Outline

- SPECT Imager Design

 Imaging optics for gamma rays
 Design tradeoffs
 Calibration techniques
- Data Acquisition Systems
 - Architectures
 - Signals and conditioning Event detection
 - De-randomization
- Communication
 A survey of small-animal SPECT imagers



Modality Sensitivity and Resolution



SPECT Imager Properties and Main "Knobs"

Electronics

- Spatial Resolution Detectors, Optics Energy Resolution Detectors, Electronics
- Temporal Resolution
- · Sensitivity
- Count-rate Capability
- Calibration
- Synchronization

Physical Limits Detectors, Electronics Counting statistics **Optics**, **Detectors Real materials Detectors**, Electronics Complexity of underlying physics Detectors, Optics







Refraction...

- A. Snigirev, V. Kohn, I. Snigireva, B. Lengeler, "A compound refractive lens for focusing high energy x-rays," *Nature*, 384:49-51, 1996.
- B. Lengeler, C. Schroer, J. Tümmler, B. Benner, M. Richwin, A. Snigirev, I. Snigirev, M. Drakopoulos, "Imaging by parabolic refractive lenses in the hard x-ray range," *J. Synchrotom Rad.*, 6: 1153-1167, 1999.
- B. Cederström, R. N. Cahn, M. Danielsson, M. Lundqvist, D. R. Nygren, "Focusing hard x-rays with old LP's," *Nature*, 404: 951, 2000.



Reflection

- The redirection of light rays about the normal at interfaces between media of different n
- Occurs when no real solution to Snell's law
 - $\sin \theta_2 = (n_1/n_2) \sin \theta_1 \ge 1$
 - $\sin \theta_1 \ge n_2/n_1 \approx 1$ $\sin \theta_1 \approx 90^\circ$
- Problem is at gamma-ray energies for most materials:

Δ δ**≈ 10**-6

So gamma-ray mirrors are physically challenging...





Polycapillary Optics

- Hollow glass rods allow propagation of x-rays/y-rays via total internal reflection
- Tapered capillaries used for focusing in the synchrotron community
 Turned into a demonstration SPECT/CT instrument by Ritman et al





Multilayer Mirror Optics

- A single mirror has low collection efficiency
- NASA has pioneered multi-shell mirrors made up of arrays of metal foils
- Practical up to 35 keV
 - Above these energies, low reflectivity and finish errors limit efficiency
 - Long focal lengths (> 1 m)



Reflection/Diffraction

P.J. Serlemitsos and Y. Soong, "Foil x-ray mirrors," Astrophysics and Space Science, vol. 239, pp. 177-196, 1996

P.J. Serlemitsos, "Conical Foil X-ray Mirrors: Performance and Projections," Applied Optics, Vol. 27, pg. 1544, 1988

G. Hildebrandt and H. Bradazek "Approaching Real X-ray Optics," The Rigaku Journal, vol. 17(1), pp. 13-22, 2000.

M.J. Pivovaroff, W.C. Barber, F.E. Christensen, W.W. Craig, T. Decker, M. Epstein, T. Funk, C.J. Halley, B.H. Hasegawa, R. Hill, J.G. Jernigan, C. Taylor, and K. Ziock, "Small Animal Radionucidie Imaging with Focusing Gamma-ray Optics", Proc. SPIE, Vol. 5199, pp. 147-161, 2004.

M.J. Pivovaroff, T. Funk, W.C. Barber, B.D. Ramsey, B.H. Hasegawa, "Progress of focusing x-ray and gamma-ray optics for small animal imaging", Proc. SPIE, Vol. 5923, pp. 65-78, 2005.



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|---|--|---|---------------------------|----------------------------------|--|
| Element | Z | Absorption Coeff (cm ⁻¹) | Absorption Length (mm) | Transmittance (1/8" material) | |
| H₂O | 8 | .15 | 67 | 95/100 | |
| Pb | 82 | 26.823 | .373 | 1/5,000 | |
| W | 74 | 36.333 | .275 | 1/100,000 | |
| Au | 79 | 42.629 | .235 | 1/1,000,000 | |
| - | 70 | 42 921 | 228 | 1/1 000 000 | |



Absorption

Other techniques



- Scanning slit and camera



S.D. Metzler, R. Accorsi, A.S. Ayan, and R.J. Jaszczak "Silt-Slat and Multi-Silt-Slat Collimator Design and Experimentally Acquired Phantom Images" From a Rotating Prototype IEEE Trans. Nucl. Sci., 57(1), 125-134, 2010

G.L. Zeng and D. Gagnon "CdZnTe strip detector SPECT imaging with a slit collimator" Phys. Med. Biol., 49, 2257–2271, 2004.

