

MULTI-MODAL IMAGE INTEGRATION



*CARLO CAVEDON
MEDICAL PHYSICS UNIT
VERONA UNIVERSITY HOSPITAL - ITALY*

*ICTP SCHOOL ON MEDICAL PHYSICS
FOR RADIATION THERAPY
TRIESTE – ITALY – 16 APR 2015*



The Abdus Salam
**International Centre
for Theoretical Physics**
50th Anniversary 1964–2014

MULTIMODAL IMAGE INTEGRATION vs. REGISTRATION

- **image integration** = the use of two or more image sets in the process of (i.e.) treatment planning
- **image registration** = the process of making two or more image sets spatially coherent to each other
- **image fusion** = the simultaneous visualization of two or more image sets, previously coregistered

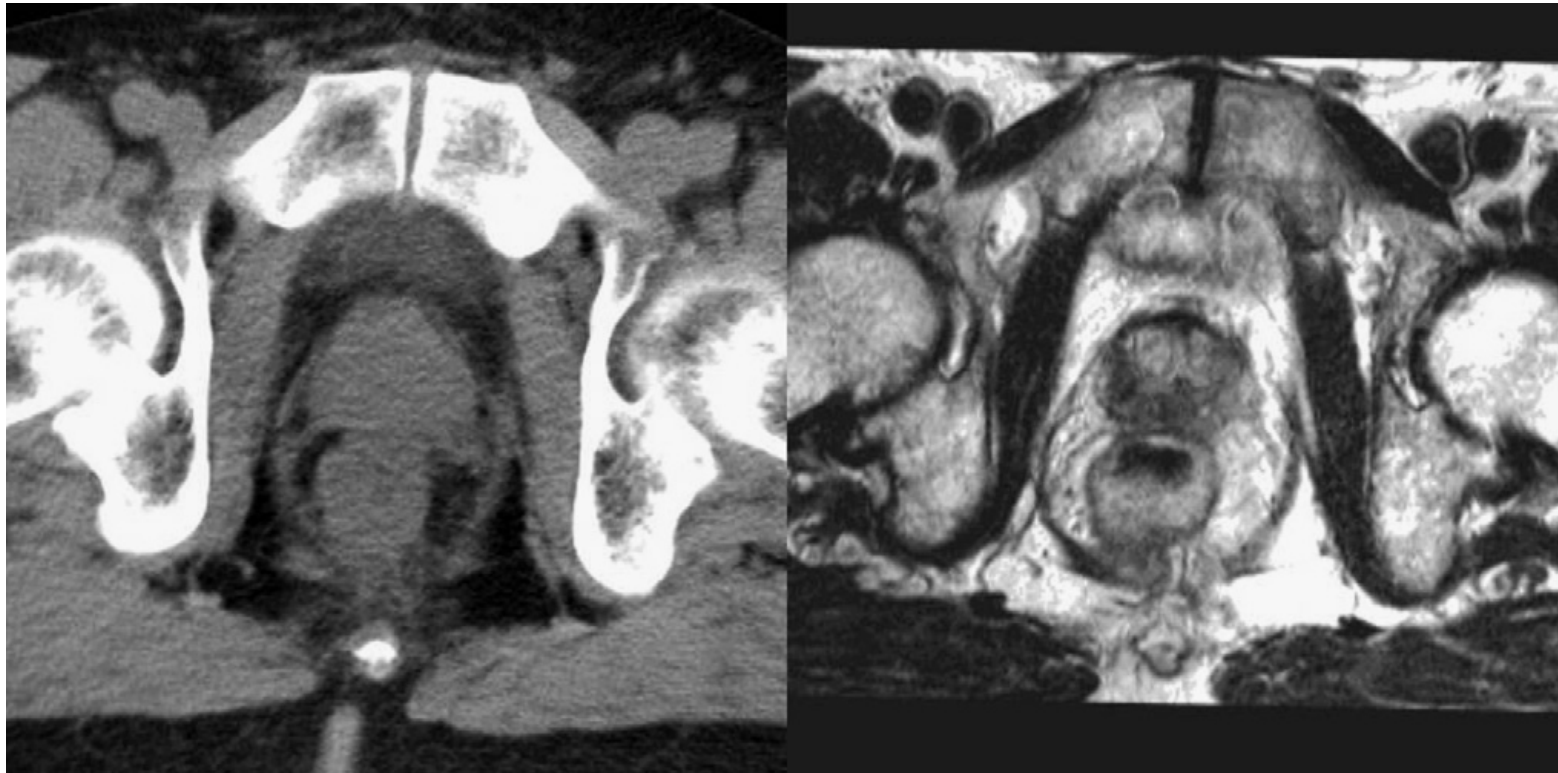
IMAGING MODALITIES RELEVANT TO TREATMENT PLANNING

- **computed tomography (CT)**
 - basic modality for treatment planning
- **magnetic resonance imaging (MRI)**
 - multimodality imaging technique
 - morphological and functional information
- **PET-CT**
 - low resolution datasets
 - CT inherent to modality – easy spatial reference
- **ultrasound (US)**
- **emerging modalities (PET-MR etc.)**

