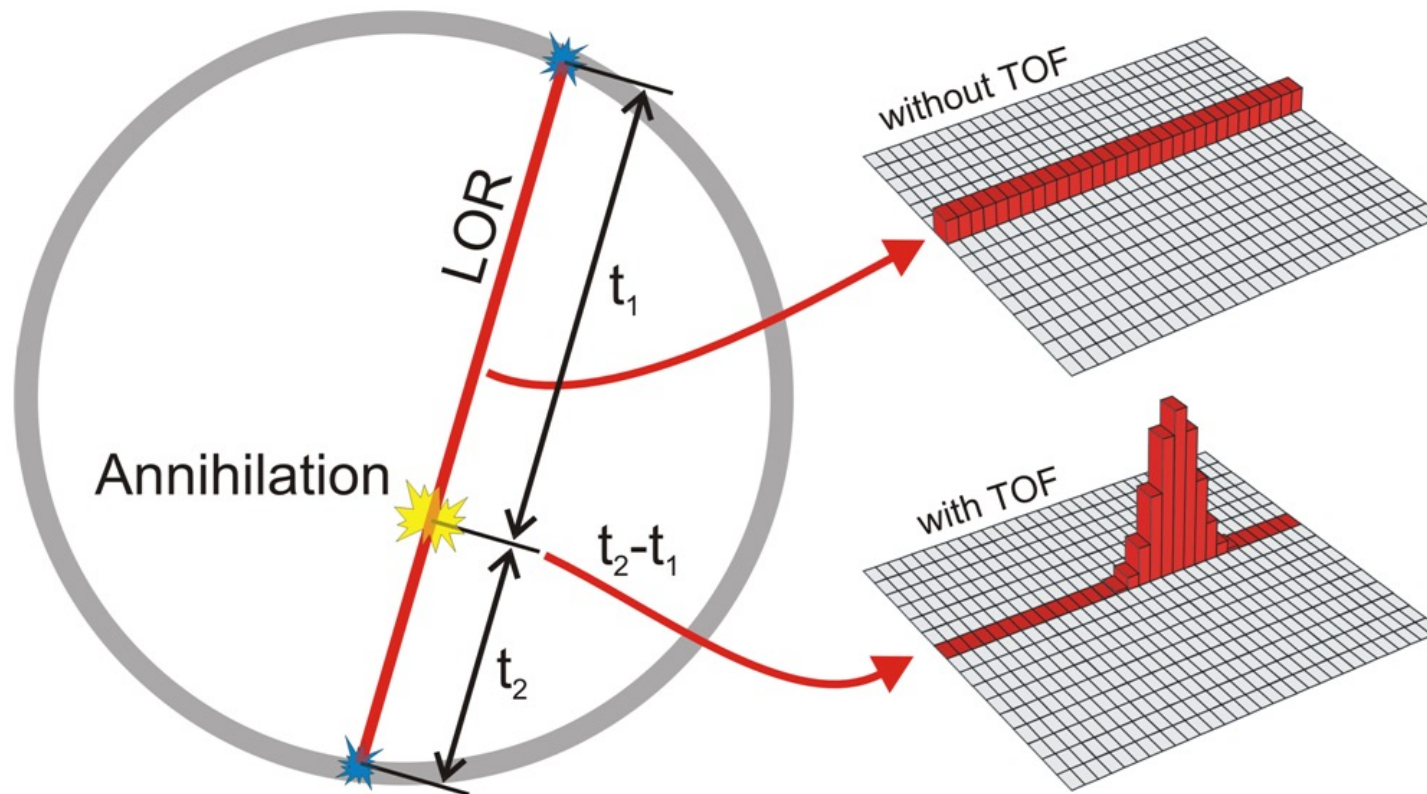


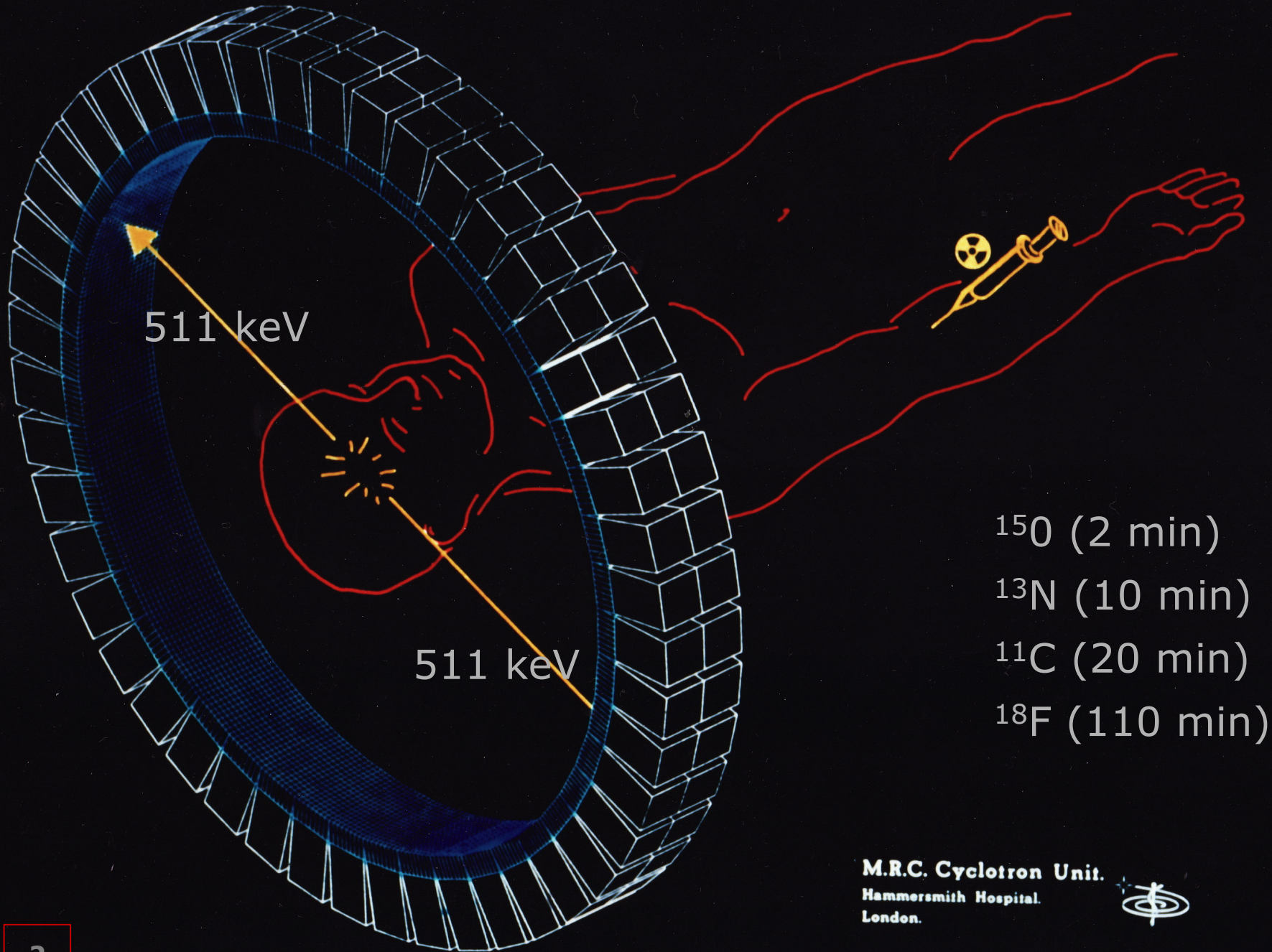
# 10 ps: a Challenge for the Future of Positron Emission Tomography

Christian Morel

Aix-Marseille Univ, CNRS/IN2P3, CPPM, Marseille, France



# Positron Emission Tomography (PET)

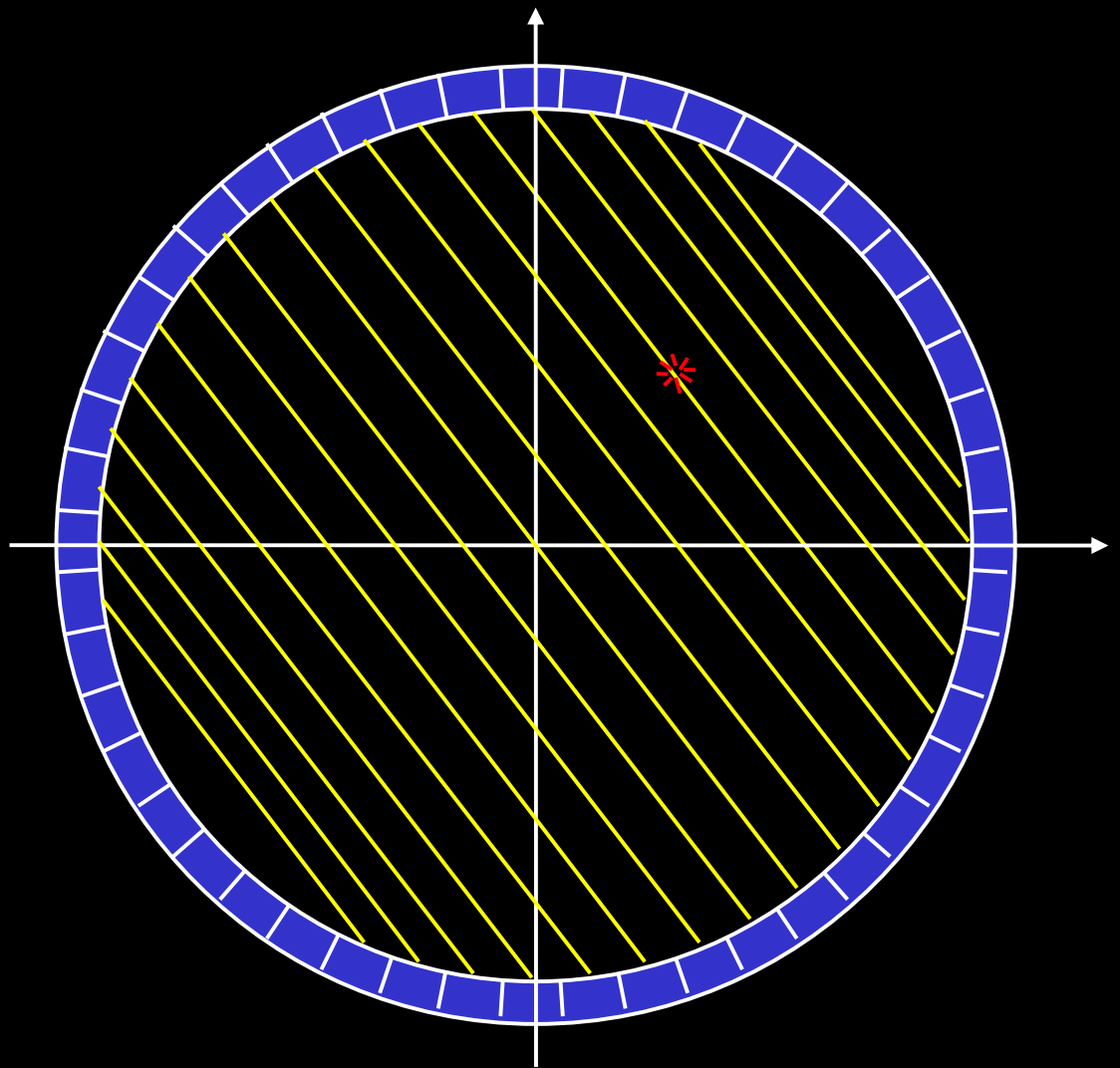
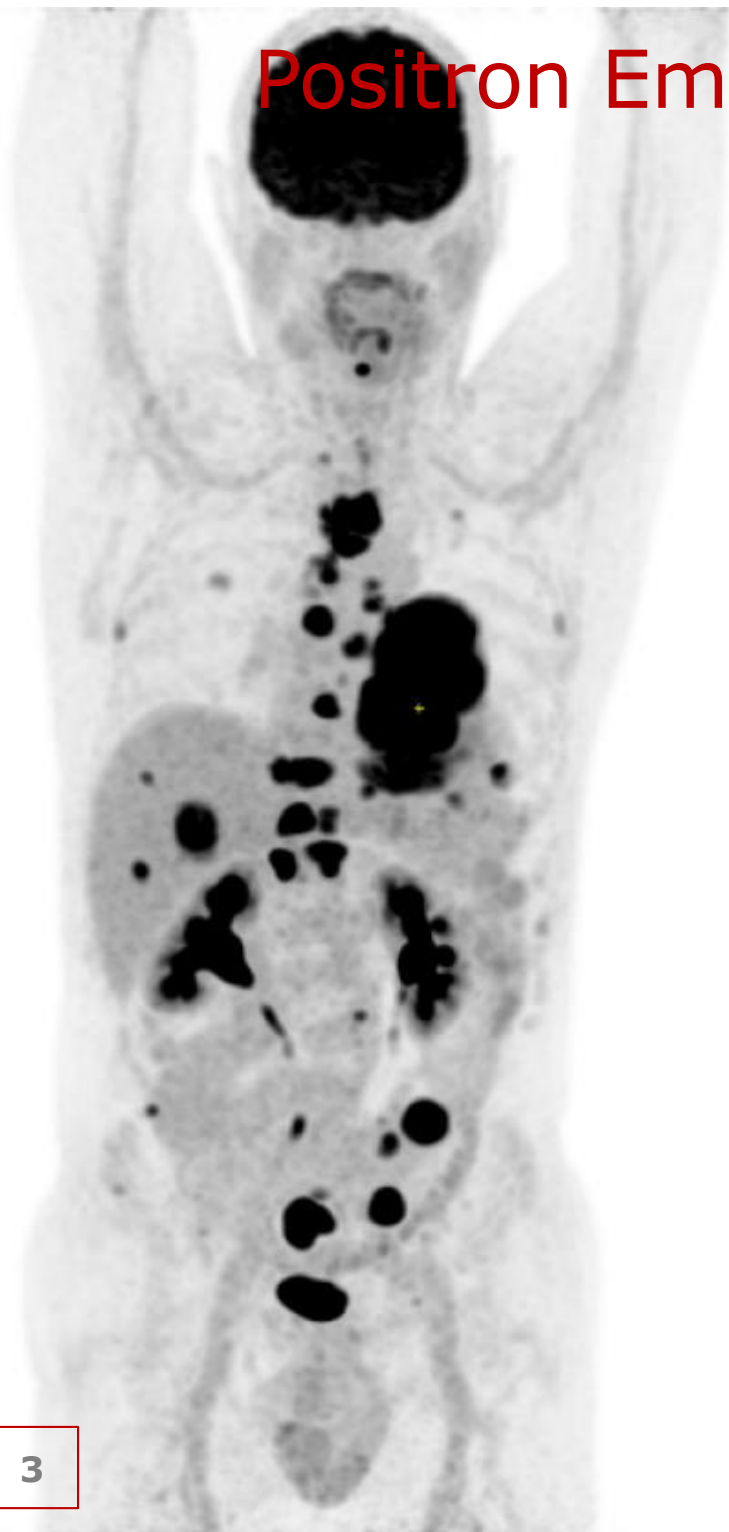


