



Software challenges and opportunities in multimodality PET/CT and PET/MR imaging

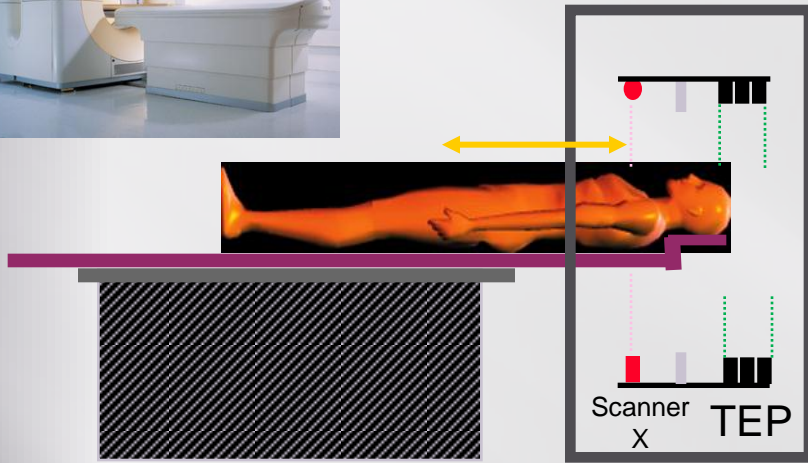
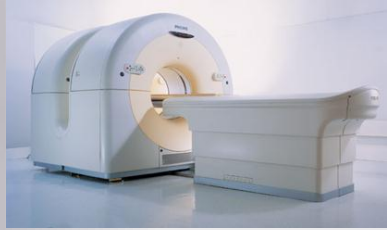
Dimitris Visvikis

Director of Research, INSERM

National Institute of Health and Medical Sciences

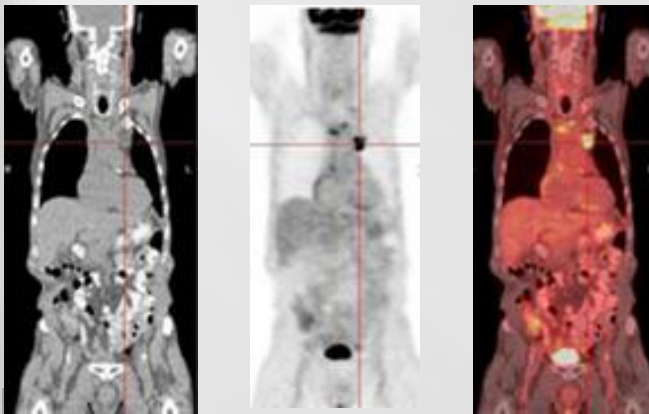
LaTIM (Medical Information Processing Lab), Brest, France

Multimodality imaging: PET/CT

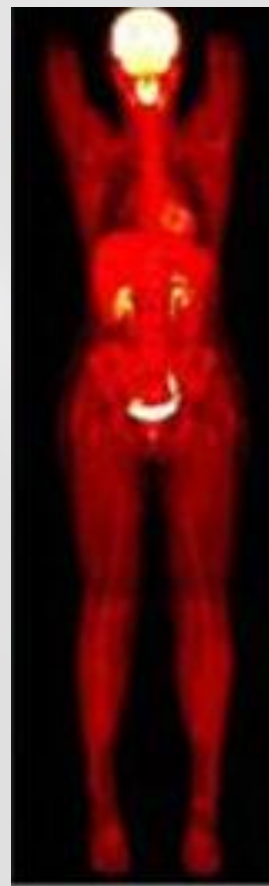
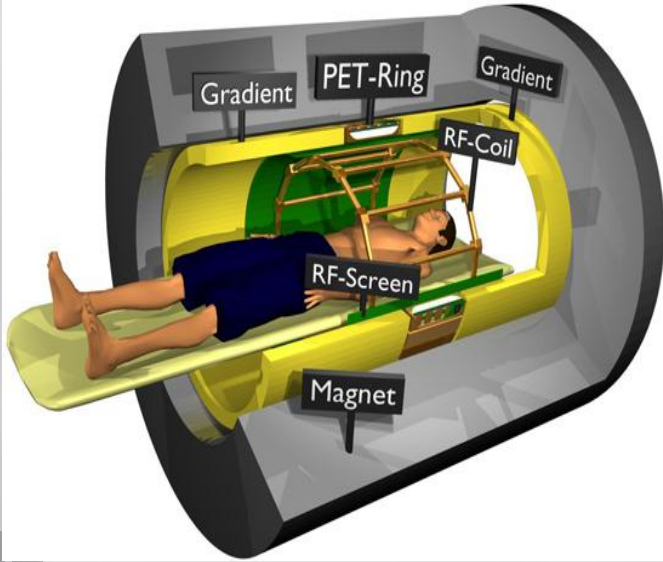


Medical Imaging revolution

- ✓ Combination of anatomy and function
- ✓ Increased patient throughput



Multimodality imaging: PET/MR



Question Combining anatomy and function: challenge or opportunity?

From a software point of view: a little bit of both

1. Corrections:

- physics principles of the detection process (scatter, attenuation, partial volume effects, noise)
- association of anatomical and functional images (physiological organ motion)

2. New image processing approaches allowing:

- combination of different multimodality image derived parameters and their associated temporal evolution
- development of predictive and prognostic models of disease outcomes

Multimodality imaging: software developments

Question **Combining PET and CT**: challenge or **opportunity**?

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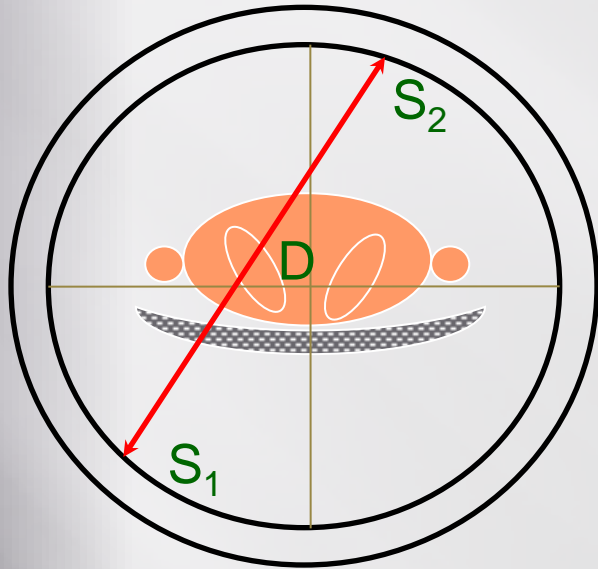
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PET attenuation correction



- Need to know the distribution of the attenuation coefficients
- Attenuation independent of the position of the annihilation along the line of response
- No need to know the activity distribution for the calculation of attenuation as it is the case in SPECT

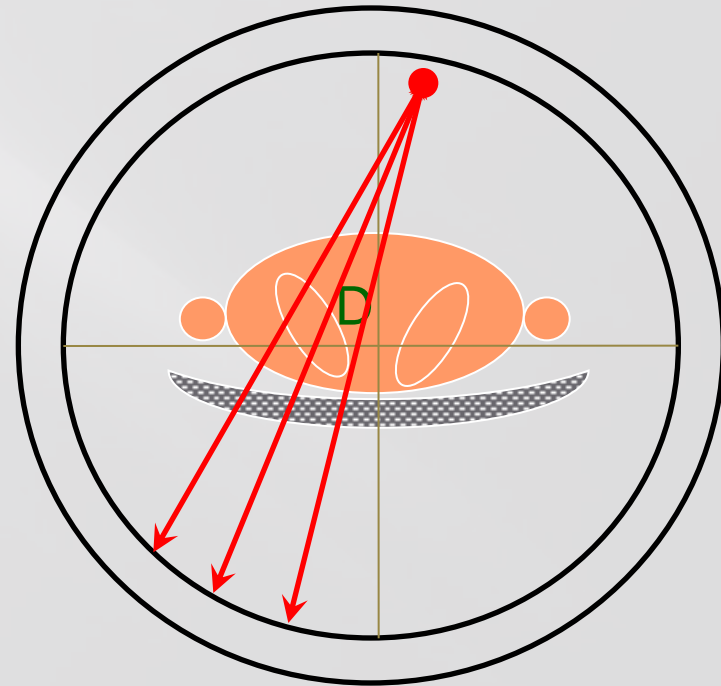
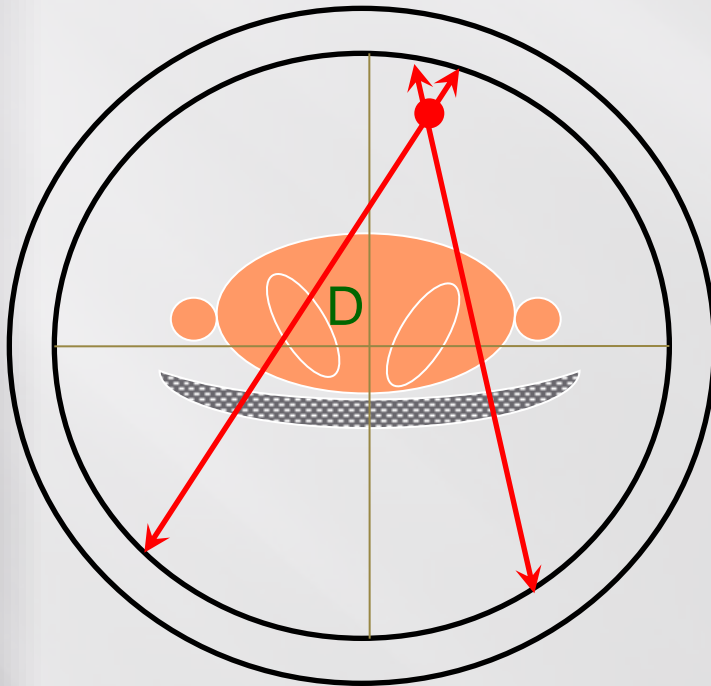


PET attenuation correction



b^+ emitter
(Ga-68/Ge-68)

