

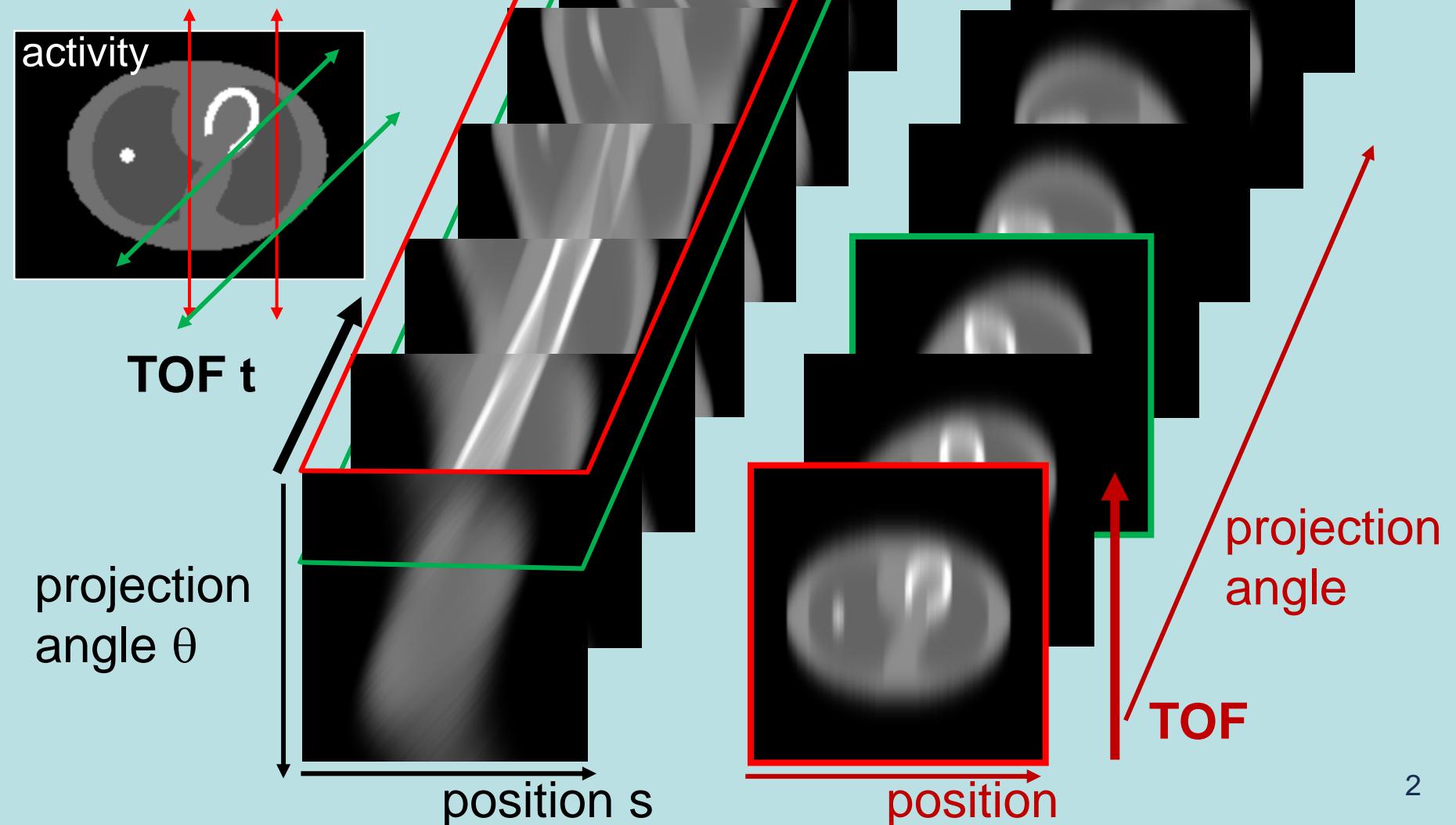


# 10 ps TOF-PET

- TOF sinograms
- no more (iterative) reconstruction?
- attenuation, scatter, randoms
- resolution:
  - image resolution and angular sampling
- SNR
- estimate attenuation from emission with MLAA
  - relax or eliminate need for attenuation measurement
  - illustrates need for accurate system calibration

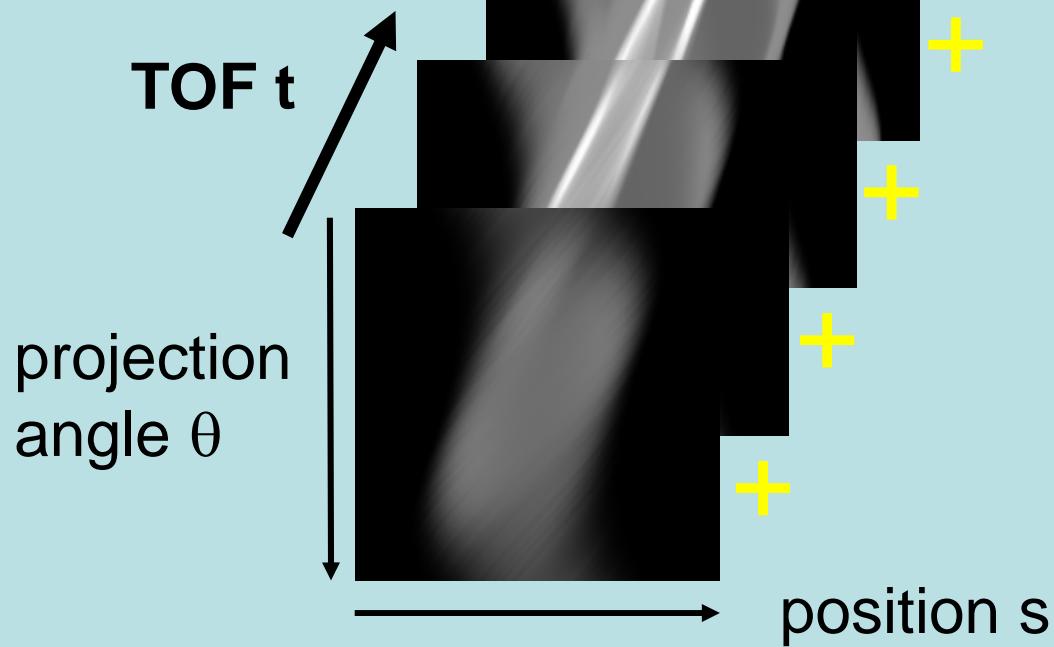


# TOF SPECT



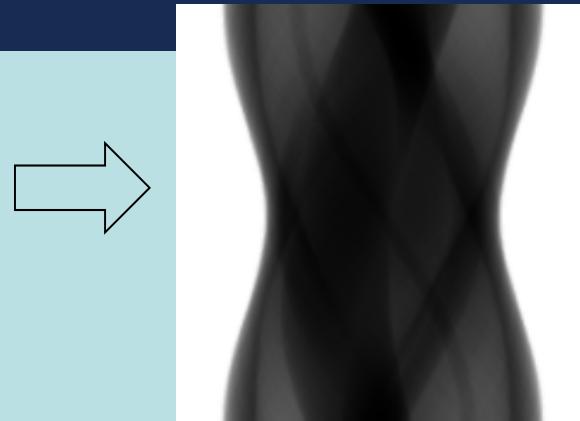
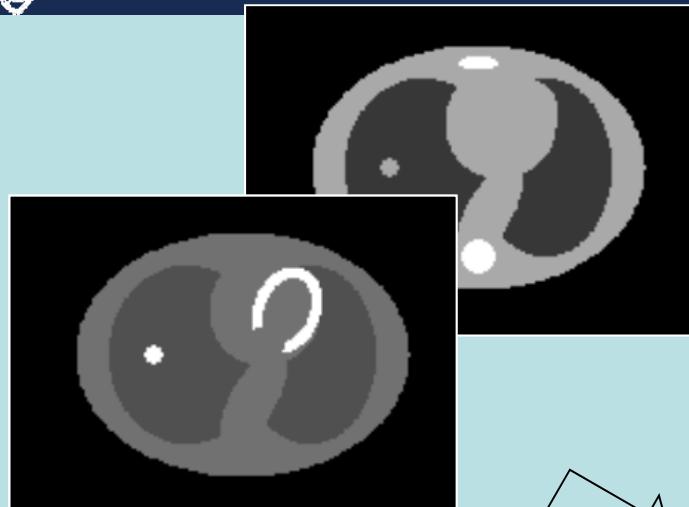


## TOF tomograms

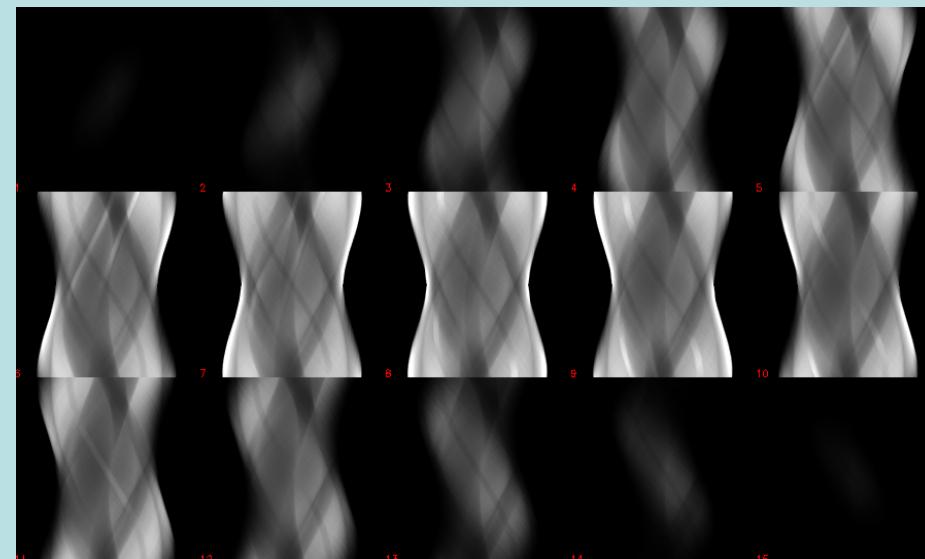
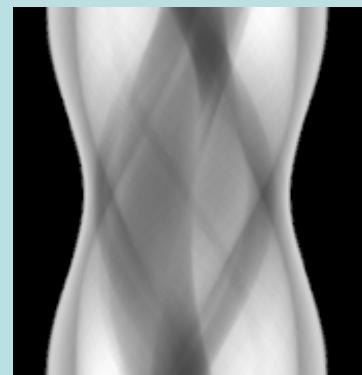




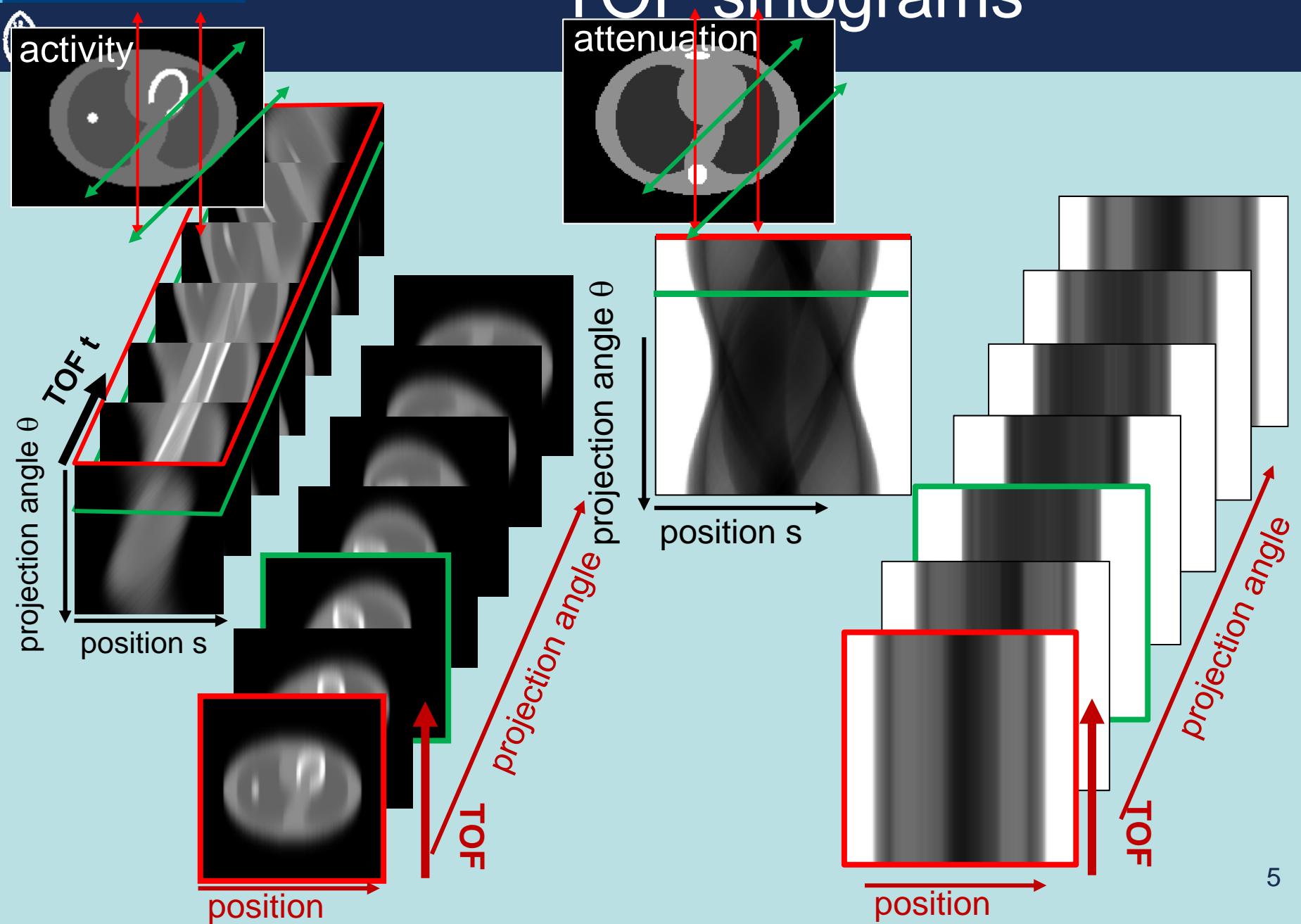
# TOF sinograms



TOF



## TOF sinograms



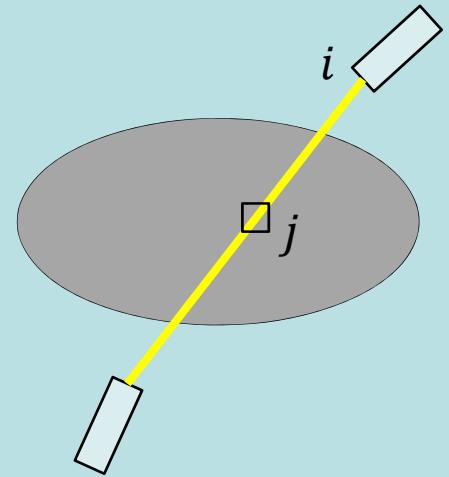


# 10 ps TOF-PET

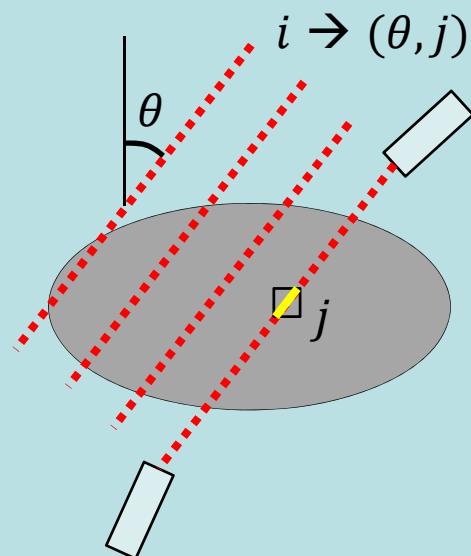
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# no more (iterative) reconstruction?



no TOF  
or  
 $10^2$  ps TOF



10 ps TOF

projection:  $y_i = \sum_j a_{ij} \lambda_j$

backprojection:  $b_j = \sum_i a_{ij} y_i$

MLEM:  $\lambda_j^{(n+1)} = \frac{\lambda_j^{(n)}}{\sum_i a_{ij}} \sum_i a_{ij} \frac{y_i}{\sum_\xi a_{i\xi} \lambda_\xi^{(n)}}$

projection:  $y_{\theta j} = a_{\theta j} \lambda_j$

backprojection:  $b_j = \sum_\theta a_{\theta j} y_{\theta j}$

MLEM:  $\lambda_j^{(n+1)} = \frac{\lambda_j^{(n)}}{\sum_\theta a_{\theta j}} \sum_\theta a_{\theta j} \frac{y_{\theta j}}{a_{\theta j} \lambda_j^{(n)}}$

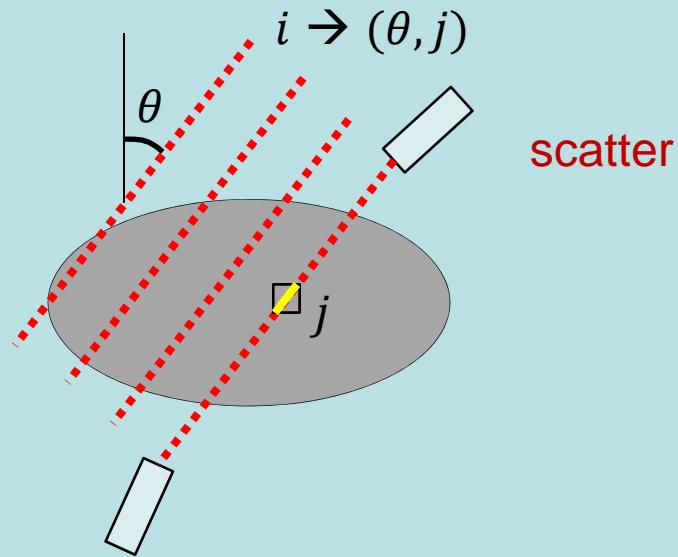
non-iterative!

$$= \frac{\sum_\theta y_{\theta j}}{\sum_\theta a_{\theta j}}$$



# no more (iterative) reconstruction?

10 ps TOF



detector resolution

$$\text{projection: } y_{\theta j} = a_{\theta j} \lambda_j + s_{\theta j}$$

$$\text{backprojection: } b_j = \sum_{\theta} a_{\theta j} y_{\theta j}$$

$$\text{MLEM: } \lambda_j^{(n+1)} = \frac{\lambda_j^{(n)}}{\sum_{\theta} a_{\theta j}} \sum_{\theta} a_{\theta j} \frac{y_{\theta j}}{a_{\theta j} \lambda_j^{(n)} + s_{\theta j}}$$