THE CENTRAL ROLE OF CT IN TREATMENT PLANNING

- CT is the tomographic modality that offers the best **spatial accuracy** (freedom from significant distortion etc.)
- CT information can be directly transformed into a **map of attenuation coefficients** => useful in dose calculation
- modern in-room verification systems are based on **x-ray transmission imaging** (e.g. CBCT) => easily registered to CT

MR FOR TREATMENT PLANNING

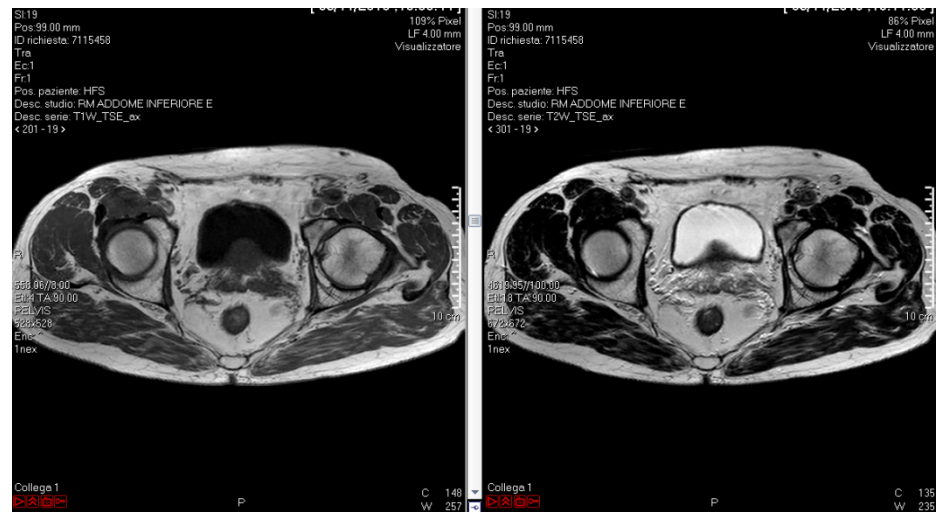


- example: comparison between CT and MR – prostate
- better visualization of soft tissue
- no direct correspondence between “gray levels” => may complicate automatic image registration

MORPHOLOGICAL T1- AND T2-BASED IMAGING

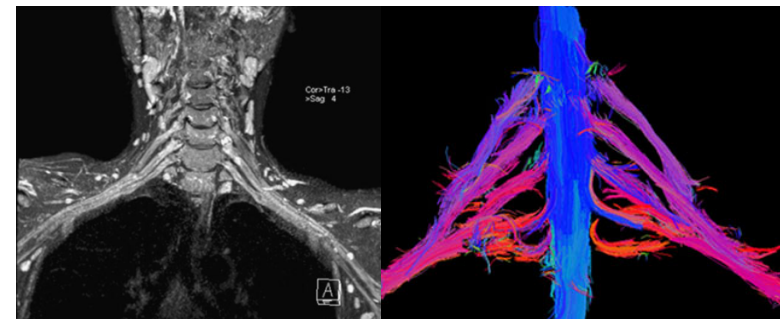
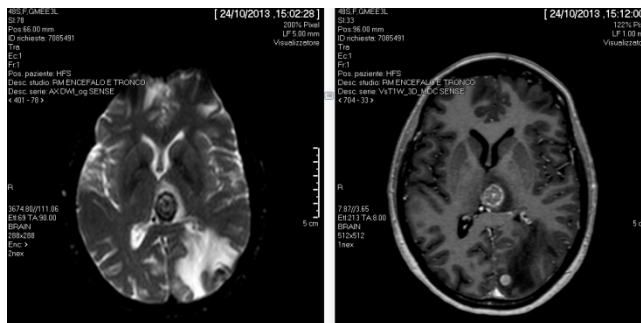
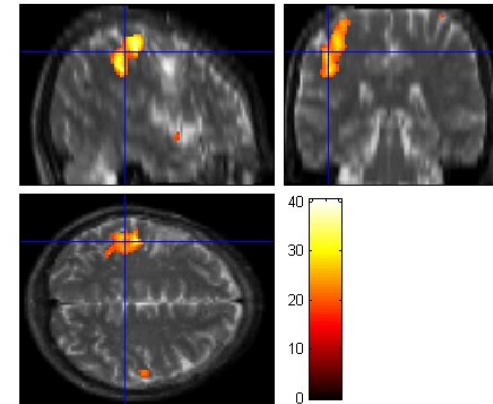
- **T1 and T2** weighting corresponds to imaging with different “modalities”
- **T1** enhances **muscle-fat** - **T2** enhances **water** (fluids)
- Paramagnetic **contrast agents** have more effect on **T1-weighted** images

