



Imaging using ionizing radiations

Positron Emission Tomography

Ziad El Bitar <u>ziad.elbitar@iphc.cnrs.fr</u> Institut Pluridisciplinaire Hubert Curien, Strasbourg

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Slides are courtesy of David Brasse



PET instrument

Positron mean range



Acolinearity

This effect depends on the energy of the positon

Angular distribution is almost gaussian With an LTMH around 0,5°

 $R_{AC} = 0,0022 \times D$

D = System diameter



\Box β^+ in a magnetic field

Objective : Limitation of the positon range (flight) in presence of an axial magnetic field – Combination PET/IRM

- Resolution phantom: Ø 1 mm 2,5 mm
- Simulation of a high resolution animal PET system : 1.8 mm

¹⁸F *B* = 0 Tesla ¹⁵O B = 0 Tesla ¹⁵O B = 15 Tesla







S. Jan, SHFJ - CEA

Physical principles: short remindr

Question:

Calculate the minimal energy of a 511 keV photon after undergoing a Compton scatter. What is the energy of the recoil electron ?

$$E_c(keV) = \frac{E_0}{1 + \frac{E_0}{m_e c^2} (1 - \cos\theta)}$$

The minimum energy is obtained when the maximum of energy is transferred to the electron. This happens for $\theta = 180^{\circ}$

$$E_c(keV) = \frac{511}{2 - \cos\theta} \qquad E_e(keV) = E - E_c = 511\frac{1 - \cos\theta}{2 - \cos\theta}$$

Numerical application: find the energy of the scattered photon and the recoil electron

$$E_c(keV) = \frac{511}{2 - \cos 180^\circ} = 170keV$$

$$E_e = E - E_c = 511 - 170 = 340 \text{keV}$$

Interaction cross section

$$I(x) = I(0) \exp(-\mu x)$$

$\mu \approx \mu_{compton} + \mu_{photoelectric}$

@ 511 keV

	$\mu_{Compton} \ (\mathrm{cm}^{-1})$	$\mu_{Photoelectric}\ ({ m cm}^{-1})$	μ (cm ⁻¹)
Soft tissues	~0,096	~ 0,00002	~ 0,096
Bone	~ 0,169	~ 0,001	~ 0,17
BGO	0,51	0,40	0,96
Lead	0,76	0,89	1,78
Tungsten	1,31	1,09	2,59

Question:

What is the probability that a 511 keV photon undergoes a Compton scatter @ 7.5 cm inside the brain?

What would be the probability value for a photon coming from the liver @ 20 cm from the body surface?

$$\frac{I(x)}{I(0)} = \exp(-\mu x) \qquad \mu_{Compton} = 0,096 \text{ cm}^{-1}$$

Brain: x = 7,5 cm; I(x)/I(0) = 0,49 (49 % of photons are unscattered) Liver: x = 20 cm; I(x)/I(0) = 0,15 (15 % of photons are unscattered)



The objective of the detection system is to detect photons coming from the body and that did not scatter.

Need for a dense material Optimize the rate of photolectric/compton cross sections.

Question:

Calculate the thickness of BGO required to stop 90% of 511 keV photons.

$$\frac{I(x)}{I(0)} = 0,1 = \exp(-0.96 \times x)$$

x = 2,4cm

Used scintillators

Scintillators (Density g/cm ³)	Yield (ph/511keV)	Decay (ns)	Refraction index	μ @511 keV (cm ⁻¹)	μ _{Ph} /μ _C @511 keV
Nal:TI BGO LSO:Ce GSO:Ce Ba F_2 YAP:Ce LaB r_3 LaC I_3 LuAP LYSO	3,67 7,13 7,40 6,71 4,89 5,37 5,29 3,86 8,34 7,11	19400 4200 ~ 13000 ~ 4600 700; 4900 ~ 9200 32000 23000 5110 17300	230 300 ~ 47 ~ 56 0,6; 630 ~ 27 16 25 18 41	1,85 2,15 1,82 1,85 1,56 1,95 1,9 1,9 1,9	0,34 0,96 0,88 0,70 0,45 0,46 0,45 0,36 0,95 0,83	0,22 0,78 0,52 0,35 0,24 0,05