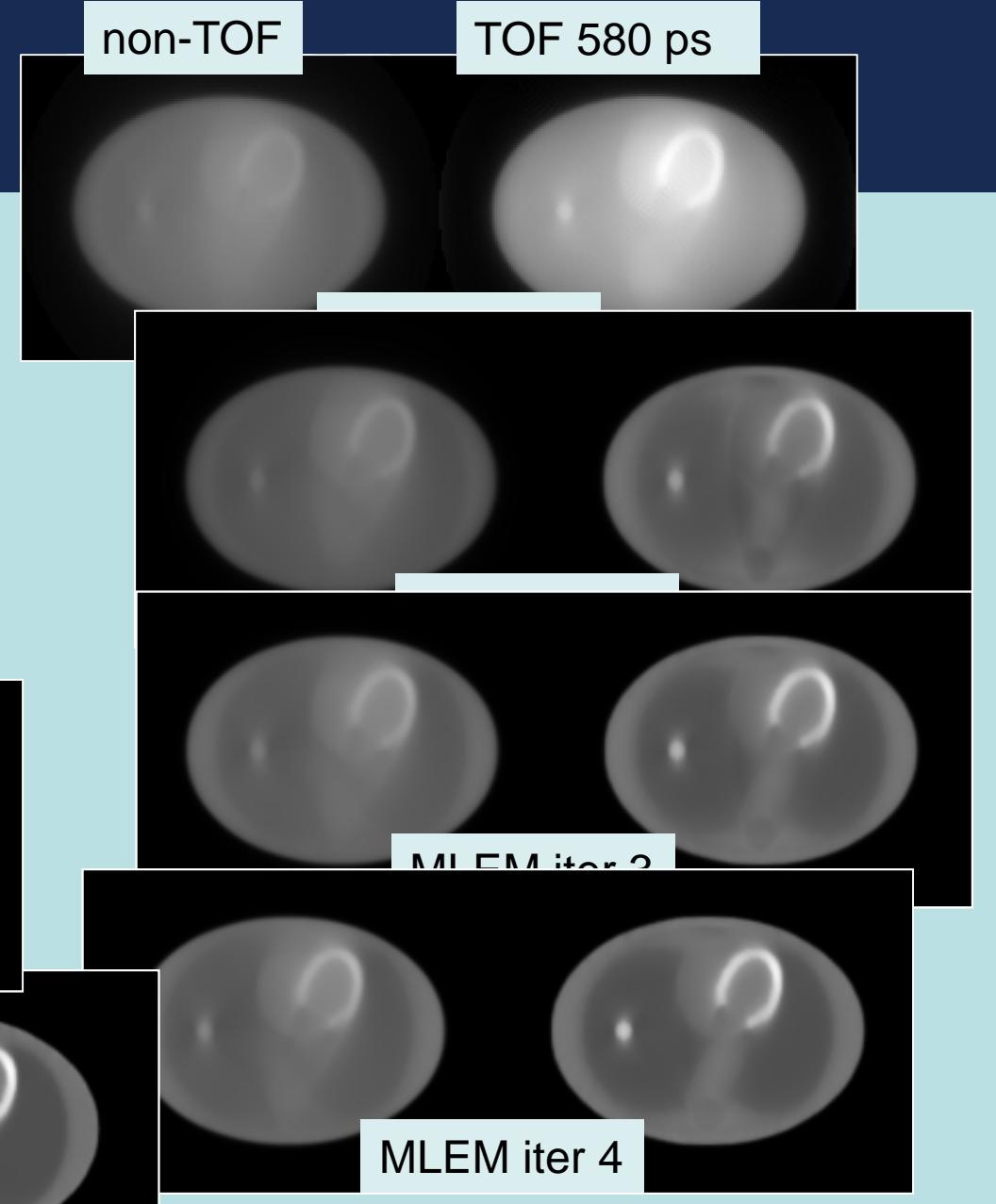
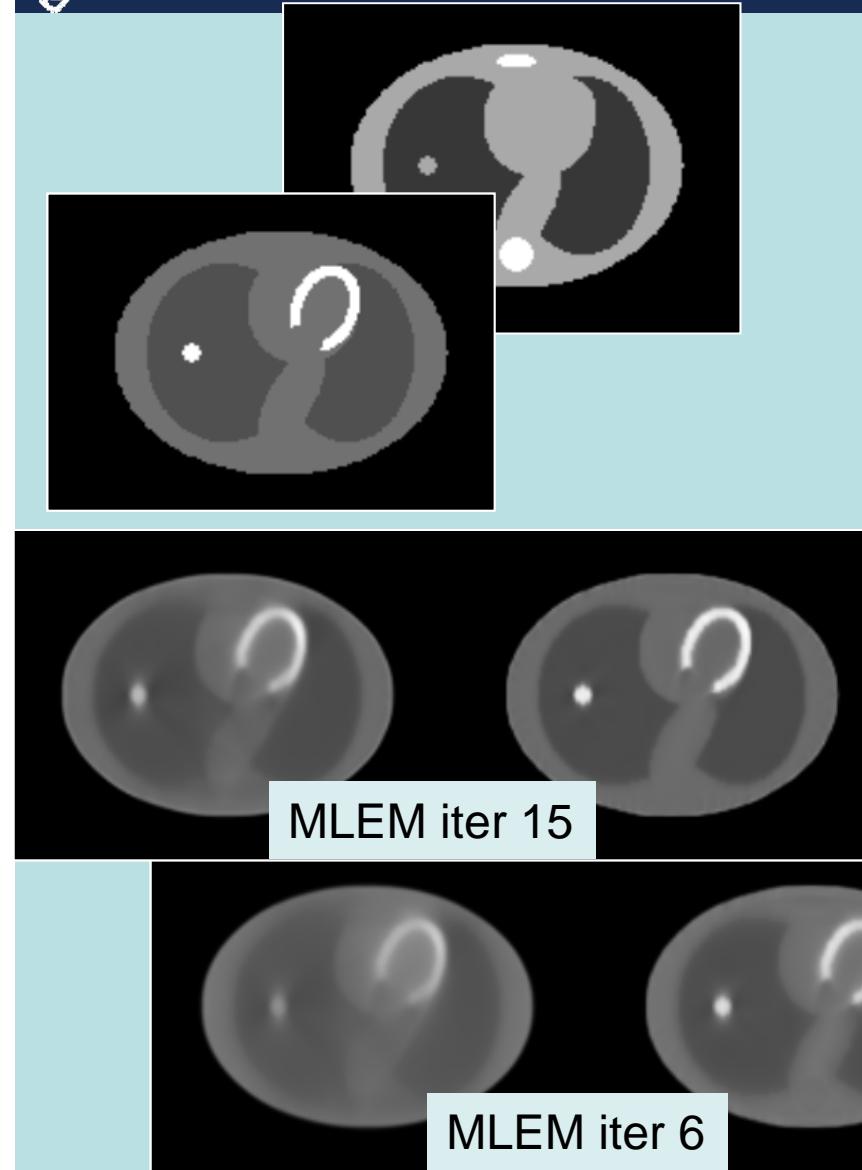
iterative!

$$\text{projection: } y_{\theta k} = \sum_j d_{\theta k j} a_{\theta j} \lambda_j$$

$$\text{backprojection: } b_j = \sum_{\theta k} d_{\theta k j} a_{\theta j} y_{\theta k}$$

iterative!

$$\text{MLEM: } \lambda_j^{(n+1)} = \frac{\lambda_j^{(n)}}{\sum_{\theta k} d_{\theta k j} a_{\theta j}} \sum_{\theta k} d_{\theta k j} a_{\theta j} \frac{y_{\theta k}}{\sum_{\xi} d_{\theta k \xi} a_{\theta \xi} \lambda_{\xi}^{(n)}}$$

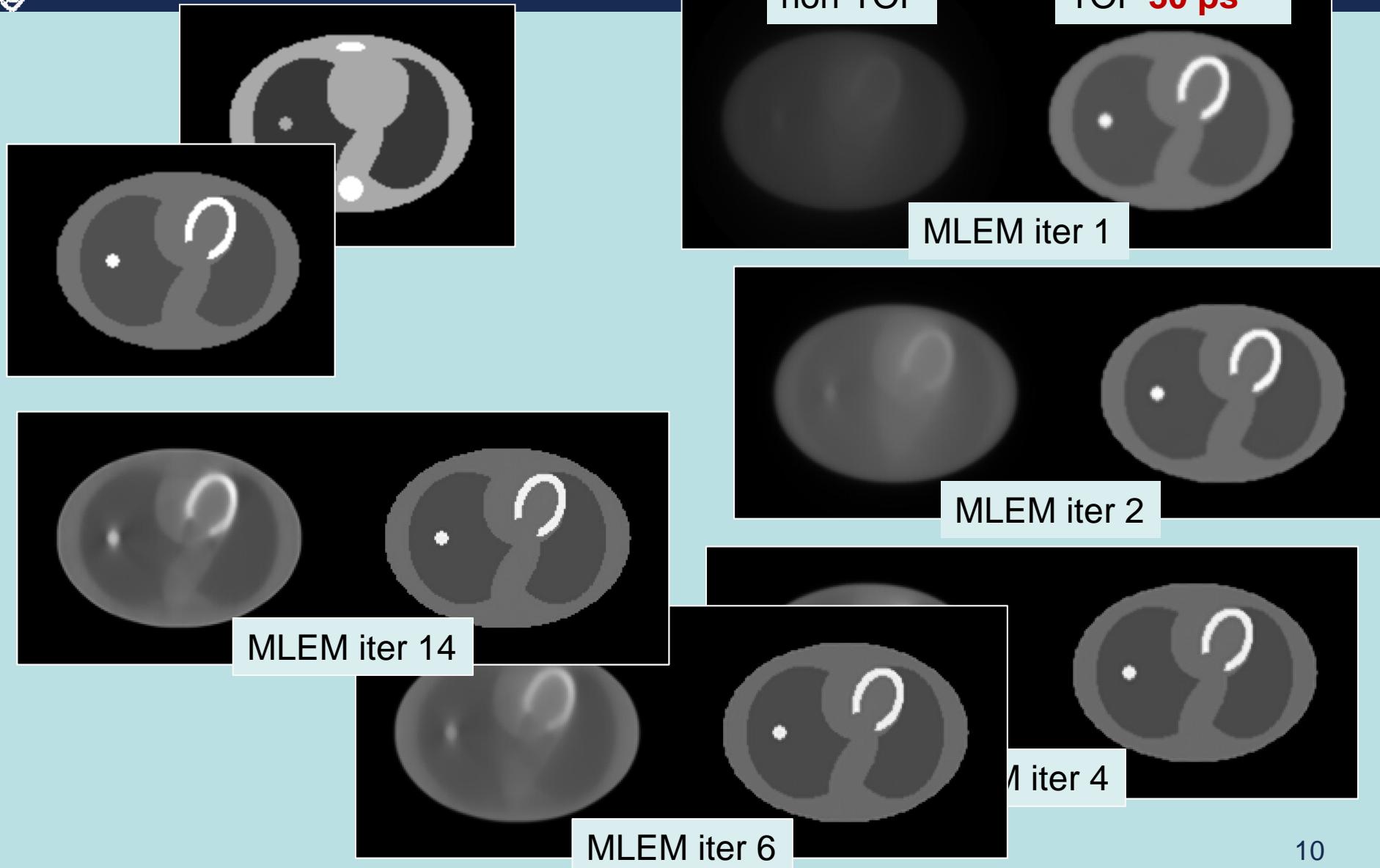




# TOF-PET

non-TOF

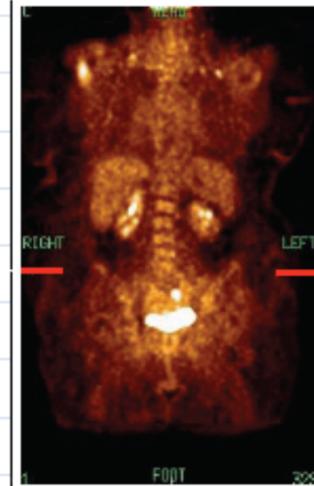
TOF **30 ps**



# Improvement in lesion detectability with TOF

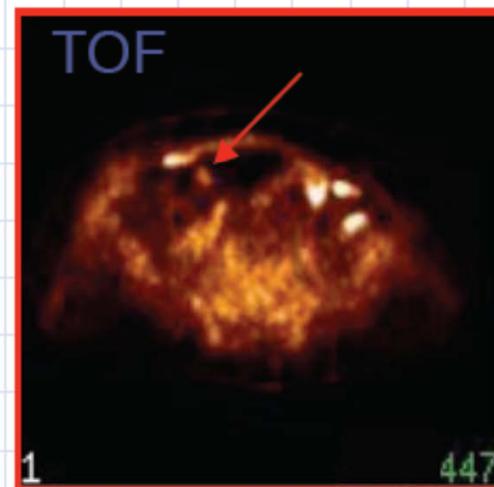
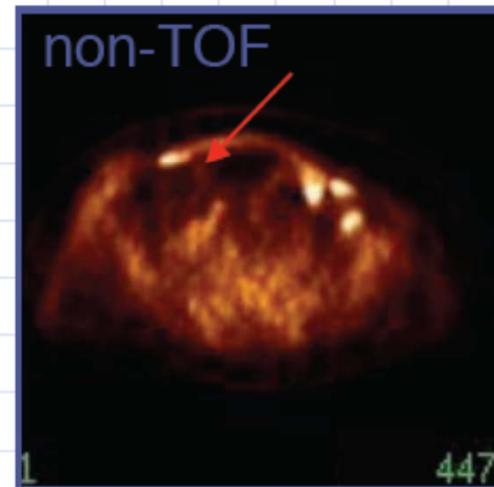
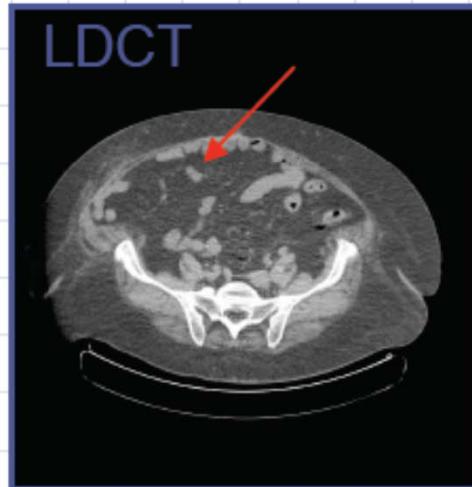


Heavy-weight  
patient study



Colon cancer  
119 kg, BMI = 46.5  
13 mCi, 2 hr post-inj  
3 min/bed

\*Gemini TF



Department of Radiology

UNIVERSITY OF PENNSYLVANIA





# 10 ps TOF-PET

- TOF sinograms
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  - relax or eliminate need for attenuation measurement
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## attenuation

activity

attenuation

no attenuation corr.

non-TOF

TOF 580 ps

TOF 10 ps

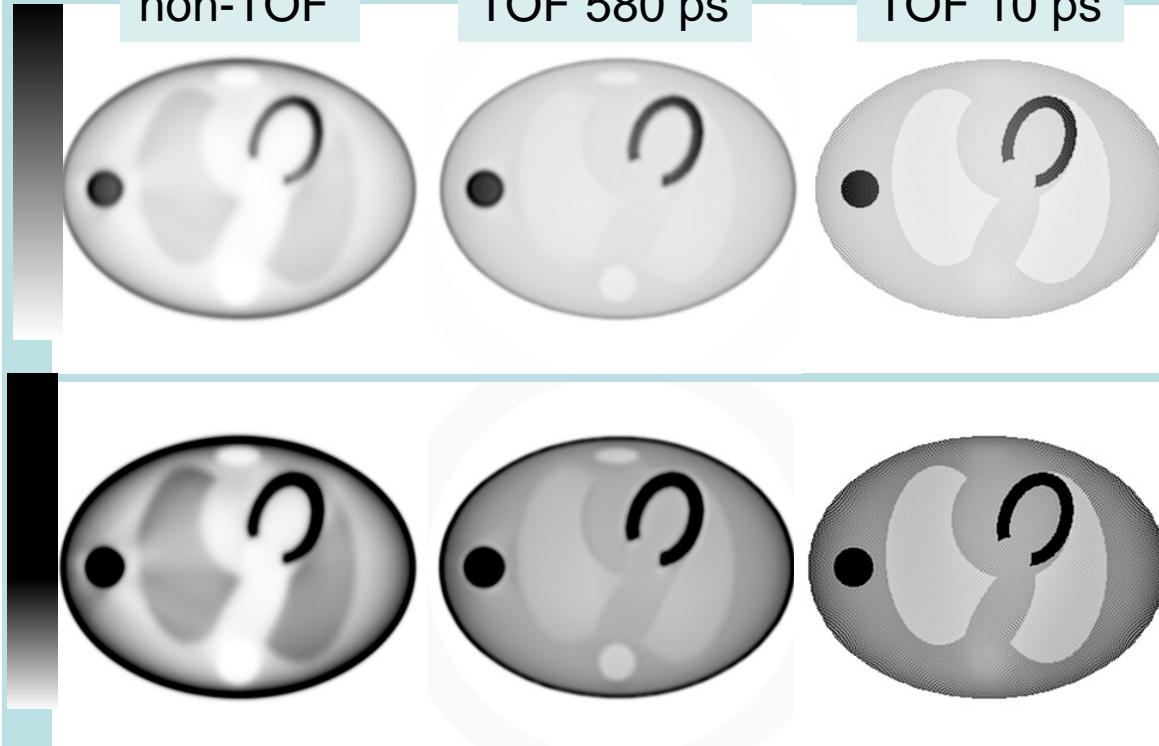
with attenuation corr.

non-TOF

TOF 580 ps



TOF 10 ps



Conti M 2011 Why is TOF PET reconstruction a more robust method in the presence of inconsistent data? *Phys Med Biol* 2011, 56: 155-168



# scatter and randoms

randoms fraction ~ total coincidence window ~ diameter of the FOV

- independent of TOF
- “easily” and accurately estimated
  - from delayed coincidences
  - or from singles
- randoms fraction up to 50% in clinical scans

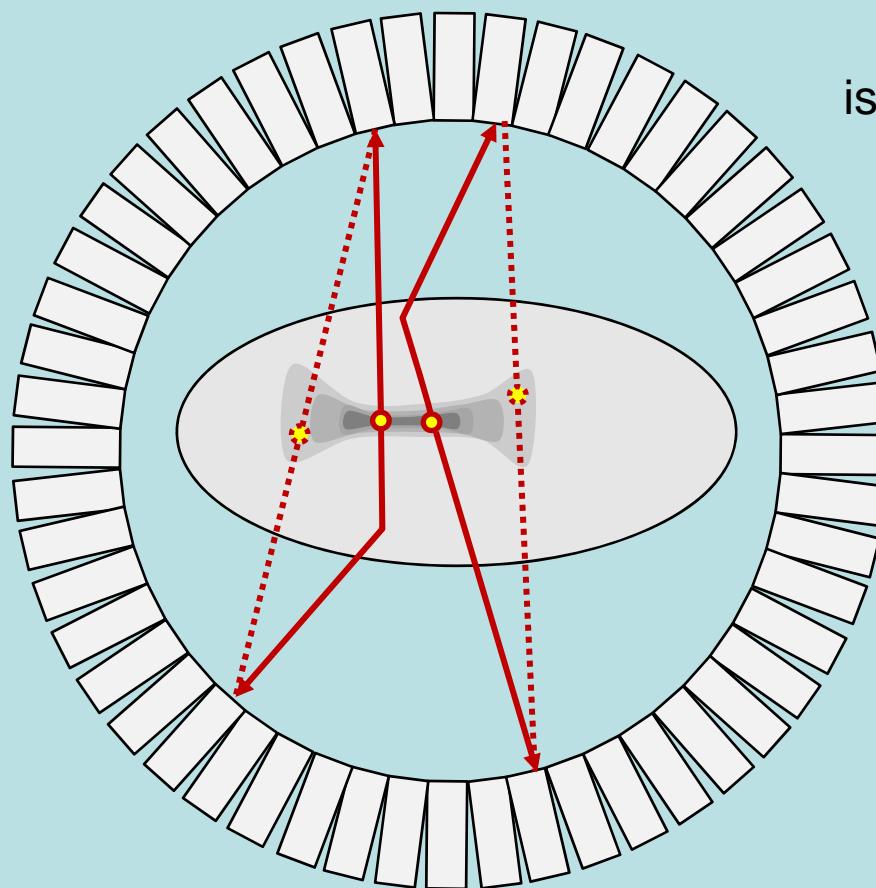
scatter fraction ~ energy resolution

- dependent on TOF
- very difficult to estimate
  - accelerated Monte Carlo
  - energy measurement?
- scatter fraction 40 .. 50% in clinical scans



# scatter and randoms

scatter estimation software usually assumes that scatter is smooth



is it smooth in TOF direction?