*left: T1-weighted MR image
right: T2-weighted MR image*



FUNCTIONAL INFORMATION FROM MRI

- MRI can provide valuable **functional information** by means of:
 - *diffusion-weighted imaging (**DWI**) – including maps of apparent diffusion coefficient (**ADC**) and diffusion tensor imaging (**DTI**) – tractography*
 - ***fMRI** based on the **BOLD** effect*
 - *arterial spin labeling (**ASL**)*
 - ...

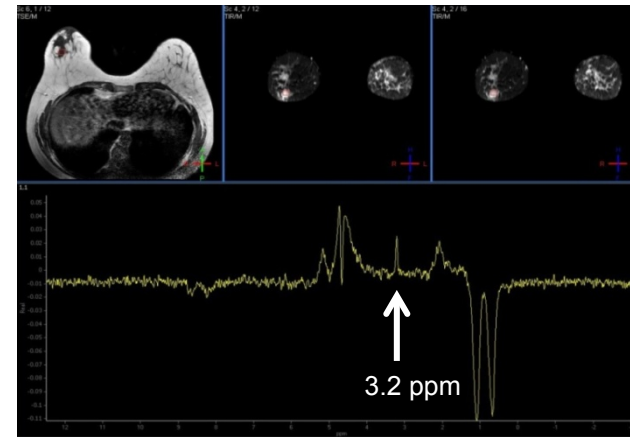
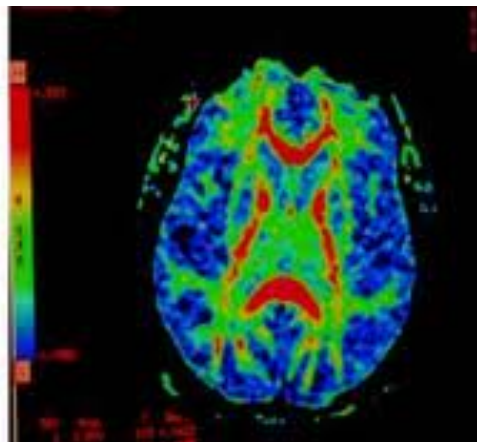
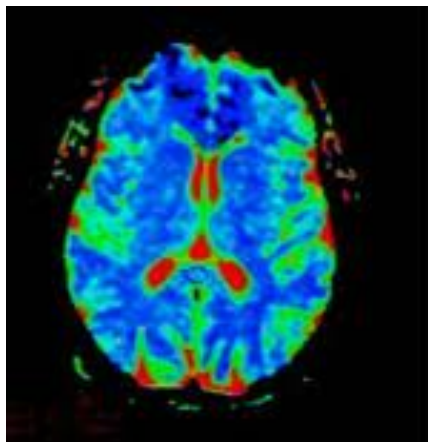


FUNCTIONAL INFORMATION FROM MRI

- functional MRI is characterized by **low spatial resolution (low SNR)**
 - fMRI is often reported on **anatomical atlases** for reference
- => registration to CT might be difficult because of **poor “common information”**