Used photodetectors

Photomultiplier HPD MCP

Solid detector Photodiode Avalanche photodiode

Important parameters to consider:

Quantum efficiency Gain Signal to noise ratio Speed Geometry Cost/readout channel

Photodetector/Crystal coupling

« Block detector »



$$= \frac{(D + B) - (C + A)}{\Sigma}$$
$$= \frac{(A + B) - (C + D)}{\Sigma}$$



Photodetector/Crystal coupling

Extension of « Block detector »: « Quadrant sharing »





4 PMT for 64 pixels Pixel size / 2 Cost reducing

Adapted for large detectors Dead zones edges

Photodetector/Crystal coupling

Continuous detector

Large FOV: 40-50 cm Dead zone at edges

Curved detector possible



Barycenter calculation of the interaction position The spatial resolution depends on the number of detected photons-> Nal:TI etc...

25 mm = Compromise between resolution and efficiency

Thickness	R _{intrinsèque}
10 mm	3 mm
25 mm	4-5 mm

Should have the capability of generating many events, unless increasing the dead time ...



System configuration

Detection in coïncidence The time resolution of the system depends on the constant decay of the used crystal used electronic readout Typically $\tau = 5-6$ ns for BGO-NaI:TI $\tau = 2-3$ ns for LSO



Different registered events



Resolution: Response function

The intrinsic detector resolution can be derived in 2 components

Geometry (Considers a perfect detector) Physics (Physical properties of the detector)



Parallax problem



Two important consequences Broadening of the response (Assymetry) False positionning

The resolution of a detector system the quadratic sum of the « elementary » resolutions

Positron mean range Acolinearity Geometrical factors Intrinsic spatial resolution (continuous detector) Physical factors

Question:

Hypothesis: Different responses can be approximated by gaussian functions

Calculate the system resolution for a ¹⁸F source centred in the field of view of a clinical scanner which diameter is 80 cm and the size of a detection element is 6 mm

 $R_i = 6mm / 2 = 3mm$ $R_{AC} = 0,0022 \times 800 = 1,76mm$ $R_{positron} = 0,102mm$

 $R_{système} = \sqrt{3^2 + 1,76^2 + 0,102^2} = 3,48mm$

Detection efficiency

The number of registered events is given by

The amount of injected radioactivity Fraction of radioactivity targeting the Region Of Interest (ROI) Duration of the exam Detection efficiency of the PET system

> Unit Cps/Bq/ml

The system efficiency if the product of several factors

Detector efficiency @ 511 keV Detection solid angle Source positionning with respect to detector The width of the energy window The width of the time window

Detection efficiency

The detection efficiency of an elementary detector is the product of

The probability to detect an incident photon Times

The fraction of events selected in the energy window $\mathcal{E} = \left(1 - e^{-\mu d}\right) \times \Phi$

 μ = linear attenuation coefficient of the material d = material thickness

Detection efficiency in coïncidence

$$\varepsilon = \left(1 - e^{-\mu d}\right)^2 \times \Phi^2$$

System geometrical efficiency

The product of

Detection solid angle covered by the detector for a given source position (Ω) Filling rate (Materials volume / total volume) (φ)

$$\Omega = 4\pi \sin \left[\tan^{-1} (A/D) \right]$$

Point source centred in a circular system With diameter D And axial coverage A

 $\phi = \frac{\text{section} \times \text{height}}{\left(\text{section} + \text{deadzone}\right) \times \left(\text{height} + \text{deadzone}\right)}$

Detection efficiency of the system

$$\eta \approx 100 \times \frac{\varepsilon^2 \varphi \Omega}{4\pi}$$

Question:

Calculate the detection efficiency of a point source centred in the field of view of the system. TEP: simple detector ring of 80 cm diameter composed of detector elements whom dimensions are $4,9 \times 6 \times 30 \text{ mm}^3$.