γ emitter (Cs-137)



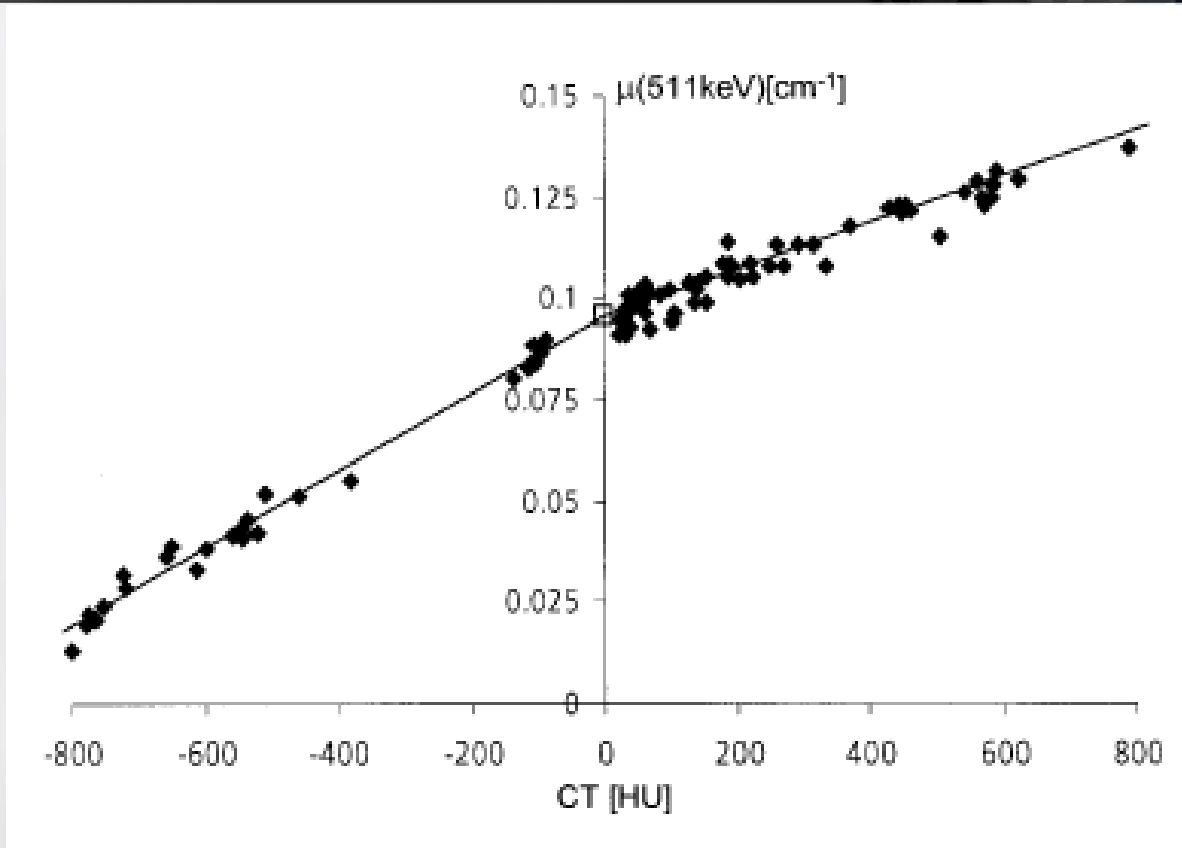
PET attenuation correction



- Noisy
- Need for segmentation
- Long acquisition times (15-20mins)



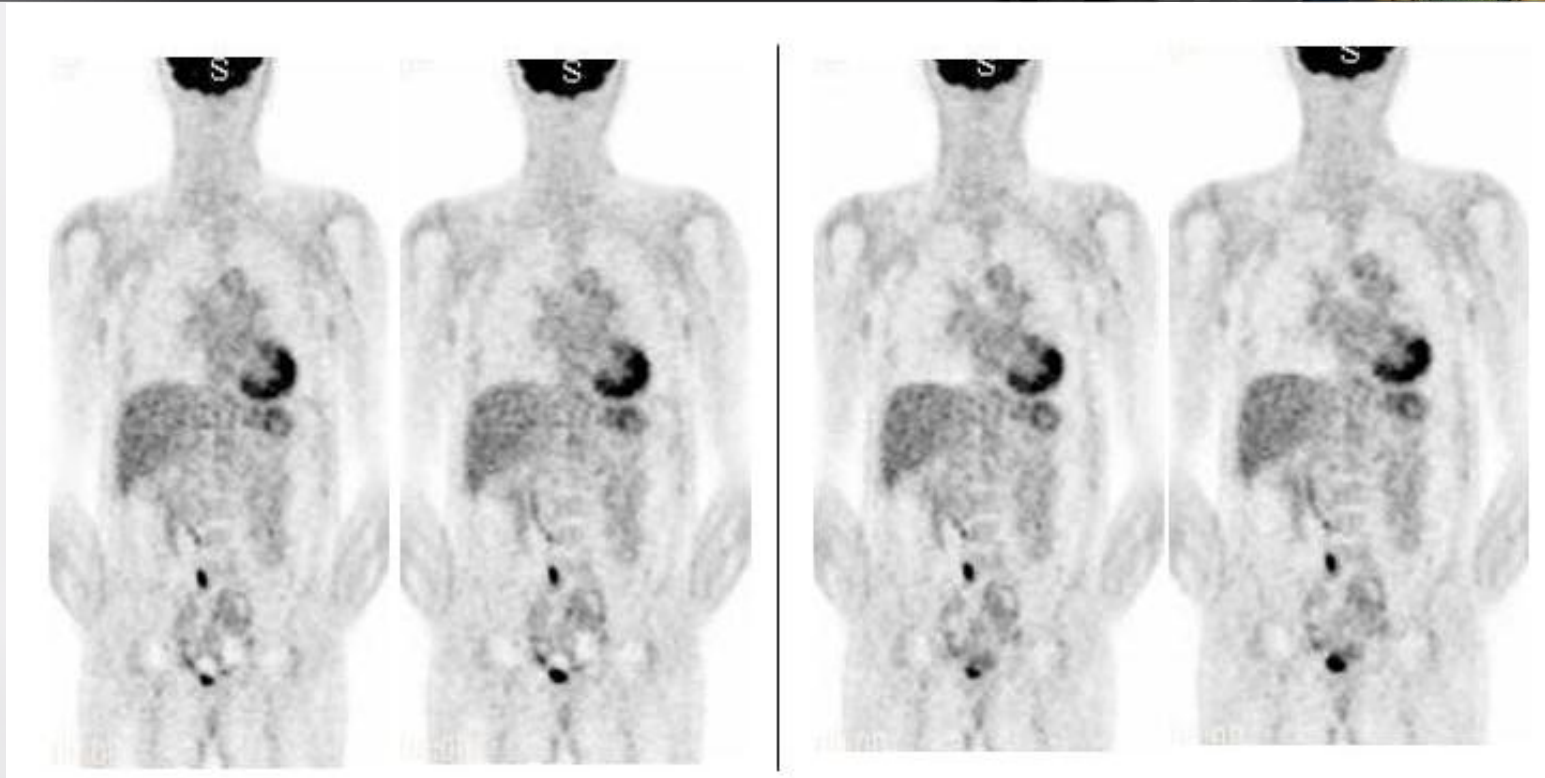
PET/CT attenuation correction



$$\mu = \mu_{H2O} \frac{(HU + 1000)}{1000}$$

$$\mu = \mu_{H2O}^{PET} + HU \frac{\mu_{H2O}^{CT} (\mu_{bone}^{PET} - \mu_{H2O}^{PET})}{1000(\mu_{bone}^{CT} - \mu_{H2O}^{CT})}$$

PET/CT attenuation correction



Standard AC

CT based AC

- Equivalent quantitative accuracy
- > 20% superior contrast

Question **Combining PET and MR: challenge** or opportunity?

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Tissue classification based on 2 point Dixon's MR sequence



Water



Fat



Classify
and combine

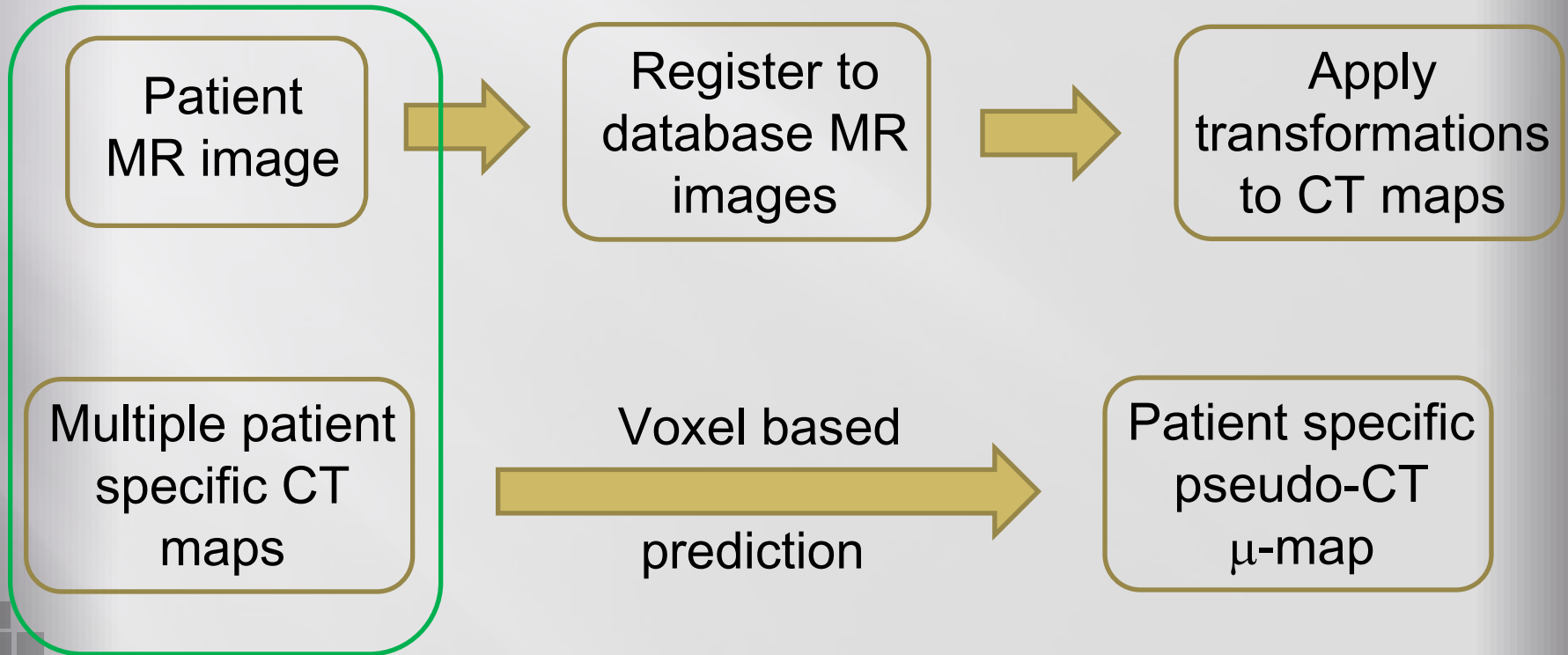


Corresponding CT



Attenuation correction: atlas based

Machine learning approach: Atlas registration and pattern recognition
 (Atlas: Database of registered CT and MR acquisitions)



