## **Part 1: From Physics to Statistics**

or "What quantity is reconstructed?" (in emission tomography)

#### Outline

- Decay phenomena and fundamental assumptions
- Idealized detectors
- Random phenomena
- Poisson measurement statistics
- State emission tomography reconstruction problem

## What Object is Reconstructed?

In *emission imaging*, our aim is to image the *radiotracer distribution*.

#### The what?

At time t = 0 we inject the patient with some *radiotracer*, containing a "large" number N of metastable atoms of some radionuclide.

Let  $\vec{X}_k(t) \in \mathbb{R}^3$  denote the position of the *k*th *tracer atom* at time *t*. These positions are influenced by blood flow, patient physiology, and other unpredictable phenomena such as Brownian motion.

The ultimate imaging device would provide an exact list of the spatial locations  $\vec{X}_1(t), \ldots, \vec{X}_N(t)$  of all tracer atoms for the entire scan.

Would this be enough?

## **Atom Positions or Statistical Distribution?**

Repeating a scan would yield different tracer atom sample paths  $\left\{ \vec{X}_k(t) \right\}_{k=1}^{N}$ .

: statistical formulation

**Assumption 1.** The spatial locations of individual tracer atoms at any time  $t \ge 0$  are *independent* random variables that are all *identically distributed* according to a common probability density function (pdf)  $f_{\vec{X}(t)}(\vec{x})$ .

This pdf is determined by patient physiology and tracer properties.

Larger values of  $f_{\vec{X}(t)}(\vec{x})$  correspond to "hot spots" where the tracer atoms tend to be located at time *t*. Units: inverse volume, *e.g.*, atoms per cubic centimeter.

The *radiotracer distribution*  $f_{\vec{X}(t)}(\vec{x})$  is the quantity of interest.

(Not 
$$\left\{ \vec{X}_k(t) \right\}_{k=1}^N$$
!)

### **Example: Perfect Detector**



True radiotracer distribution  $f_{\vec{X}(t)}(\vec{x})$  at some time *t*.

A realization of N = 2000 i.i.d. atom positions (dots) recorded "exactly."

Little similarity!

# **Binning/Histogram Density Estimator**



Estimate of  $f_{\vec{X}(t)}(\vec{x})$  formed by histogram binning of N = 2000 points. Ramp remains difficult to visualize.

### **Kernel Density Estimator**



Gaussian kernel density estimator for  $f_{\vec{X}(t)}(\vec{x})$  from N = 2000 points.



Horizontal profiles at  $x_2 = 3$  through density estimates.

#### **Poisson Spatial Point Process**

Assumption 2. The number of injected tracer atoms N has a Poisson distribution with some mean

$$u_N \stackrel{\triangle}{=} E[N] = \sum_{n=0}^{\infty} nP[N=n].$$

Let N(B) denote the number of tracer atoms that have spatial locations in any set  $B \subset \mathbb{R}^3$  (VOI) at time  $t_0$  after injection.

 $N(\cdot)$  is called a *Poisson spatial point process*.

Fact. For any set B, N(B) is Poisson distributed with mean:

$$E[N(B)] = E[N]P[\vec{X} \in B] = \mu_N \int_B f_{\vec{X}(t_0)}(\vec{x}) \, \mathrm{d}\vec{x}.$$

Poisson *N* injected atoms + i.i.d. locations  $\Rightarrow$  Poisson point process

## Illustration of Point Process ( $\mu_N = 200$ )



### **Radionuclide Decay**

Preceding quantities are all unobservable. We "observe" a tracer atom only when it decays and emits photon(s).

The time that the *k*th tracer atom decays is a random variable  $T_k$ .

Assumption 3. The  $T_k$ 's are statistically *independent* random variables, and are independent of the (random) spatial location.

Assumption 4. Each  $T_k$  has an exponential distribution with mean  $\mu_T = t_{1/2}/\ln 2$ .

Cumulative distribution function:  $P[T_k \le t] = 1 - \exp(-t/\mu_T)$ 



#### **Statistics of an Ideal Decay Counter**

Let K(t, B) denote the number of tracer atoms that decay by time t, and that were located in the VOI  $B \subset \mathbb{R}^3$  at the time of decay.