- $^{15}\text{O}$  (2 min)
- $^{13}\text{N}$  (10 min)
- $^{11}\text{C}$  (20 min)
- $^{18}\text{F}$  (110 min)

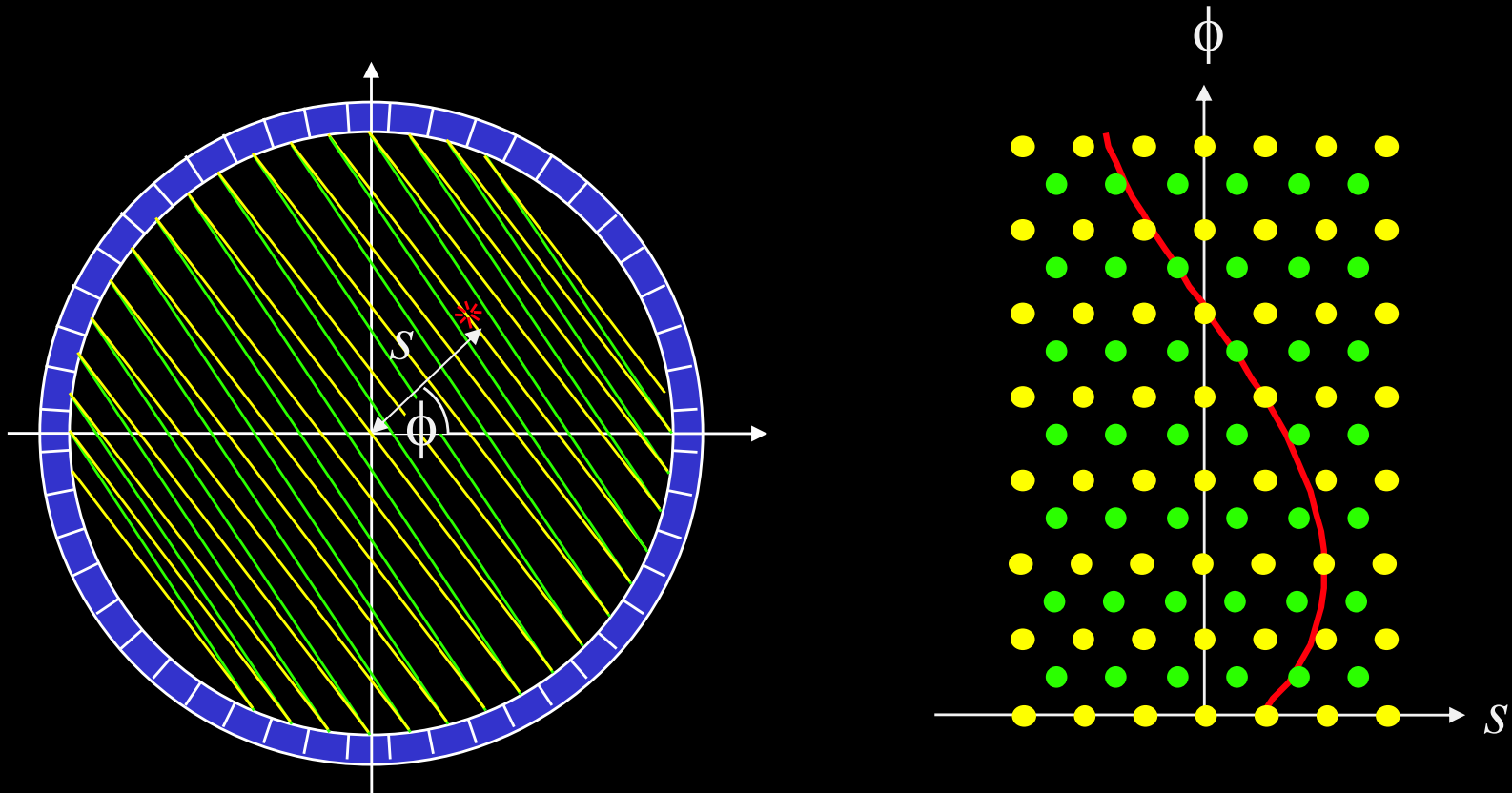
M.R.C. Cyclotron Unit.  
Hammersmith Hospital.  
London.



# Positron Emission Tomography (PET)

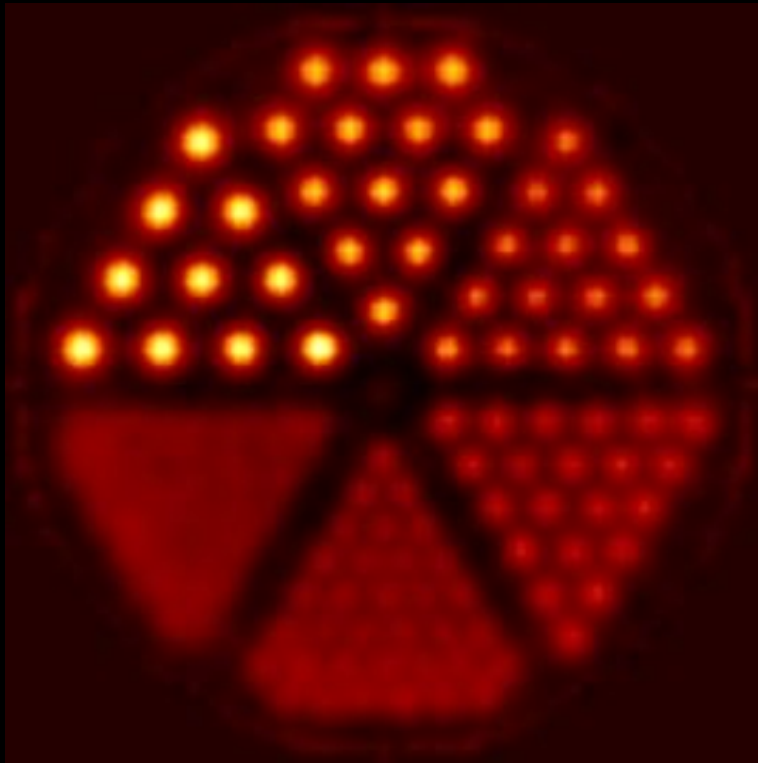


# Positron Emission Tomography (PET)

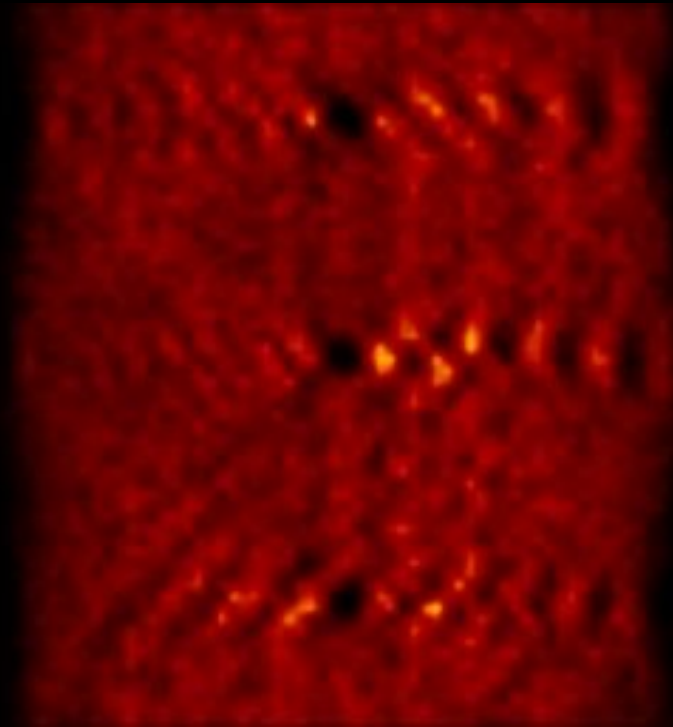


Sinogram

# Positron Emission Tomography (PET)



Reconstructed image



Sinogram

# Radial spatial resolution

$$R = a \sqrt{\left(\frac{d}{2}\right)^2 + b^2 + r^2 + (0.0022 D)^2}$$

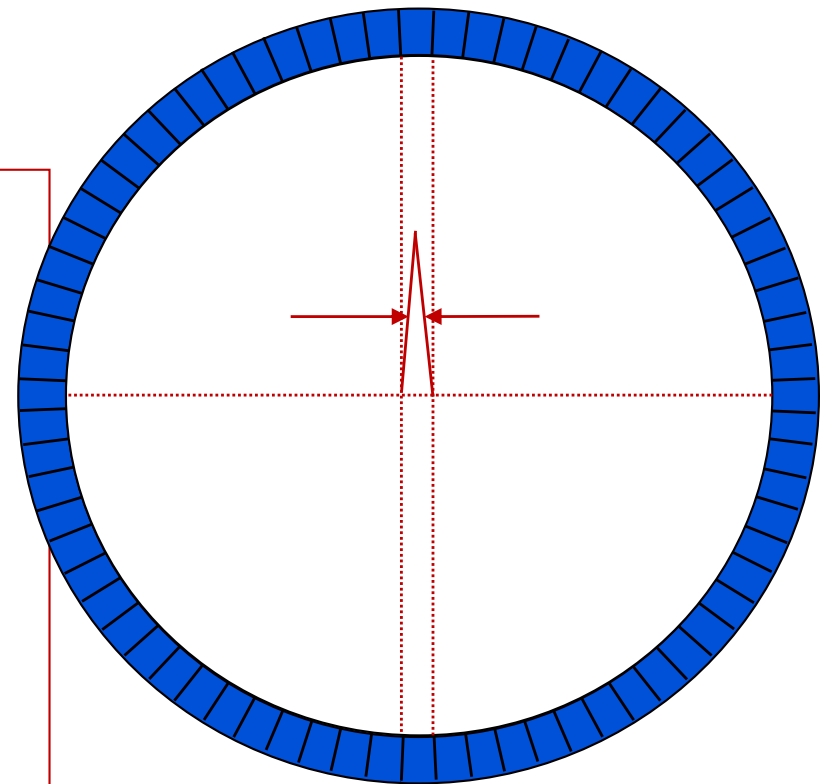
$d$  Pixel size

$D$  Ring diameter

$r$  Positron range

$b$  Detector crosstalk

$a$  Image reconstruction algorithm  
(1,1 - 1,3)