Attenuation correction: comparison

Tissue classification vs Atlas based

Average Relative VOI PET Quantification Errors for Mean SUVs; Results Compared with Those Achieved Using CTAC

Region	CT _{nobone}	CT _{seg}	PsCT _{MRSEG}	PsCT _{AT&PR}
Thorax	1.0% ± 0.8%	10.2% ± 7.5%	13.5% ± 10.7%	14.0% ± 11.4%
Abdomen	5.7% ± 8.9%	7.8% ± 10.3%	12.7% ± 10.3%	5.2% ± 4.3%
Pelvis	3.7% ± 4.2%	5.5% ± 5.9%	15.8% ± 9.6%	4.9% ± 5.1%
Average	3.6% ± 6.0%	7.6% ± 8.2%	14.1% ± 10.2%	7.7% ± 8.4%
Lesions	2.8% ± 4.5%	4.0% ± 5.2%	7.5% ± 7.9%	5.7% ± 4.7%

Errors are averaged across all patients for body regions and lesions. Data are mean ± SD.

Account for bone structures

Use of Ultra-short Echo Time Triple Echo MR sequence

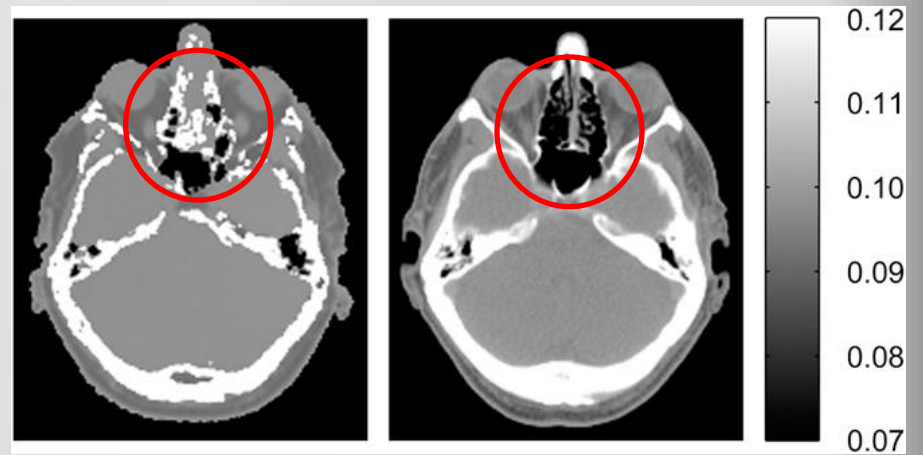
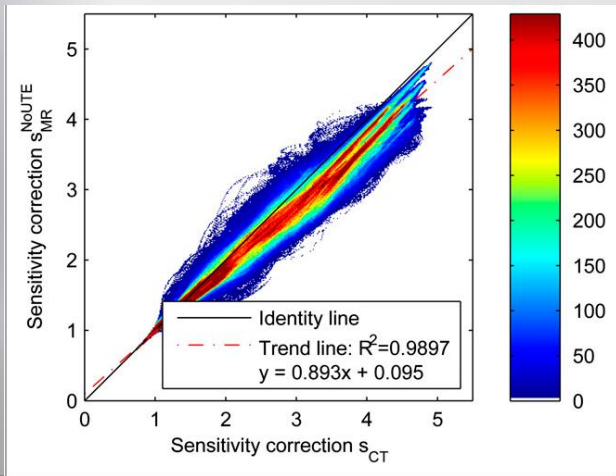
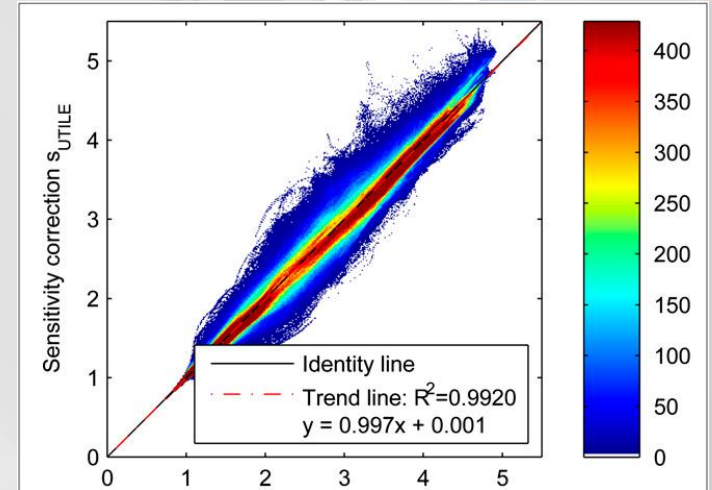
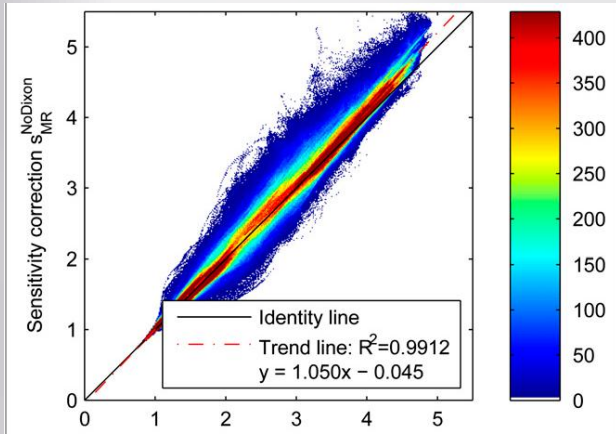
- ✓ Ultra-short echo time sampling for bone segmentation based on a dual-echo technique (Keereman et al JNM 2010, Catana et al JNM 2010)

+

- ✓ Gradient echos for Dixon water-fat separation

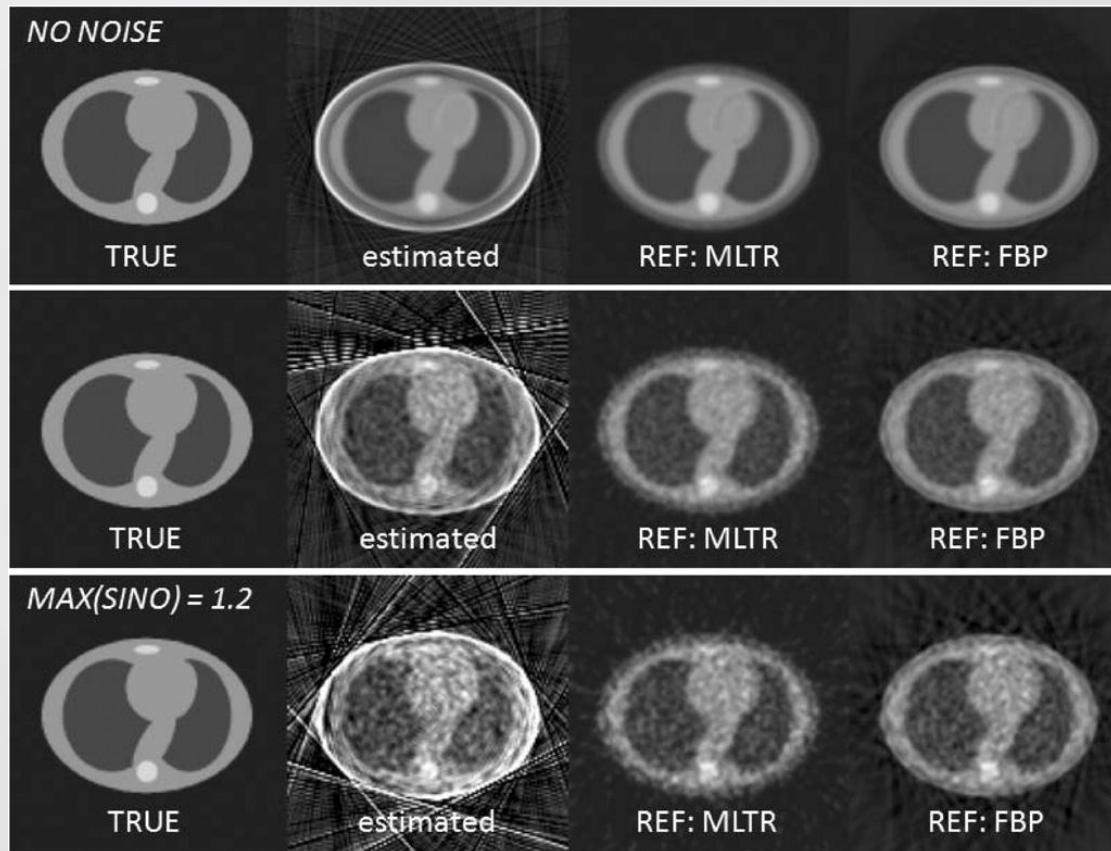
4 tissue classification (air, fat, soft tissue and bone)
using predefined attenuation factors