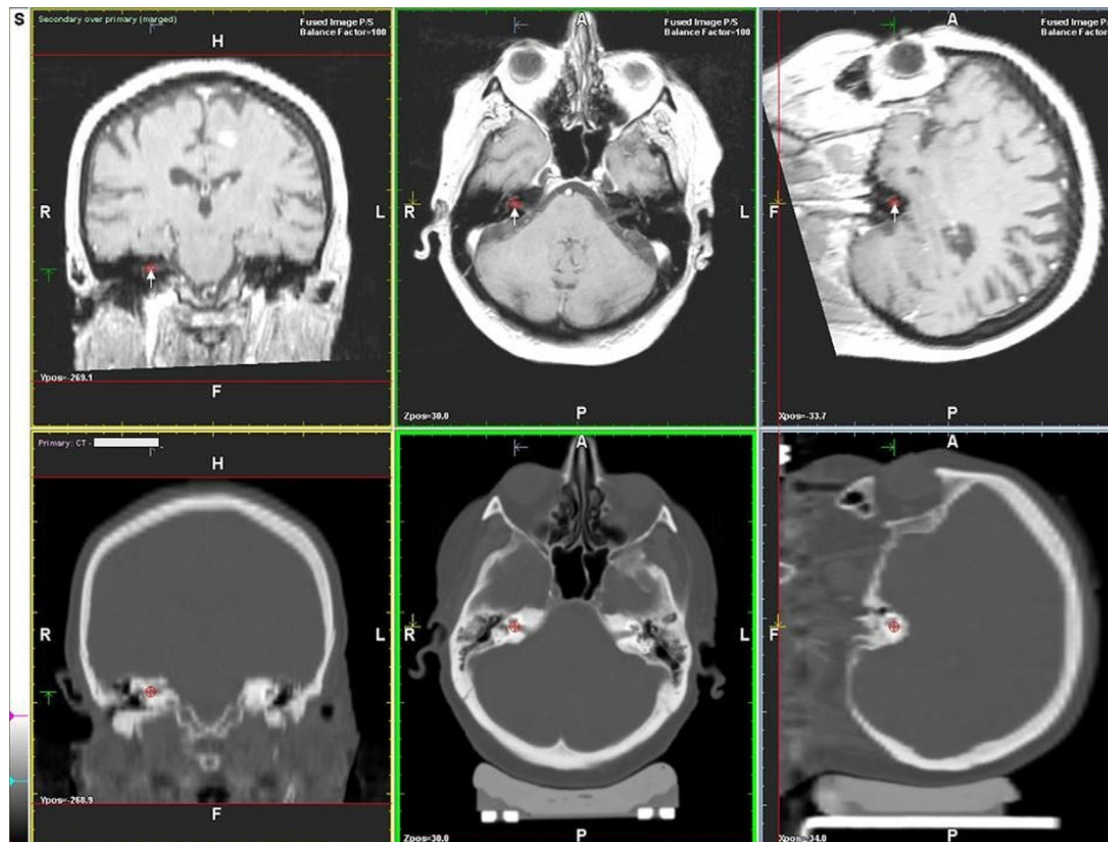
MULTIPARAMETRIC MR IMAGING

- Special MRI modalities such as **DWI** (ADC) and **spectroscopy** may be integrated for diagnostic purposes (multi-parametric imaging)
- Multi-parametric datasets are **usually not employed in the treatment planning process**; special attention needed