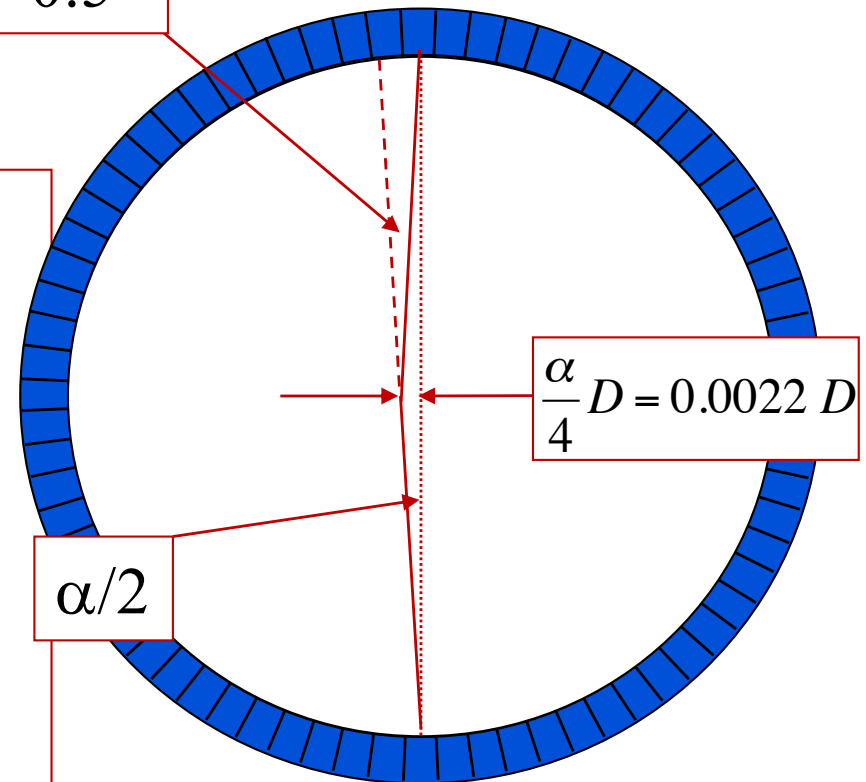


# Spatial resolution at image centre

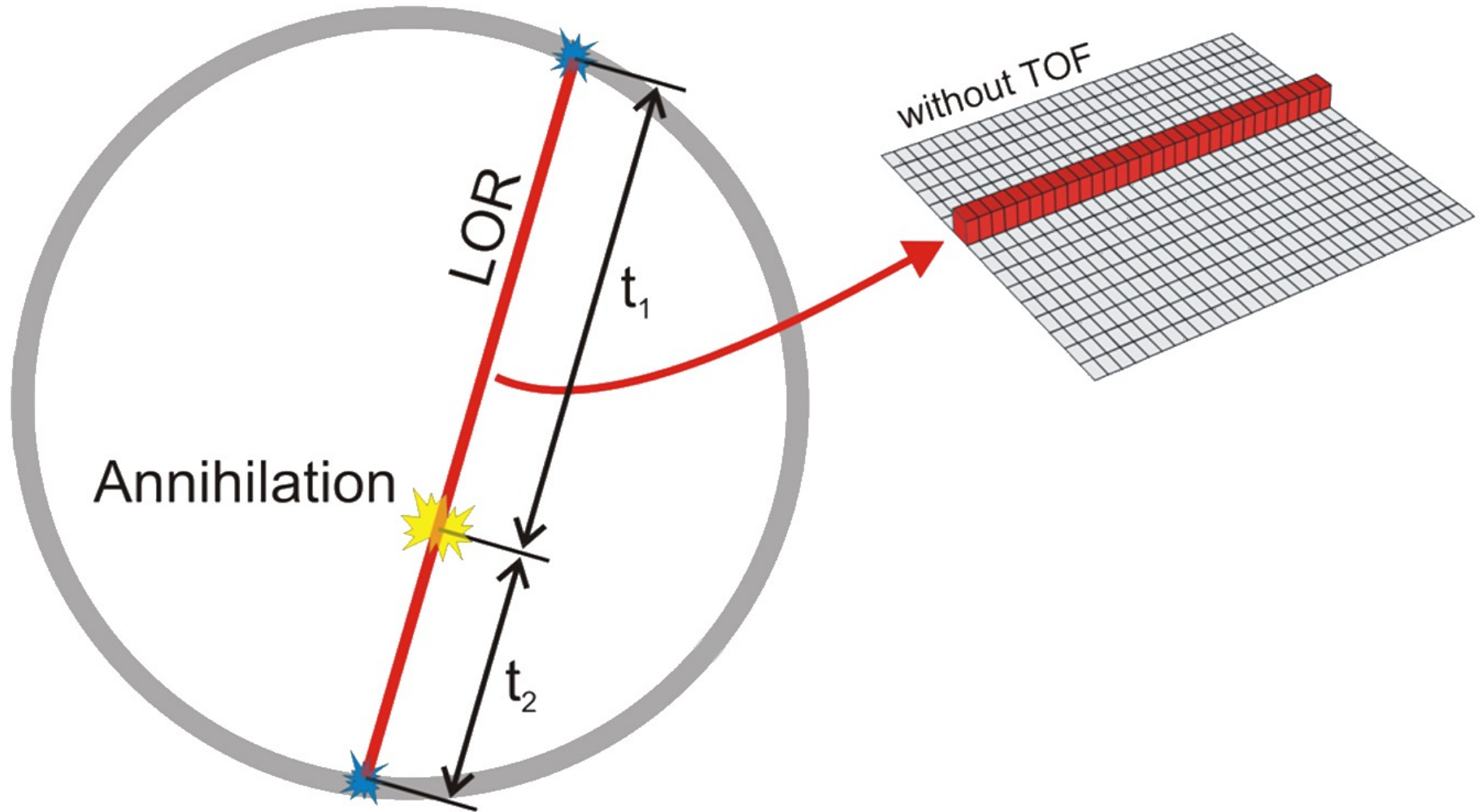
$$R = a \sqrt{\left(\frac{d}{2}\right)^2 + b^2 + r^2 + (0.0022 D)^2}$$

- $d$  Pixel size
- $D$  Ring diameter
- $r$  Positron range
- $b$  Detector crosstalk
- $a$  Image reconstruction algorithm  
(1,1 - 1,3)

$$\alpha \sim 0.5^\circ$$

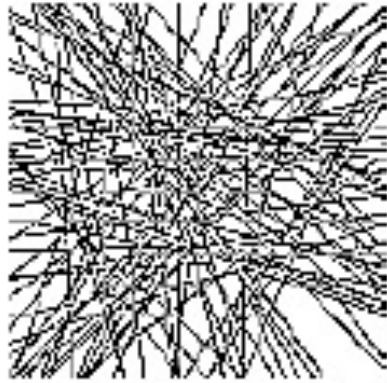


# Backprojection

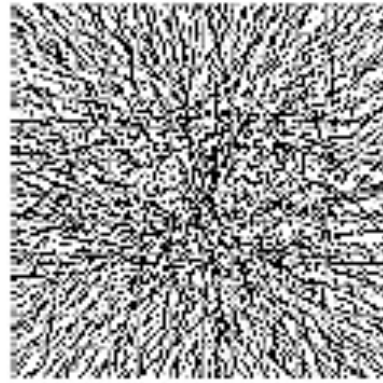




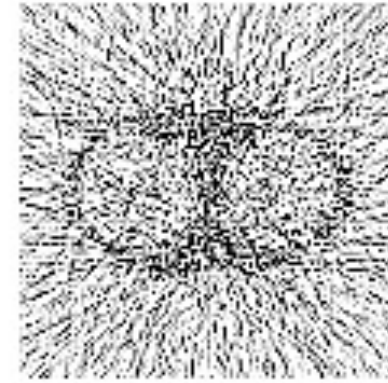
# Tomography and counting statistics



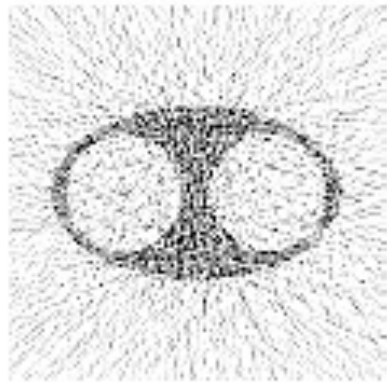
$10^2$



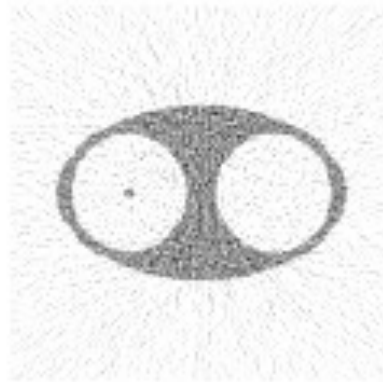
$10^3$



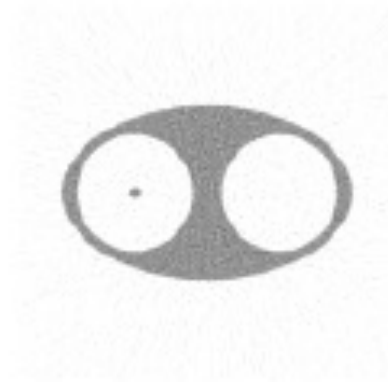
$10^4$



$10^5$



$10^6$



$10^7$

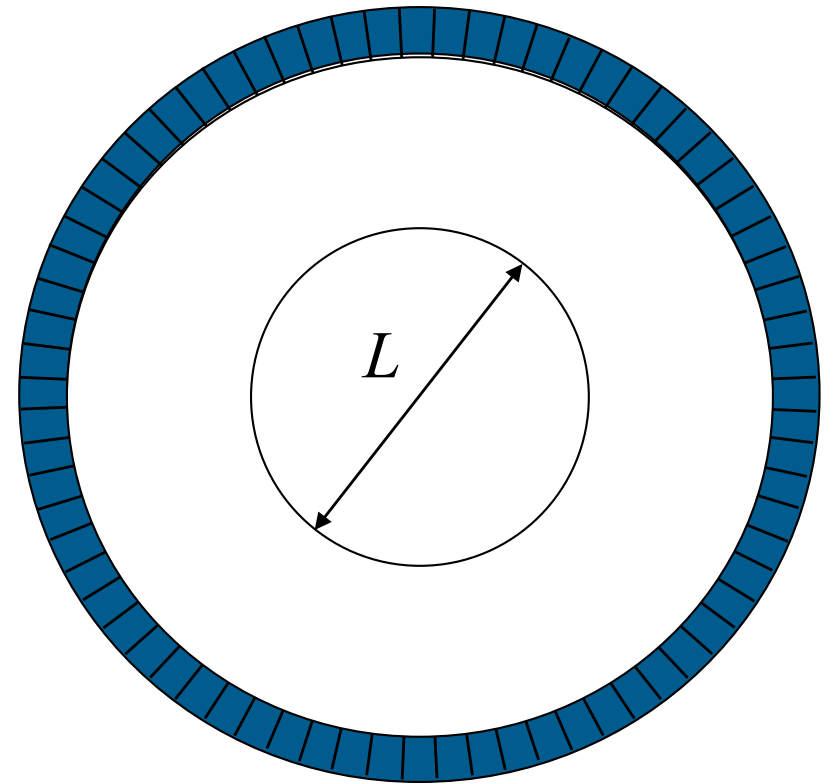
Courtesy: C. Comtat, CEA-SHFJ

# SNR and counting statistics

$$\text{SNR} = \frac{A}{\Delta A} = \frac{N_{\beta^+}}{\sqrt{N_{\beta^+}}} = \sqrt{N_{\beta^+}}$$

$$\Rightarrow N_{\beta^+} = \text{SNR}^2$$

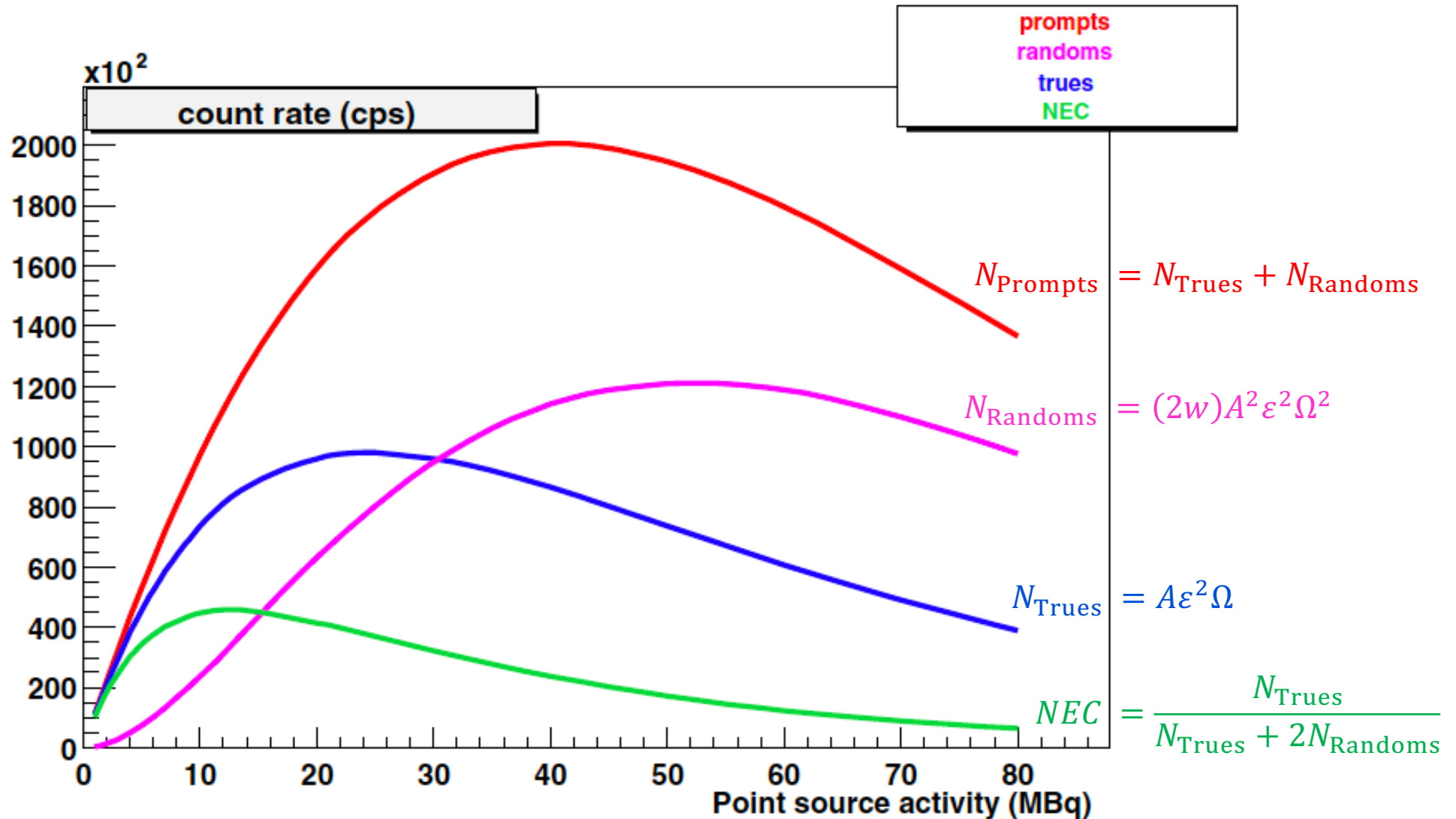
$$N_{\text{evt}} = \left(\frac{L}{d}\right)^3 \times \text{SNR}^2 \times \left(\frac{L}{d}\right)$$



Improving spatial resolution  $\times 2 \Rightarrow$  increasing counting statistics  $\times 16$  to get unchanged SNR in the reconstructed image voxels