Maybe not

- harder to estimate
- may contain more information

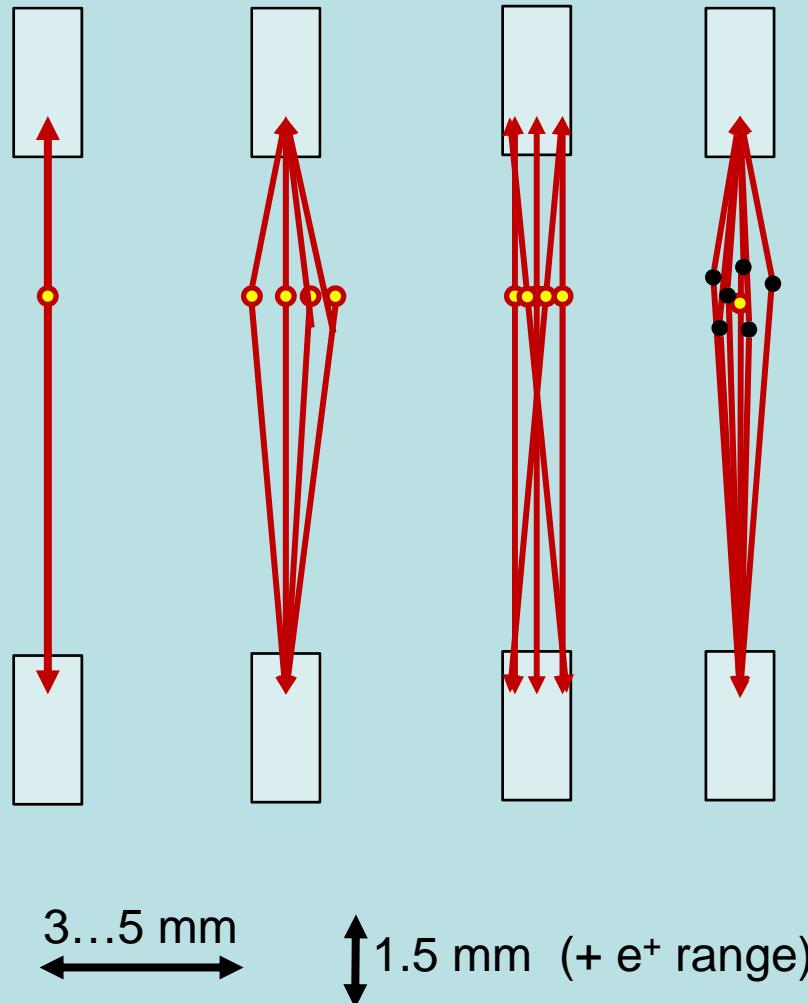


# 10 ps TOF-PET

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# resolution



Depth of interaction:  
measured to achieve 10 ps TOF

acollinearity:  
2D blurring perpendicular to TOF-axis  
 $\sim 0.0022 \text{ mm/mm} * \text{diameter}$

detector resolution:  
2D blurring perpendicular to TOF-axis  
 $\sim \text{crystal\_width} / 2$

positron range:  
3D blurring!

<i>isotope</i>	<i>mean [mm]</i>	<i>max [mm]</i>
<sup>11</sup> C:	1.1	4.1
<sup>18</sup> F:	0.6	2.4
<sup>68</sup> Ga:	2.9	8.2
<sup>82</sup> Rb:	5.9	14.1



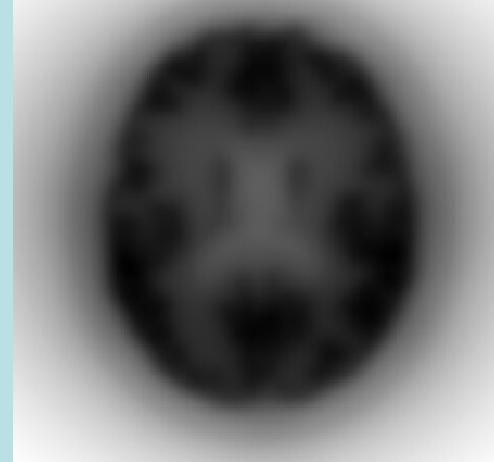
# resolution

resolution in TOF-direction ~1.5 mm, in detector direction 5 mm

true activity



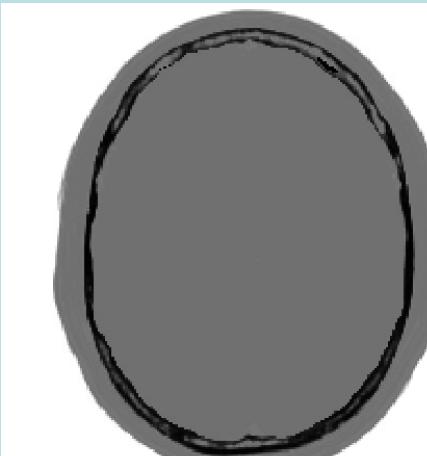
non-TOF backproj



TOF backproj



true attenuation



non-TOF OSEM



TOF OSEM





# resolution

same when only vertical parallel projection is used

true activity



non-TOF backproj

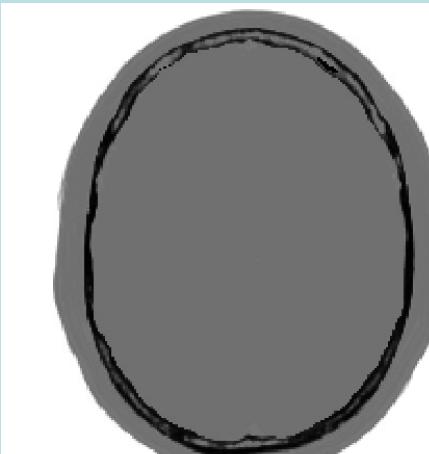


TOF backproj

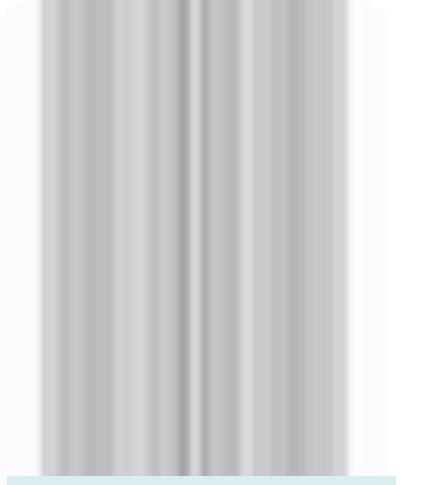


*any angular sampling is fine*

true attenuation



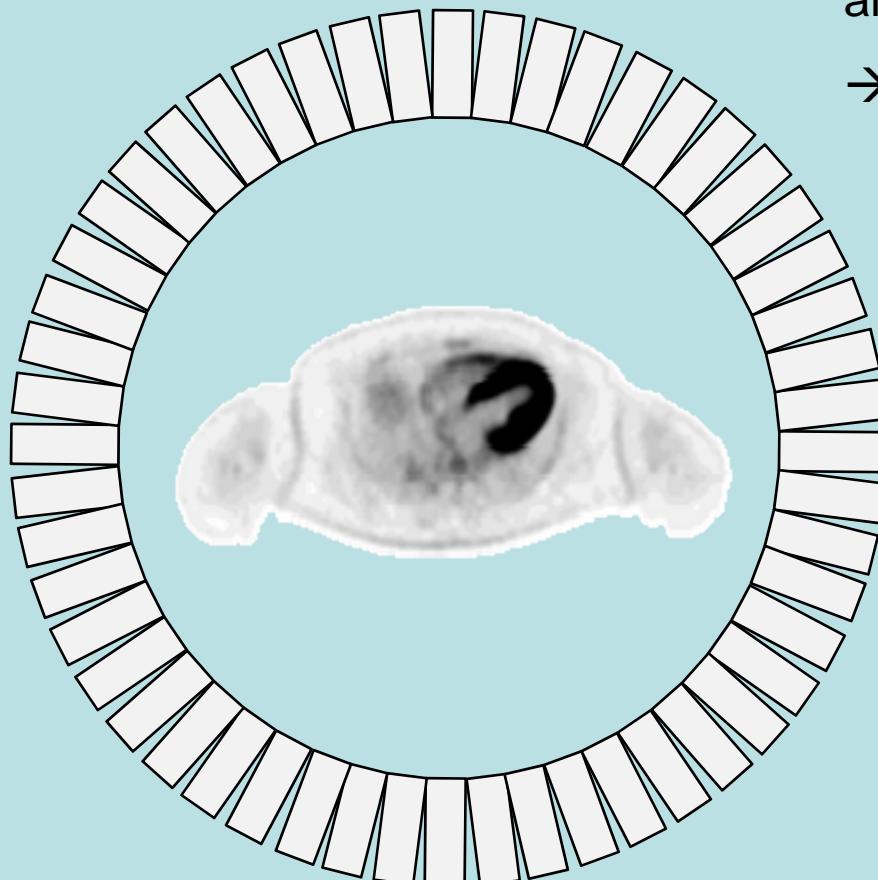
non-TOF OSEM



TOF OSEM



# resolution



angular sampling ± irrelevant  
→ optimize image SNR (or NEC)





# resolution

resolution dominated by TOF → use poor detector resolution

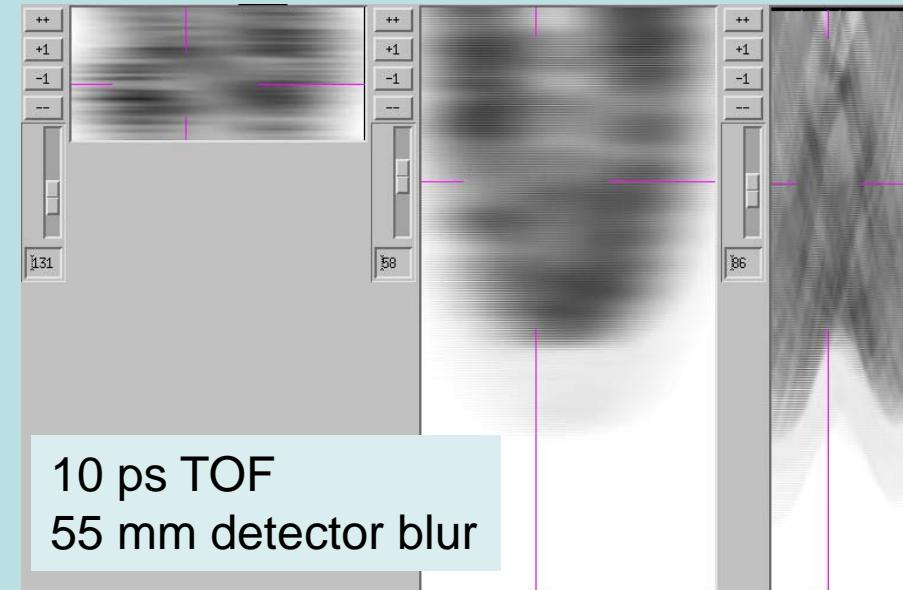
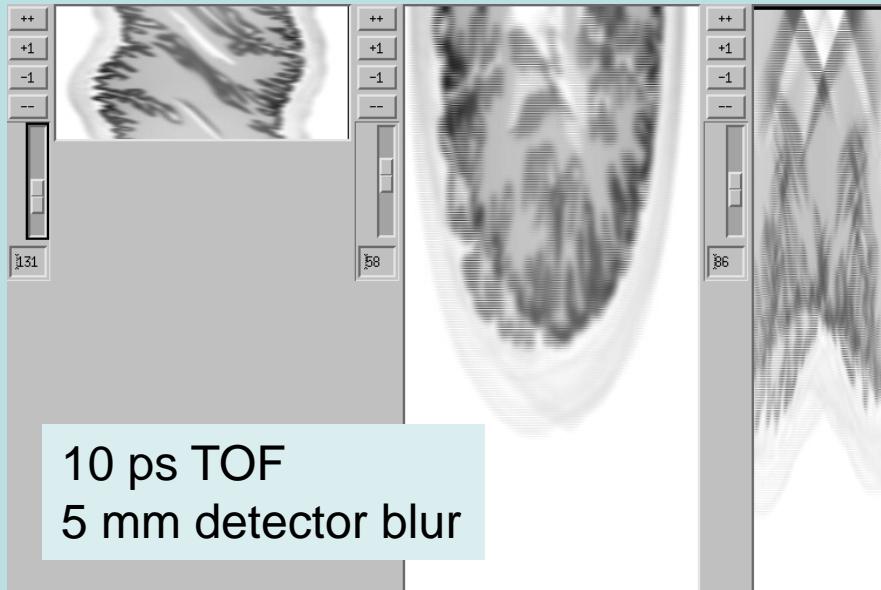
5 mm detector resol



55 mm detector resol



non-TOF  
sinogram





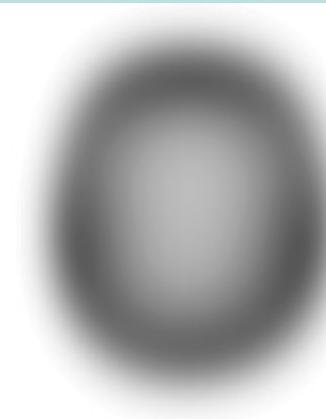
55 mm detector resol



true activity



non-TOF OSEM



TOF OSEM



### M-20-6 - Can Time-of-Flight Information be Used to Mitigate Detector Induced Blur in PET Reconstruction?

M. Toussaint<sup>1,2</sup>, J. - P. Dussault<sup>1</sup>, R. Lecomte<sup>2</sup>

**Session:** M-20 - Image Reconstruction II

**Date:** Saturday, October 28, 2017, 10:20 am

**Room:** Centennial III ()

*IEEE NSS MIC 2017*



# 10 ps TOF-PET

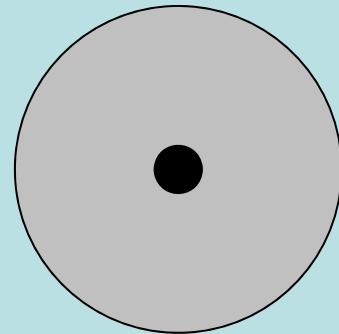
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Improvement in variance with FBP [1]:

for center of uniform disk:

$$\frac{\text{var}_{\text{nonTOF}}}{\text{var}_{\text{TOF}}} \simeq \sqrt{\frac{2 \ln 2}{\pi}} \frac{D}{\Delta x} = 0.66 \frac{D}{\Delta x} \quad (\text{if } \Delta x \ll D)$$



$$D$$



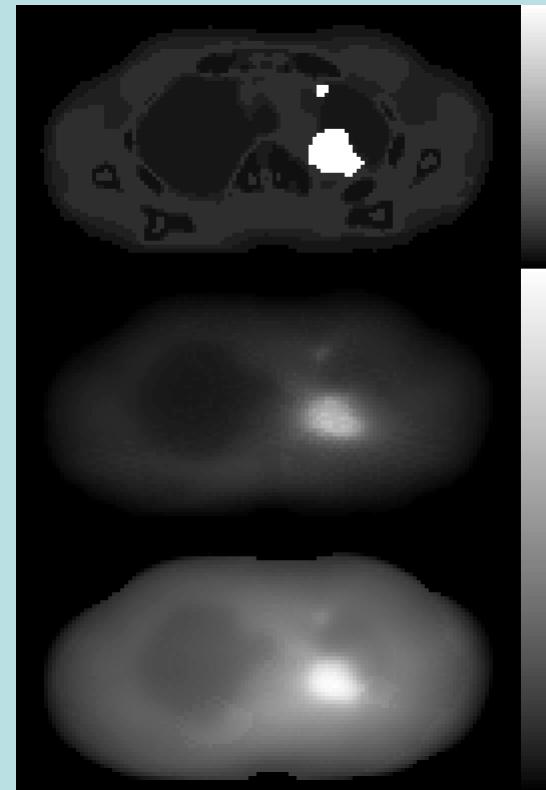
$$\frac{\text{var}_{\text{TOF}1}}{\text{var}_{\text{TOF}2}} \simeq \frac{\Delta x_2}{\Delta x_1}$$

Similar findings for MLEM

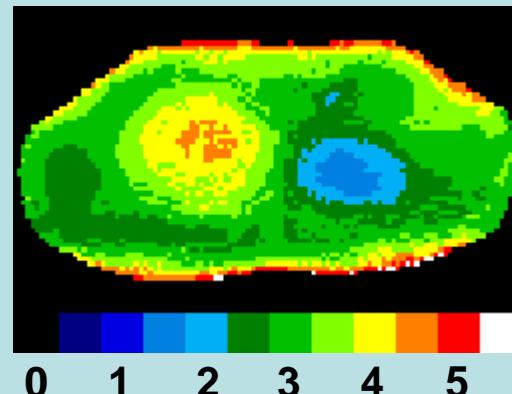


- 2D realistic thorax phantom
- No scatter, no randoms
- Attenuation modeled
- Intrinsic resolution 0.5 cm
- Pixel size 0.3375 cm
- FOV of 64.8 cm
- Target resolution 1.2 cm FWHM