Reflector size: 0,25 mm

80% of the events are selected by the energy window

$$\varepsilon = (1 - e^{-(0.96 \times 3)}) \times 0.8 = 0.755$$

$$\Omega = 4\pi \sin[\tan^{-1}(0.6/80)] = 0.094$$

$$\varphi = (4.4 \times 5.5)/(4.9 \times 6) = 0.823$$

$$\eta = 100 \times 0.755^{2} \times 0.094 \times 0.823/4\pi = 0.35\%$$

Data representation



 $r = x\cos\phi + y\sin\phi$





Acquisition 2D



Direct slices

Crossed Slices

Acquisition 3D





[11C]flumazenil images of benzodiazapene receptor distribution



2D PET

3D PET

Question:

Calculate the detection efficiency of a point source centred in the field of view of the system. TEP: 16 detector ring of 80 cm diameter composed of 4,9 x 6 x 30 mm³ detector elements. Reflector size: 0,25 mm

80% of the events are selected by the energy window.

3D Acquisition mode.

 $\varepsilon = (1 - e^{-(0.96 \times 3)}) \times 0.8 = 0.755$ unchanged $\Omega = 4\pi \sin \left[\tan^{-1} (0.6 \times 16/80) \right] = 1.50$ $\phi = (4.4 \times 5.5) / (4.9 \times 6) = 0.823$ uchanged $\eta = 100 \times 0.755^2 \times 1.50 \times 0.673 / 4\pi = 5.59\%$ To compare with 0.35 %

Question:

BGO -> LSO

$$\varepsilon = (1 - e^{-(0,88\times3)}) \times 0,9 = 0,836$$

$$\Omega = 4\pi \sin[\tan^{-1}(0,6\times16/80)] = 1,50 \quad \text{unchanged}$$

$$\varphi = (4,4\times5,5)/(4,9\times6) = 0,823 \quad \text{unchanged}$$

$$\eta = 100 \times 0,836^2 \times 1,50 \times 0,673/4\pi = 6,19\% \quad \text{To compare with 5,59\%}$$

Acquisition protocole



Static or dynamic acquisition in order to cover the whole body.

Data Correction



$$f_a(s,\theta) = \int_{-\infty}^{M} f(x,y) dt \, \mathrm{e}^{-\int_{-\infty}^{M} \mu(x,y) dt'} \times \int_{M}^{+\infty} f(x,y) dt \, \mathrm{e}^{-\int_{M}^{+\infty} \mu(x,y) dt'}$$

$$g_a(s,\theta) = \int_{-\infty}^{+\infty} f(x,y) dt \times e^{-\int_{-\infty}^{+\infty} \mu(x,y) dt'}$$

The attenuation •

- Do not depend on the localisation of the point of emission in the LOR.
- Depends only on the attenuation integral $\mu(x,y)$ different of 0.
- Depends on the function $\mu(x,y)$
 - \rightarrow required measurement.
- Example @ 511 keV:
 - Soft tissues: μ = 0,096 cm⁻¹
 - Muscle: μ = 0,1 cm⁻¹
 - Bone: μ = 0,134 cm⁻¹
 - Water: μ = 0.097cm⁻¹

Attenuation artifact

- Loss of an important number of photons
 - Around 17 % of photons pairs emitted in the centre of the brain.
 - Around 5 % of photons emitted in centre of the thorax.
- Bias in quantification.
- Unequal attenuation depending on the depth.
- Error in diagnosis.



Measurement of the map of the linear attenuation coefficients.