Fact. K(t, B) is a *Poisson counting process* with mean

$$E[K(t,B)] = \int_0^t \int_B \lambda(\vec{x},\tau) \, \mathrm{d}\vec{x} \, \mathrm{d}\tau,$$

where the (nonuniform) emission rate density is given by

$$\lambda(\vec{x},t) \stackrel{ riangle}{=} \mu_N rac{e^{-t/\mu_T}}{\mu_T} \cdot f_{\vec{X}(t)}(\vec{x}).$$

Ingredients: "dose," "decay," "distribution"

Units: "counts" per unit time per unit volume, *e.g.*,  $\mu$ Ci/cc.

"Photon emission is a Poisson process"

What about the actual measurement statistics?

#### **Idealized Detector Units**

A nuclear imaging system consists of  $n_d$  conceptual *detector units*.

Assumption 5. Each decay of a tracer atom produces a recorded count in at most one detector unit.

Let  $S_k \in \{0, 1, ..., n_d\}$  denote the index of the incremented detector unit for decay of *k*th tracer atom. ( $S_k = 0$  if decay is undetected.)

**Assumption 6.** The  $S_k$ 's satisfy the following conditional independence:

$$P\left(S_1,\ldots,S_N | N, T_1,\ldots,T_N, \vec{X}_1(\cdot),\ldots,\vec{X}_N(\cdot)\right) = \prod_{k=1}^N P\left(S_k | \vec{X}_k(T_k)\right).$$

The recorded bin for the *k*th tracer atom's decay depends only on its position when it decays, and is independent of all other tracer atoms.

(No event pileup; no deadtime losses.)

# **PET Example**



$$n_d \leq (n_{\text{crystals}} - 1) \cdot n_{\text{crystals}}/2$$

### **SPECT Example**



 $n_d = n_{\text{radial\_bins}} \cdot n_{\text{angular\_steps}}$ 

### **Detector Unit Sensitivity Patterns**

**Spatial localization:** 

 $s_i(\vec{x}) \stackrel{\triangle}{=}$  probability that decay at  $\vec{x}$  is recorded by *i*th detector unit.

### Idealized Example. Shift-invariant PSF: $s_i(\vec{x}) = h(\vec{k}_i \cdot \vec{x} - r_i)$

- $r_i$  is the radial position of *i*th ray
- $\vec{k}_i$  is the unit vector orthogonal to *i*th parallel ray
- $h(\cdot)$  is the shift-invariant radial PSF (*e.g.*, Gaussian bell or rectangular function)



### **Example: SPECT Detector-Unit Sensitivity Patterns**



Two representative  $s_i(\vec{x})$  functions for a collimated Anger camera.

### **Example: PET Detector-Unit Sensitivity Patterns**



## **Detector Unit Sensitivity Patterns**

 $s_i(\vec{x})$  can include the effects of

- geometry / solid angle
- collimation
- scatter
- attenuation
- detector response / scan geometry
- duty cycle (dwell time at each angle)
- detector efficiency
- positron range, noncollinearity
- ...

System sensitivity pattern:

$$s(\vec{x}) \stackrel{\triangle}{=} \sum_{i=1}^{n_d} s_i(\vec{x}) = 1 - s_0(\vec{x}) \le 1$$

(probability that decay at location  $\vec{x}$  will be detected at all by system)

### **System Sensitivity Pattern** $s(\vec{x})$



Example: collimated 180° SPECT system with uniform attenuation.

# **Detection Probabilities** $s_i(\vec{x}_0)$ (vs det. unit index *i*)



## **Summary of Random Phenomena**

- Number of tracer atoms injected N
- Spatial locations of tracer atoms  $\{\vec{X}_k\}_{k=1}^N$
- Time of decay of tracer atoms  $\{T_k\}_{k=1}^N$
- Detection of photon  $[S_k \neq 0]$
- Recording detector unit  $\{S_k\}_{i=1}^{n_d}$

### **Emission Scan**

Record events in each detector unit for  $t_1 \le t \le t_2$ .