- ↗ scan duration
- ↗ crystal thickness (crystal efficiency)
- ↗ injected activity
- ↗ solid angle

# Noise Equivalent Count Rate (NECR)



# Inorganic scintillators for PET

	NaI	BGO	BaF <sub>2</sub>	L(Y)SO	LuAP	LaBr <sub>3</sub>
Density (g/cm <sup>3</sup> )	3.67	7.13	4.88	7.40	8.34	5.3
Effective Z	51	74	53	66	65	52
Photofraction (%)	18	42	19	33	32	14
Decay time (ns)	230	300	0.6 620	35-45	17	16
Light yield (ph/MeV)	43000	8200	1430 9950	28000	11400	60000
Peak emission (nm)	415	480	220 310	420	365	300
Refraction index	1.85	2.15	1.47	1.82	1.97	1.9

# Radial spatial resolution

$$R(s) = a \sqrt{\left(\frac{d}{2}\right)^2 + \frac{(w^2 - d^2)}{D^2} s^2 + b^2 + r^2 + (0.0022 D)^2}$$

$d$  Pixel size

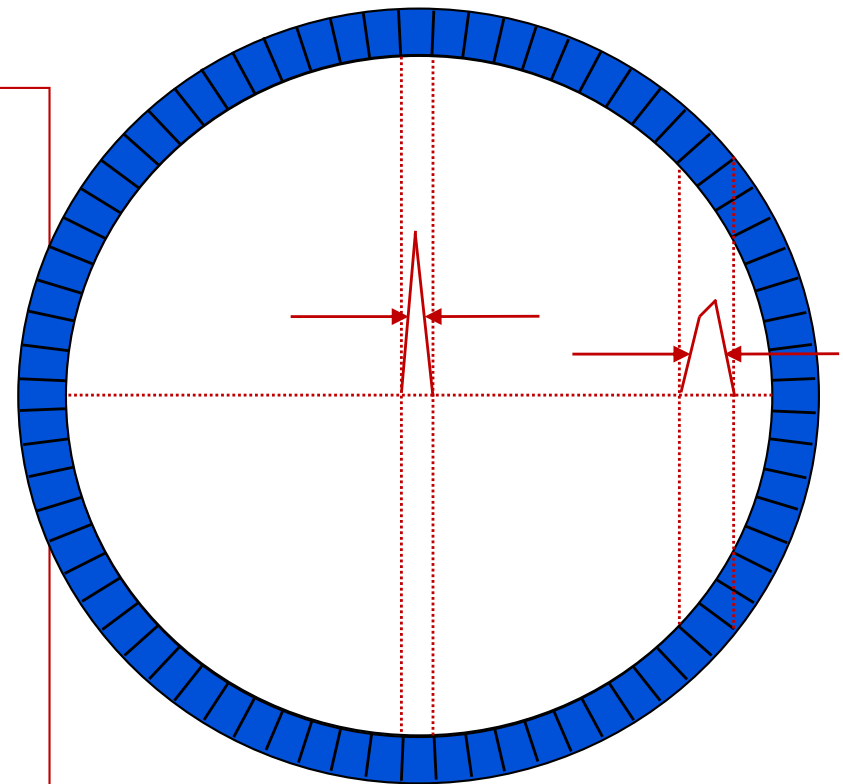
$w$  Pixels thickness

$D$  Ring diameter

$r$  Positron range

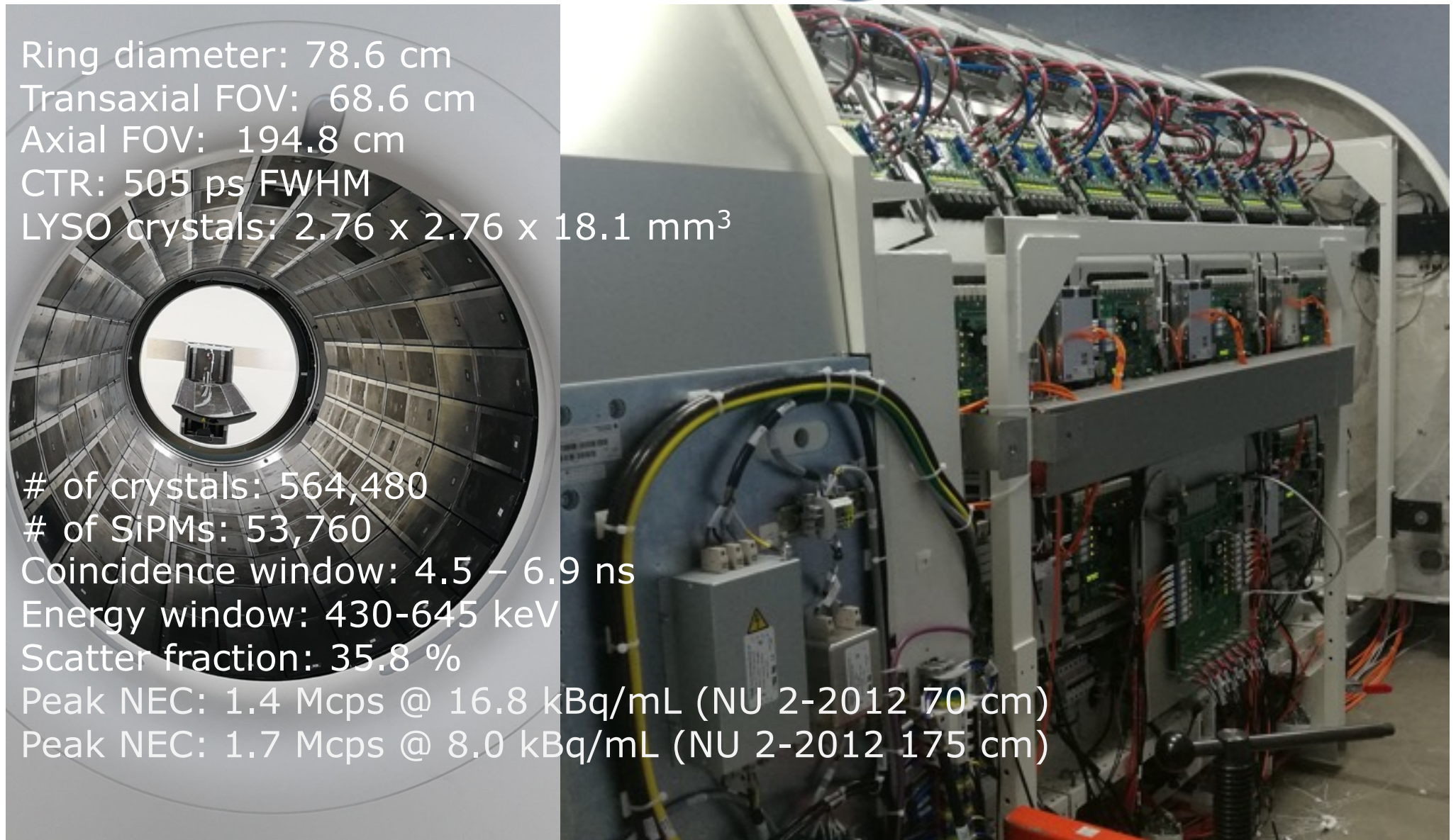
$b$  Detector crosstalk

$a$  Image reconstruction algorithm  
(1,1 - 1,3)



# Total Body-PET **EXPLORER** scanner

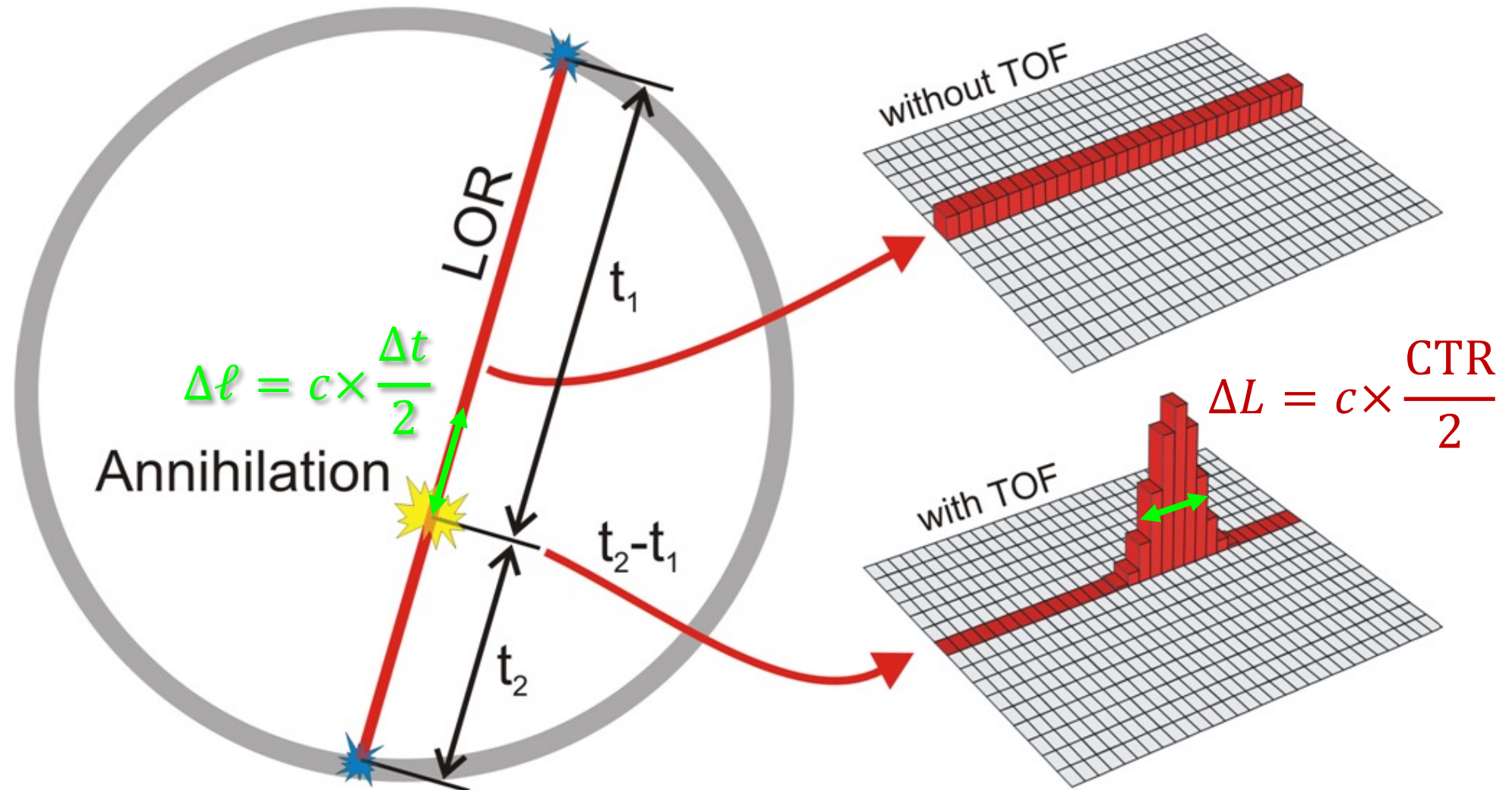
14



Ring diameter: 78.6 cm  
Transaxial FOV: 68.6 cm  
Axial FOV: 194.8 cm  
CTR: 505 ps FWHM  
LYSO crystals:  $2.76 \times 2.76 \times 18.1 \text{ mm}^3$

# of crystals: 564,480  
# of SiPMs: 53,760  
Coincidence window: 4.5 – 6.9 ns  
Energy window: 430-645 keV  
Scatter fraction: 35.8 %  
Peak NEC: 1.4 Mcps @ 16.8 kBq/mL (NU 2-2012 70 cm)  
Peak NEC: 1.7 Mcps @ 8.0 kBq/mL (NU 2-2012 175 cm)

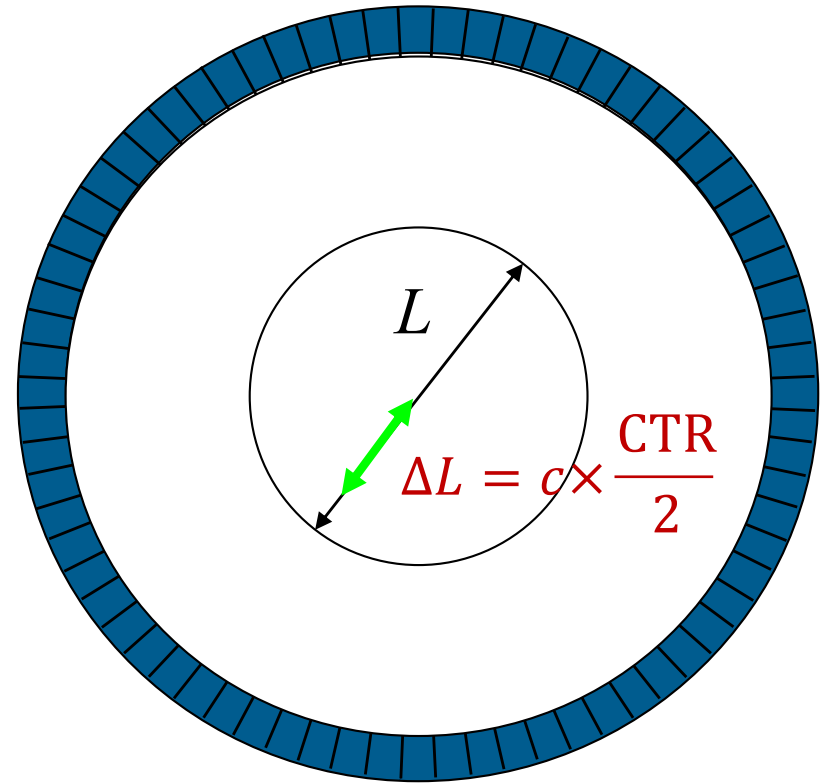
# Time-Of-Flight (TOF)-PET



# Impact of TOF-PET on image SNR

$$N_{\text{evt}} = \left(\frac{L}{d}\right)^3 \times \text{SNR}^2 \times \left(\frac{L}{d}\right)$$

$$N_{\text{TOF}} = \left(\frac{L}{d}\right)^3 \times \text{SNR}^2 \times \left(\frac{\Delta L}{d}\right)$$