Attenuation correction: improvements



The solution may come from PET

Explore consistency conditions and TOF information to derive the attenuation maps from the non-attenuation corrected emission datasets



Multimodality imaging: software developments

Question **Combining anatomy and function:** challenge or **opportunity?**

From a software point of view: a little bit of both

1. Corrections:

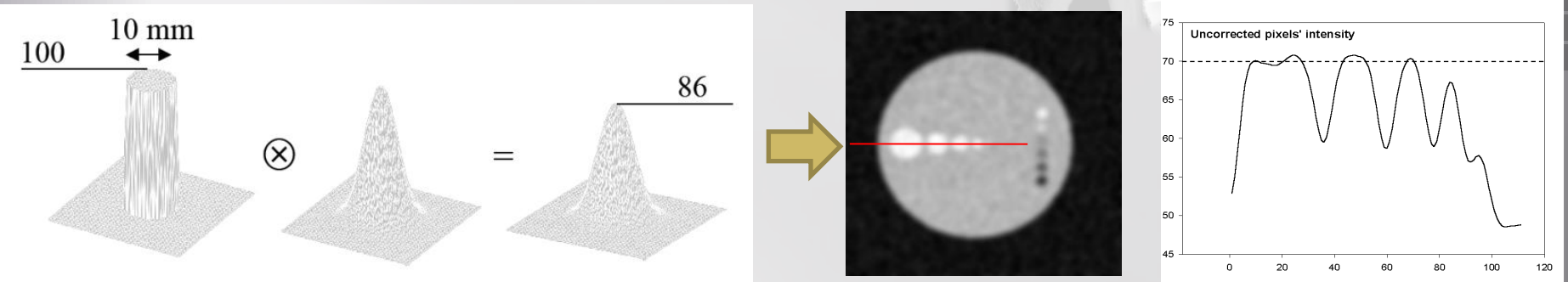
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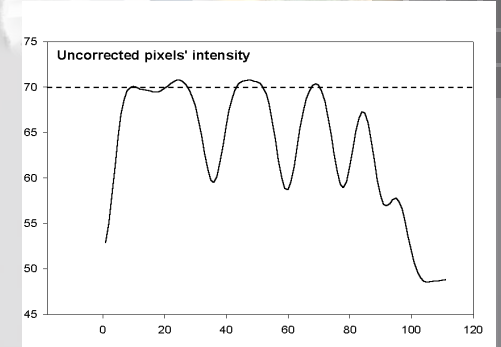
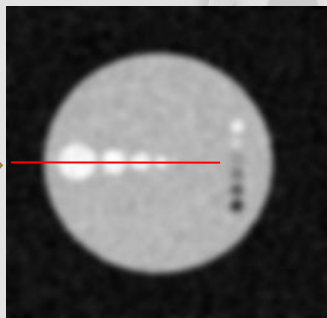
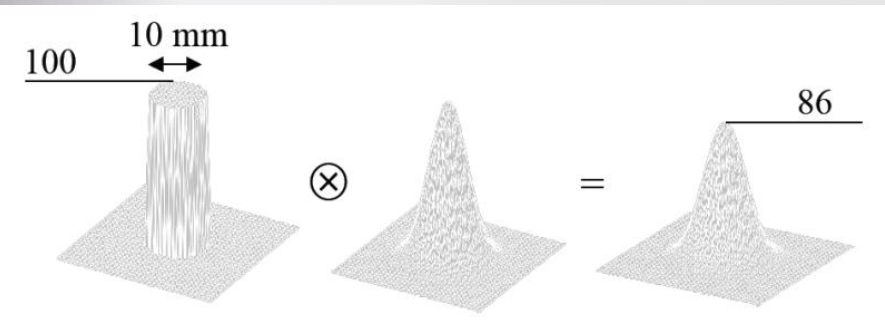
Partial volume effects correction



Different classifying categories may be identified

- (i). Emission only or anatomical detail based
- (ii). Region of interest or voxel-wise based
- (ii). Segmentation needed or not
- (iii). Assumptions considering regions homogeneity
- (iv). Post-reconstruction vs reconstruction based

Partial volume effects correction



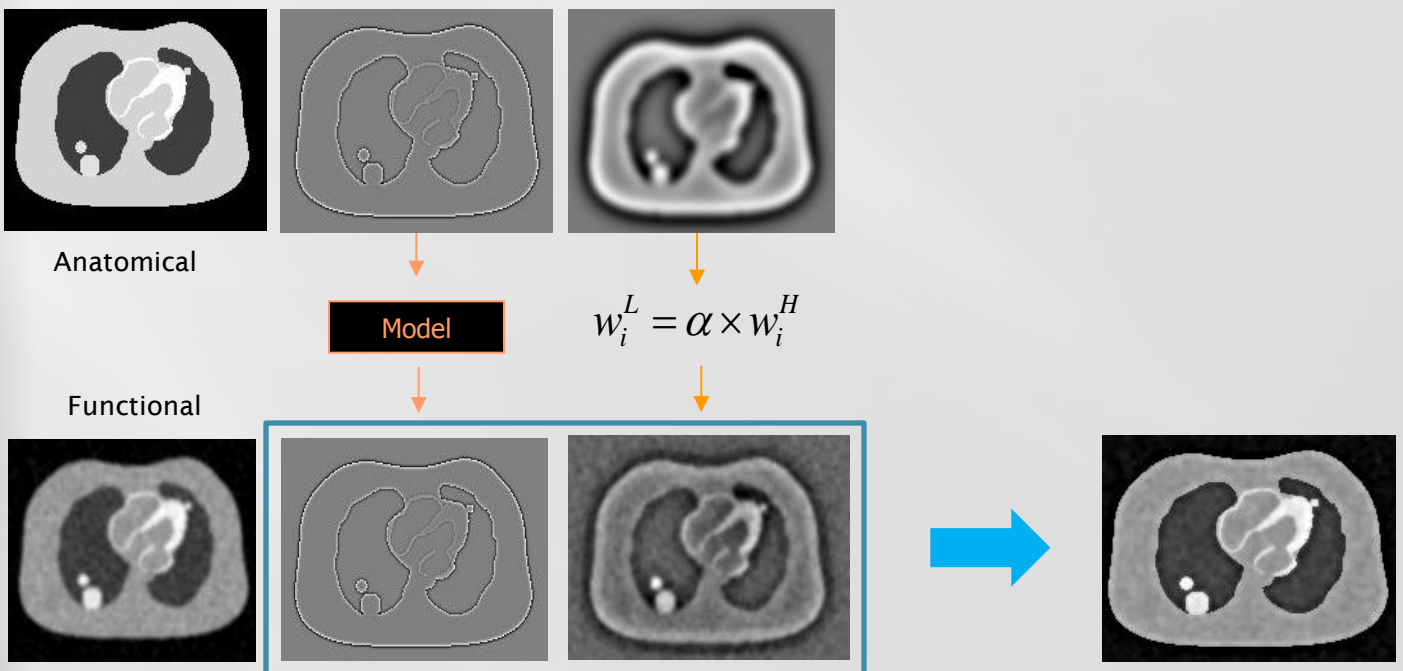
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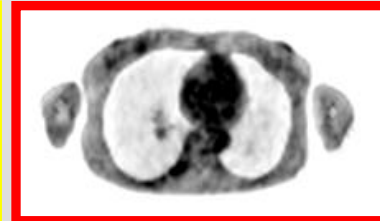
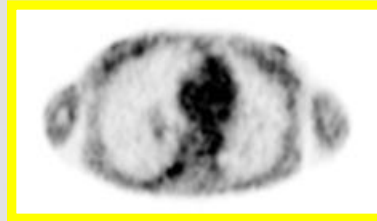
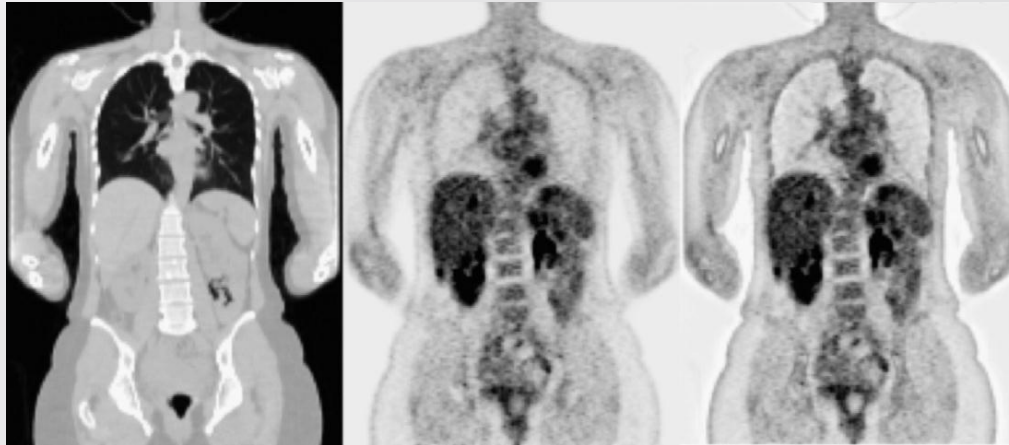
Partial volume effects correction

Post-processing voxelwise approaches (overall image improvement)

Multi-resolution Analysis

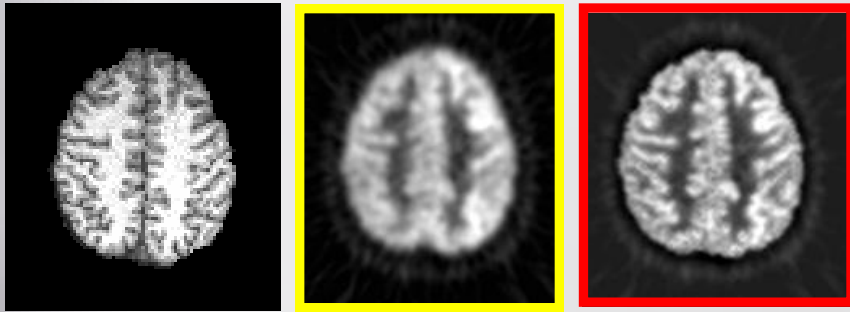


Oncology:

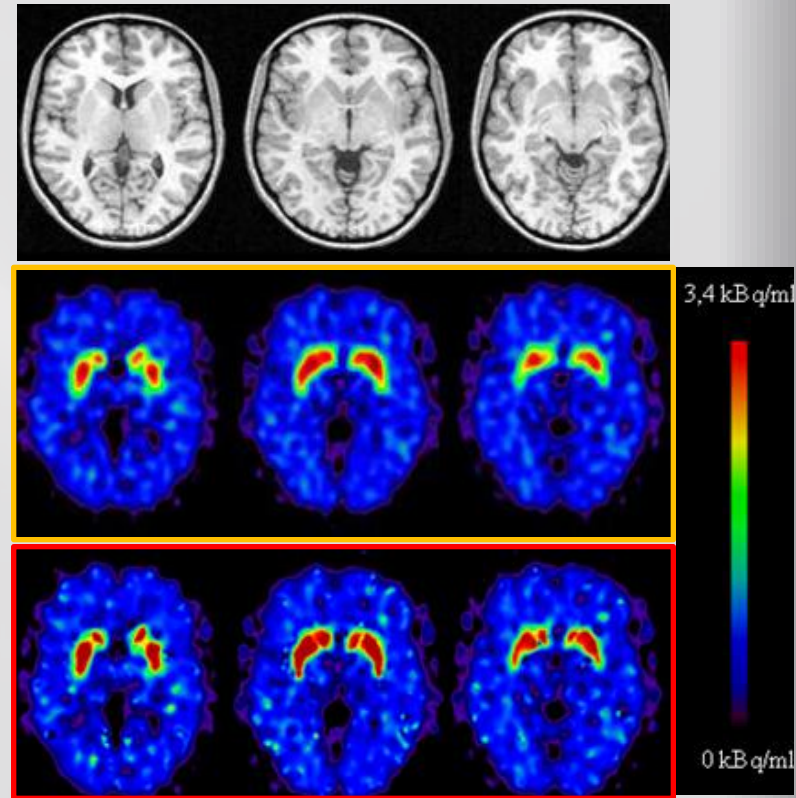


Lung backgd activity +2%
Tumour/Lung activity +20%

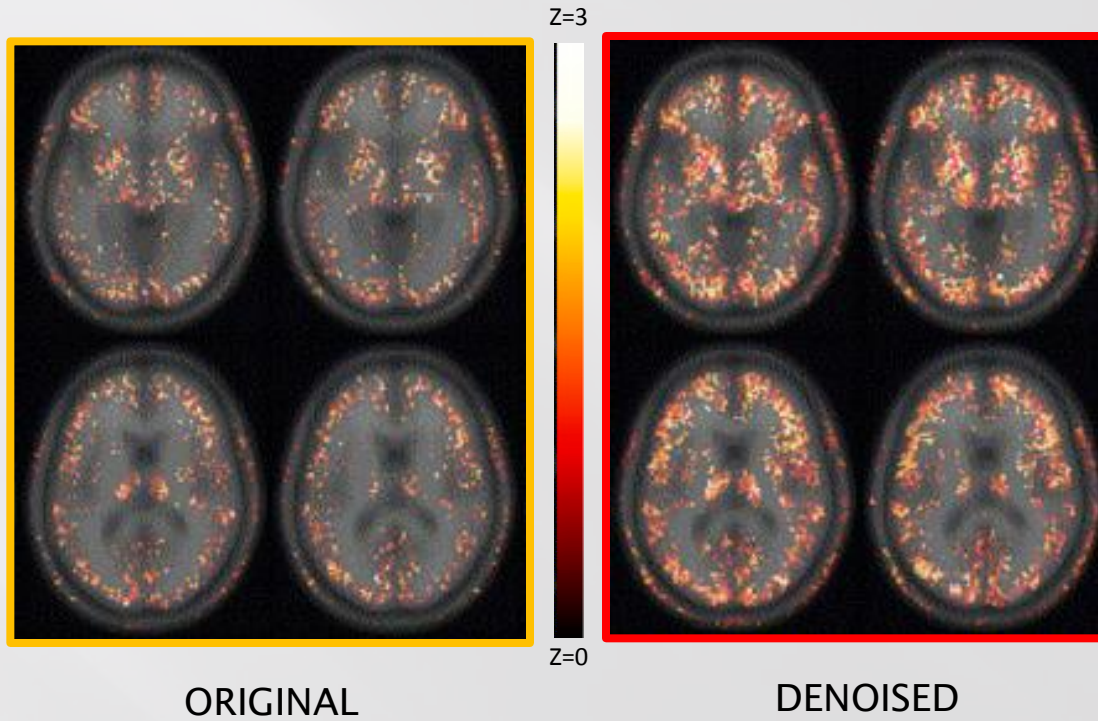
Neurology:



White matter -20%
Grey matter +30%



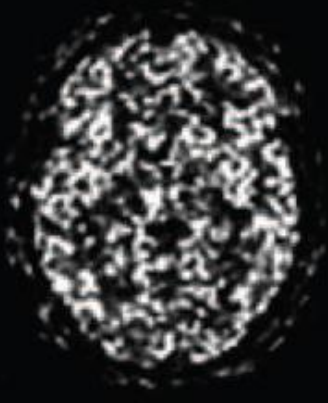
Multi-resolution Analysis: denoising



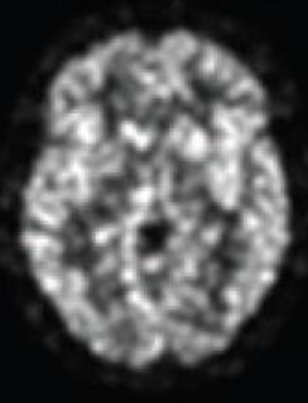
Reconstruction based PVC using MR based priors



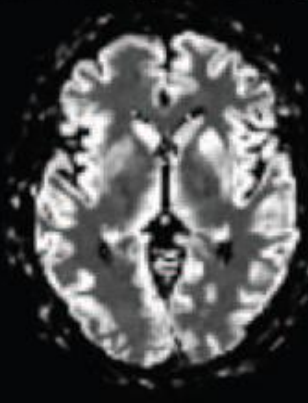
MLEM



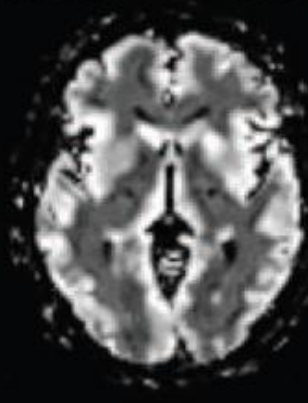
Post-sm. MLEM



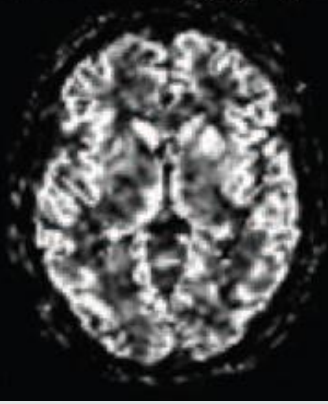
A-MAP perf. segm.



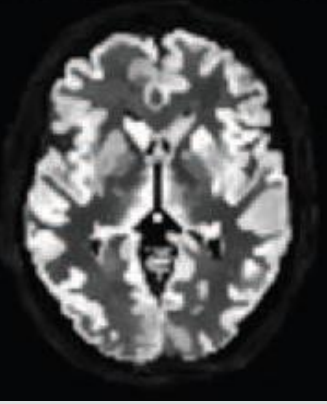
A-MAP SPM segm.



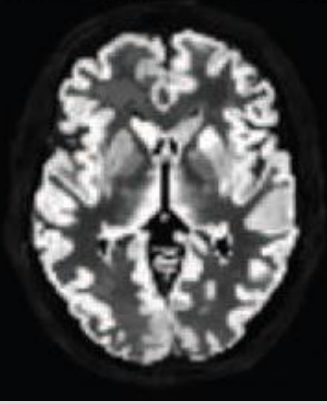
Joint entropy (JE)



Bowsher 4 of 18



Bowsher 9 of 80



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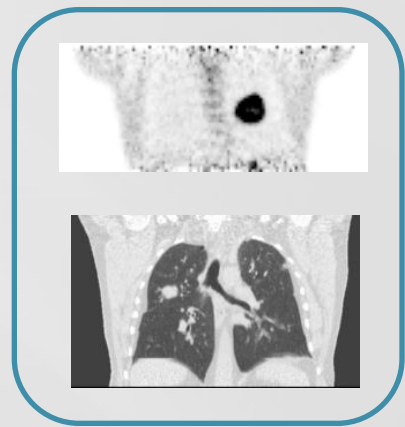
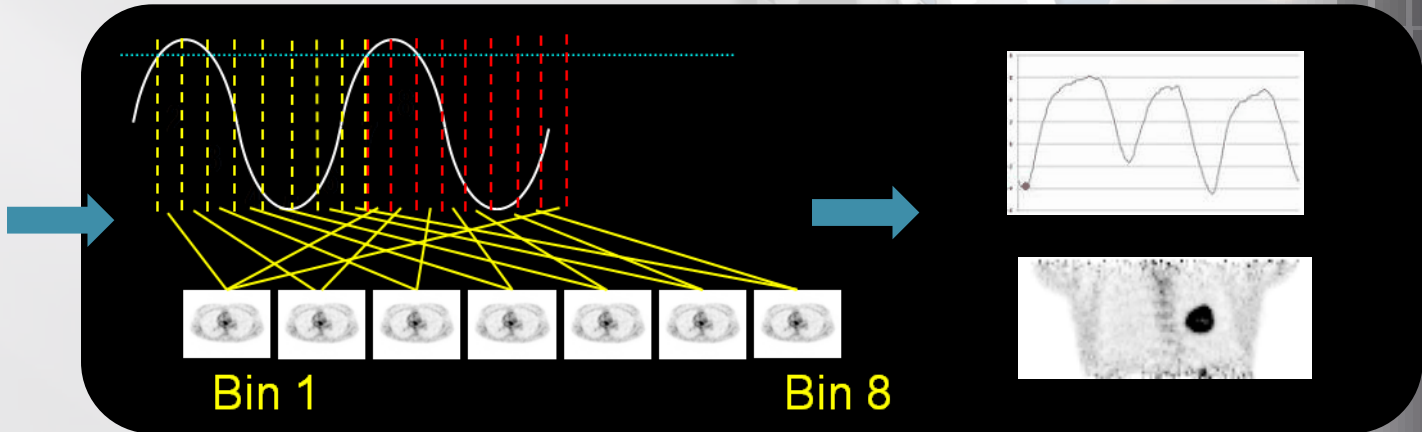
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Respiratory motion: PET/CT