 $Y_i \stackrel{\triangle}{=}$  number of events recorded by *i*th detector unit during scan, for  $i = 1, ..., n_d$ .  $Y_i \stackrel{\triangle}{=} \sum_{k=1}^N \mathbf{1}_{\{S_k = i, T_k \in [t_1, t_2]\}}$ .

The collection  $\{Y_i : i = 1, ..., n_d\}$  is our *sinogram*.

Note  $0 \leq Y_i \leq N$ .

Fact. Under Assumptions 1-6 above,

$$Y_i \sim \text{Poisson}\left\{\int s_i(\vec{x})\lambda(\vec{x})\,\mathrm{d}\vec{x}\right\}$$
 (cf "line integral")

and  $Y_i$ 's are statistically independent random variables, where the *emission density* is given by

$$\lambda(\vec{x}) = \mu_N \int_{t_1}^{t_2} \frac{1}{\mu_T} e^{-t/\mu_T} f_{\vec{X}(t)}(\vec{x}) \, \mathrm{d}t.$$

(Local number of decays per unit volume during scan.)

Ingredients:

- dose (injected)
- duration of scan
- decay of radionuclide
- distribution of radiotracer

## **Poisson Statistical Model (Emission)**

Actual measured counts = "foreground" counts + "background" counts.

Sources of background counts:

- cosmic radiation / room background
- random coincidences (PET)
- scatter not account for in  $s_i(\vec{x})$
- "crosstalk" from transmission sources in simultaneous T/E scans
- anything else not accounted for by  $\int s_i(\vec{x})\lambda(\vec{x}) d\vec{x}$

#### Assumption 7.

The background counts also have independent Poisson distributions.

Statistical model (continuous to discrete)

$$Y_i \sim ext{Poisson}\left\{\int s_i(\vec{x})\lambda(\vec{x})\,\mathrm{d}\vec{x}+r_i
ight\}, \qquad i=1,\ldots,n_d$$

 $r_i$ : mean number of "background" counts recorded by *i*th detector unit.

## **Emission Reconstruction Problem**

Estimate the emission density  $\lambda(\cdot)$  using (something like) this model:

$$Y_i \sim \text{Poisson}\left\{\int s_i(\vec{x})\lambda(\vec{x})\,\mathrm{d}\vec{x}+r_i\right\}, \qquad i=1,\ldots,n_d.$$

Knowns:

- $\{Y_i = y_i\}_{i=1}^{n_d}$ : observed counts from each detector unit
- $s_i(\vec{x})$  sensitivity patterns (determined by system models)
- *r*<sub>*i*</sub>'s : background contributions (determined separately)

Unknown:  $\lambda(\vec{x})$ 

### **List-mode acquisitions**

Recall that conventional sinogram is temporally binned:

$$Y_i \stackrel{\triangle}{=} \sum_{k=1}^N \mathbf{1}_{\{S_k=i, T_k \in [t_1, t_2]\}}.$$

This binning discards temporal information.

List-mode measurements: record all (detector,time) pairs in a list, *i.e.*,

$$\{(S_k,T_k): k=1,\ldots,N\}.$$

List-mode dynamic reconstruction problem:

Estimate  $\lambda(\vec{x},t)$  given  $\{(S_k,T_k)\}$ .

# **Emission Reconstruction Problem - Illustration**

 $\lambda(\vec{x})$ 



 $\{Y_i\}$ 



 $x_1$ 

r

### **Example: MRI "Sensitivity Pattern"**



Each "k-space sample" corresponds to a sinusoidal pattern weighted by:

- RF receive coil sensitivity pattern
- phase effects of field inhomogeneity
- spin relaxation effects.

$$y_i = \int f(\vec{x}) c_{\rm RF}(\vec{x}) \exp(-\iota \omega(\vec{x})t_i) \exp(-t_i/T_2(\vec{x})) \exp\left(-\iota 2\pi \vec{k}(t_i) \cdot \vec{x}\right) d\vec{x} + \varepsilon_i$$

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