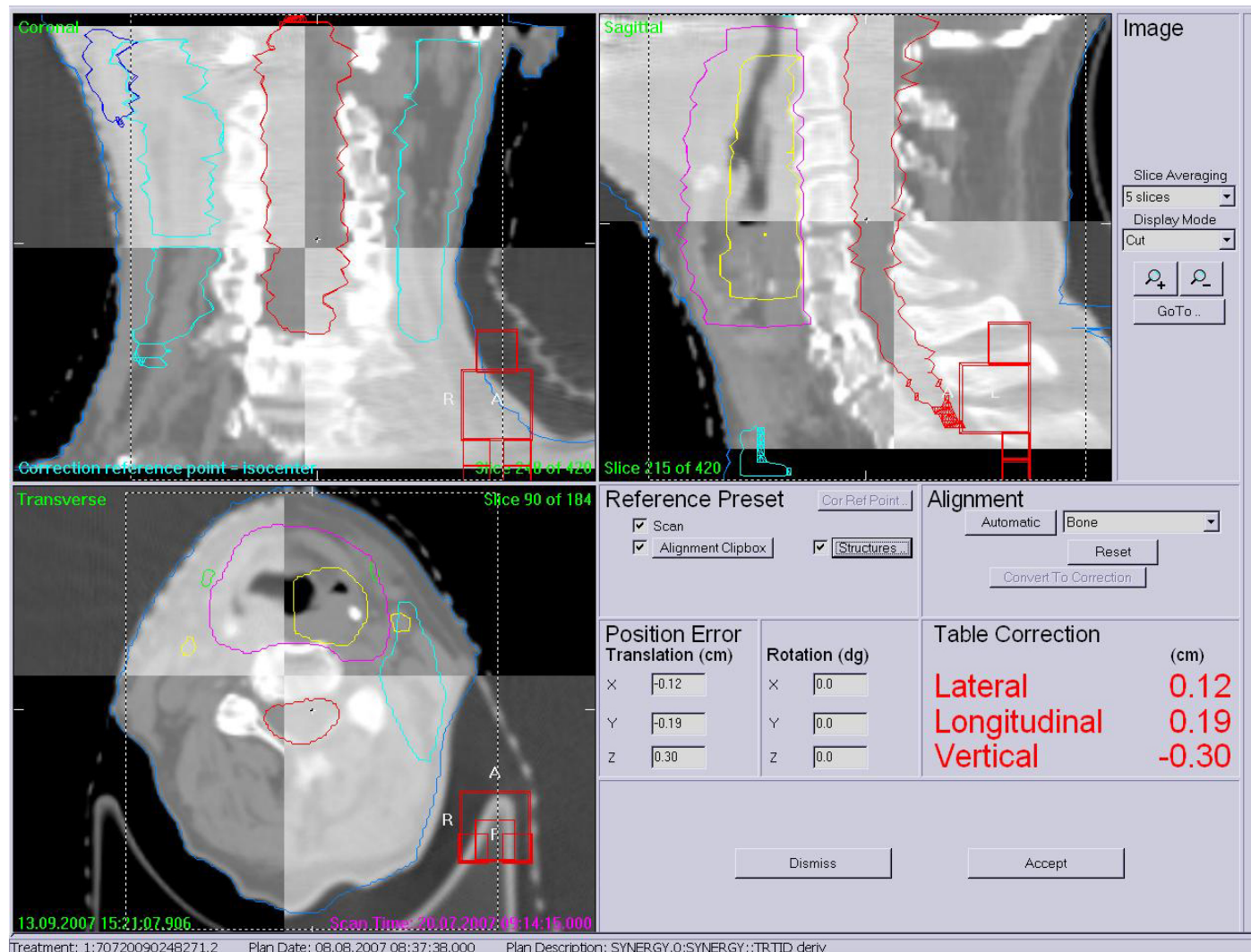
COREGISTRATION BETWEEN MRI AND CT

- Strictly **rigid transformation** in the brain
- 3 translations+3 rotations => 6 parameters



- Diagnostic MRI is usually **rotated around the L-R axis** compared to CT
- **Correction needed** – might not be evident on axial orientation
- Inferior regions might introduce **deformations**

COREGISTRATION BETWEEN MRI AND CT



COREGISTRATION BETWEEN MRI AND CT

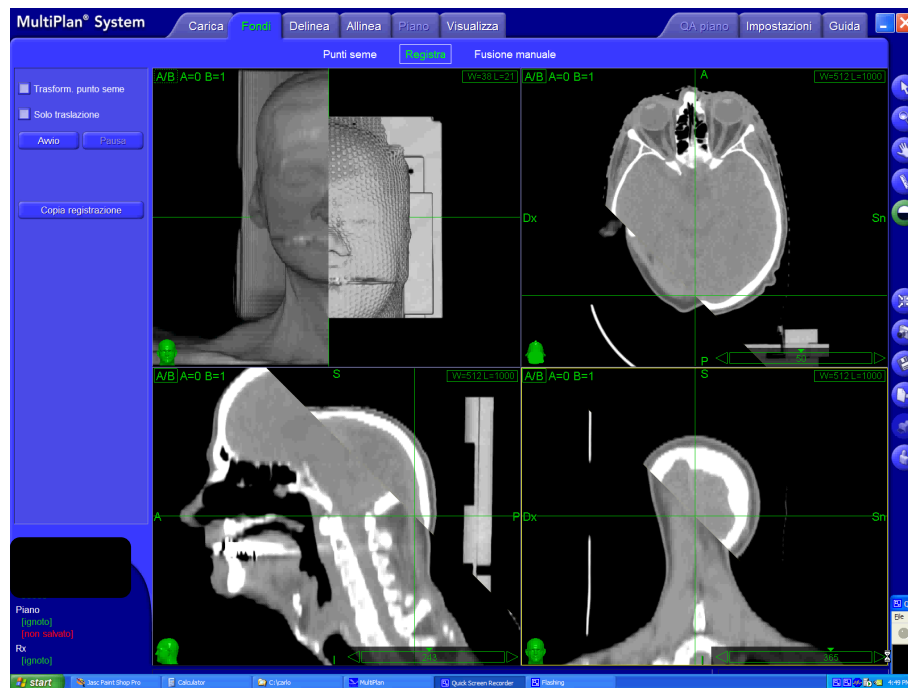
- Use of “**clip-boxes**” in case of deformations to disregard in the registration process
- Commercially available treatment planning systems and 3rd part software **may offer** this functionality
- Privilege the anatomical region that has to be coregistered – leave any uncontrolled region free