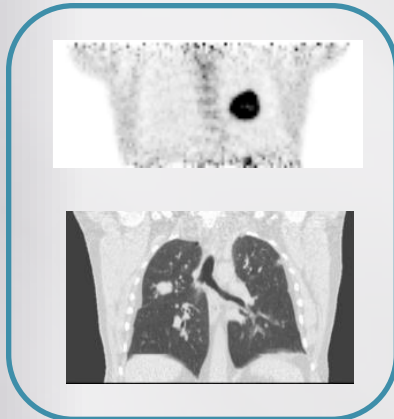
Synchronisation
External detectors

This block illustrates the synchronization process. It shows two photographs of external detectors used for respiratory monitoring. Below the photos is a graph of a respiratory waveform, which is used to time-align the PET and CT scans.

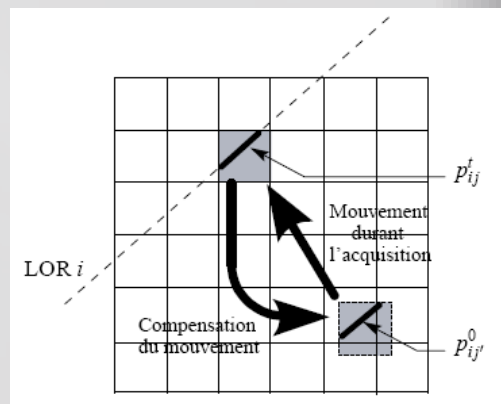
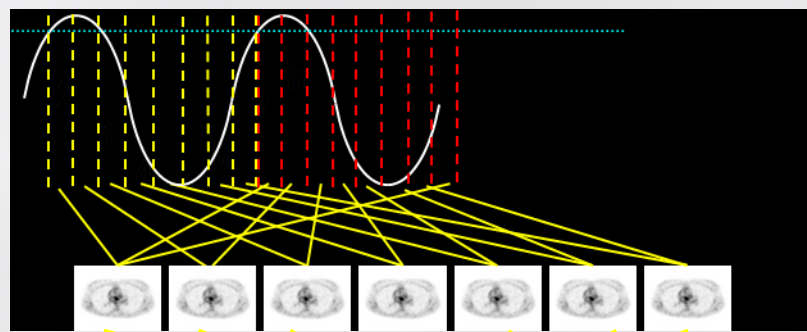
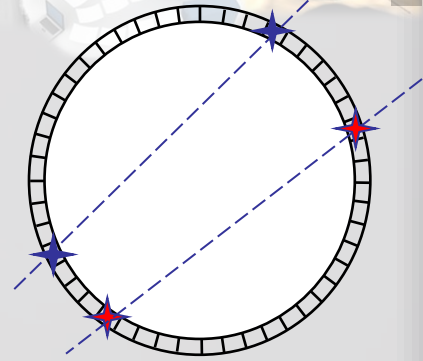


Affine or
deformable
motion model

Respiratory motion: PET/CT



Affine or deformable motion model





Can we use the same workflow for motion correction in PET/MR?

YES!

Differences that need to be taken into consideration?

- ✓ Temporal sampling
- ✓ MR sequences that measure motion

Are there any **advantages** / **disadvantages** in using MR?

- ✓ No ionising radiation
- ✓ Simultaneous acquisition
- ✓ MR incompatibility of different external motion measurement devices

Different protocols for motion estimation in MR

- ❑ **Dynamic 3D** T1/T2 acquisitions
- ❑ **Dynamic 2D** T1/T2 acquisitions binned into 4D images (using normalised mutual information or 1D navigators)

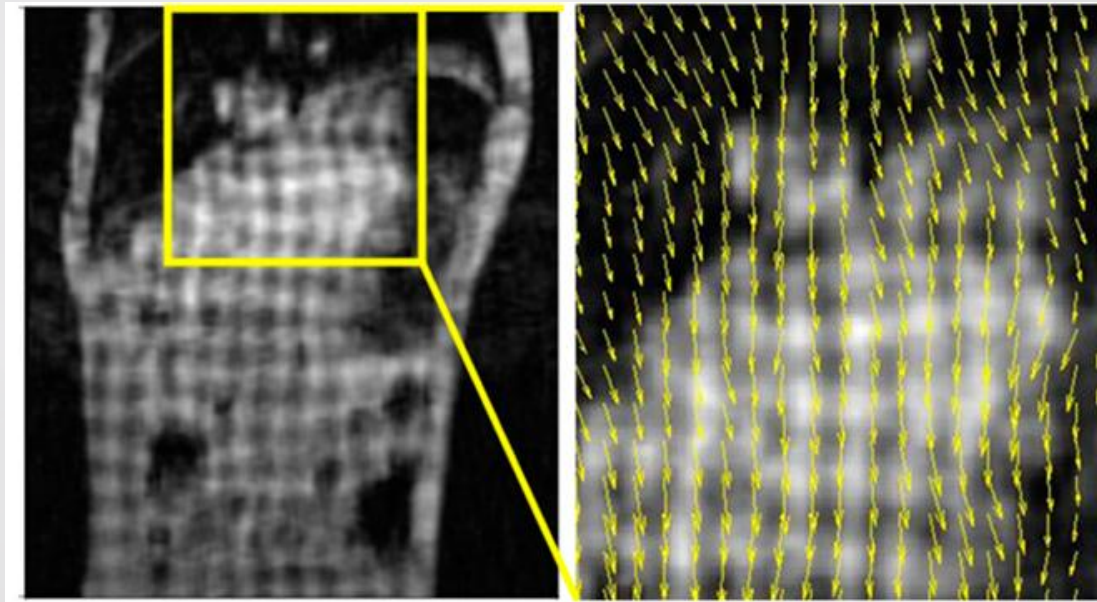
Combination with affine or deformable image registration for the derivation of 3D vector fields

Limitation: only applicable to sequences providing anatomical contrast

Respiratory motion: MR sequences

Different protocols for motion estimation in MR

- ❑ **Tagged MRI acquisition:** superimpose a regular tagging pattern on the object magnetisation distribution (1D, 2D or 3D in the phase or amplitude domains)



Different protocols for motion estimation in MR

- ❑ **Tagged MRI acquisition:** superimpose a regular tagging pattern on the object magnetisation distribution (1D, 2D or 3D in the phase or amplitude domains)

Advantage: Does not require the use of deformable image registration for the derivation of 3D vector fields

Limitation: Does not allow estimation of motion in the lungs.
MR images are usually affected by artefacts.

Different protocols for motion estimation in MR

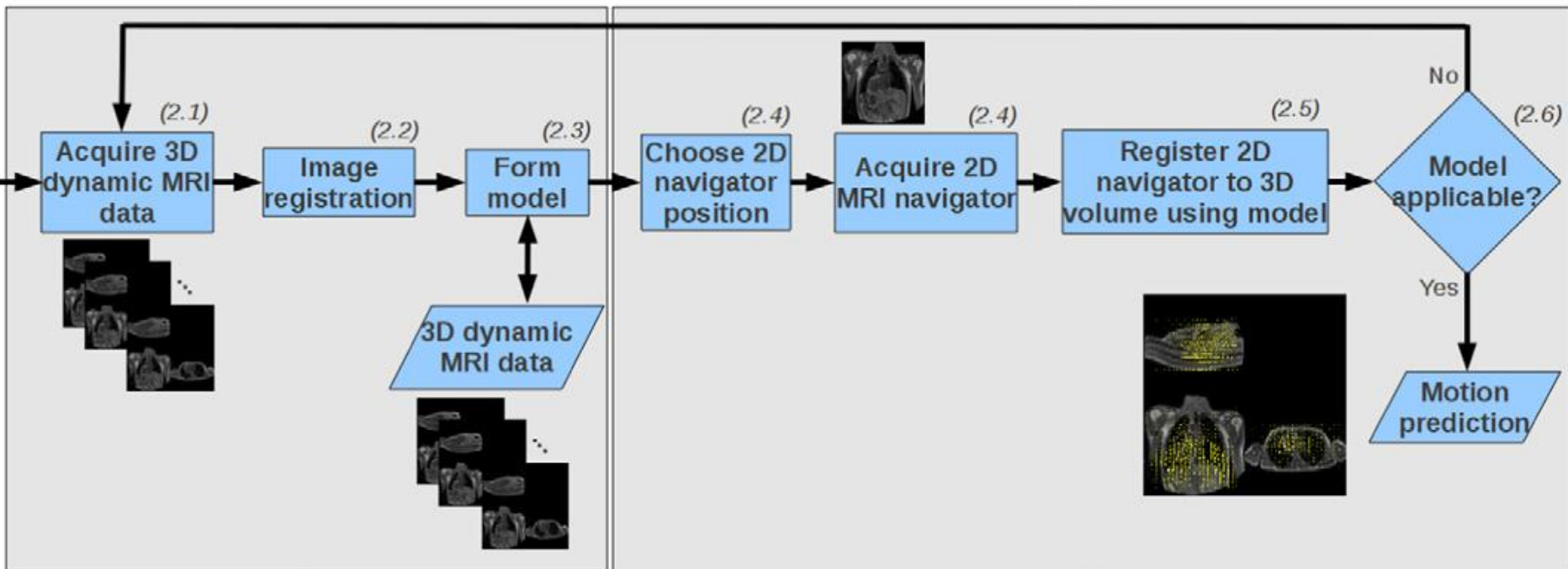
- ❑ **Patient specific motion model** based on establishing a relationship between 4D internal structures displacement and diaphragmatic 1D/2D navigators

