, up, down in 2D; 26 neighbors in 3D)
ψ called the *potential function*



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

Regularization parameters: Dramatic effects on IQ

"q generalized gaussian" potential function with tuning parameters: β , δ , p, q: $\beta \psi(t) = \beta \frac{\frac{1}{2}|t|^p}{1+|t/\delta|^{p-q}}$. (Thibault *et al.*, Med. Phys., Nov. 2007)



p = q = 2 $p = 2, q = 1.2, \delta = 10 \text{ HU}$ p = q = 1.1

noise:	11.1	10.9	9	10.8
(#lp/cm):	4.2	7.2		8.2

Which parameter choice enables the lowest dose scan?

Piecewise constant phantoms



Phantom:

MLEM:

Lee *et al.*, IEEE T-NS, 2002, 300K counts non-convex "broken parabola" potential function and deterministic annealing

Summary of statistical image reconstruction choices

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

Reconstruction Method \triangleq **Models** + **Cost Function** + **Algorithm**

- 5. Minimization algorithms: $\hat{x} = \underset{x}{\operatorname{arg\,min}} \Psi(x)$
- The minimization algorithm has *no affect on image quality / dose if* we iterate "until convergence."
- $\circ\,$ For convergent algorithms, IQ determined entirely by cost function Ψ including the models within it.
- What happens we stop iterating "before convergence?" Affects dose? (Depends on algorithm and initial image...)
- Finding fast algorithms for arg min is key to ubiquitous low-dose CT. e.g., alternating direction method of multipliers (ADMM) (Ramani & Fessler, IEEE T-MI, Mar. 2012)

Summary: Factors that affect dose reduction

In conjectured order from most important to least:

- 1. Regularizer
- 2. Statistical model / data-mismatch term
- 3. System model
- 4. Object model / non-negativity constraint
- 5. Iterative minimization algorithm

Other considerations that affect dose reduction

- 2D versus 3D reconstruction
 - 8 immediate neighbors in 2D versus 26 neighbors in 3D
 - transaxial versus axial relationships
- axial versus helical scans
- angular sampling: fine versus sparse
- dual energy / spectral CT versus single kVp

Recent dose figures for statistical image reconstruction

Yadava *et al.*, AAPM 2010, AAPM 2011 "average-probability-of-detection and average-detection-score of full dose FBP and 1/4th dose Veo [MBIR] are comparable."

Cohen et al., AAPM 2011

[Phantom] "measurement differences between full dose FBP and 1/4th dose Veo were found to be statistically insignificant, indicating similar image quality."

Katsura *et al.*, Eur. Radiol. Aug. 2012 (hot off the press) Compared FBP, ASIR, and "MBIR" (with no almost description); concluded MBIR can work with "80% less radiation."

How low (dose) can you go?



http://laidoffinnyc.wordpress.com/2009/03/04/

- Depends on where you start
- Depends on many reconstruction method choices
- all of which affect IQ (which is harder to assess for nonlinear reconstruction methods)
- Conjecture: sub-mSv exams are achievable

Contrast-dependent edge resolution: 1D



Challenge: Shape of edge response depends on contrast when "preserving edges."

Contrast-dependent edge resolution: 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 in statistical image reconstruction

Modeling

- Statistical modeling for very low-dose CT
- Resolution effects
- Spectral CT
- Object motion
- Dynamic imaging (perfusion CT)
- Parameter selection / performance characterization
 - Performance prediction for nonquadratic regularization
 - Effect of nonquadratic regularization on detection tasks
 - Choice of regularization parameters for nonquadratic regularization

• Algorithms

- $\circ\,$ optimization algorithm design
- software/hardware implementation
- Moore's law alone will not suffice

(dual energy, dual source, motion, dynamic, smaller voxels ...)

Clinical evaluation

• ...

Many research opportunities to aid this CT revolution...

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