COREGISTRATION BETWEEN MRI AND CT

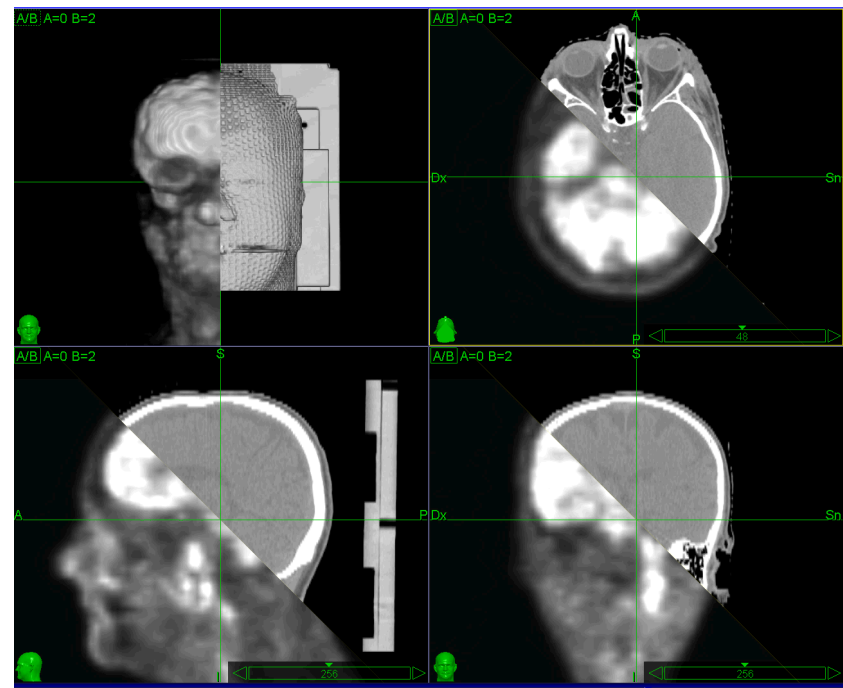
- Obtaining **similar (consistent) initial orientation** is often essential even in case of automatic transformation – robustness of algorithms to different initial orientation is an issue in general
- Use of **patient positioning devices** recommended in case of multimodality imaging – example: PET-to-CT
- Pay attention to **MR compatibility - safety!**

COREGISTRATION: examples

CT-to-CT



PET-to-CT



PET-CT FOR TREATMENT PLANNING

- **^{18}F -FDG PET-CT** imaging is increasingly growing since the introduction of clinical PET-CT scanners (ca. 2000)
- Applications to Radiation Oncology: **PET-based volumes of reference** (BTV=biological target volume)
- Clinical decisions (including “BTV” delineation) generally based on the **Standardized Uptake Volume (SUV)**

PET-CT FOR TREATMENT PLANNING

$$SUV = \frac{c(t)}{A(t)} \cdot bw$$

c = activity concentration (MBq/kg), A = injected activity (MBq), bw =body weight (kg)

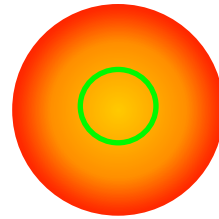
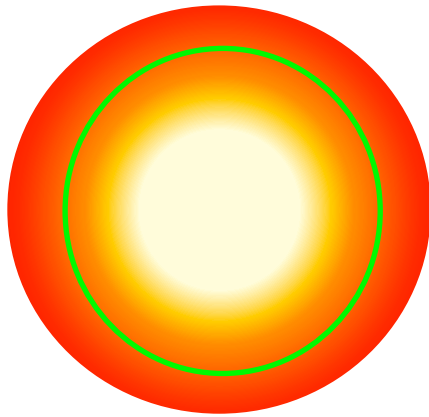
- Importance of **standardization** (patient weight, uptake time, injected activity and correction for decay in the uptake time ...)
- **Lesion motion** might have negative (even destructive) effects on SUV quantification (see specific module)