Respiratory motion: motion modelling

Different protocols for motion estimation in MR

Model formation

Model application



Different protocols for motion estimation in MR

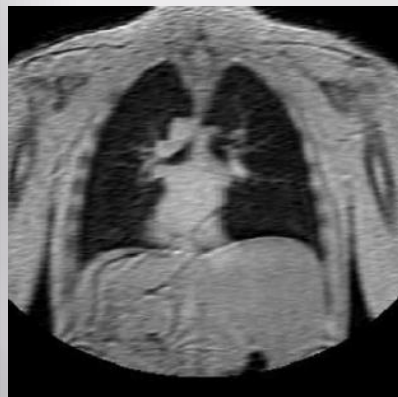
- ❑ **Patient specific motion model** based on establishing a relationship between 4D internal structures displacement and diaphragmatic 1D/2D navigators

Limitation: A dynamic 3D T1 acquisition is necessary for each patient in order to derive the motion model



Different protocols for motion estimation in MR

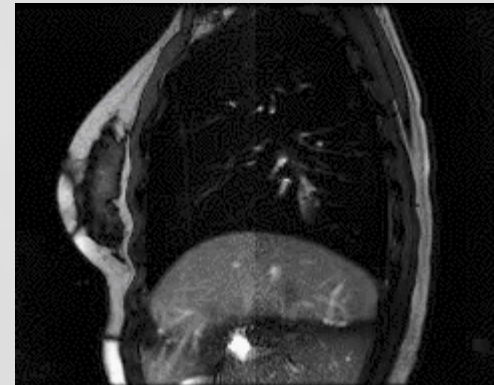
- ❑ **Generic motion model** based on using a patient database of 4D internal structures displacements and associated diaphragmatic 1D/2D navigators



2D Navigator

+

$$U(t) = B s(t)$$



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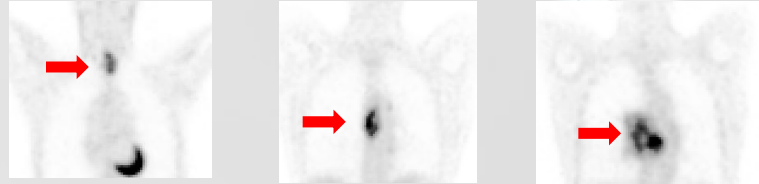
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PET image derived parameters

✓ Predictive and prognostic value of functional volume for different cancer models

Initial FDG PET scan

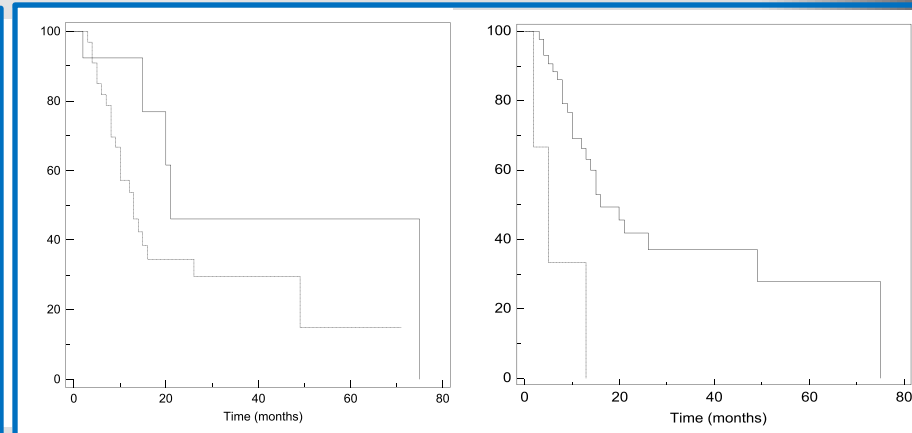
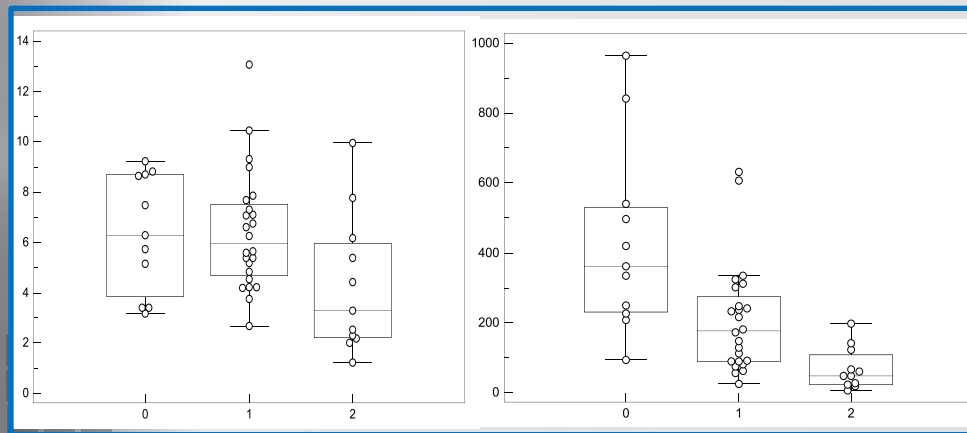


Better patient stratification and management (survival, therapy response)

Example: 45 esophageal cancer patient

Therapy response

Survival



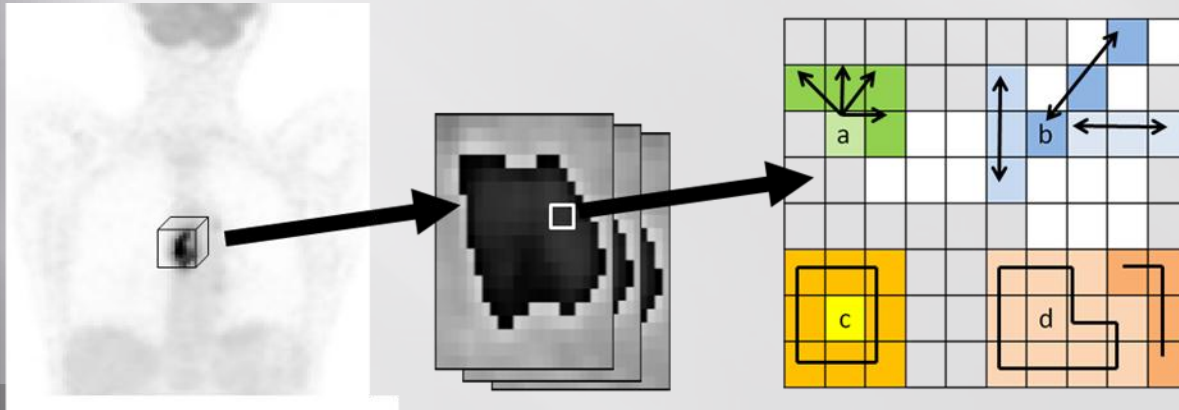
Characterising tracer distribution

✓ Characterisation of tumour activity distribution

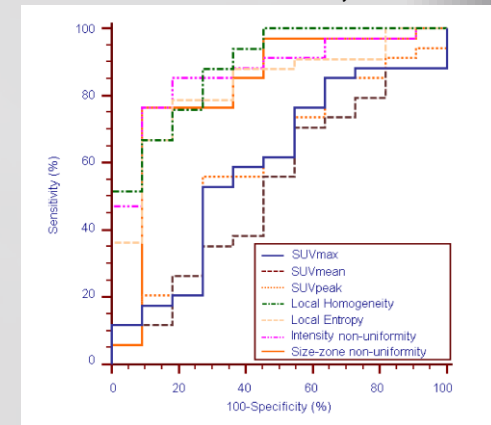
Hypothesis: Intra-tumour tracer distribution variability represents a challenge in molecular imaging. Is there any additional information that can be extracted from PET images?