True activity

TOF PET variance  
 $\Delta t = 500 \text{ ps}$ 

Non-TOF PET variance

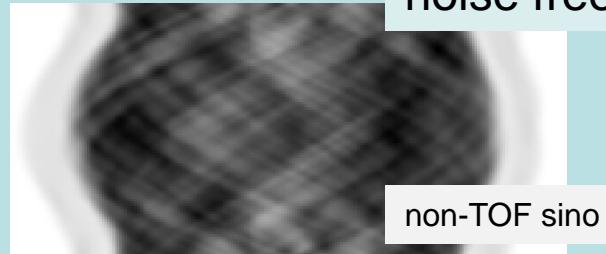


Variance improvement  
due to TOF information



# SNR

noise free



non-TOF sino



TOF sino (1 of 440)



true

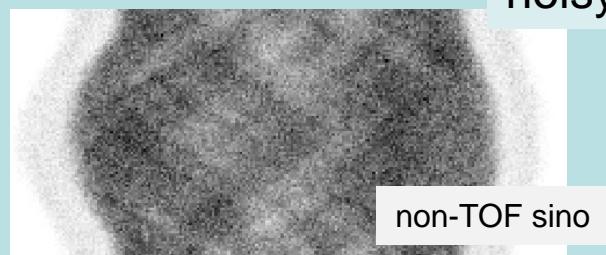


non-TOF



TOF

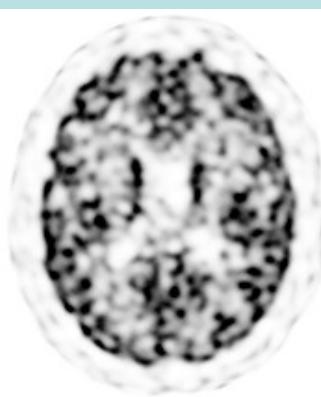
noisy



non-TOF sino

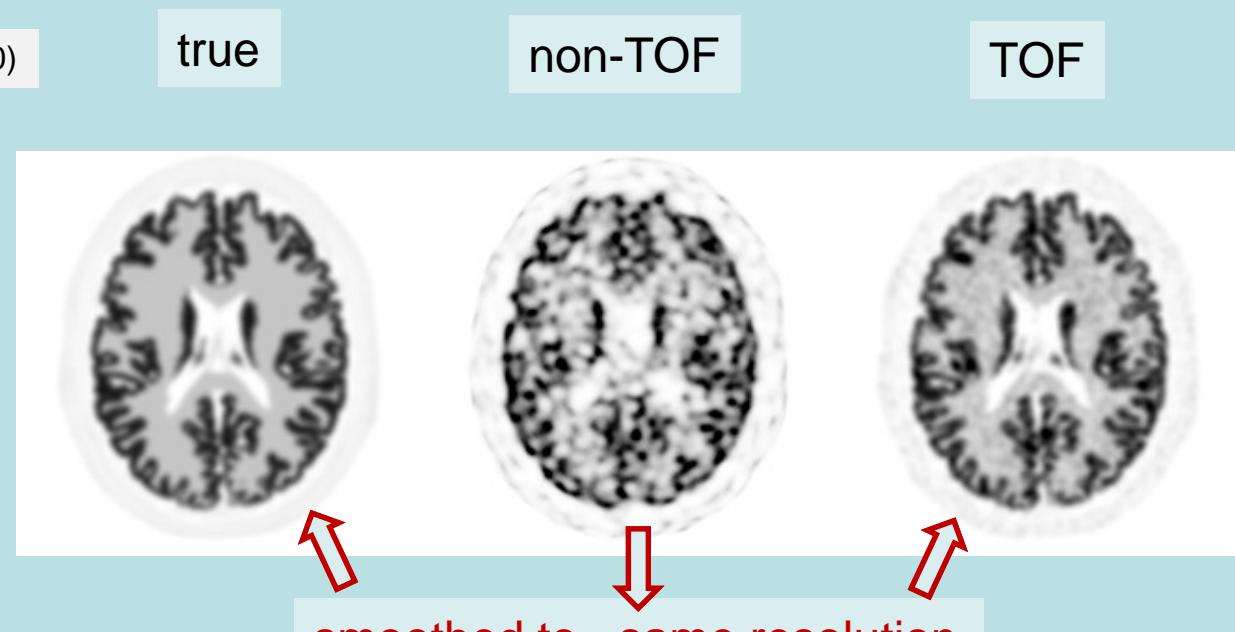
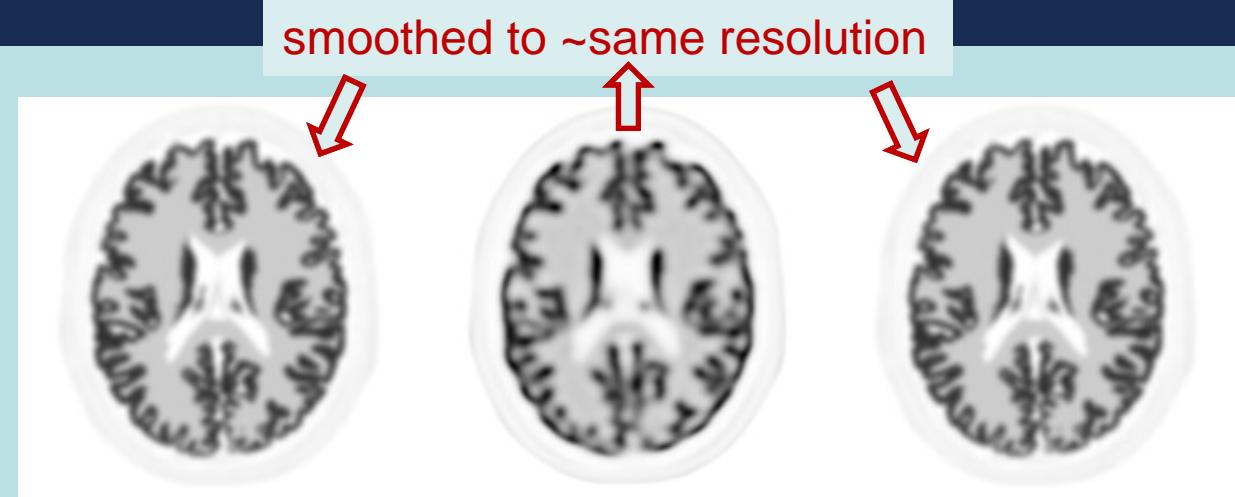
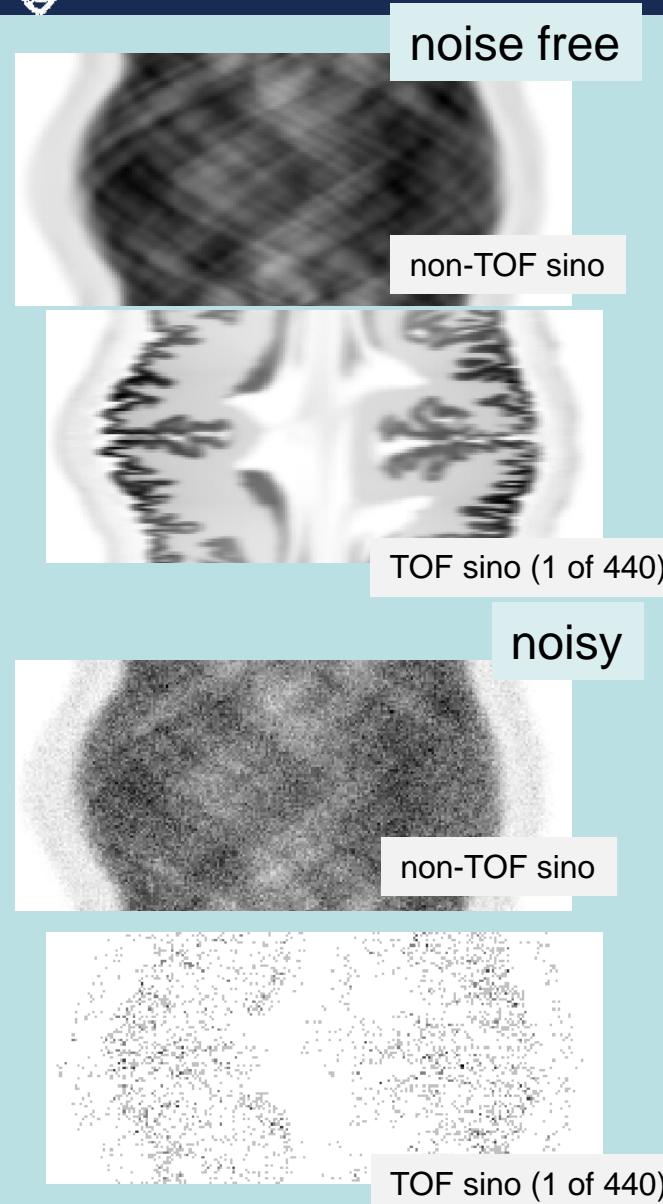


TOF sino (1 of 440)





## SNR





SNR

non-TOF OSEM

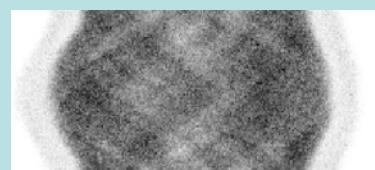
TOF OSEM



10 ps TOF and  
**55 mm** detector  
blur



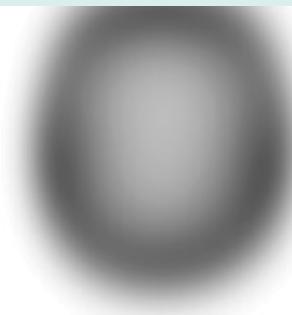
10 ps TOF and  
**5 mm** detector  
blur



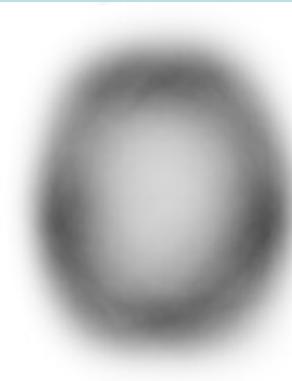
true activity



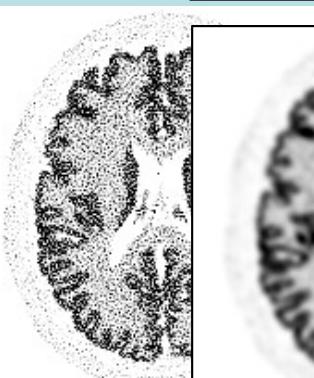
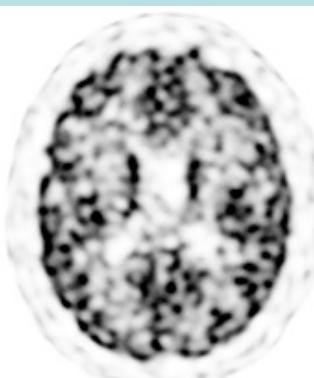
non-TOF OSEM



TOF OSEM



smoothed





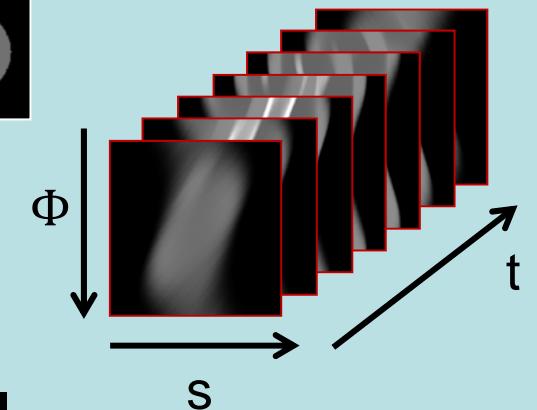
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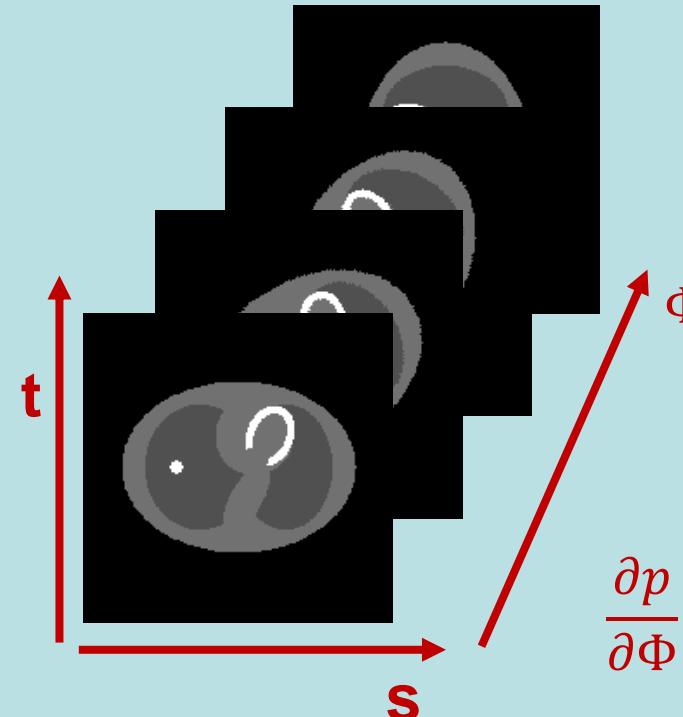
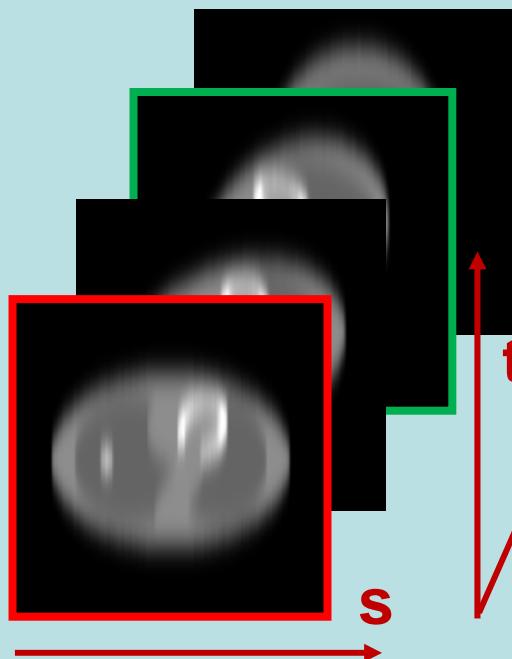


# TOF consistency

$p$  is a consistent “2D”-TOF-PET sinogram



$$\rightarrow Dp = t \frac{\partial p}{\partial s} + \frac{\partial p}{\partial \Phi} - s \frac{\partial p}{\partial t} + \sigma^2 \frac{\partial^2 p}{\partial s \partial t} = 0$$

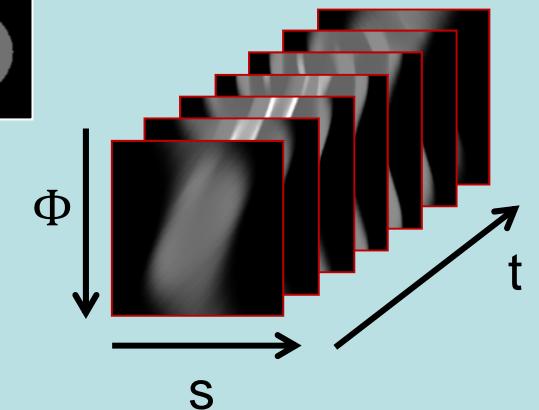


$$\frac{\partial p}{\partial \Phi} = s \frac{\partial p}{\partial t} - t \frac{\partial p}{\partial s}$$



# TOF consistency

$p$  is a consistent “2D”-TOF-PET sinogram



$$\rightarrow Dp = t \frac{\partial p}{\partial s} + \frac{\partial p}{\partial \Phi} - s \frac{\partial p}{\partial t} + \sigma^2 \frac{\partial^2 p}{\partial s \partial t} = 0$$

set  $p_{it} = \frac{y_{it}}{a_i}$

$$a_i = e^{-\sum_j l_{ij} \mu_j}$$

$$D \frac{y_{it}}{a_i} = t \frac{\partial(y_{it}/a_i)}{\partial s} + \frac{\partial(y_{it}/a_i)}{\partial \Phi} - s \frac{\partial(y_{it}/a_i)}{\partial t} + \sigma^2 \frac{\partial^2(y_{it}/a_i)}{\partial s \partial t} = 0$$

$$t \frac{\partial y_{it}}{\partial s} \frac{1}{a_i} - t \frac{y_{it}}{a_i^2} \frac{\partial a_i}{\partial s} + \dots = 0$$

$$\times a_i \quad t \frac{\partial y_{it}}{\partial s} - t \frac{y_{it}}{a_i} \frac{\partial a_i}{\partial s} + \dots = 0$$

$$Dy_{it} = y_{it} \left( t \frac{\partial \ln(a_i)}{\partial s} + \frac{\partial \ln(a_i)}{\partial \Phi} \right) + \sigma^2 \frac{\partial y_{it}}{\partial t} \frac{\partial \ln(a_i)}{\partial s} \quad t = 1..T$$



# TOF consistency

$$Dy_{it} = \left( t y_{it} + \sigma^2 \frac{\partial y_{it}}{\partial t} \right) \frac{\partial \ln(a_i)}{\partial s} + y_{it} \frac{\partial \ln(a_i)}{\partial s} \quad t = 1..T$$

if  $y_{it} > 0$ : least squares fit,  $T$  equations

2 unknowns  $\frac{\partial \ln(a_i)}{\partial s}$  and  $\frac{\partial \ln(a_i)}{\partial \Phi}$

$$a_i = e^{-\sum_j l_{ij}\mu_j} \rightarrow \ln(a_i) = -\text{proj}(\mu)$$



attenuation sinogram  $\text{proj}(\mu)$  determined up to a constant

=

attenuation factor  $a$  determined up to a factor

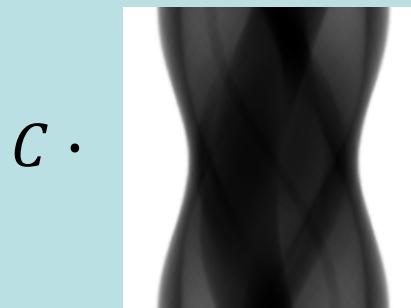
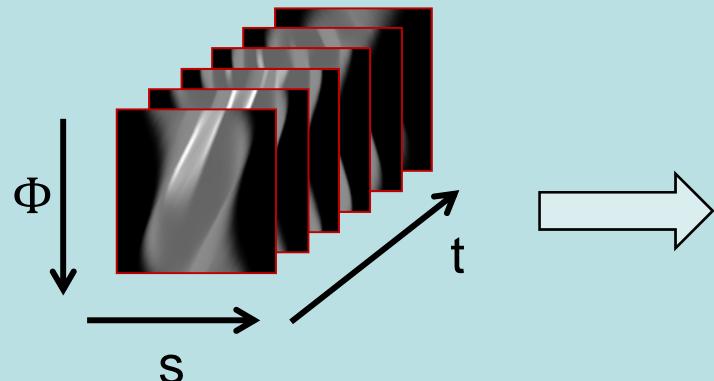


## TOF consistency

$$Dy_{it} = \left( t y_{it} + \sigma^2 \frac{\partial y_{it}}{\partial t} \right) \frac{\partial \ln(a_i)}{\partial s} + t y_{it} \frac{\partial \ln(a_i)}{\partial s} \quad t = 1..T$$

if  $y_{it} > 0$ : least squares fit,  $T$  equations

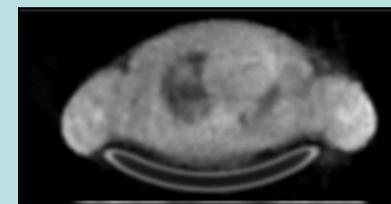
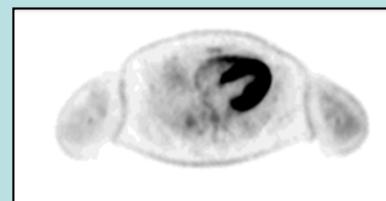
2 unknowns  $\frac{\partial \ln(a_i)}{\partial s}$  and  $\frac{\partial \ln(a_i)}{\partial \Phi}$





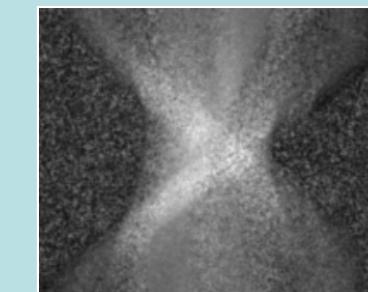
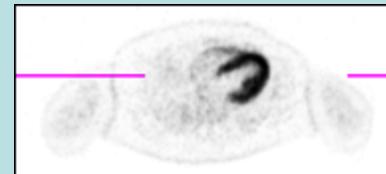
# attenuation in TOF-PET

- TOF-PET data



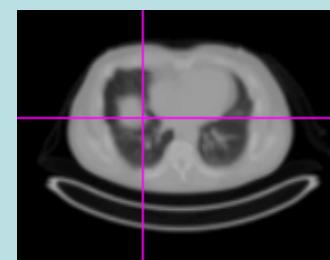
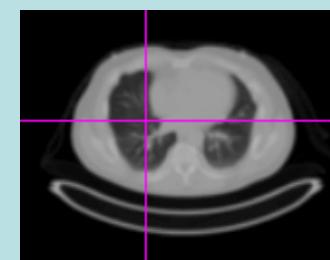
- MLAA

*Maximum Likelihood reconstruction of Activity and Attenuation*



- MLACF

*Maximum Likelihood reconstruction of Activity and attenuation Correction Factors*



- MLRR

*Maximum Likelihood Reconstruction of activity and Registration of attenuation image*



maximize likelihood  $L(y, \lambda, \mu)$

$$L = \sum_i \sum_t y_{it} \ln \widehat{y}_{it} - \widehat{y}_{it}$$

$$\widehat{y}_{it} = e^{-\sum_k l_{ik} \mu_k} \sum_j c_{itj} \lambda_j + s_{it}$$

Alternated parameter update:

constant  $\mu$        $\widehat{y}_{it} = a_i \sum_j c_{itj} \lambda_j + s_{it}$       (TOF-) MLEM

constant  $\lambda$        $\widehat{y}_{it} = b_{it} e^{-\sum_k l_{ik} \mu_k} + s_{it}$       (TOF-) MLTR