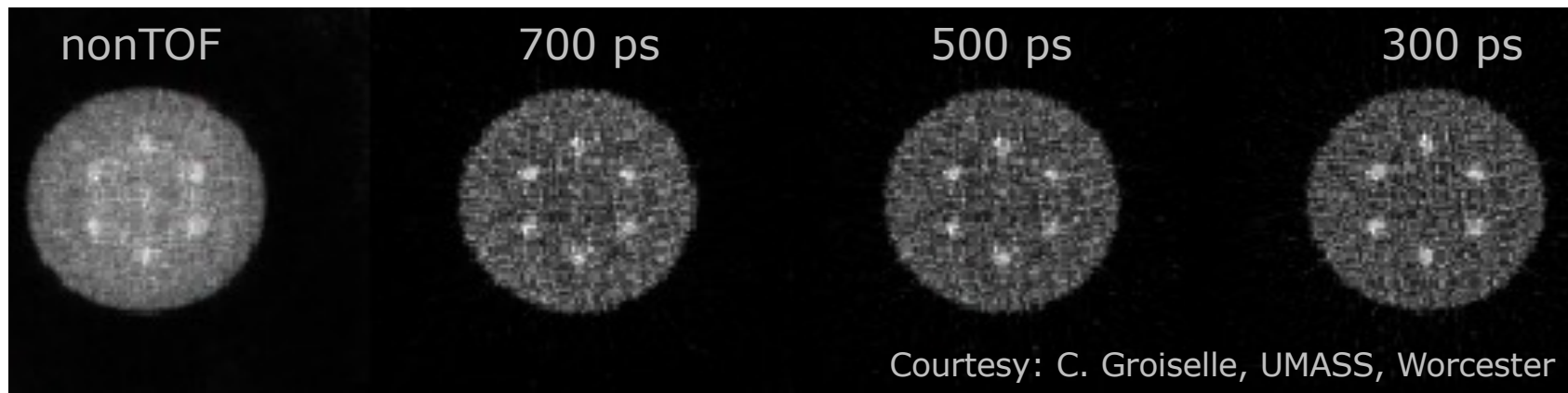
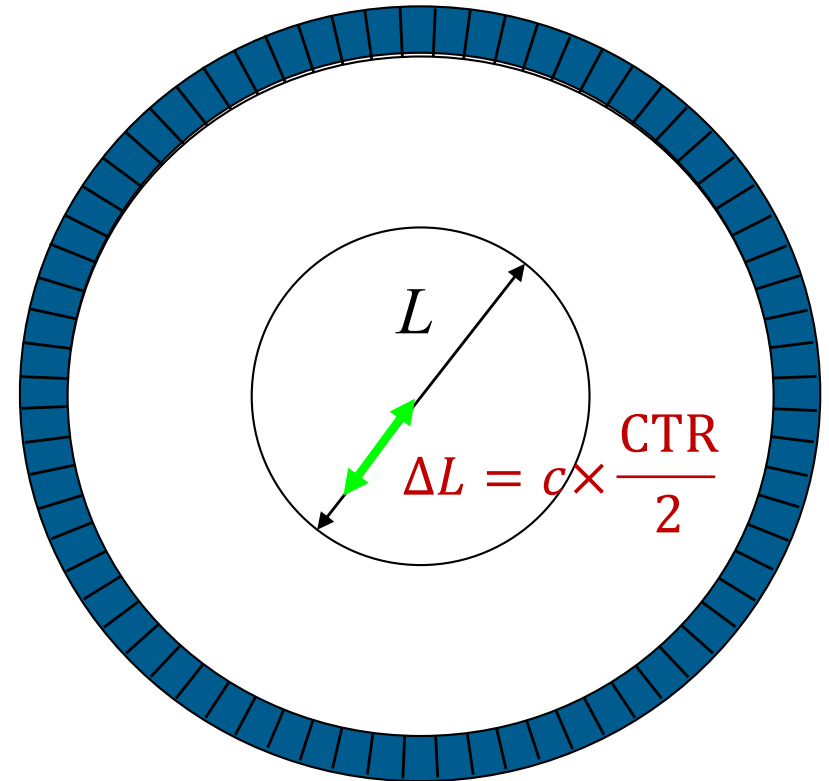


# Impact of TOF-PET on image SNR

$$N_{\text{evt}} = \left(\frac{L}{d}\right)^3 \times \text{SNR}_{\text{nonTOF}}^2 \times \left(\frac{L}{d}\right)$$

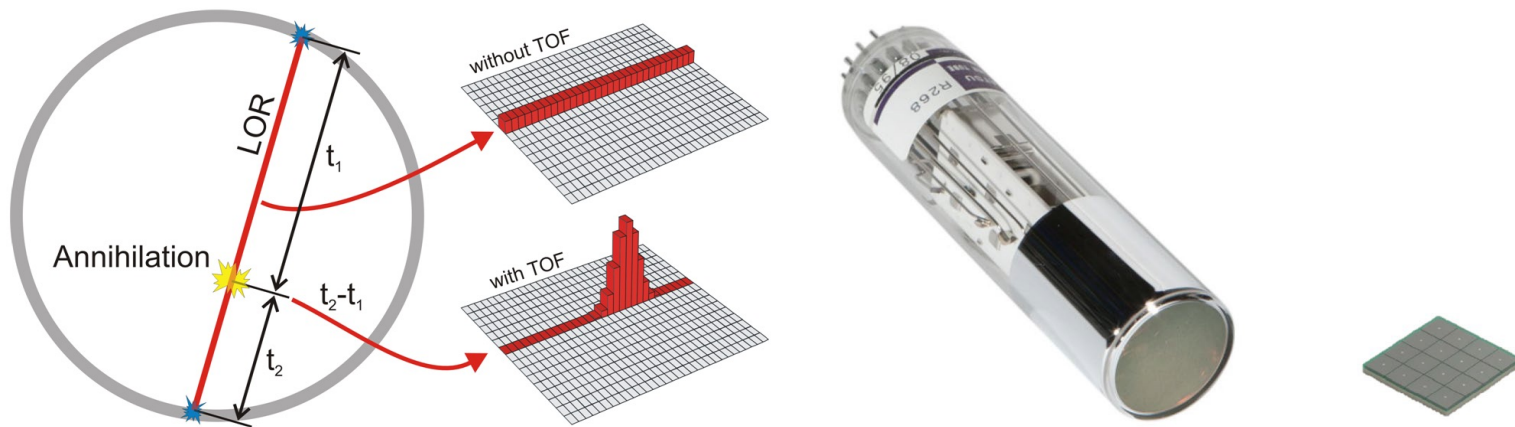
$$N_{\text{evt}} = \left(\frac{L}{d}\right)^3 \times \text{SNR}_{\text{TOF}}^2 \times \left(\frac{\Delta L}{d}\right)$$

$$\left(\frac{\text{SNR}_{\text{TOF}}}{\text{SNR}_{\text{nonTOF}}}\right)^2 = \frac{2L}{c \times \text{CTR}}$$

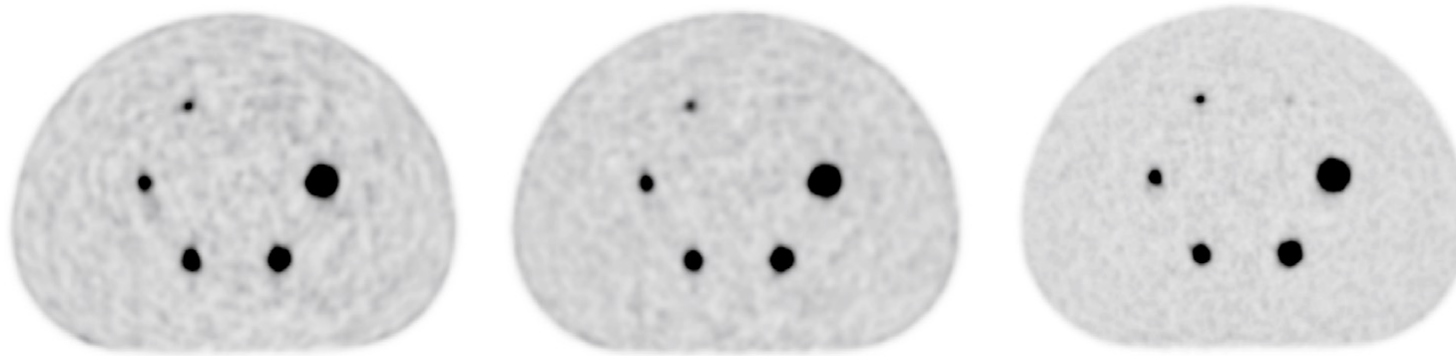


Courtesy: C. Groiselle, UMASS, Worcester

# State-of-the-art TOF-PET



$$\left( \frac{SNR_{TOF}}{SNR_{nonTOF}} \right)^2 = \frac{2D}{c \times CTR}$$

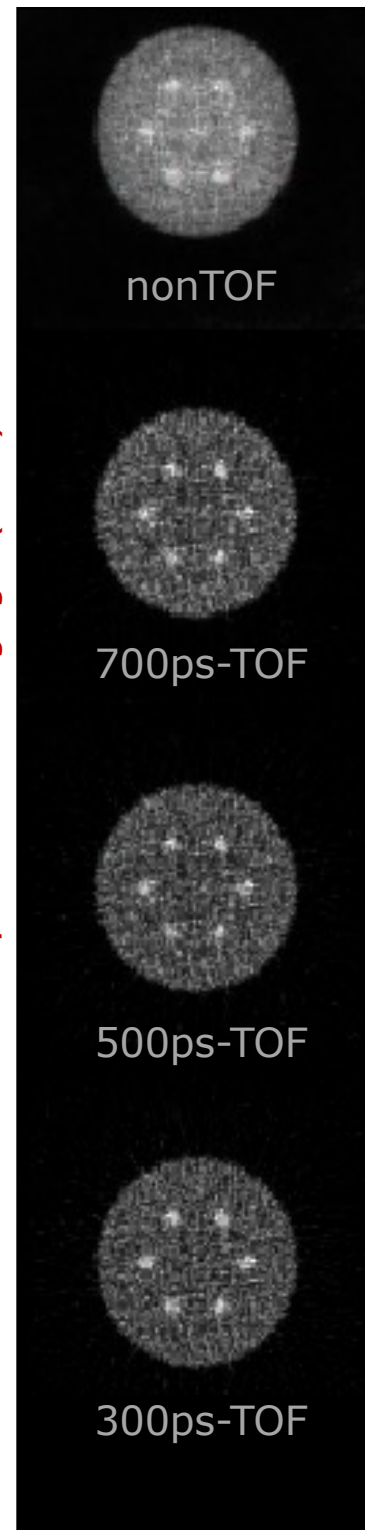


nonTOF

527ps-TOF

210ps-TOF

M. Conti and B. Bendriem, Clin Transl Imaging 7 (2019) 139–147



nonTOF

700ps-TOF

500ps-TOF

300ps-TOF

# Brief history of (TOF&TB-)PET scanners

- 60s: TOF-PET advocated by Anger (LBL), Brownell (MGH) and Budinger (LBL)
  - 80s: First TOF-PET scanners for  $^{15}\text{O}$  and  $^{13}\text{N}$  imaging using CsF/PMT and BaF<sub>2</sub>/PMT (PETT (St-Louis), TTV01-3 (Grenoble), TOFPET-I (Houston), SP3000/UW (Seattle))  
**CTR: 450-750 ps FWHM**
  - 90s: 3D-PET imaging with nonTOF-PET scanners using BGO/PMT
  - Mid-90s: nonTOF-PET/CT scanner using LSO/PMT (4-6 ns coincidence time window)
  - Mid-00s: TOF-PET scanners using L(Y)SO/PMT or LaBr<sub>3</sub>/PMT  
**CTR: 450-550 ps FWHM**
  - Mid-10s: TOF-PET scanners using L(Y)SO/SiPM  
**CTR: 300-400 ps FWHM**
  - End-10s: TOF-PET scanner using LSO/SiPM  
Biograph Vision **CTR: 210 ps FWHM**
- Total Body (TB)-PET using L(Y)SO/SiPM