PET-CT FOR TREATMENT PLANNING

- Use of SUV to define biological volumes of reference suffers from **several limitations**
- **Fixed threshold** (e.g. 2.2): different behaviour for small and large lesions
- **Percentage of SUV_{max}**: underestimation in case of inhomogeneous uptake and reconstruction artifacts (e.g. Gibbs artifact in resolution-modeling reconstruction - PSF)
- **Tumor motion** is an additional bias

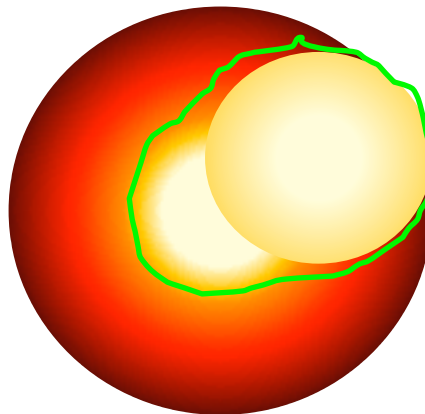
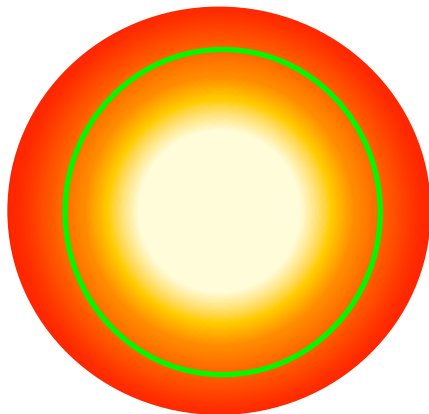
PET-CT FOR TREATMENT PLANNING

- threshold-based contouring (e.g. $SUV=2.2$)



- small lesions might be underestimated due to small SUV values – large lesions might be overestimated

- percentage-based contouring (e.g. 40% of SUV_{max})



- inhomogeneous lesions tend to be underestimated because of high SUV spots

PET-CT FOR TREATMENT PLANNING

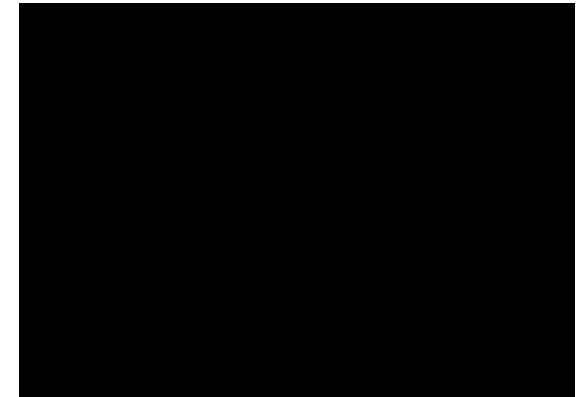
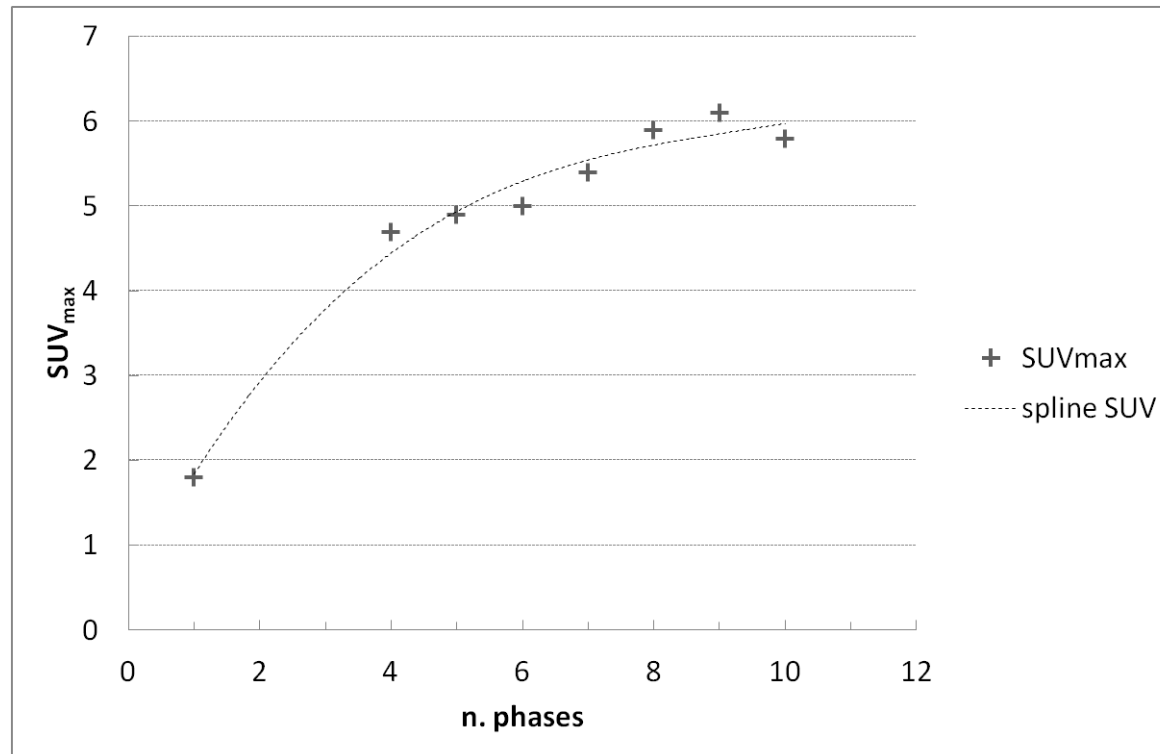
- more refined algorithms are based e.g. on the maximum gradient (gradient-based) or on *object-recognition* or *classification* algorithms
- there is no recognized “best-in-class” algorithm so far – a critical approach is always necessary when using commercially-available systems
- new algorithms (especially based on object recognition/classification methods) might be more robust with respect to motion artifacts etc. – more research needed