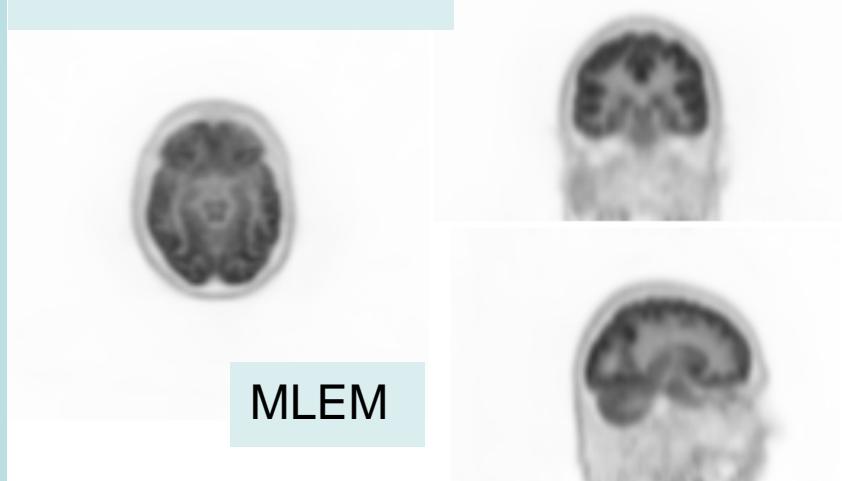
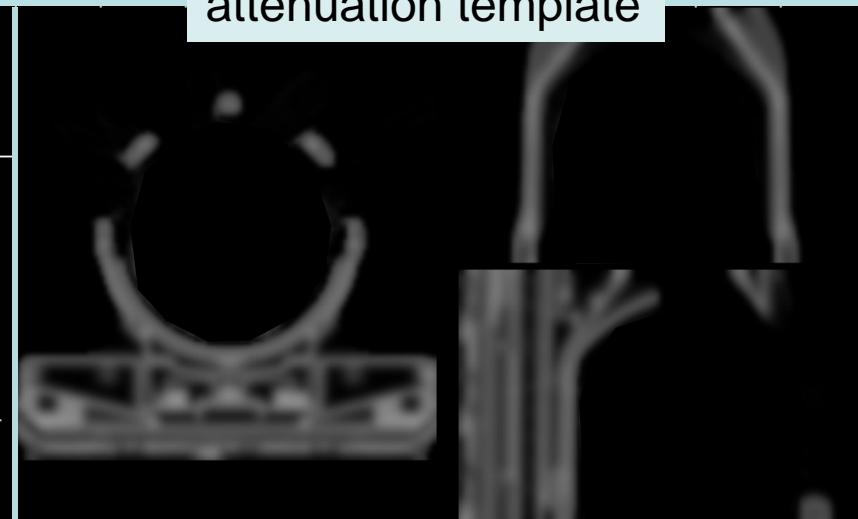
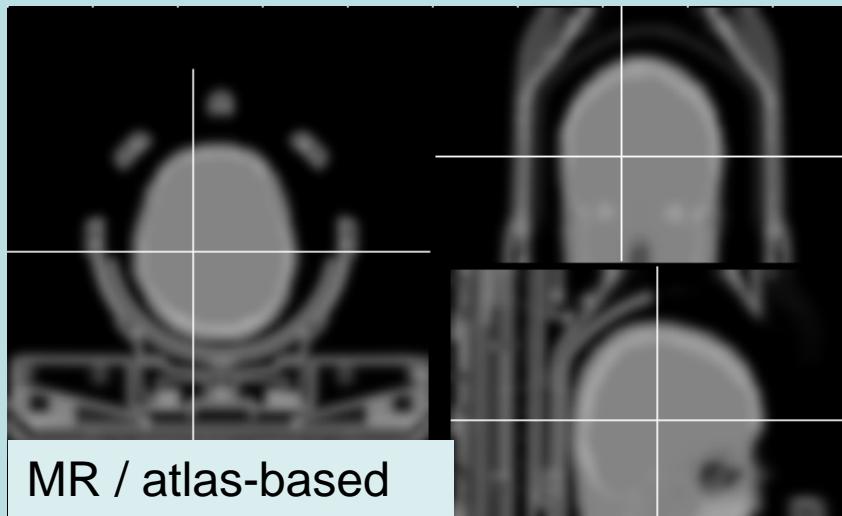
if  $s_{it} \sim \widehat{y}_{it}$   
or  $s_{it} \simeq 0$        $\rightarrow$        $\widehat{y}_i = b_i e^{-\sum_k l_{ik} \mu_k} + s_i$       MLTR

a few MLTR sub-iterations per MLEM (OSEM) sub-iteration



# TOF-PET MLAA

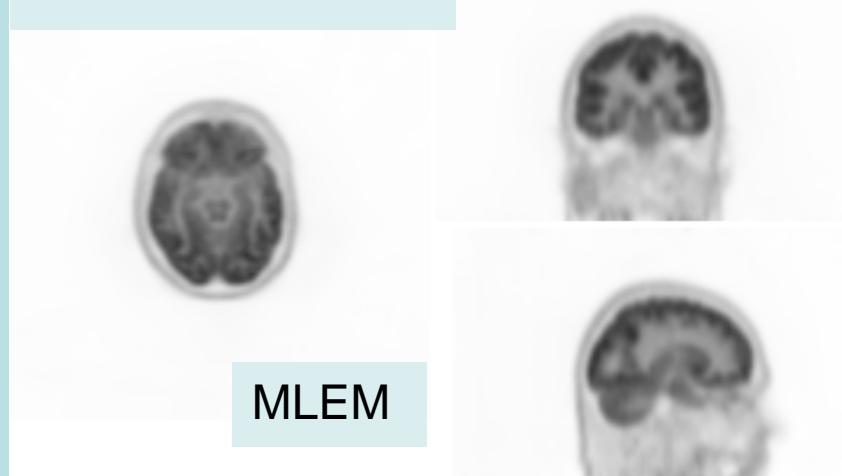
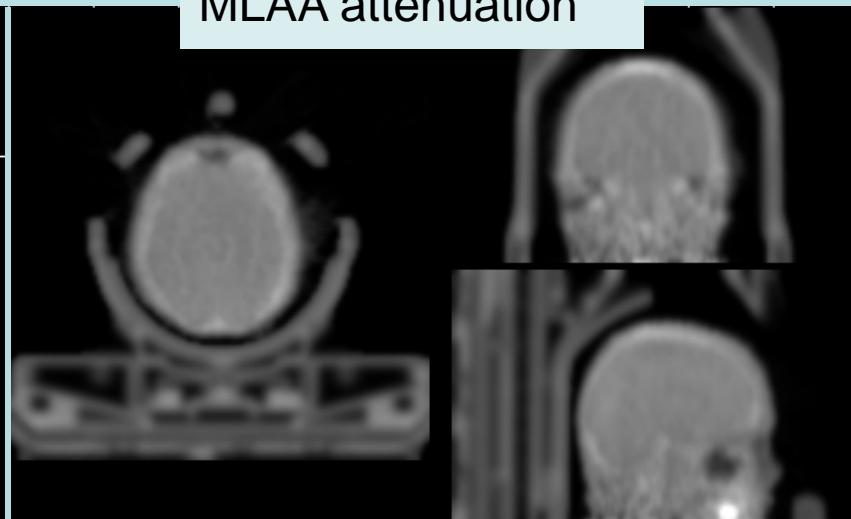
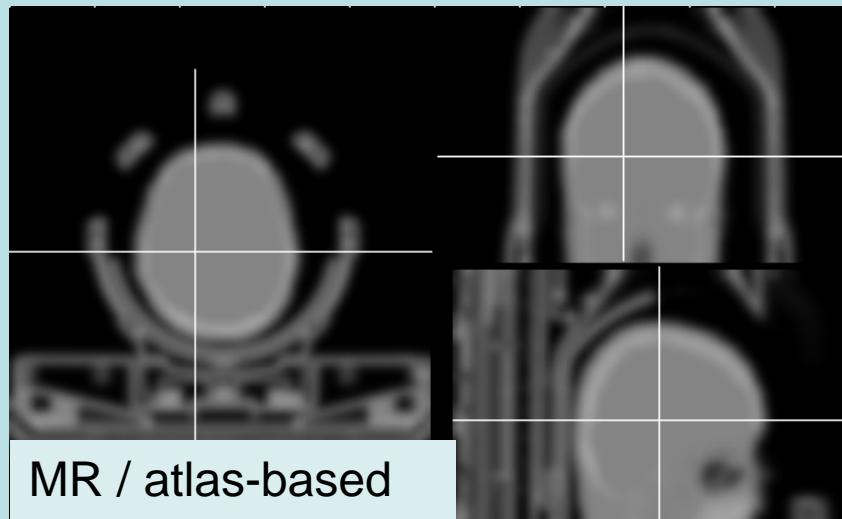
TOF: 380 ps





# TOF-PET MLAA

TOF: 380 ps





# TOF-PET MLAA

true

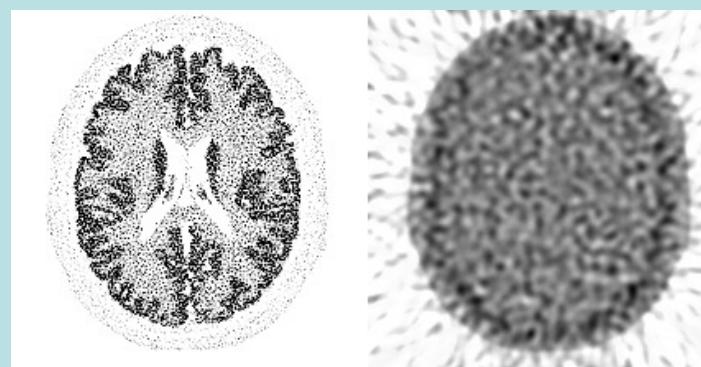


MLAA, using  
known body  
contour

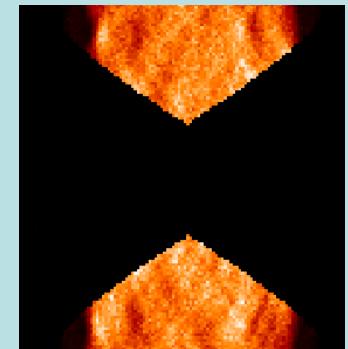
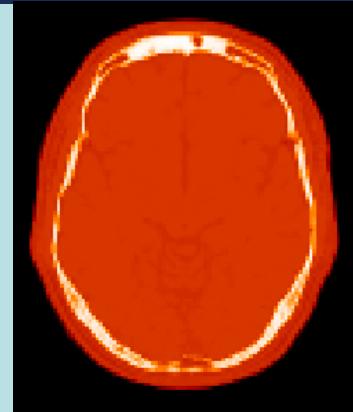
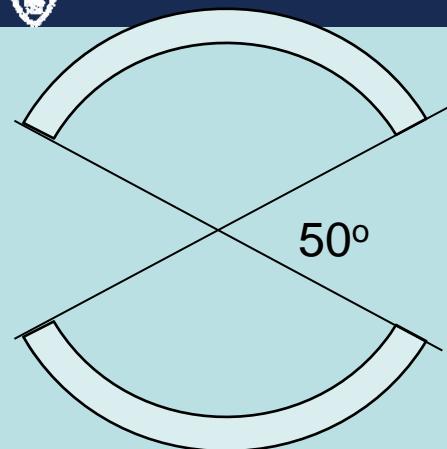


noise free

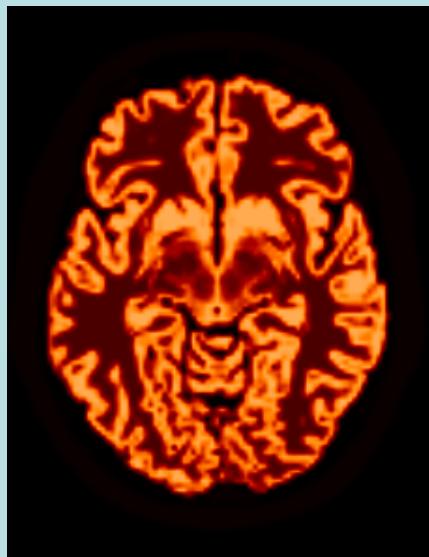
MLAA, using  
known body  
contour



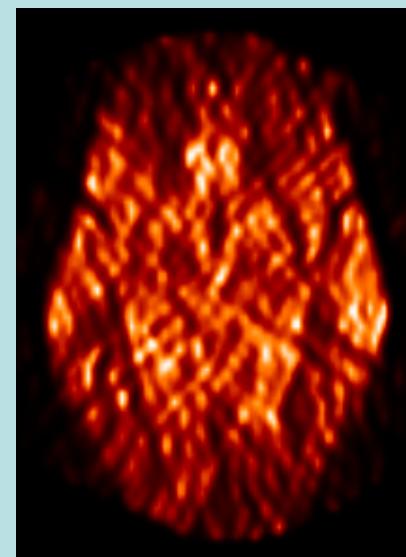
noisy



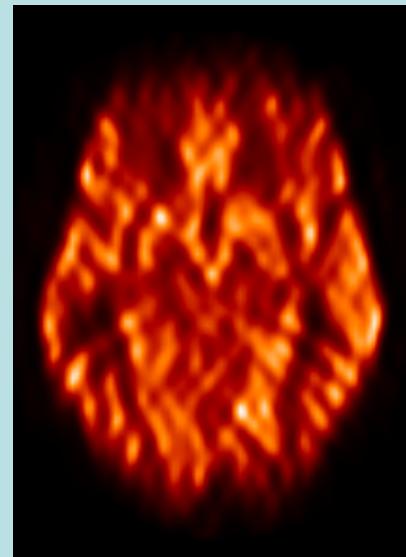
sinogram



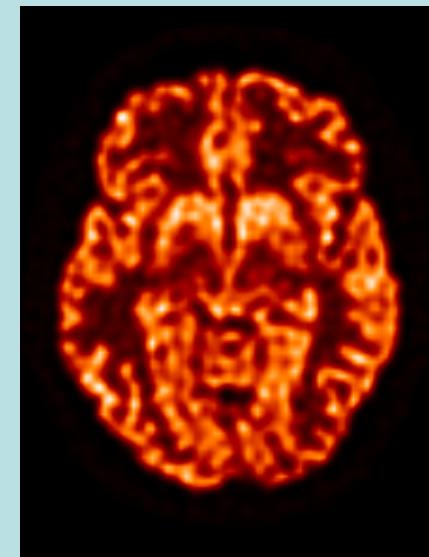
TRUE



non-TOF



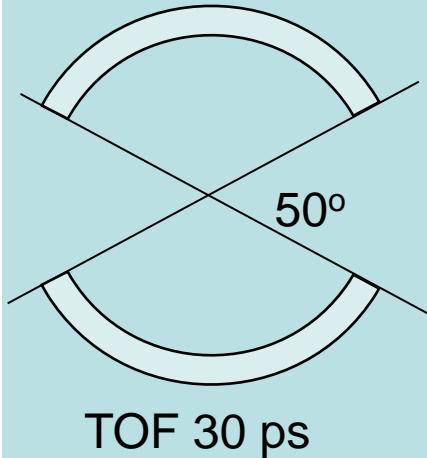
TOF 400 ps



TOF 30 ps

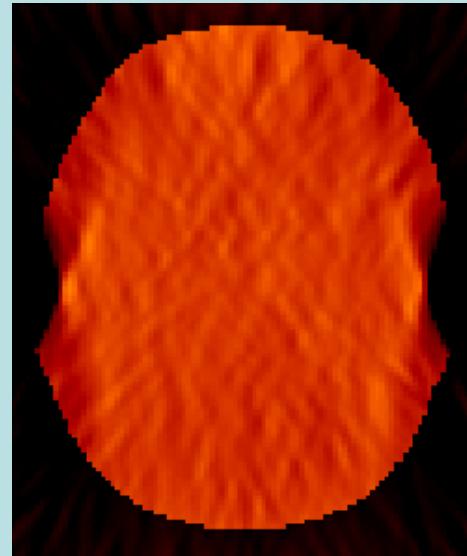
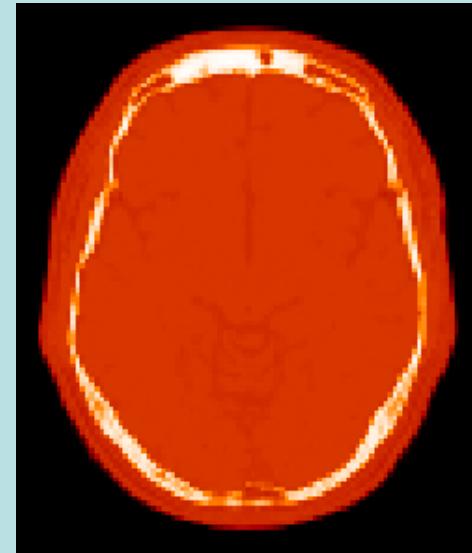


# TOF-PET: MLAA

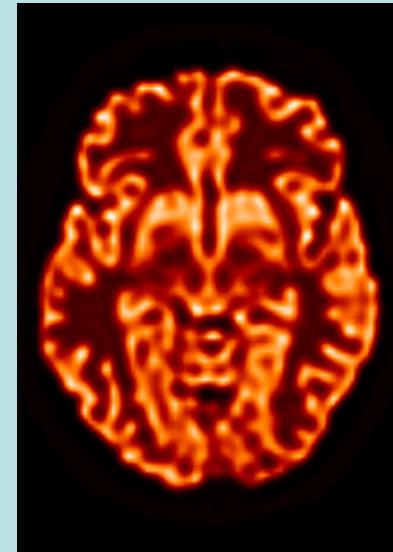
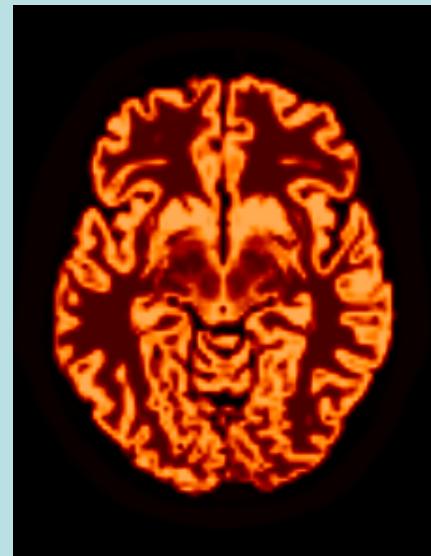