# Brief history of (TOF&TB-)PET scanners

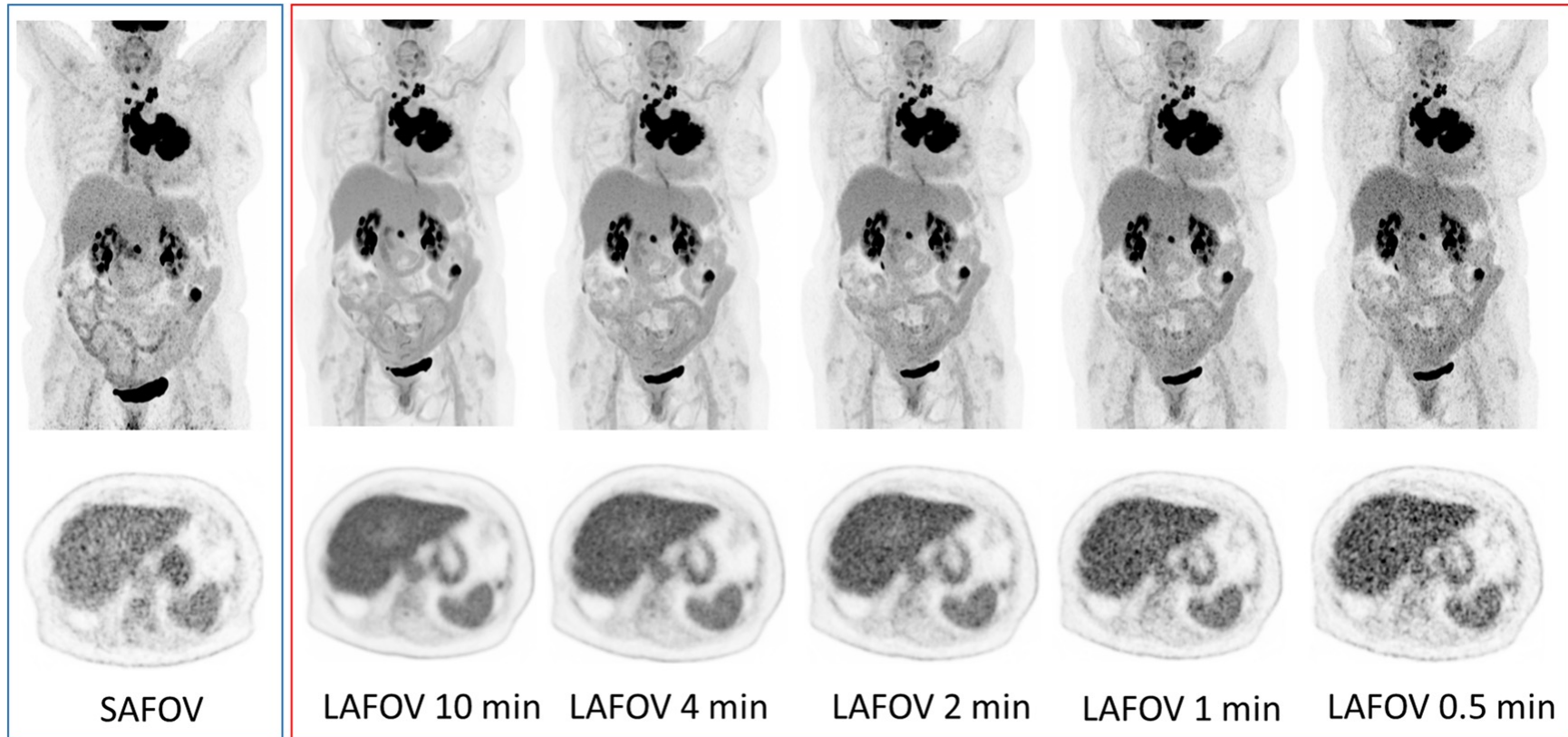
- 60s: TOF-PET advocated by Anger (LBL), Brownell (MGH) and Budinger (LBL)
- 80s: First TOF-PET scanners for  $^{15}\text{O}$  and  $^{13}\text{N}$  imaging using CsF/PMT and BaF<sub>2</sub>/PMT (PETT (St-Louis), TTV01-3 (Grenoble), TOFPET-I (Houston), SP3000/UW (Seattle))  
**CTR: 450-750 ps FWHM**
- 90s: 3D-PET imaging with nonTOF-PET scanners using BGO/PMT
- Mid-90s: nonTOF-PET/CT scanner using LSO/PMT (4-6 ns coincidence time window)
- Mid-00s: TOF-PET scanners using L(Y)SO/PMT or LaBr<sub>3</sub>/PMT  
**CTR: 450-550 ps FWHM**
- Mid-10s: TOF-PET scanners using L(Y)SO/SiPM  
**CTR: 300-400 ps FWHM**
- End-10s: TOF-PET scanner using LSO/SiPM  
Biograph Vision **CTR: 210 ps FWHM, AFOV: 26.3 cm**
- Total Body (TB)-PET using L(Y)SO/SiPM  
uEXPLORER **CTR: 505 ps FWHM, AFOV: 194.8 cm**  
PennPET EXPLORER **CTR: 250 ps FWHM, AFOV: 140 cm**  
Biograph Vision Quadra **CTR: 230 ps FWHM, AFOV: 106 cm**

# Total Body-PET scanner Biograph Vision Quadra

Ring diameter: 82 cm  
 Transaxial FOV: 78 cm  
 Axial FOV: 106 cm  
**CTR: 230 ps FWHM**  
 LSO crystals:  
 3.2 x 3.2 x 20 mm<sup>3</sup>

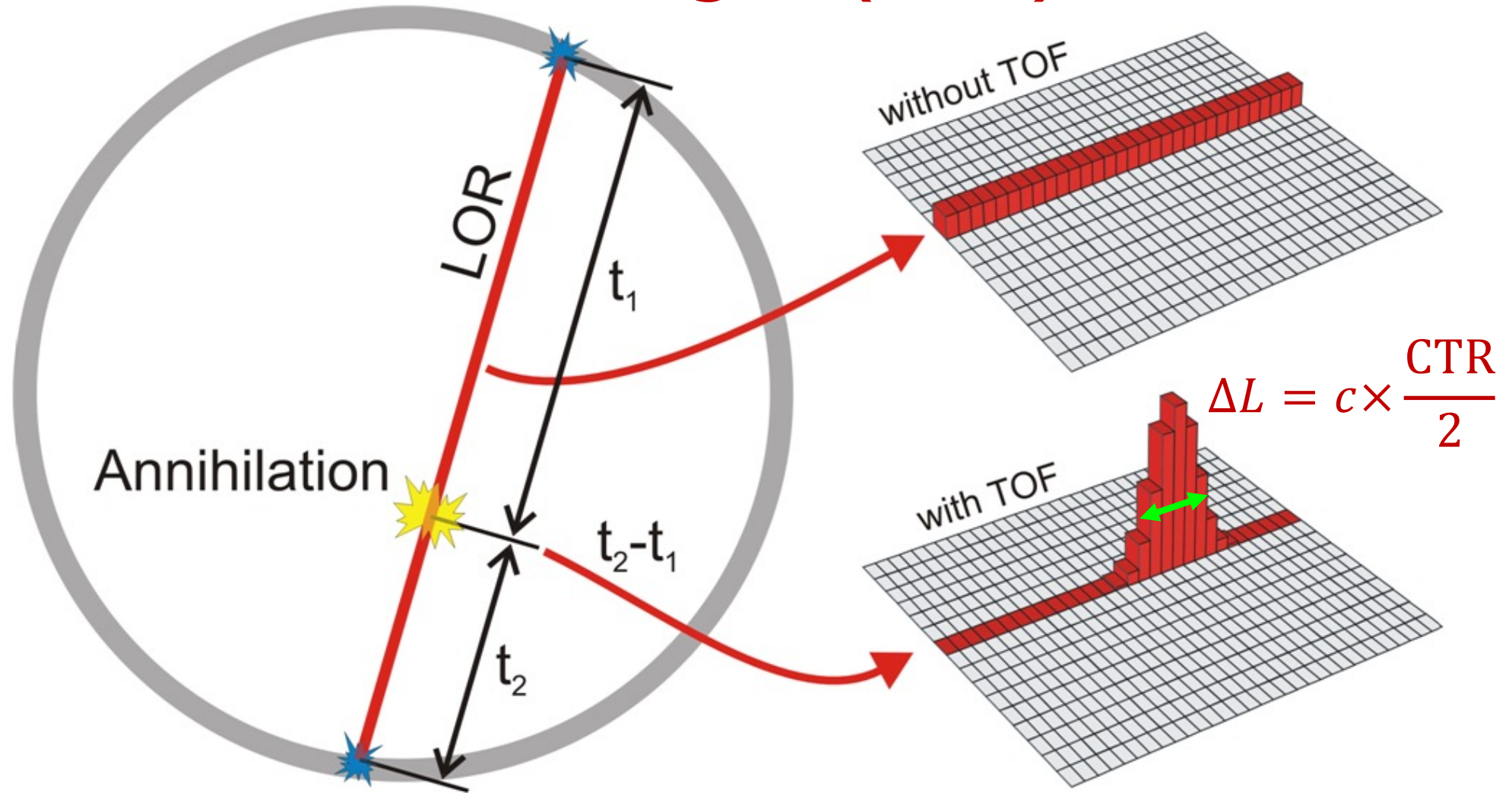
# of crystals: 243,200  
 # of SiPM arrays (16 x 16): 9,728  
 Coincidence window: 4.7 ns  
 Energy window: 435-585 keV  
 Scatter fraction: 37 %  
 Peak NEC: 3.0 Mcps @ 27.5 kBq/mL (NU 2-2018)