Objectives: Analysis of the activity distribution at local and regional levels

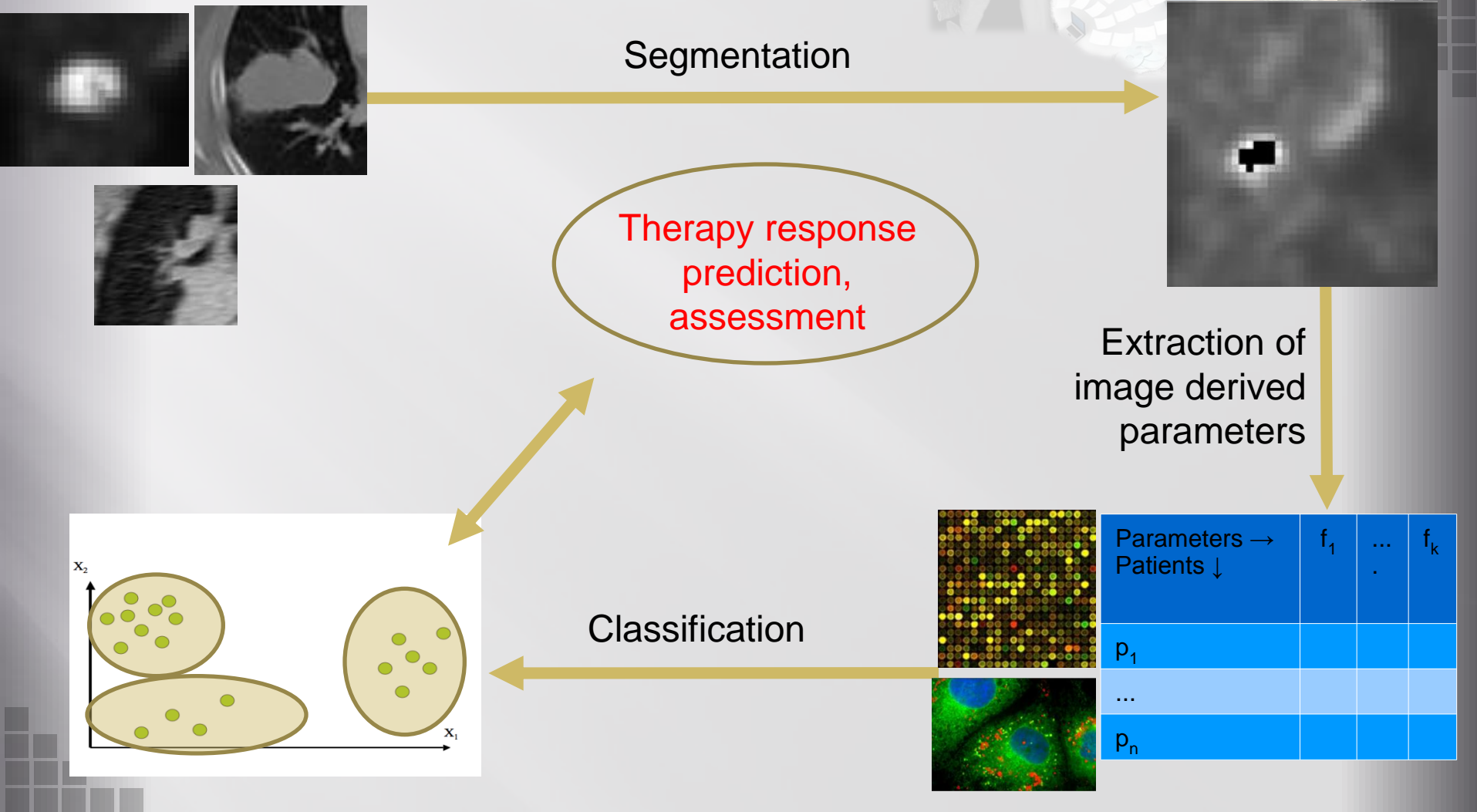
1. Stratification: therapy response, survival, distinction benign / malignant lesions
2. Correlation with molecular and biological measures (genomics)
3. Multi-tracers and multi-modality analysis (ex: $\alpha\beta 3$ Expression, glucose métabolisme, diffusion)



Entropy, Local Homogeneity, Regional Homogeneity

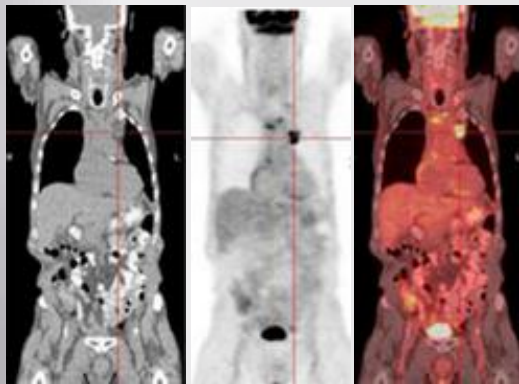


Multi-modality information paradigm

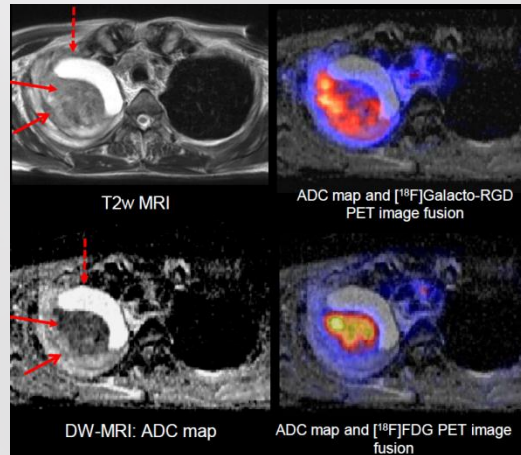


Multi-modality information paradigm

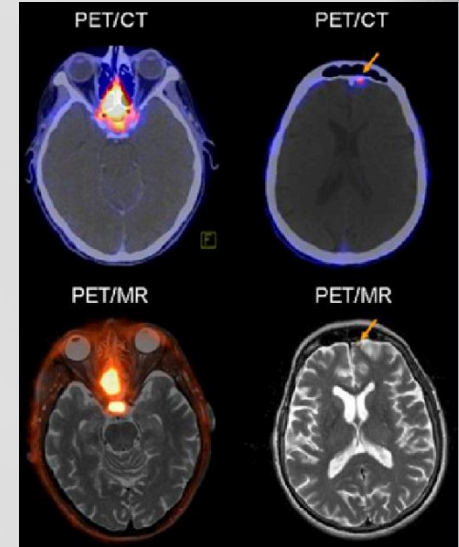
Multi modality / multi tracer imaging: PET/CT, SPECT/CT, PET/MRI...



CT PET/CT PET/CT fusion



A.J. Beer, et al. *SNM annual meeting 2010*



B.J. Pichler, et al. *Journal of Nuclear Medicine 2010*

- ✓ Combining function and anatomy
- ✓ Multi modal imaging PET/MRI, multi-tracers, 4D imaging, etc.
- ✓ Towards increasingly large datasets with heterogeneous but complementary information

Multi-modality and temporal evolution

Extraction of pertinent information:

FDG

CT



FLT

MRI



- Information fusion
- Segmentation
- Classification

	Temps					
Modalité 1 Traceur 1			...			Suivi thérapeutique / temporel
Modalité 1 Traceur 2			...			
...	
...	
Modalité i Traceur j			...			
			

Analyse multi-traceur multi-modale

Before treatment

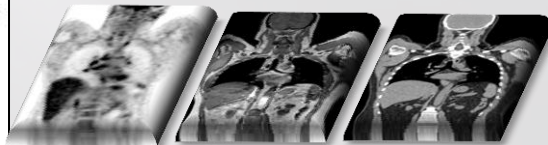
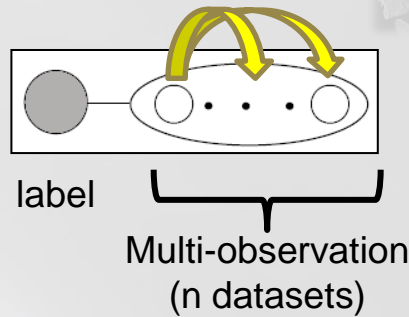
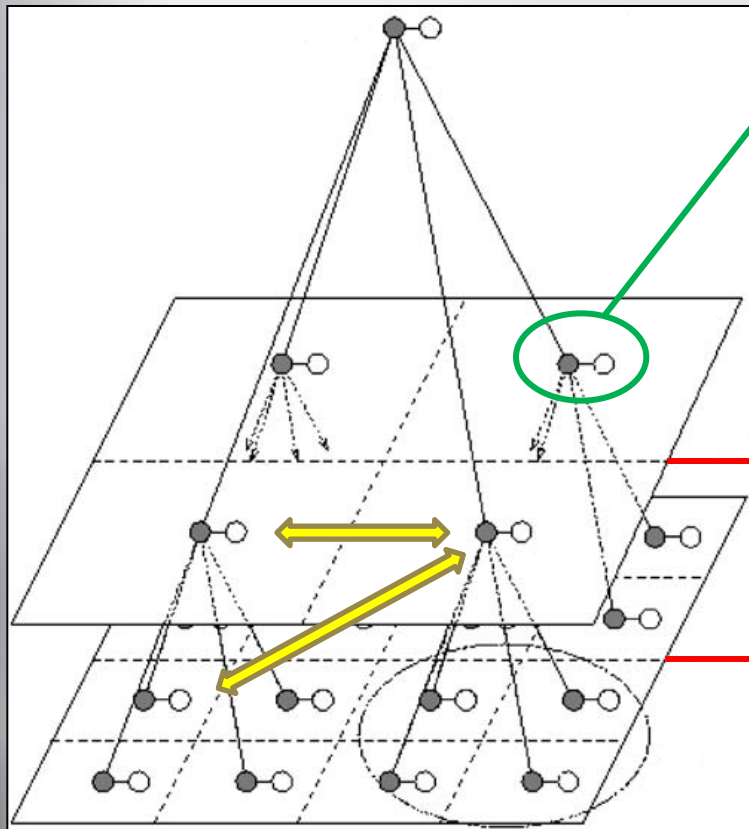


During treatment



Multi-resolution and multi-observation hierarchical models

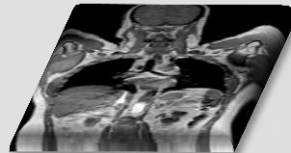
Bayesian inference



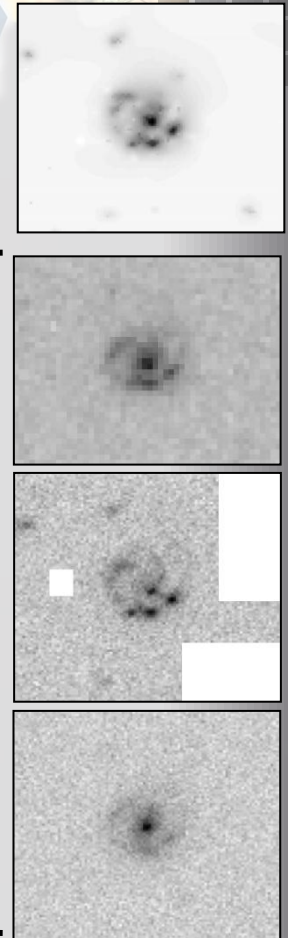
resolution i



resolution $i \times 4$



Fusion



Ex. Astronomy images
(3 observations)

- Handling missing data
- Handling data of variable resolution levels
- Possibility of complex models (correlated noise, fuzzy modeling, etc.)



Multimodality imaging is producing challenges in terms of software development but also plenty of opportunities.

MR sequences is the key for providing answers to a number of challenges associated with PET/MR.

Existing approaches previously developed in PET/CT multimodality imaging may be explored within the PET/MR field.

New software developments are necessary to efficiently explore the multitude of information that may be available from multimodality imaging.