TOF 30 ps

TRUE



joint estimation

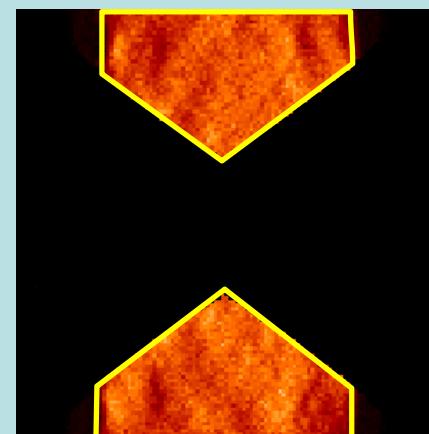
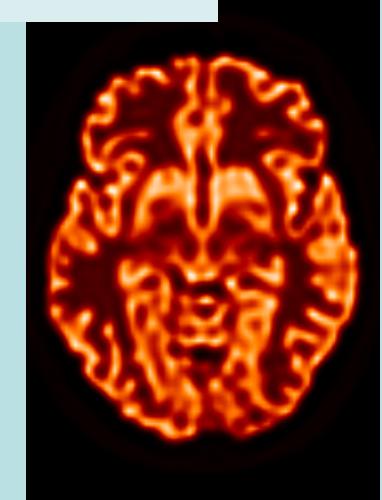
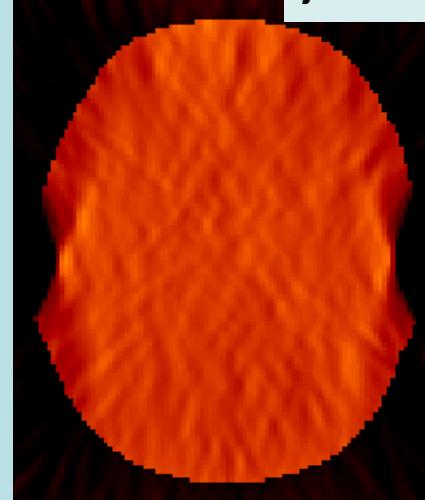
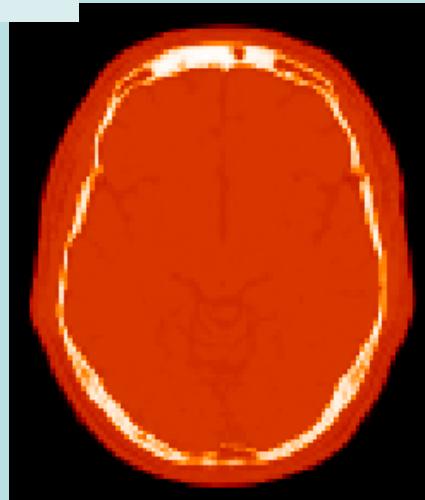




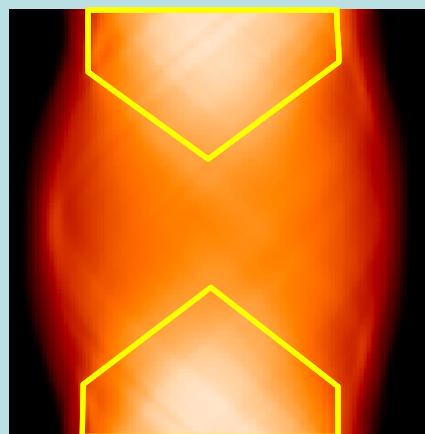
## TOF-PET: MLAA

TRUE

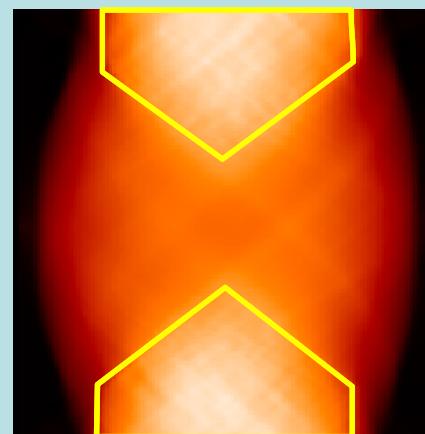
joint estimation



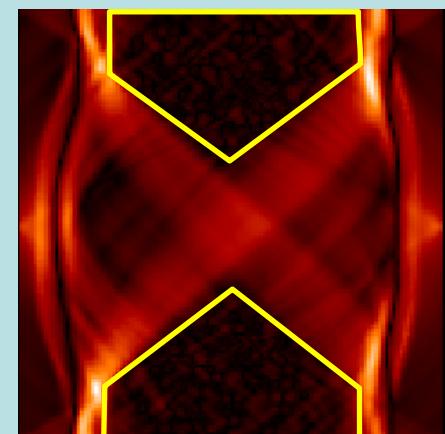
measured activity  
sinogram



true attenuation



estimated  
attenuation



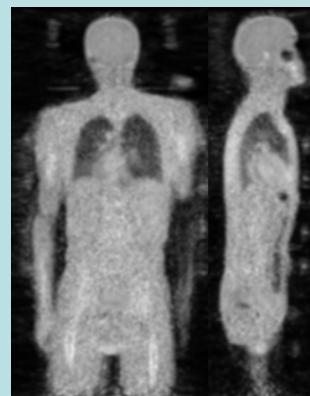
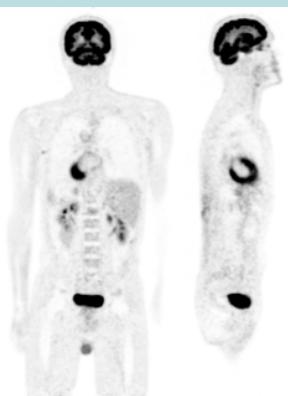
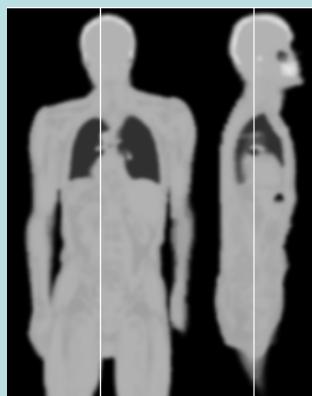
difference true  
and estimated  
attenuation



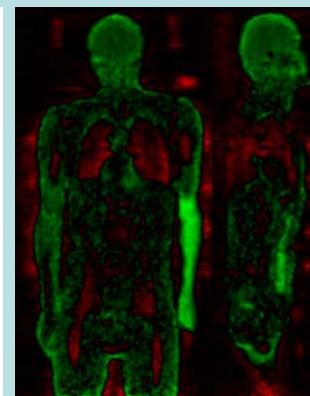
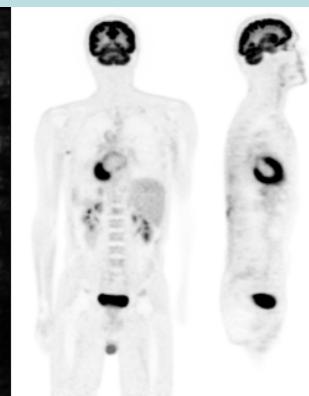
## TOF-PET: MLAA

MLEM (CT-based atten)

patient 1



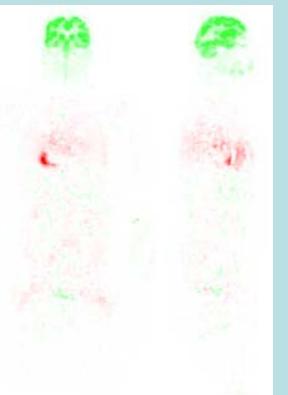
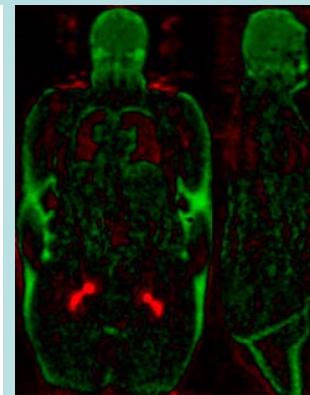
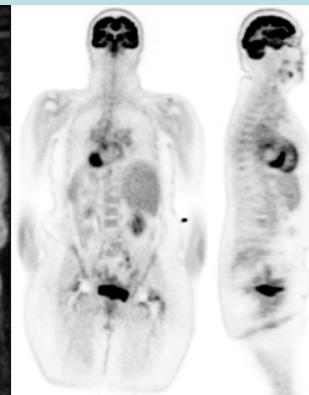
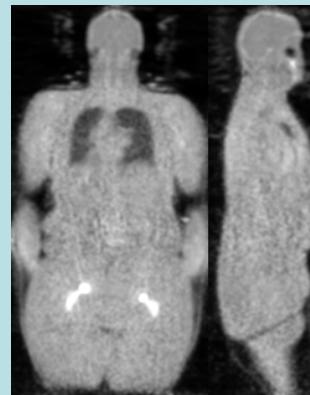
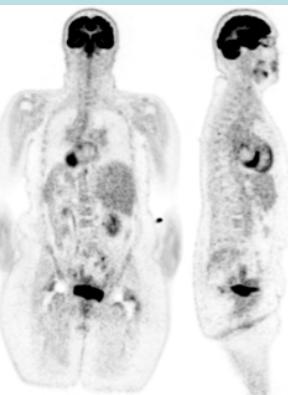
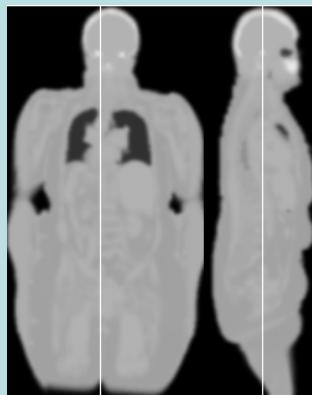
MLAA

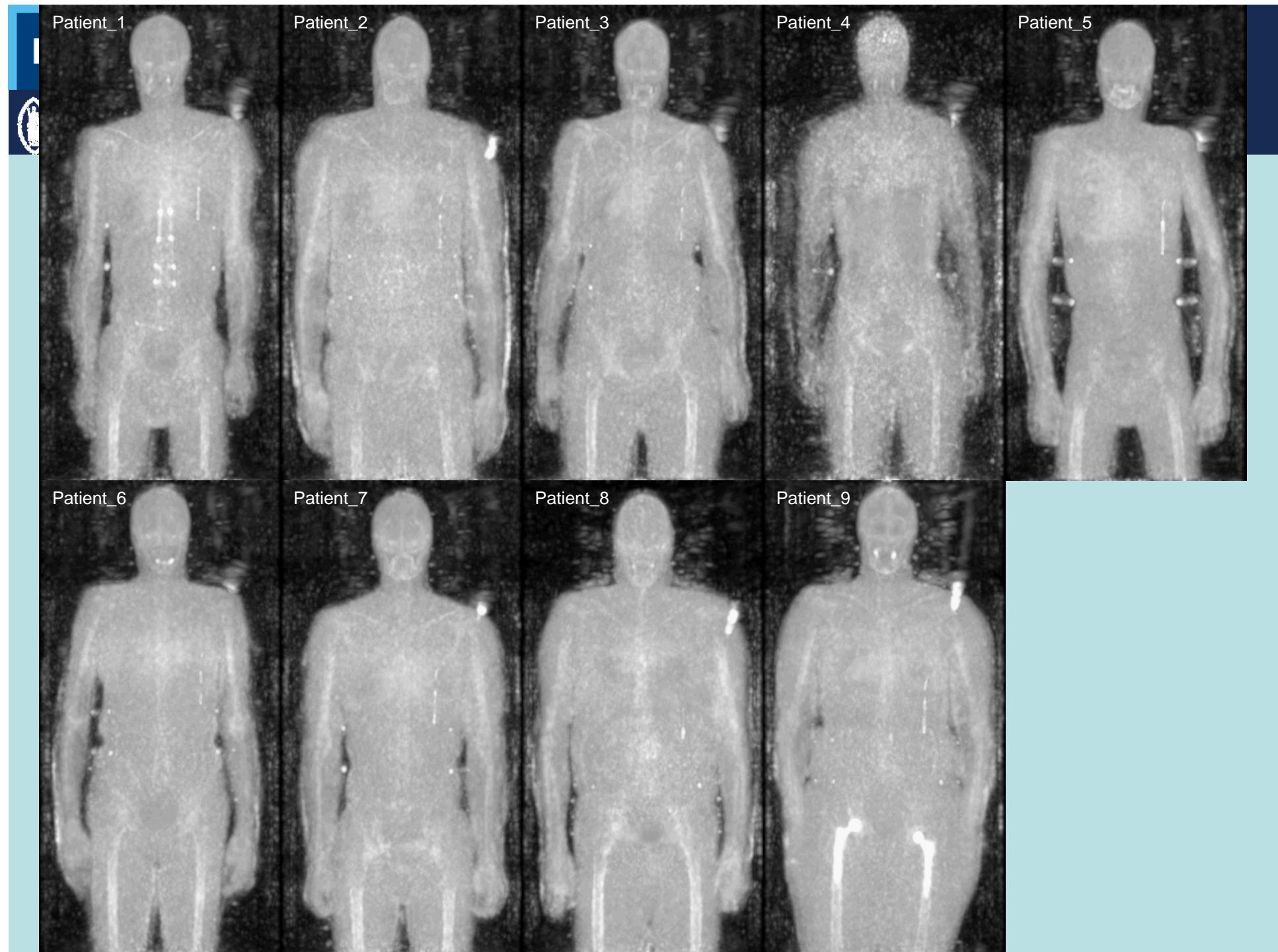


MLAA - MLEM



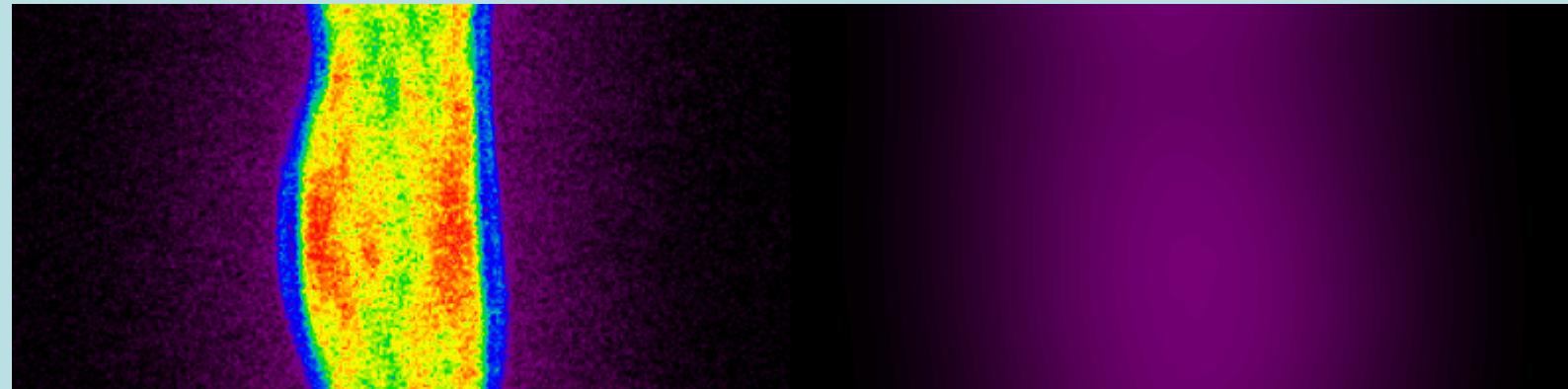
patient 9







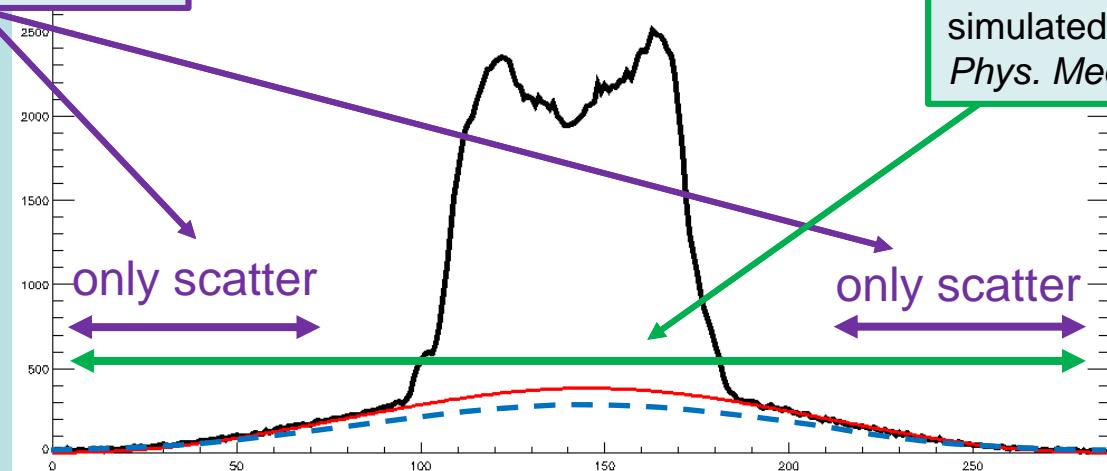
## scatter estimation



sinogram

estimated scatter

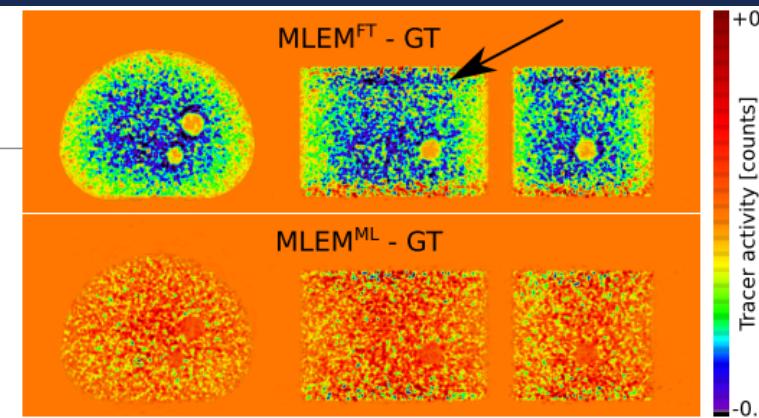
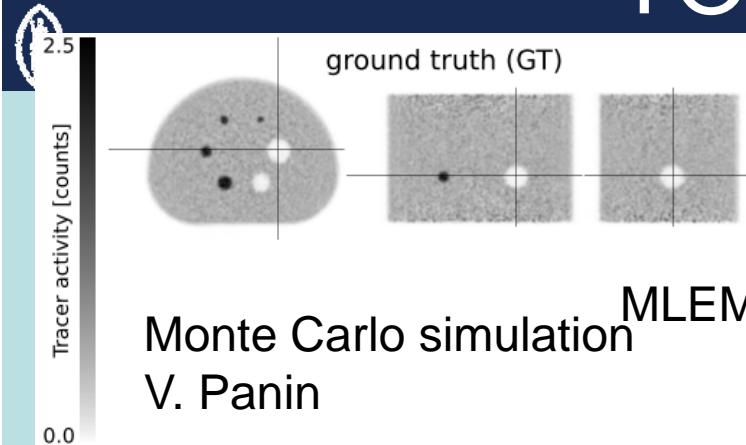
“scatter tail fitting”



A. Rezaei et al. Plane-dependent ML scatter scaling: 3D extension of the 2D simulated single scatter (SSS) estimate.  
*Phys. Med. Biol.*, 62, pp.6515-6531.