G.A. Prenosil et al., JNM 2021



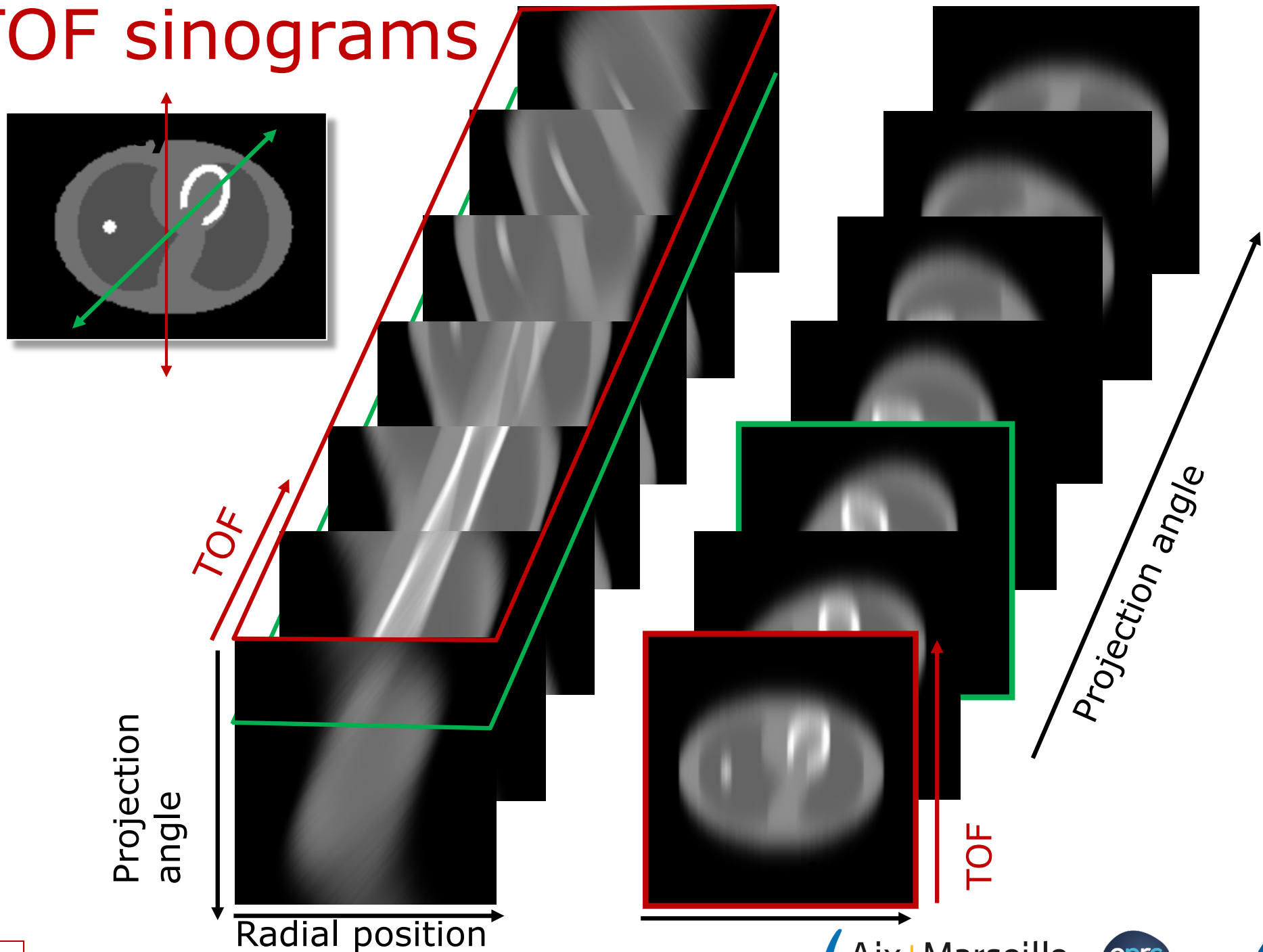
I. Alberts et al., EJNMMI 2021

# Time-Of-Flight (TOF)-PET

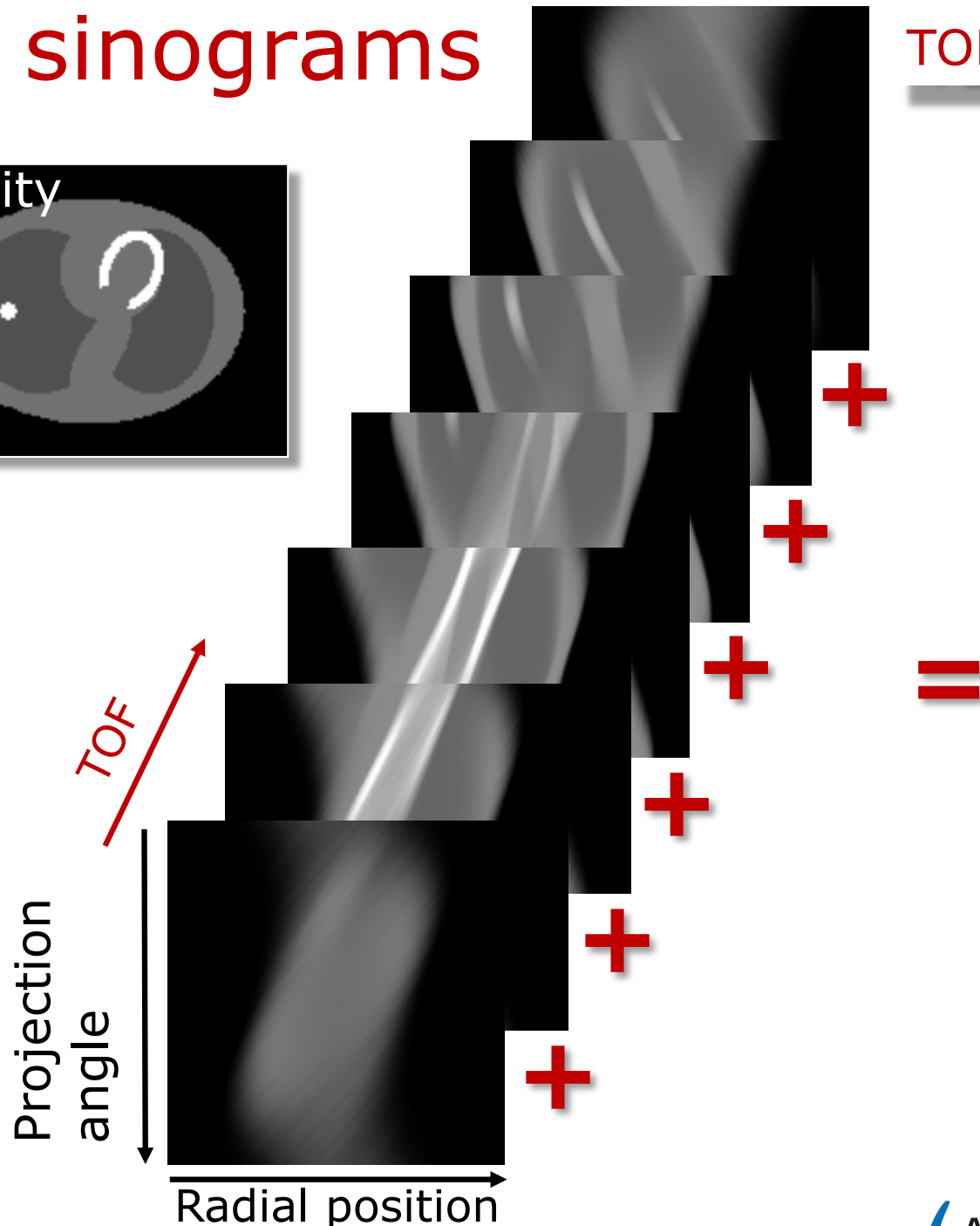


$$30 \text{ cm/ns} \times \frac{10 \text{ ps}}{2} = 1.5 \text{ mm}$$

# TOF sinograms



# TOF sinograms



TOF:  $(s, \theta, z, \delta, t)$

nonTOF:  $(s, \theta, z, \delta)$



nonTOF

=

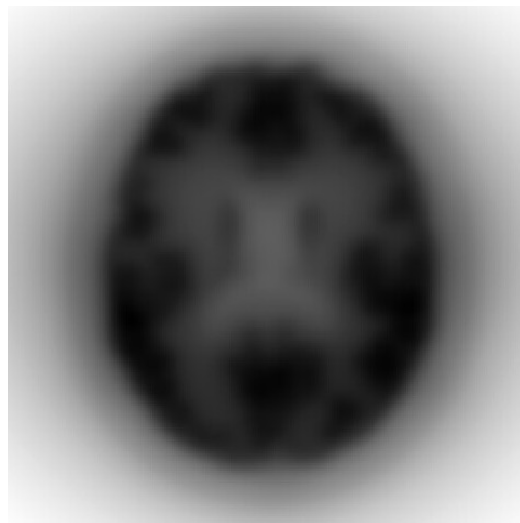


Resolution in TOF-direction  $\sim 1.5$  mm  
Resolution in detector direction 5 mm

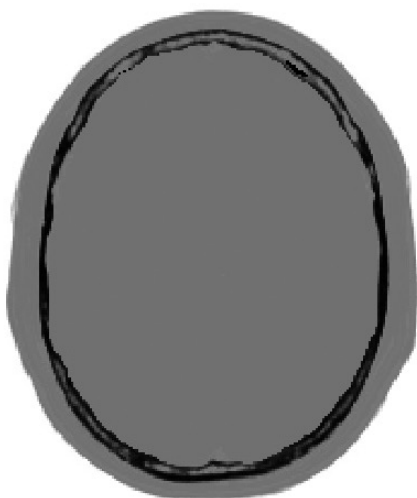
true activity



nonTOF backproj



TOF backproj



true attenuation



nonTOF OSEM



TOF OSEM

# Using only the vertical projection angle

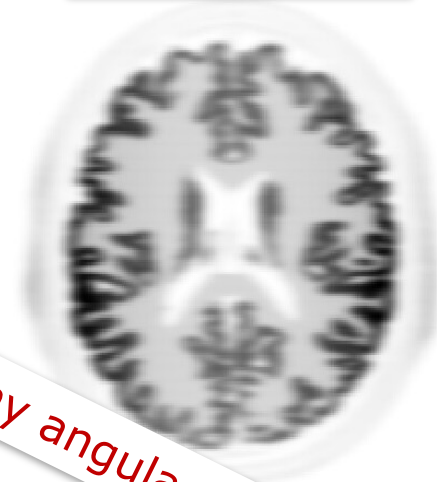
true activity



nonTOF backproj

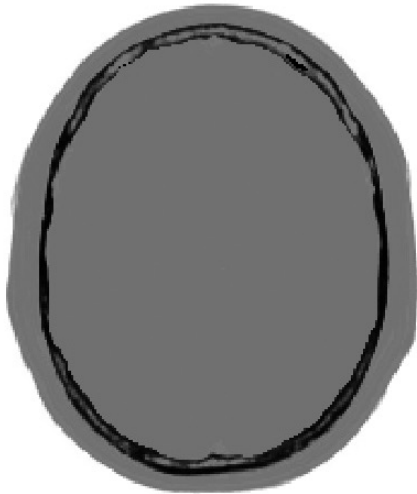


TOF backproj

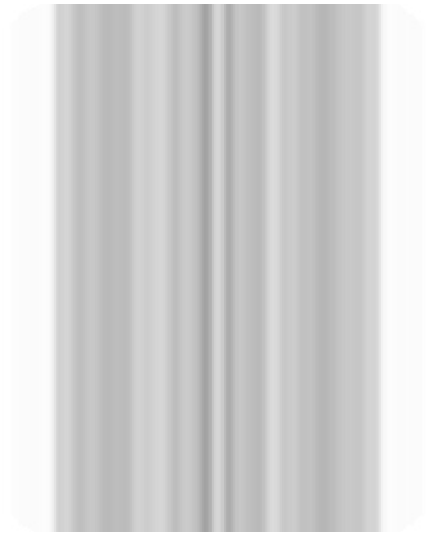


any angular sampling is fine

true attenuation



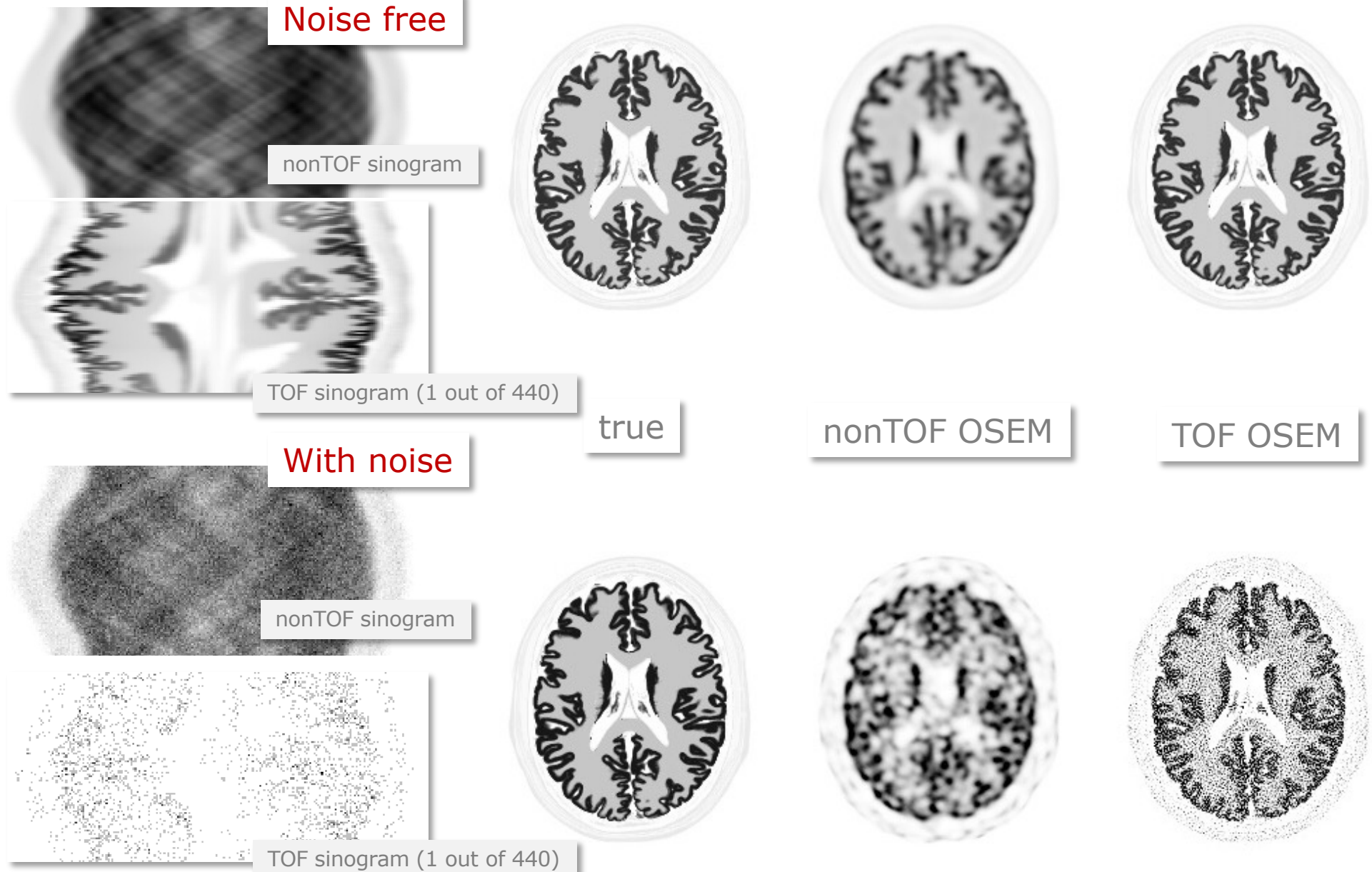
nonTOF OSEM



TOF OSEM



# Signal to Noise Ratio (SNR)



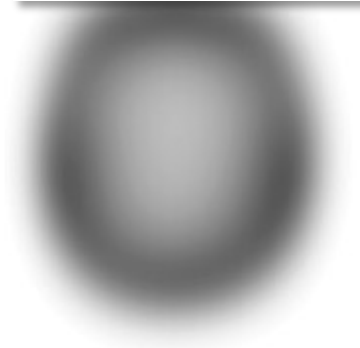
Courtesy: J. Nuyts, Univ Leuven



true activity



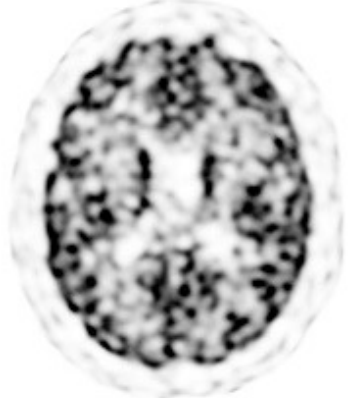
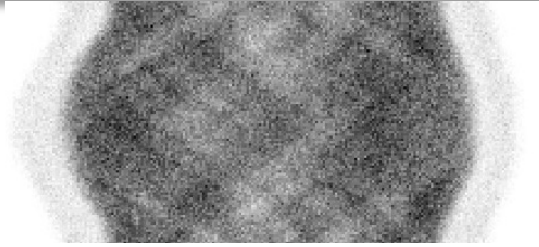
nonTOF OSEM



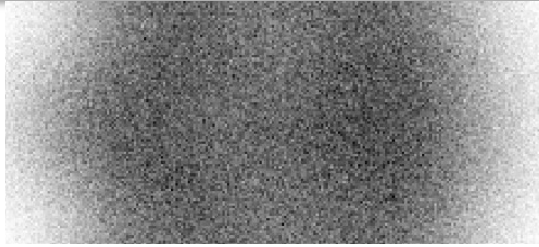
TOF OSEM



10 ps CTR  
5 mm detector blur



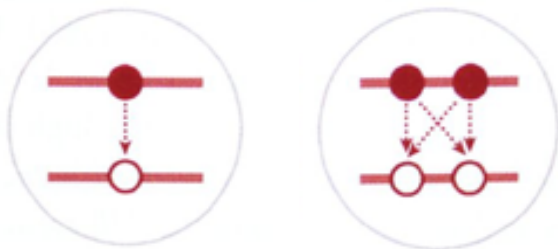
10 ps CTR  
55 mm detector blur



Courtesy: J. Nuyts, Univ Leuven

# Sources possibles de photons prompts (< 1 ns)

Exciton/bi-exciton  
stable at 300 K  
(e.g. CdSe CQwells)



Activators  $\text{Ce}^{3+}$  : 5d-4f  
 $\text{Ca}^{2+}$  &  $\text{Mg}^{2+}$  co-doping  
 $\tau_r \sim 20$  ps  $\tau_d < 16$  ns

Hot intraband  
luminescence  
0.1-10 ps  
(e.g.  $\text{PbWO}_4$ ,  
 $\text{CaWO}_4$ )