S. Walrand, F. Jamar, M. de Jong, and S. Pauwels "Evaluation of Novel Whole-Body High-Resolution Rodent SPECT (Linoview) Based on Direct Acquisition of Linogram Projections" *J. Nucl. Med.*, 46,1872–1860, 2005.

P. Edholm, G.T. Herman, and D.A. Roberts "Image Reconstruction from Linograms : Implementation and Evaluation" IEEE Trans. Med. Imag., 7 (3), 239-246, 1988.



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Multiple Pinholes: To Multiplex or Not to Multiplex

- Definitely don't want pinhole arrangement symmetric with respect to rotation axis
 Mutiple lobes in PSF can give some reconstruction artifacts false lesions
 More iterations of reconstruction algorithm typically required
- Benefit depends on object and tracer distribution



S.R. Meikle, P. Kench, A.G. Weisenberger, R. Wojcik, M.F. Smith, S. Majews S. Eberl, R.R. Fulton, A.B. Rosenfeld, and M.J. Fulham, "A prototype coded aperture detector for small animal SPECT" IEEE NSS/MIC Cont. Rec. vols 1–4, pp 1580–4, 2002. Center ter Gamma-Ray Imir



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Adaptive, Anamorphic Projection

- Crossed-slit collimator decouples axial and transaxial
- magnifications
 Make full use of detector area regardless of object shape



Pinhole Clusters

Replace a single pinhole with an array of smaller pinholes
 Keep angles shallower to reduce keel edge penetration
 Good for higher energies (511 keV)





Absorption

- Collimators 4 basic flavors
- We use for our solid-state detectors:
 - Parallel hole
 - Laminated photoetched W
 - 100 layers 7 mm thick
 380 µm bore spacing, 120 µm septa

 - Efficiency = 5 × 10⁻⁵
 Manufactured to our spec by Tecomet of Woburn, MA



www.harpellassociates.com



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Absorption

There is ample literature on the design and analysis of single and multiple pinhole systems. Good entry points are the following papers and the literature they cite.

F.P. DiFilippo, "Design and performance of a multi-pinhole collimation device for small animal imaging with clinical SPECT and SPECT-CT scanners" Phys. Med. Biol., 53, 4185-4201, 2008.

S.R. Meikle, P. Kench, M. Kassiou, and R. B. Banati, "Small animal SPECT and its place in the matrix of molecular imaging technologies" *Phys. Med. Biol.*, 50, R45-61, 2005.

R. Accorsi and S. D. Metzler, *Analytic Determination of the Resolution-Equivalent Effective Diameter of a Pinhole Collimator* *IEEE Trans. Med. Imag.*, 23 (6), 750-763, 2004.

N. U. Schramm, G. Ebel, U. Engeland, T. Schurrat, M. Béhé, and T. M. Behr, "High-Resolution SPECT Using Multi-pinhole Collimation" *IEEE Trans. Nuc. Sci.*, 50(3), 315-320, 2003.

R.J. Jaszczak, J. Li, H. Wang, M.R. Zulutsky, R.E. Coleman "Pinhole collimation for ultra-high-resolution small-field-of-view SPECT," *Phys. Med. Biol.*, 39, 425-437, 1994.



Conterner To Conterner Ray Insigning













Key Points

 Magnification: as large as the required field-of-view and camera size permits.

Obliquity: if the object to pinhole distance is very short, then vignetting by the pinhole and depth-of-interaction effects become problematic at the edges of the detector.

Design needs to consider the imaging experiment to be performed – and there will in general be a need for different pinhole sizes and locations for different imaging tasks.