Source of ⁶⁸Ge

- Utilisable for a large period (T = 271 jours)
- Important dead time for detector near to the source.
- Acquisition: 15 to 30 min.
- Noisy image
- Small bias compared to true value (measured @ 511 keV)
- Source of simple photons (¹³⁷Cs)
 - Utilisable for a large period (T = 30,2 ans)
 - Acquisition: 5 to 10 min.
 - Biais with respect to true value
 - − Mesurement @ 662 keV: requirement of value conversion 662 keV \rightarrow 511 keV
 - Possible simultaneous Emission/Transmission acquisition









Acquisition protocole

• Projections acquisition in the presence of an object:

$$I(s,\theta) = I_0(s,\theta) \cdot e^{-\int_{-\infty}^{+\infty} \mu(x,y)dt}$$

- Full flux projections acquisition I_0
- Linear attenuation coefficient reconstruction μ(x,y) if required.
- Scaling coefficients if required.
- Calculation of attenuation correction factors (ACF):

$$e^{\int_{-\infty}^{+\infty} \mu(x,y)dt} = \frac{I_0(s,\theta)}{I(s,\theta)}$$

Attenuation correction

Two possible approaches

Projections correction

Projections multiplication

$$g_{c}(s,\theta) = \int_{-\infty}^{+\infty} f(x,y) dt \times e^{-\int_{-\infty}^{+\infty} \mu(x,y) dt'} \times ACF$$



Reconstruction of corrected projections



Correction during reconstruction

Iterative reconstruction with the modeling of the attenuation into the projector



Problems related to attenuation correction

- Patient motion between emission and transmission
 - Data scaling
 - Emission/Transmission simultaneous acquisition
- Noise propagation in images corrected from attenuation
 - Filtering the attenuation map
 - Segmentation of attenuation map
 - Usage of low noise attenuation map



Compton scatter

• In patient \implies Mispositioned coïncidence



• Dans le cristal

Détérioration de la résolution intrinsèque Rejet d'événements

Scatter artifact in PET

- Mispositioned coïncidences
 - Blurr
 - Loss in image contrast
 - Activity outside of the object
 - Quantitative bias
- More artifact in 3D artifact



Corrections de diffusion en TEP

- Estimation of the number of the scattered photons by energy spectra study:
 - Double energy window
 - True coïncidence estimation
- Estimation of the scattered photons from the projections:
 - Convolution
 - Profile approximation from outside the object
- Estimation of the scatterd by calculating the distribution:
 - Analytical calculation
 - Simulation de Monte Carlo



Scattered profile approximation

- Hypothesis:
 - Events outside the object
 - \rightarrow scatter distribution
 - Image of scatter = Low frequency image



Distribution calculation

- Hypothesis:
 - Known true events distribution
 - Known transmission map
- Algorithm

Possible iteration

- A Analytical calculation or Monte Carlo simulation of the scattered photons distribution.
 - \rightarrow Scattered sinogram
 - B Sinogram acquired estimation of scattered sinogram
 - \rightarrow Sinogram corrected from scatter
 - *C* Image reconstruction
 - \rightarrow Estimation of the true events distribution

3D methods comparison

PET field of view (FOV)



No activity outside the FOV

 \bigcirc

Figure of merit	Absolute c	Absolute concentration (kBq/ml)		Contrast (%) SNR			
Case/compartment	В		D		С	A	-
Calibration concentration	5.88		4.86		100	-	_
AC	7.66±0.28	30%	5.31±0.17	9%	63.82±1.15	21.91±5.17	No correction
DEW	6.05±0.23	3%	4.62±0.18	-5%	91.63±1.84	15.42±3.64	DEW
CVS	6.49±0.30		4.68±0.23		84.11±3.85	18.79±4.54	
SRBSC	6.52±0.30		4.76±0.22		86.26±3.95	19.46±4.72	
MCBSC1	6.51±0.24	11%	4.81±0.21	-1%	81.31±3.93	9.74±2.43	Monte Carlo
MCBSC2	6.55±0.27		4.78±0.15		85.02±1.76	10.32±2.05	_

Zaidi et al, Eur J Nucl Med 2000:1813-1826

3D methods comparison

PET field of view (FOV)