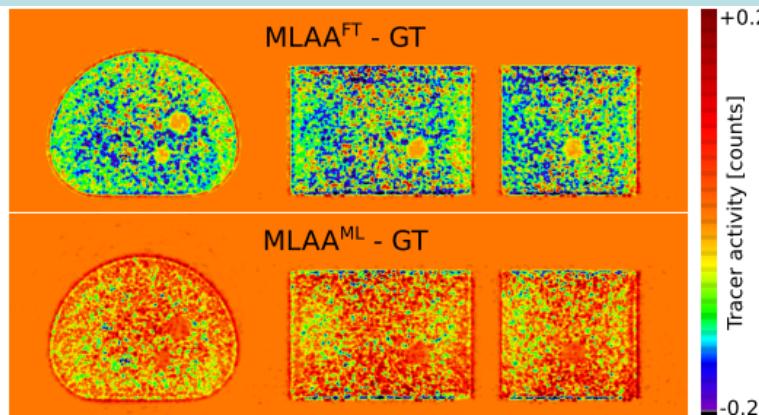
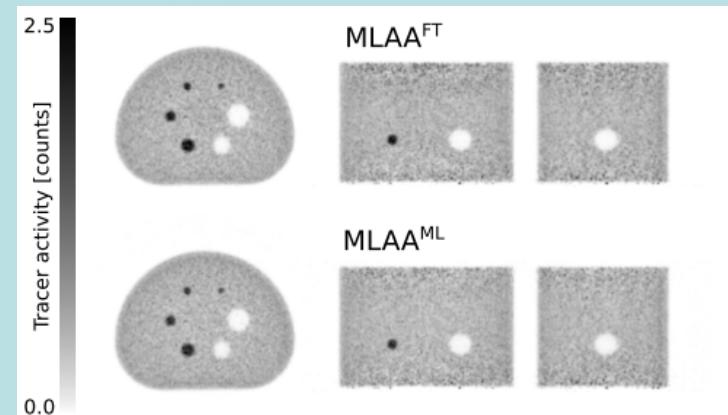
Simulated  
Single Scatter

## TOF-PET MLAA: scatter



tail fitting  
overestimated  
scatter

ML fitting



tail fitting

ML fitting

tail fitting

ML fitting

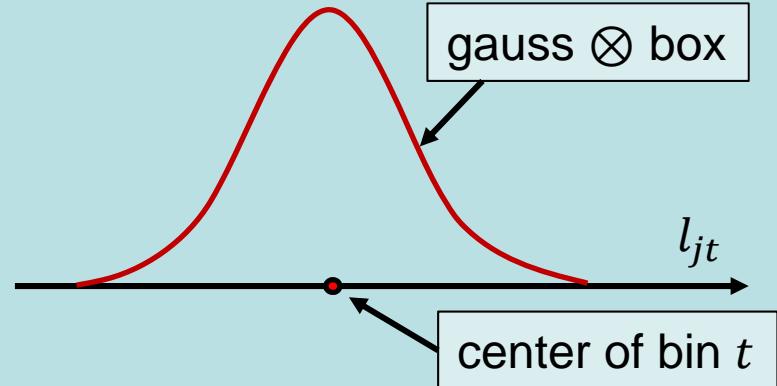


# timing and efficiency calibration

expected data given activity  $\lambda$ :

$$\bar{y}_{it} = a_i n_i \sum_j c_{ij} g_{ijt} \lambda_j + n_i s_{it}$$

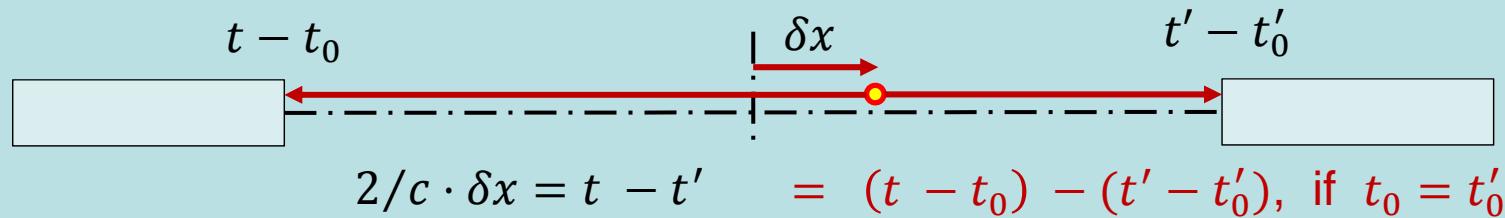
$$g_{ijt} = \frac{1}{2} \left( \operatorname{erf}\left(\frac{l_{jt} + b}{\sigma\sqrt{2}}\right) - \operatorname{erf}\left(\frac{l_{jt} - b}{\sigma\sqrt{2}}\right) \right)$$



$\sigma$ : TOF-resolution

$b$ : bin width

$l_{jt}$ : distance voxel  $j$  to center of bin  $t$



$g_{ijt}$  depends on **timing offsets** ( $l_{jt}$ ) and on **TOF resolution**  $\sigma$

$n_i$  represents LOR sensitivity



# timing and efficiency calibration

we estimate

- LOR timing offsets
- LOR TOF resolution values
- LOR efficiency corrections to vendor supplied normalization

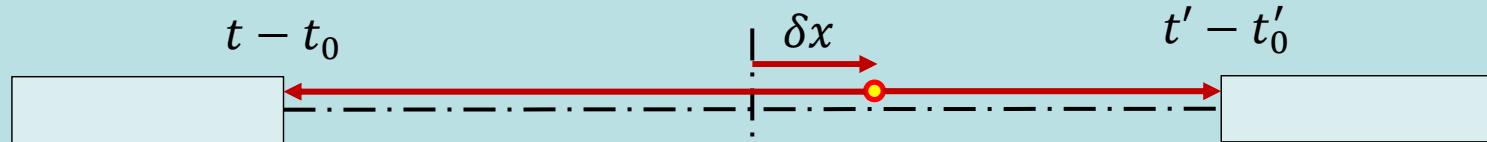
assuming LOR parameters are determined by its two crystals:

- LOR timing offset:  $\delta t_{LOR} = \delta t_{crystal1} - \delta t_{crystal2}$
- LOR TOF resolution:  $\sigma_{LOR} = \sqrt{\sigma_{crystal1}^2 + \sigma_{crystal2}^2}$
- LOR efficiency:  $\epsilon_{LOR} = \epsilon_{crystal1} \epsilon_{crystal2}$

crystal parameters determined during ML-reconstruction of activity  
using CT-based attenuation correction

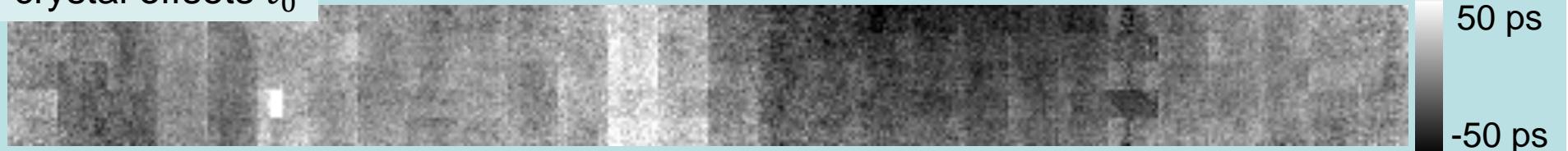


# TOF-PET MLAA: TOF offsets

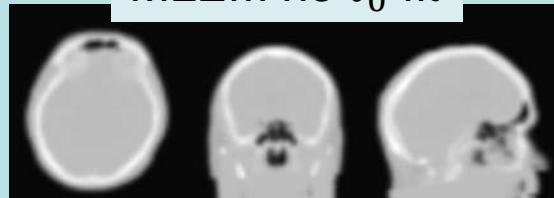


crystal offsets  $t_0$

$$2/c \cdot \delta x = t - t' = (t - t_0) - (t' - t'_0), \text{ if } t_0 = t'_0$$



MLEM no  $t_0$  fit



MLAA no  $t_0$  fit



MLAA with  $t_0$  fit

50 ps

-50 ps

+10%

-10%

MLEM\_no\_  $t_0$  – MLEM\_  $t_0$

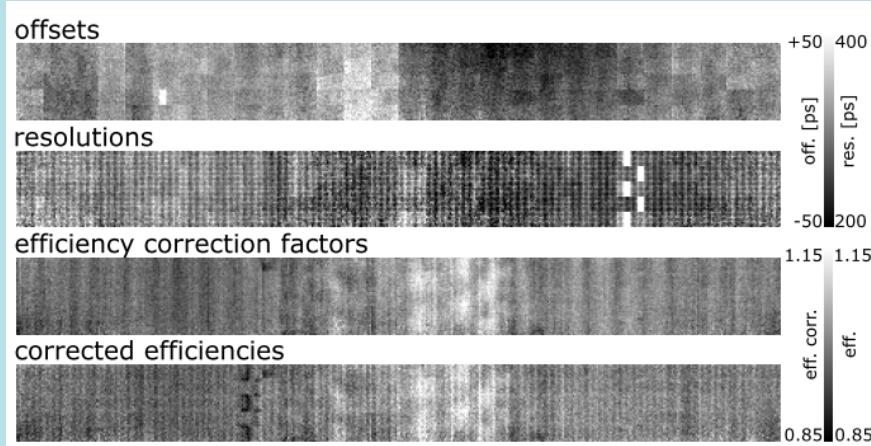
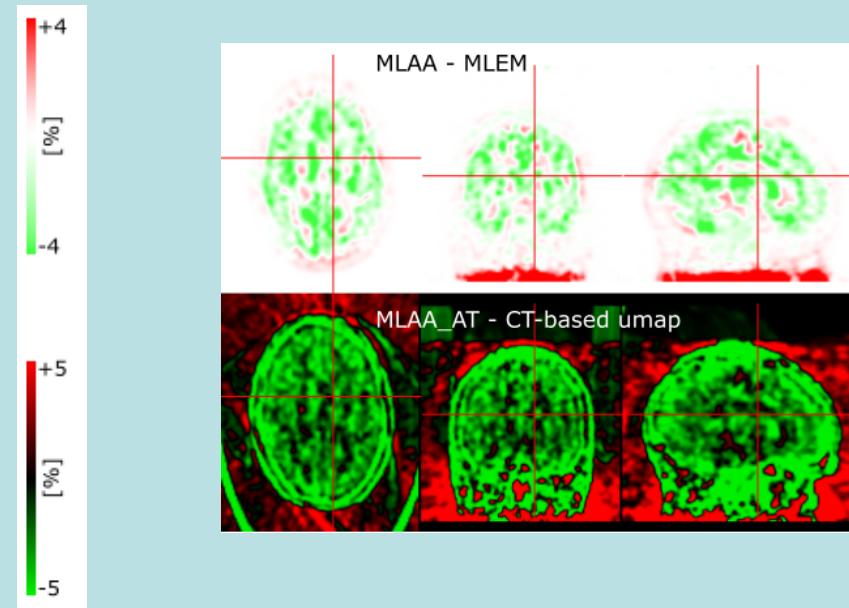
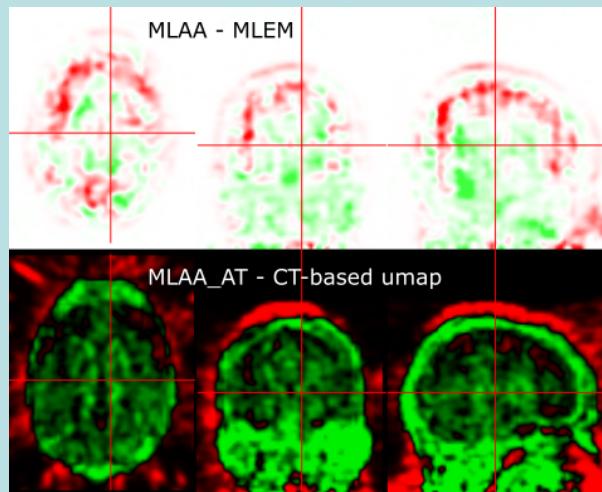
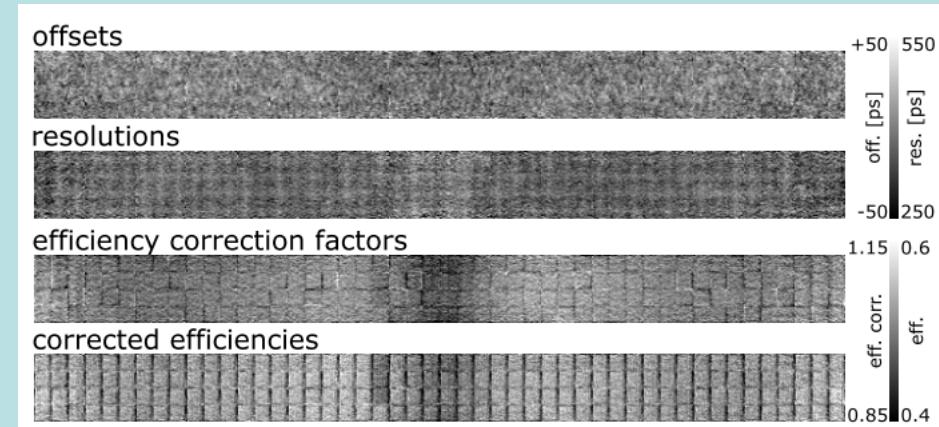
MLAA\_no\_  $t_0$  – MLEM\_no\_  $t_0$

MLAA\_  $t_0$  – MLEM\_  $t_0$

48



# timing and efficiency calibration

GE Signa ( $\pm 385$  ps)Siemens mCT ( $\pm 535$  ps)

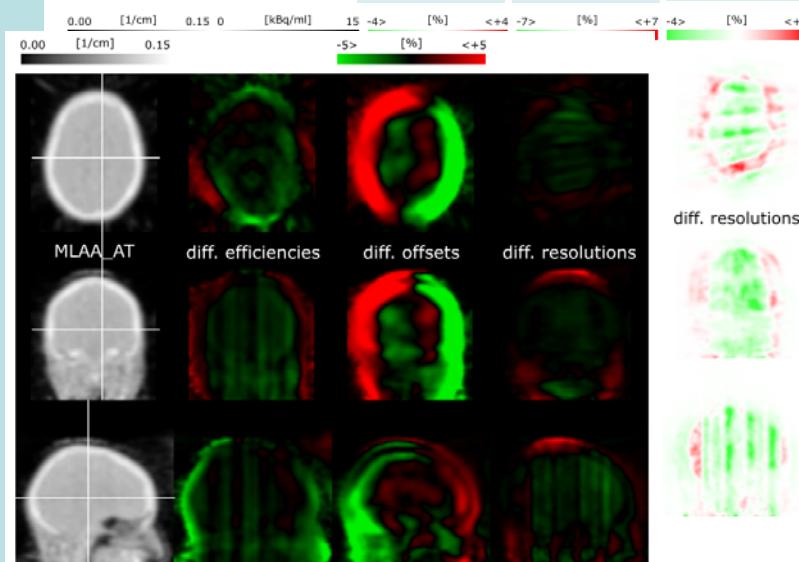


# timing and efficiency calibration

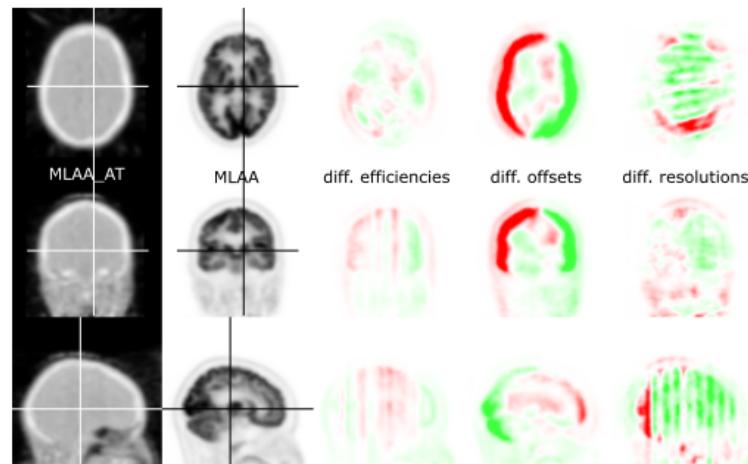
GE Signa

 $\epsilon$   
-4..4%

 $\delta t$   
-7..7%

 $\sigma$   
-4..4%


(a) SIGNA MLAA atten

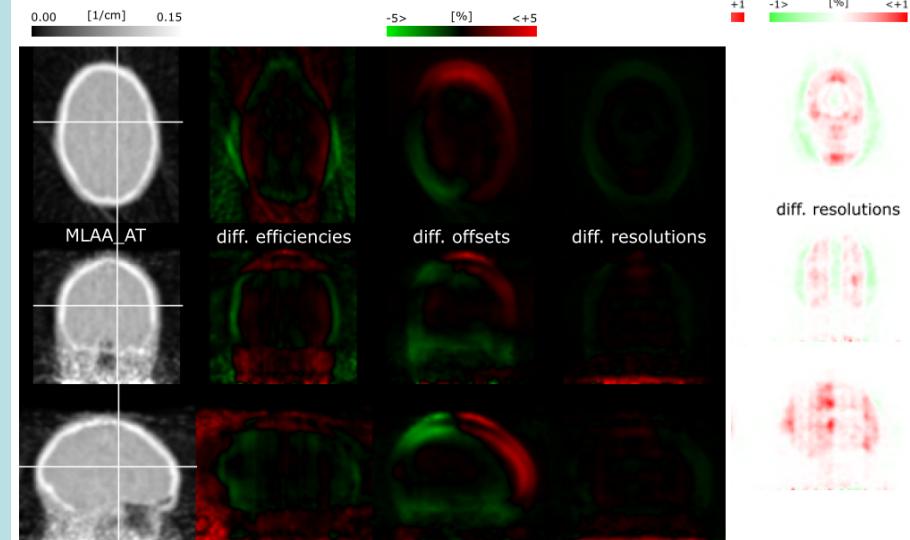


(b) MLAA

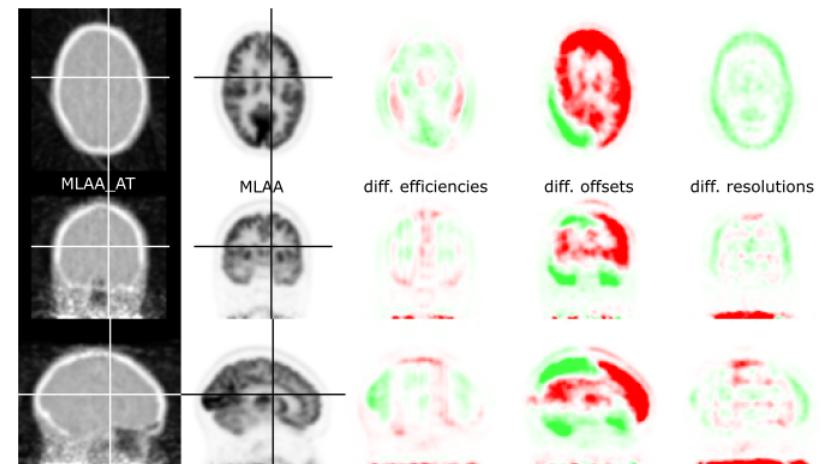
Siemens mCT

 $\epsilon$   
-2..2%

 $\delta t$   
-1..1%

 $\sigma$   
-1..1%


(b) Biograph mCT MLAA atten



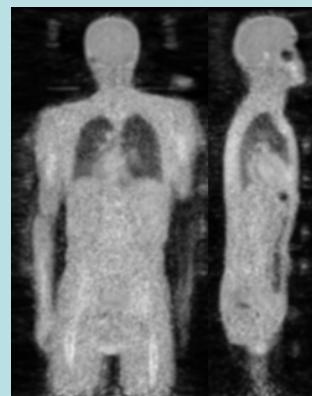
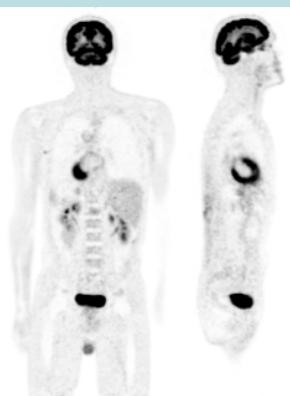
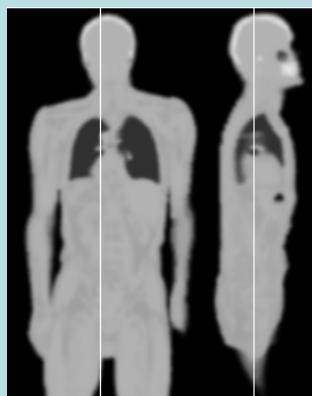
(b) MLAA



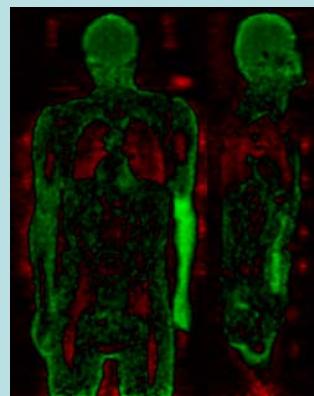
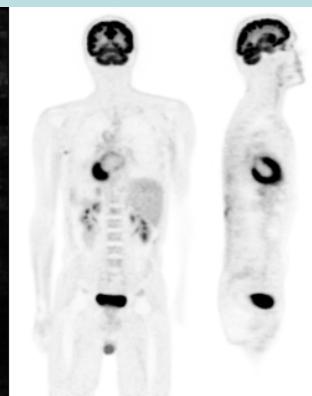
## TOF-PET: MLAA