One (or a few) Camera Design Options



- Simplest arrangement

Abnormal position for mouse
 Measured or modeled PSF must be rotated during reconstruction

• axis Pros:

Cons:



Stationary mouse

- Camera rotates about horizontal axis
- Pros:
- Mouse in normal position Cons:
- High precision motion required with possibly heavy camera(s)
 PSF probably needs to be modeled



Multi-Camera or Annular Camera Design Options Stationary cameras • Pros: House in normal position High sensitivity All data acquisition in parallel – dynamic capability PSF can be accurately measured Cons: Imager size and complexity C Inter In Essente Ray Integing

Commercial Pre-Clinical Imager Examples







Mediso/Bioscan NanoSPECT/CT

Gamma Medica FLEX Triumph

Siemens Inveon

- Scintillation detectors pixelated or monolithic
- Solid-state detectors CZT

USPECT

- Resolution from magnification
- Sensitivity from multiple pinholes

CareStream Albira



Summary of SPECT Imager Design Decisions

- Camera type
 - Technology, size, number of resolvable elements, sensitivity, energy resolution, count-rate capability
- Number of cameras
 - Acquisition time
- · What, if anything, moves and how
- Aperture type
 - Collimator or pinhole(s) or other
- · Sensitivity, magnification, and field of view
- Next:
 - Calibration - Acquisition



Calibration

Directly measure camera and imager response.

Calibrates optical performance and corrects for mechanical tolerances, electronic variations, and detector material imperfections.





Exhaustive: scan point source in regular 3-D grid throughout object volume

Parametric: scan to tune modeling/compensation parameters





Scan collimated source in regular 2-D grid across every camera face

- Shadow grid



Calibration Stages



Rotation and secondary translation makes it easy to scan the source precisely above each camera face.









Calibration References

F. van der Have, B. Vastenhouw, M. Rentmeester, and F.J. Beekman "System Calibration and Statistical Image Reconstruction for Ultra-High Resolution Stationary Prinhole SPECT" IEEE Trans. Med. Imag., 27(7), 960-971, 2008

Y-C. Chen, L.R. Furenlid, D.W. Wilson, and H.H. Barrett, 'Calibration of Scintillation Cameras and Pinhole SPECT Imaging Systems' Chap. 12 in Small-Animal SPECT Imaging, New York: Springer Science Business Media, 195–201, 2005.

And references therein.



Conter In Estimate Ray Intéging

Trends

• Data processing and acquisition benefit from the rapid pace of technology development in:

- ► Fast, low-noise op amps and A/D converters
 - Readout ASICS
 - GPUs and FPGAs
 - Networking and communications
 - > Computing power and storage capacities



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Asynchronous Solid-State Detectors

Integrating Readouts

Synchronous (clocked) devices

- Gated-integrator readouts
- CZT pixel arrays
- Charge storing detectors





B.W. Miller, H.B. Barber, H.H. Barrett, I. Shestakova, B. Singh, V.V. Nagarkar, "Single-photon spatial and energy resolution enhancement of a columnar CsI(TI)/EMCCD gamma-camera using maximum likelihood estimation *Proc.* SPIE, 2006.

G A. de Vree, A H.Westra, I. Moody, F. van der Have, K.M. Ligtvoet, and F.J. Beekman, "Photon-Counting Gamma Camera Based on an Electron-Multiplying CCD" *IEEE Trans. Nucl. Sci.*, 52(3), 580-588, 2005.



Gated Integrators

- Applicable to all kinds of detectors
- Convenient for big arrays and implementation in ASICs
- However, integrate leakage current
- kT/C noise from the reset
- Work around CDSH







Pixel Array Detectors

- 2-D array scanned and processed externally
- Can read entire raster into buffer and scan with DSP/CPU
- Double buffer to prevent loss of data
- Better: on the fly detection with gate array
 - •Need data from 2-D sub-pixel area
 - Pipeline processing

>Rastered readout must be run through multi-tap shift register to access pixels from adjacent rows



Data-Acquisition Architectures

- List-mode: raw data is maintained as an ordered list • - benefits

 - Can apply new statistical algorithms to any data
 No information loss
 Can reconstruct on different attributes image, fluence, raw list
 Can add physiological signals as entries in list
 - drawbacks
 - Data lists occupy large amounts of memory
- · Image mode: data is processed and stored as a bitmap benefits