Activity outside of the FOV

0

Figure of merit	Absolute concentration (kBq/ml)				Contrast (%)	SNR	
Case/compartment	В		D		С	A	
Calibration concentration	5.88		4.86		10	-	
AC	7.94±0.30	35%	5.47±0.15	13%	64.60±1.08	19.04±4.69	No correction
DEW	6.14±0.21	4%	4.61±0.10	-5%	95.74±2.09	12.37±3.97	DEW
CVS	6.72±0.32		4.82±0.20		84.90±3.34	16.24±4.33	
SRBSC	6.76±0.32		4.90±0.19		86.78±3.30	16.81±4.60	
MCBSC1	6.62±0.31	13%	4.72±0.24	-3%	86.23±2.64	9.78±3.37	Monte Carlo
MCBSC2	6.77±0.24		4.94±0.18	- / -	86.33±1.54	9.33±2.33	

Zaidi et al, Eur J Nucl Med 2000:1813-1826

Normalization correction

- Required correction:
 - No detection uniformity between different detectors.
 - Geometrical factors (curvature of the detection ring)
 - Dead zone in collected data

Each system line response should have the same detection efficiency





Normalisation coefficient (NC)



Normalized sinogram

Normalisation coefficients

- Full flux acquisition
 - All lines of response are irradiated by the same source.
 - Require a high number of counts (important statistics).
- Modeling
 - NC = Crystal efficiency x Geometrical factor



Obtaining the coefficients

 $g(x) \times crystal interference(x \mod 8)$

 $e(\det 1) \times e(\det 2)$





Random coïncidences in PET

Detection solid angle of unique photon



- Random coïncidences depend on: •
 - Used time window
- Consequences of the random coïncidences: •
 - Bad localisaton
 - A quantitatif bias

Correction of random coïncidences

Measurement of delayed coïncidences

- Utilisation of a delayed window



Correction of random coïncidences

- Variance reduction
 - Smoothing of the random coïncidences sinogram delayed with an appropriate methods.
 - Estimation of the sinogram of random coïncidences fortuites from the measurement of simple photons.

$$R_{ij} = 2\tau S_i \varepsilon_i S_j \varepsilon_j$$

Simples Normalisation



Noisy Sinogram





Question:

5 minutes whole body acquisition Rate of measured coïncidences = 50 000 cps Rate of random coïncidences = 20 000 cps

Acquisition in 3D mode with 10⁶ LORs

Compare the two correction methods (noisy and not noisy) in terms of % $\Delta N_{true}/N_{true}$ per LOR

Hypothesis: Scattered coïncidences are negligeables True and random coïncidences are distributed uniformly on all the LORs

> Total number of coïncidences per LOR: $50000 \times 300 / 10^6 = 15$ Number of random coïncidences per LOR: $20000 \times 300 / 10^6 = 6$

Number of true coïncidences per LOR: 15 - 6 = 9

Noisy method $\Delta N_{vraie} = \sqrt{N_{vraie} + 2 \times N_{fortuit}} = \sqrt{9 + 12} = 4,58$ $\Delta N_{vraie} = 100 \times \frac{\Delta N_{vraie}}{N_{vraie}} = 50,9\%$ $100 \times \frac{\Delta N_{vraie}}{N_{vraie}} = 50,9\%$ $100 \times \frac{\Delta N_{vraie}}{N_{vraie}} = 50,9\%$

No Noisy method

$$\Delta N_{vraie} = \sqrt{N_{vraie} + N_{fortuit}} = \sqrt{9 + 6} = 3,87$$

$$100 \times \frac{\Delta N_{vraie}}{N_{vraie}} = 43\%$$

Quality image estimation

- How to estimate the image quality?
 - Based on the concepts of:
 - Sensibility: detection of true positives
 - Specificity: détection de false positives
 - → Receiver Operating Charactéristics: represents the probability of detecting a true positive as function of detecting a false positive.
 - Difficult set up
- Estimate the signal to noise ratio in the image: Noise Equivalent Count Rate

Signal to noise ratio in the image



• A line of response registers the following signal:



Variance calculation

The variance of an image uncorrected from attenuation and dead time:



Hypothesis: Scattered and random coïncidences correction does not increase the signal

Correction factor

$$VAR_e = ac^2 \cdot VAR$$

After dead time and attenuation correction:

Simplifying the problem



Number of coïncidences in a pixel



$$VAR_e = ac^2 \cdot VAR$$

$$SNR = k \frac{t_e}{\sqrt{VAR_e}}$$

$$t_e = T \cdot moy(ac) \cdot \frac{d^2}{\frac{\pi}{4}D^2}$$

$$SNR = k \cdot \sqrt{\frac{4}{\pi}} \left(\frac{d}{D}\right)^{\frac{3}{2}} \sqrt{\frac{moy(ac)}{ac}} \sqrt{\frac{T}{1 + \alpha_{sp} + \alpha_{rp}}}$$

Noise Equivalent Count

$$NEC \approx SNR^{2}$$

$$SNR = k \cdot \sqrt{\frac{4}{\pi}} \left(\frac{d}{D}\right)^{\frac{3}{2}} \frac{moy(ac)}{ac} \sqrt{NEC}$$

$$True \ coïncidences in the image$$

$$NEC = \frac{T}{1 + \alpha_{sp} + \alpha_{rp}}$$

$$Fraction \ for \ a \ line \ of \ response$$

$$Reminder$$

$$Hypothesis: \ Corrections \ of \ random \ and \ scattered \ coïncidences \ do \ not \ introduce \ noise$$

In presence of noise,

SignalVarianceRandom coïncidences:
$$t_p + s_p + r_p - r_p$$
 $t_p + s_p + (1 + \beta) f_{FOV} r_p$

Scattered coïncidences: $t_p + s_p + r_p - r_p - \alpha_{cp}(t_p + s_p) \quad t_p + s_p + \beta r_p + \alpha_{cp}^2(t_p + s_p + (1 + \beta)f_{FOV}r_p)$ $(1 + \alpha_{cp}^2)(t_p + s_p + (1 + \beta)f_{FOV}r_p)$

Noise Equivalent Count

$$NEC = \frac{T}{1 + \alpha_{sp} + \alpha_{rp}}$$

$$NEC = \frac{T}{(1 + \alpha_{cp}^{2})(t_{p} + s_{p} + (1 + \beta)f_{FOV}r_{p})}$$

$$The X_{i} are approximations of X_{p} in the image$$

$$NEC = \frac{T^{2}}{(1 + \alpha_{ci}^{2})(T + S_{i} + (1 + \beta)f_{FOV}R_{i})}$$

The more often, $\alpha_{ci} = 0$ because the scattered correction does not introduce noise

$$NEC = \frac{T^2}{\left(T + S_i + \left(1 + \beta\right) f_{FOV} R_i\right)}$$

Application example: Random correction methods comparison

TEP HR+

Detector size (mm)	4.0 × 4.1 × 30
No. of slices	63
Slice width (mm)	2.4
Ring diameter (cm)	82.7
Axial FOV (cm)	15.2
Max. axial angle	10.2°
Mode	2D/3D

23 patients (13m, 10f) (170 \pm 10) cm (76 \pm 16) kg

Dose distribution, 1h agter injection de [18F]-FDG





Methods comparison

Count rate per bed position



Methods comparison

Relative improvement of NEC:



Improvement in NEC

In summary

TABLE 1 Methods for Estimating Random Coincidences

Method	Comments
Delayed coincidences Smoothed delayed coincidences Calculated from single photon rates	Accurate. Higher noise (Eq. 1). Lowest processing requirements. Accurate. Lower noise ($0 \le k \ll 1$, Eq. 1). Higher processing requirements. Potential for bias if scanner is not properly calibrated. Lower noise ($0 \le k \ll 1$, Eq. 1). Low processing requirements.

Variance reduction used with high $\frac{f_{FOF}R}{T+S}$

Crystal choice