MLEM (CT-based atten)

patient 1



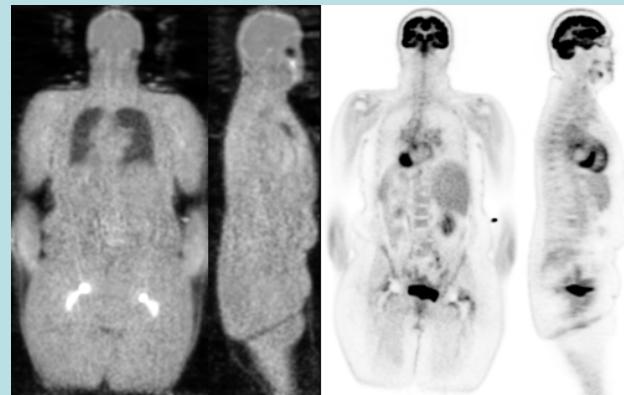
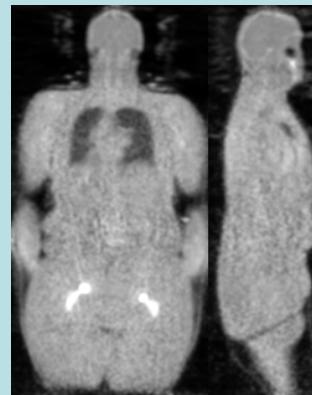
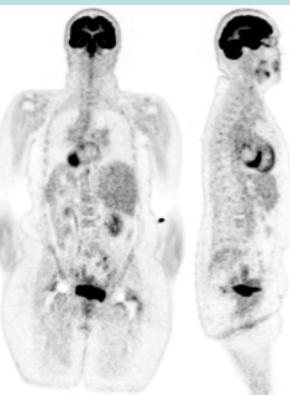
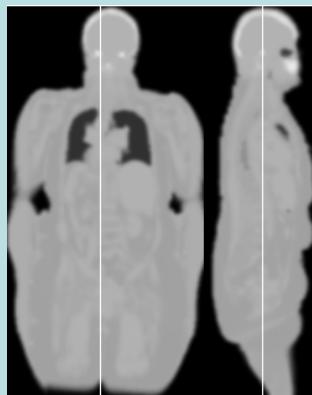
MLAA



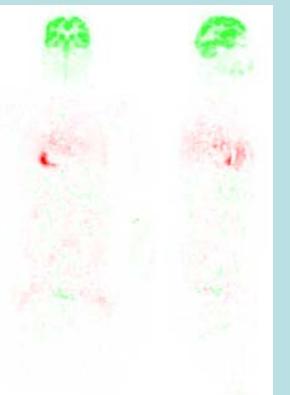
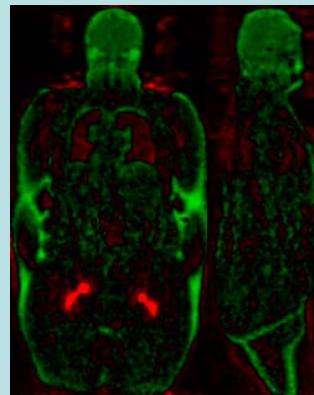
MLAA - MLEM



patient 9



before ...

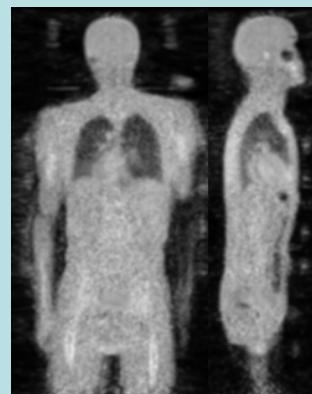
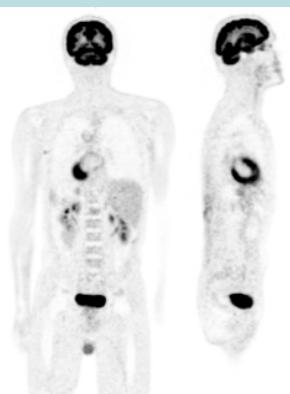
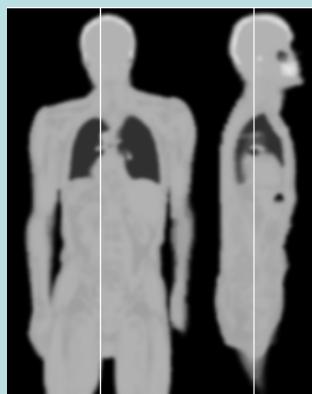




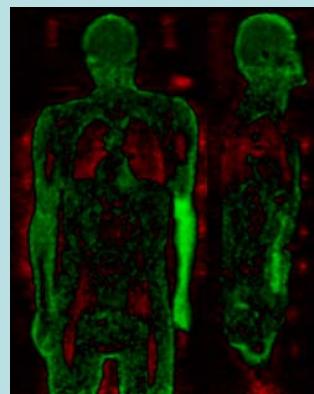
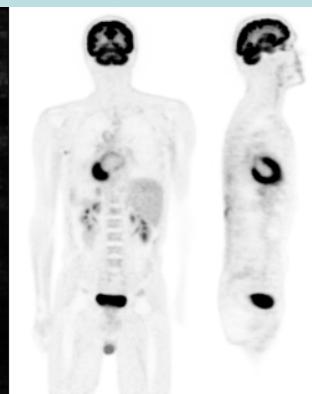
## TOF-PET: MLAA

MLEM (CT-based atten)

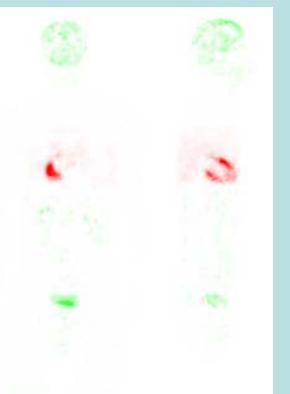
patient 1



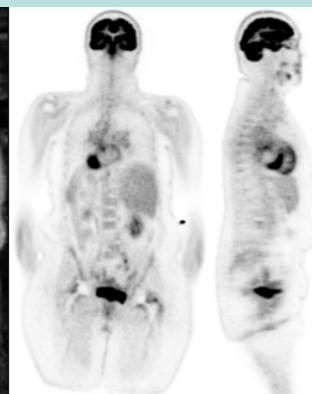
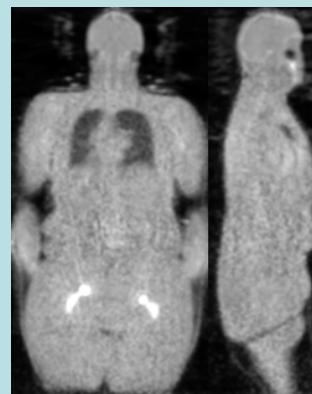
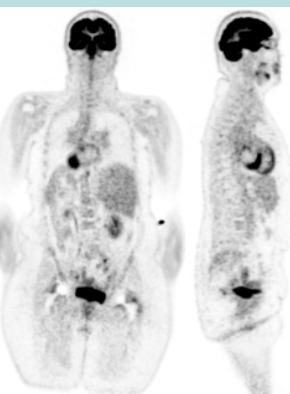
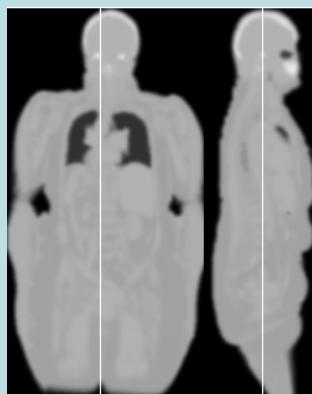
MLAA



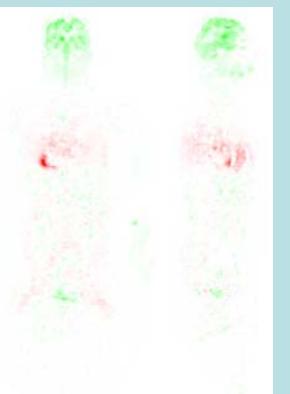
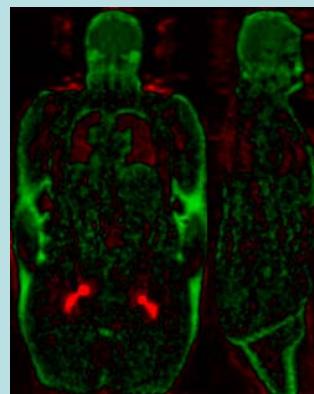
MLAA - MLEM



patient 9



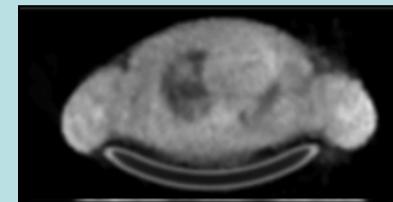
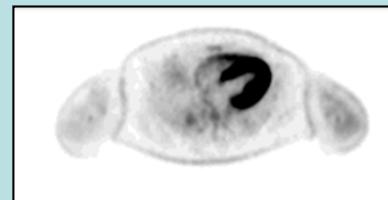
... after





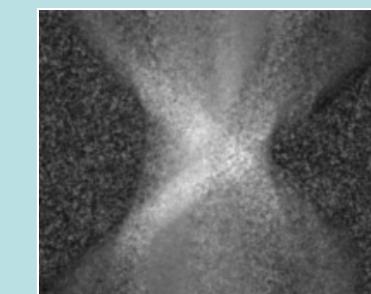
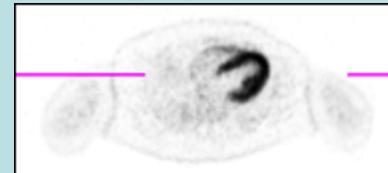
# attenuation in TOF-PET

- TOF-PET data



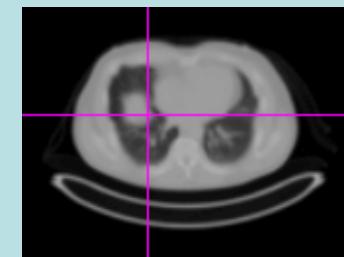
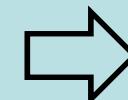
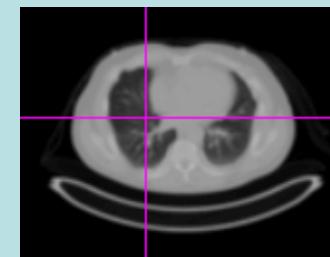
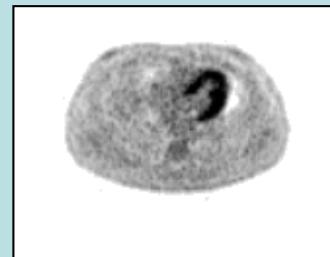
*Maximum Likelihood reconstruction of Activity and Attenuation*

- MLACF



*Maximum Likelihood reconstruction of Activity and attenuation Correction Factors*

- MLRR



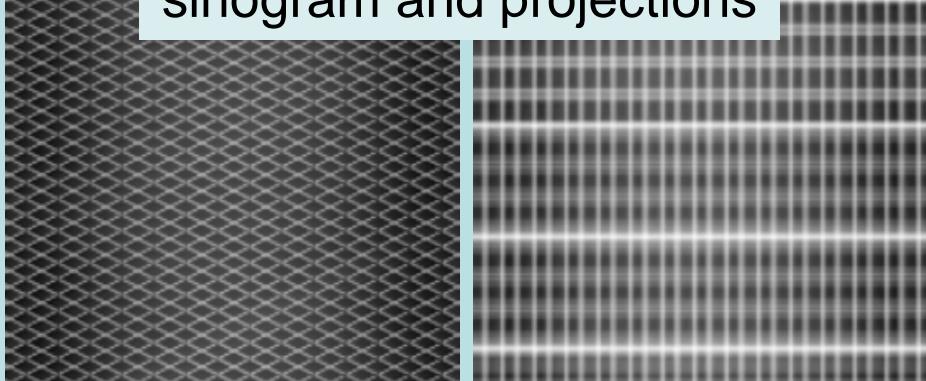
*Maximum Likelihood Reconstruction of activity and Registration of attenuation image*



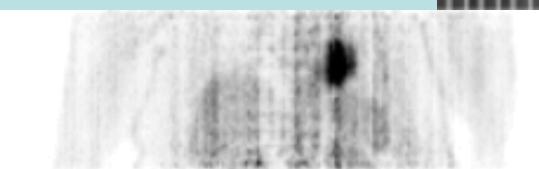
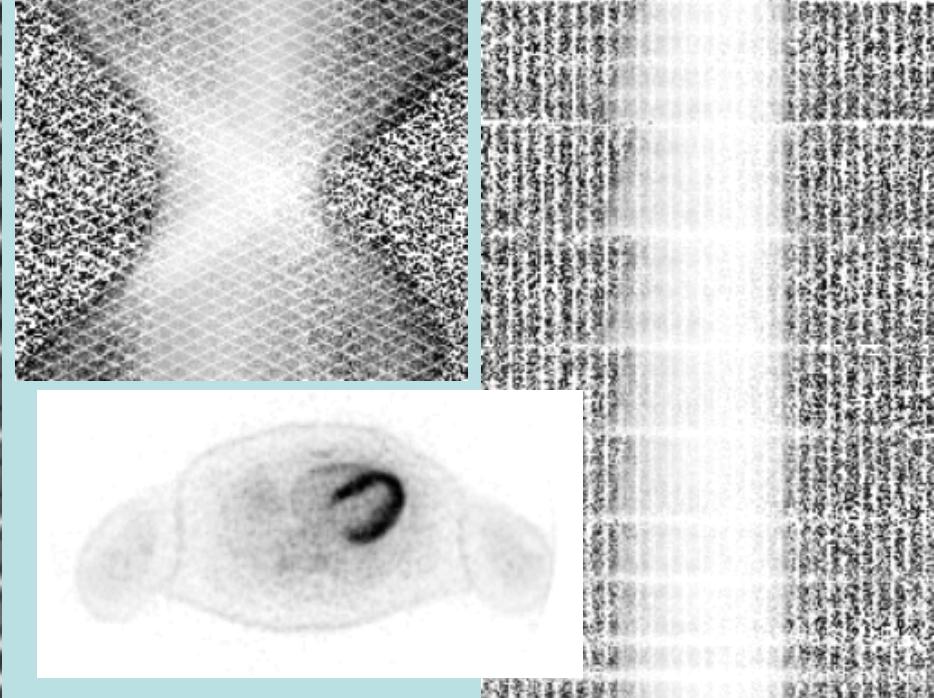
# MLACF: estimation of $\lambda$ and $a$

sensitivity

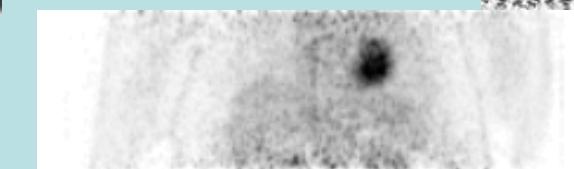
sinogram and projections



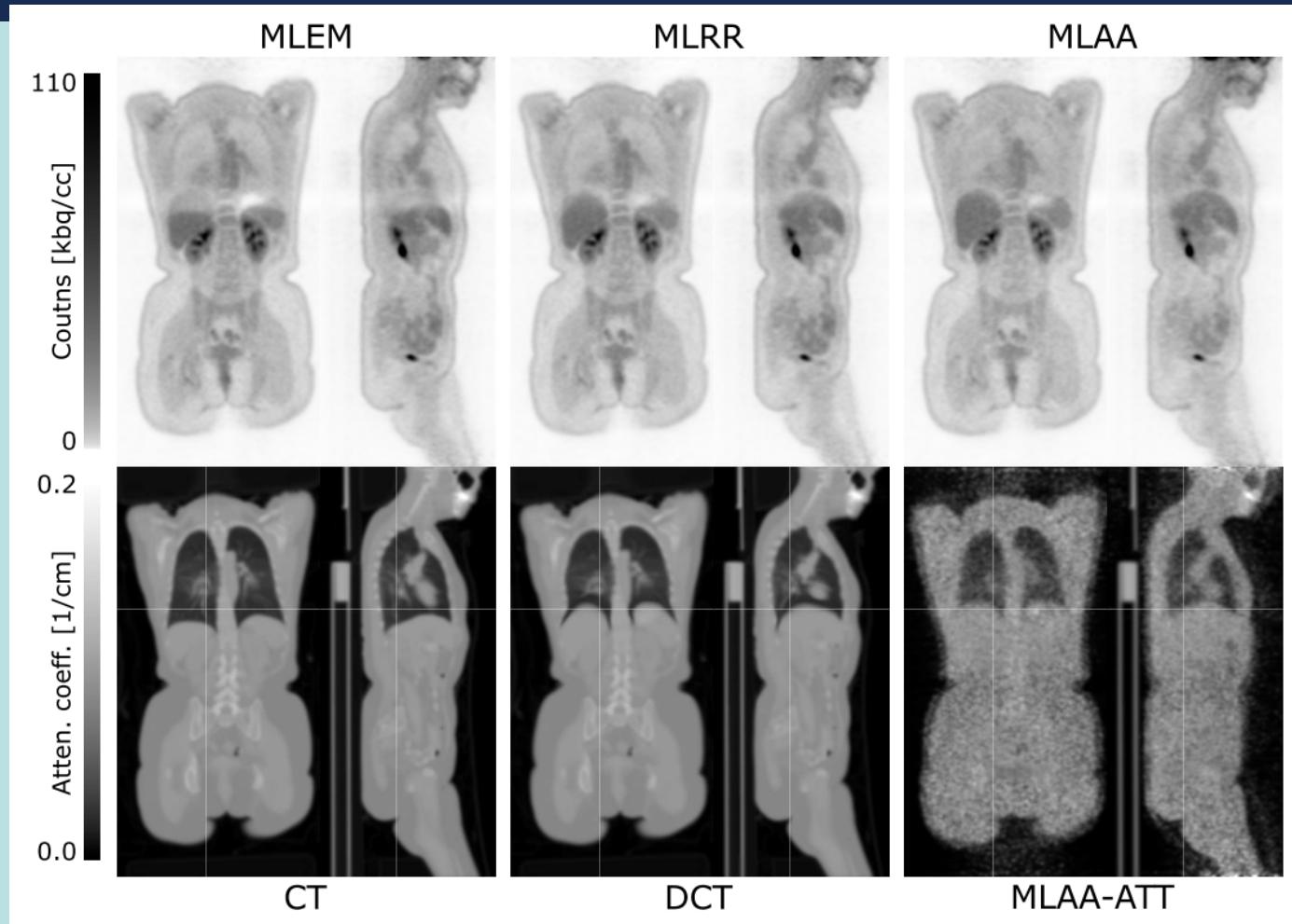
MLACF



MLEM with attenuation corr  
without sensitivity corr



MLACF estimates product of both

MLRR: estimate  $\lambda$  and CT deformation

from Rezaei et al., to be submitted to *J Nucl Med*

see also: Rezaei et al., *Phys Med Biol* 2016; 61 (4): 1852 – 1874.

Bousse et al., *Phys Med Biol* 2016 (3) L11



# 10 ps TOF-PET

## 10 ps TOF

- reconstruction
    - faster
    - more robust
  - (much) better resolution
  - limited angle problem eliminated
  - (much) better SNR
  - better joint estimation of attenuation & activity
- remaining problems
    - scatter
    - positron range
  - system calibration essential

thanks!