 - Data have fixed size, typically modest
 Data can quickly be visualized
 - drawbacks
 - Original observations not available for further processing









FastSPECT II Imager

Key Features:

• 16 cameras in 2 rings of 8 with adjustable radial position

 5 axis robotic stage for calibration and imaging subject positioning
 Exchangeable

cylindrical imaging apertures for choice of magnification/field-ofview



L. R. Furenlid, D. W. Wilson, Y. Chen, H. Kim, P. J. Pietraski, M. J. Crawford, and H. H. Barrett, "FastSPECT II: A Second-Generation High-Resolution Dynamic SPECT Imager", IEEE Trans. Nucl. Sci., 51(3), 631-635, 2004,



Modular Apertures

· Gold insert pinholes - currently 3 sizes: .1, .5, and 1 mm dia • 1/2" Pb cylinder with cast cerrobend end J cap Imaging geometry has pinholes located on lines between camera centers and imager center point $\frac{1}{4}$ Efficiency ~ 4 × 10⁻ Center In Samo Ray Integing

Fast Position Estimation



Raw list-mode data

- Cameras fully calibrated with exhaustive scanning of highly collimated source
- Maximum-likelihood
 estimation of gamma-ray interaction position and energy.

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Very Efficient ML Search Algorithm



w -

computing, pipelining and implementation in GPU or gate array



Resolution Phantom Image on FSII



1.5 mm diameter w/ 4.5 mm center to center distance
 2.0 mm diameter w/ 6 mm center to center distance



Volume Rendering

Center In Canada Ray Insigning













FSIII – Front End LM Processor

Processes events from 4 simultaneous data streams each with up to 200 640 ×480 frames per second with 7.4 μm × 7.4 μm pixels.

Frame Processing of all cameras in real time at 200 fps = 4000 fps to give ~1.23 Gpix/s

Easily handled with GPU programming



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FastSPECT III Imaging Aperture



A Survey of Systems: Triple-Head System



S. D. Metzler, R. J. Jaszczak, N. H. Patil, S. Vemulapalli, G. Akabani, and B. B. Chin "Molecular Imaging of Small Animals With a Triple-Head SPECT System Using Pinhole Collimation" *IEEE Trans. Med. Imag.*, 24(7), 853-862, 2005.

Overlapping Multipinhole System

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N. U. Schramm, G. Ebel, U. Engeland, T. Schurrat, M. Béhe, and T. M. Behr, "High-Resolution SPECT Using Multiprinhole Collimation" *IEEE Trans. Nuc. Sci.*, 50(3), 315-320, 2003.







Caster to Canno Ray Indiging

FJ, Beekman, F, van der Have, B. Vastenhouw, A.J.A. van der Linden, P.P. van Rijk, J.P.H. Burbach, and M.P. Smidt, "U-SPECT-14. Novel System for Submillimeter-Resolution Tomography with Radiolabeled Molecules in Mice" J. Nucl. Med., 48(7), 1194–1200, 2005.

U-SPECT

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Conclusions

- On-going rapid developments: imaging principles, detectors, systems and computing resources (GPUs)
- List-mode architectures have advantages, including ability to support new camera technologies without having to reinvent software and data links
- Careful calibration critical for optimal system performance
- Stationary imager designs permit dynamic and high-throughput studies
- System design process can/should be thoroughly guided by simulations with realistic digital phantoms.



Path to Higher Resolution Small-Animal SPECT

- 2-3 mm intrinsic-resolution Anger camera + 1-2 mm bore parallel-hole collimator
 + filtered back projection reconstruction
- Move to pinhole aperture with magnification
- Develop analytical forward model and switch to statistical reconstruction (ML-EM, etc...)
- Add additional pinholes and cameras to acquire projections in parallel
- Calibrate system for more accurate forward model measured H matrix
- Make pinholes smaller and move closer to object, trading FOV for magnification
- Improve camera intrinsic resolution, add DOI to reduce parallax errors
 Gate acquisition to reduce respiratory and cardiac motion
- Eliminate events with non-local energy deposition in detector
- Combine with high resolution anatomical modality
- Incorporate scatter and attenuation in object-specific H matrix
- Reconstruct in photon-by-photon list-mode form