Cross luminescence  
< 1 ns (e.g.  $\text{BaF}_2$ )  
< 300 nm  
small light yields

$$\frac{1}{\tau} = \frac{4e^2}{3\hbar c^3} \omega_{21}^3 |\vec{r}_{21}|^2$$

High donor band  
semiconductors  
< 1 ns (e.g.  $\text{ZnO}$ )  
deactivated at  
room temperature

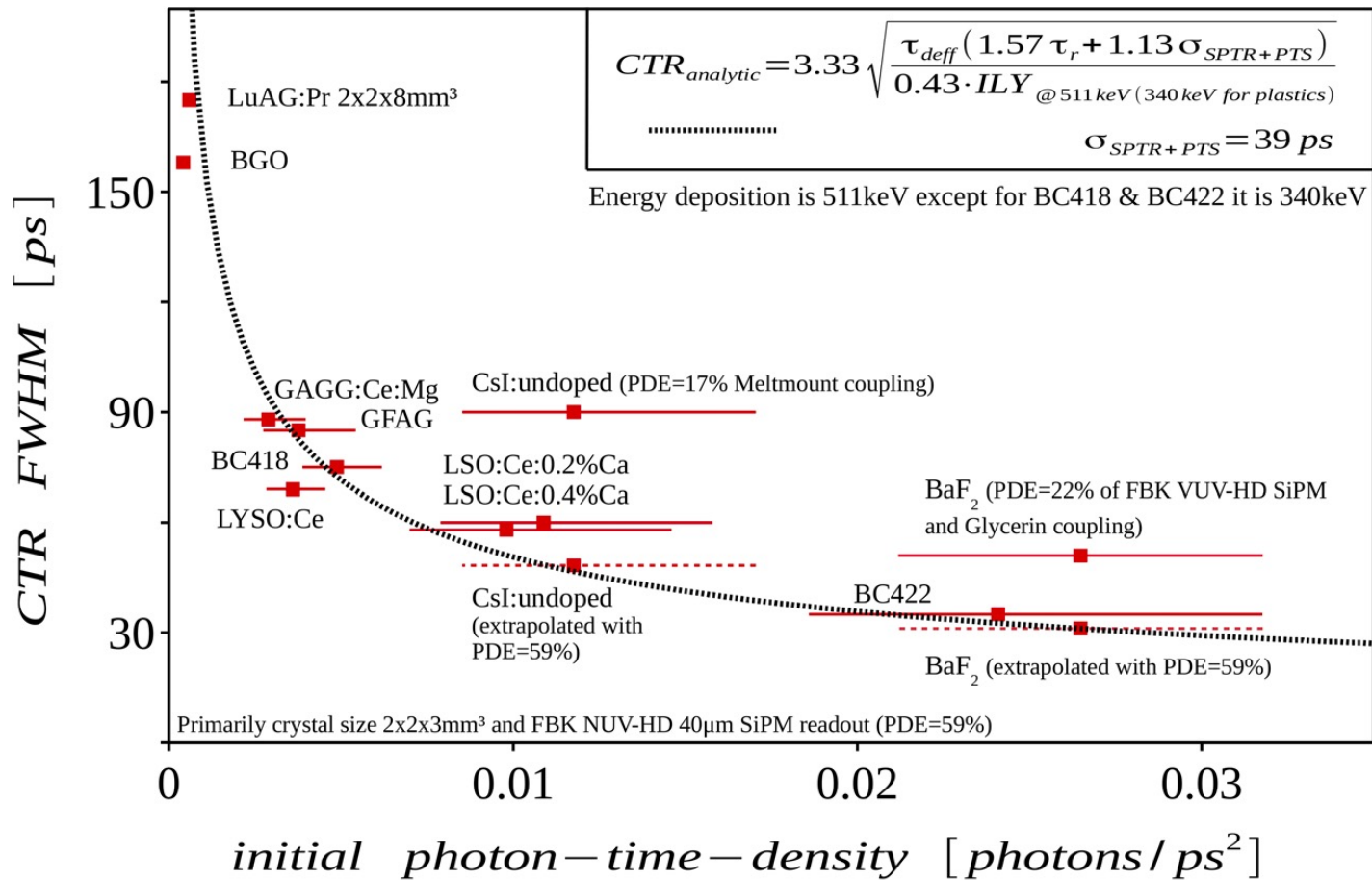
Cherenkov emission  
 $\tau \sim 5-10$  ps

Courtesy: P. Lecoq, CERN



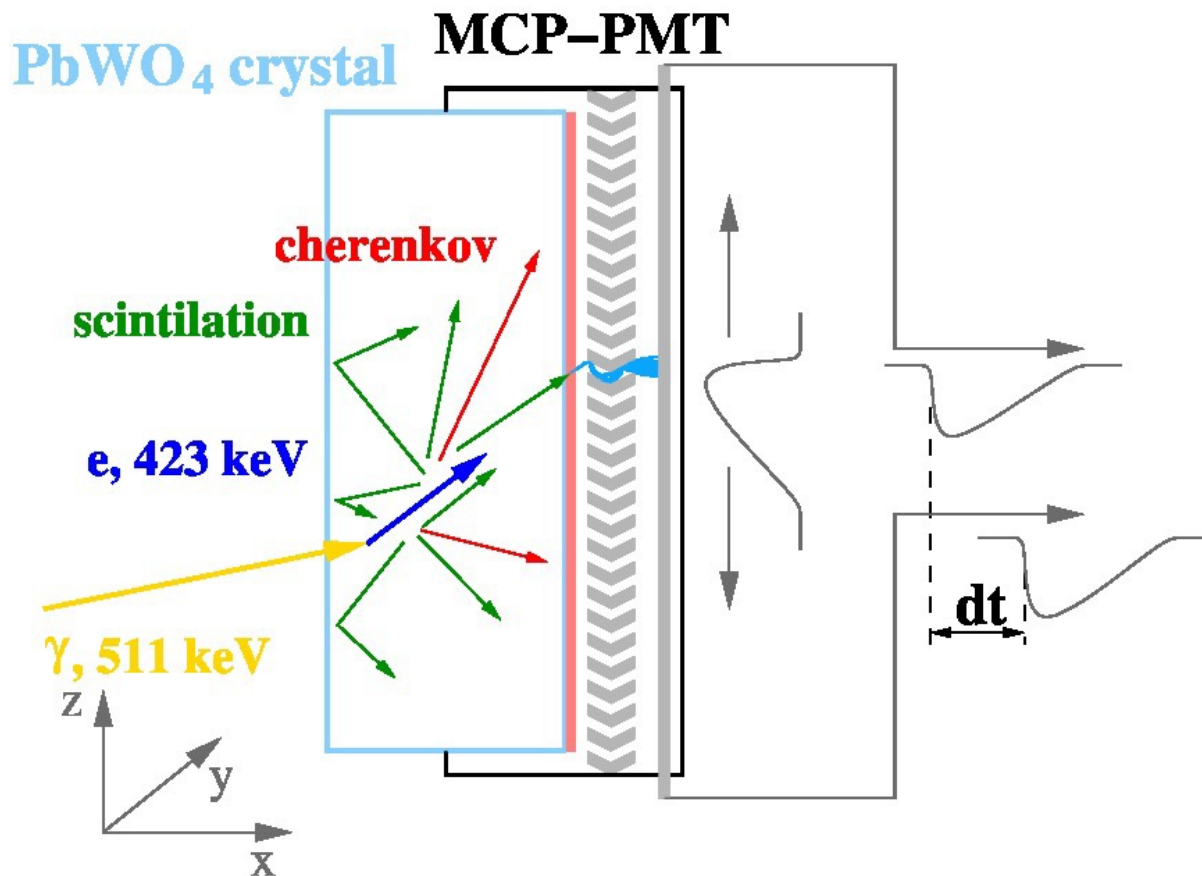
# State-of-the-art CTR measurements

$$CTR = 2.35\sqrt{2} / \sqrt{\underbrace{PDE}_{\sim 59\%} \times \underbrace{LTE}_{\sim 73\%} \times \underbrace{IPTD}}_{= \frac{LY_{@energy}}{\tau_d (1.57\tau_r + 1.13\sigma_{SPTR+PTS})}}$$



S. Gundacker et al., Phys. Med. Biol. 65 (2020) 025001  
 S. Vinogradov et al., Nucl. Inst. Meth. A 912 (2018) 149-153

- Detection of scintillation and **Cherenkov** photons emitted in PWO
- Direct deposition of a photocathode ( $n \sim 2,7$ ) on the crystal surface ( $n \sim 2,3$ )
- Encapsulation within a **Micro-Channel Plate Multiplier Tube (MCP-MT)**
- Coincidence Time resolution (CTR)  $\sim 20$  ps FWHM (exluding MCP-MT)

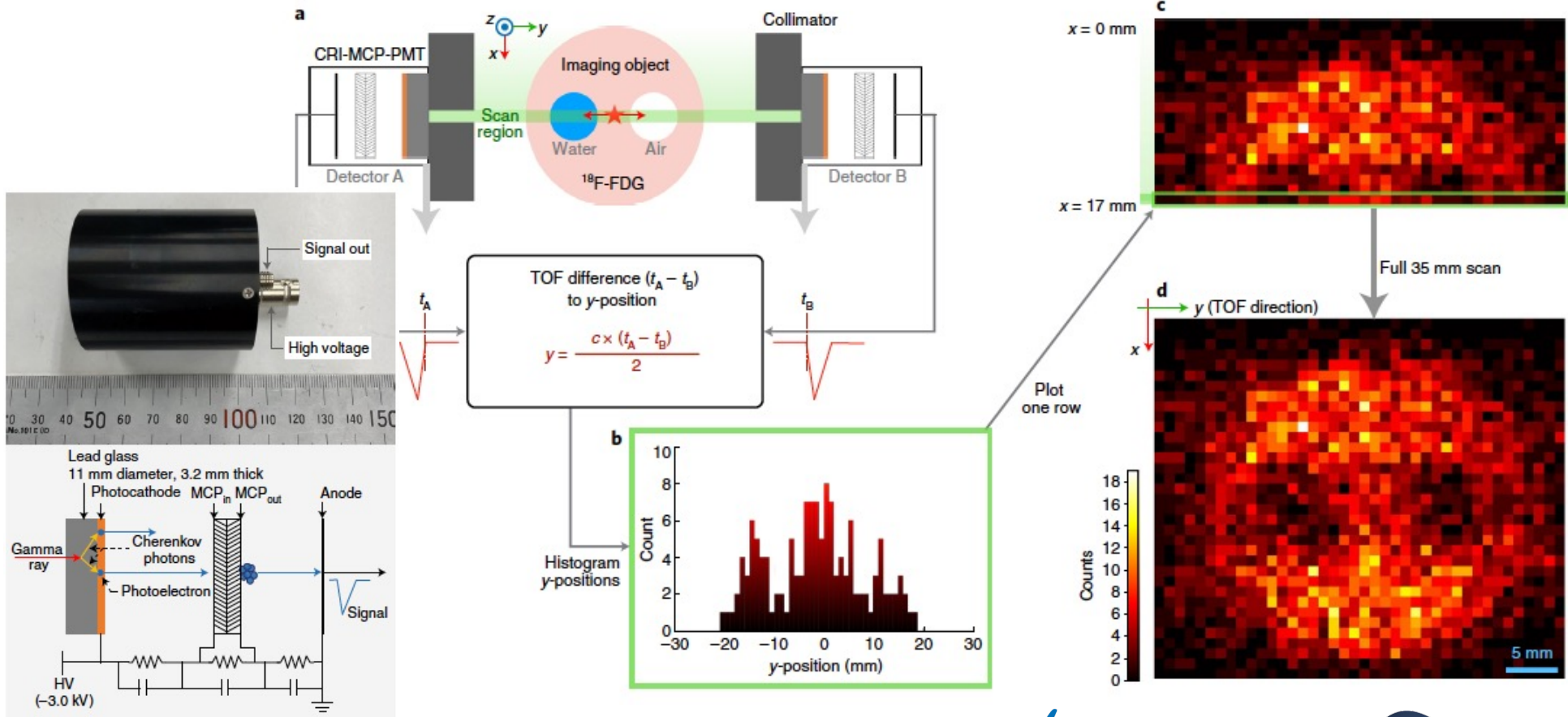
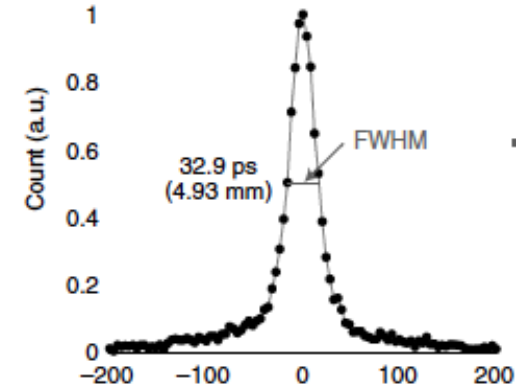


D. Yvon et al., JINST 15 (2020) P07029



# Reconstruction-free positron emission tomography

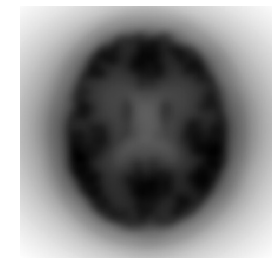
- Use of Cherenkov light for timing
- CTR 32.9 ps FWHM (4.93 mm)
- **direct Positron Emission Imaging (dPEI)**



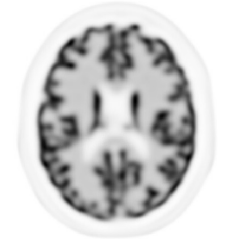
S.I. Kwon et al., Nat. Photon. 2020

# The 10 ps challenge: a step toward reconstruction-less TOF-PET

Thank you



nonTOF  
backproj



nonTOF  
OSEM



10ps TOF  
backproj



10ps TOF  
OSEM