PET-CT FOR TREATMENT PLANNING

- example of gradient-based algorithm

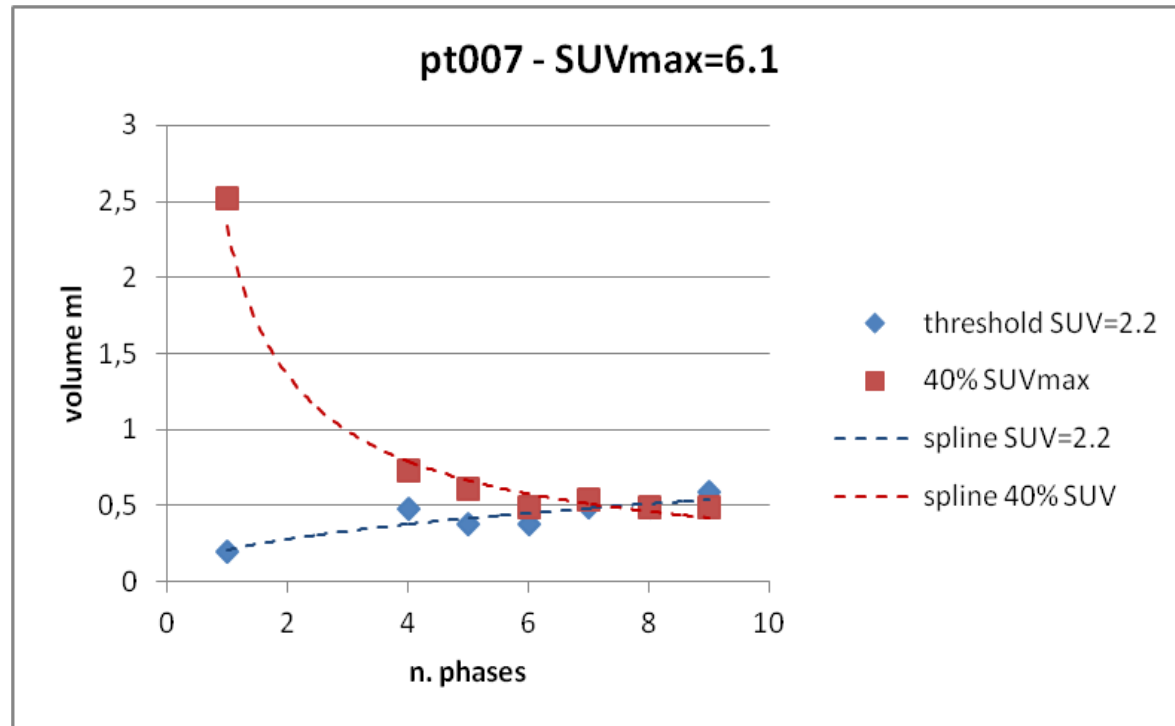


THE EFFECT OF MOTION ON SUV VALUES



- excursion 19 mm
 - $SUV_{max}=1.8$ non-gated
 - $SUV_{max}=6.1$ @9ph
- SUV_{max} in expiration as a function of the number of phase-bins

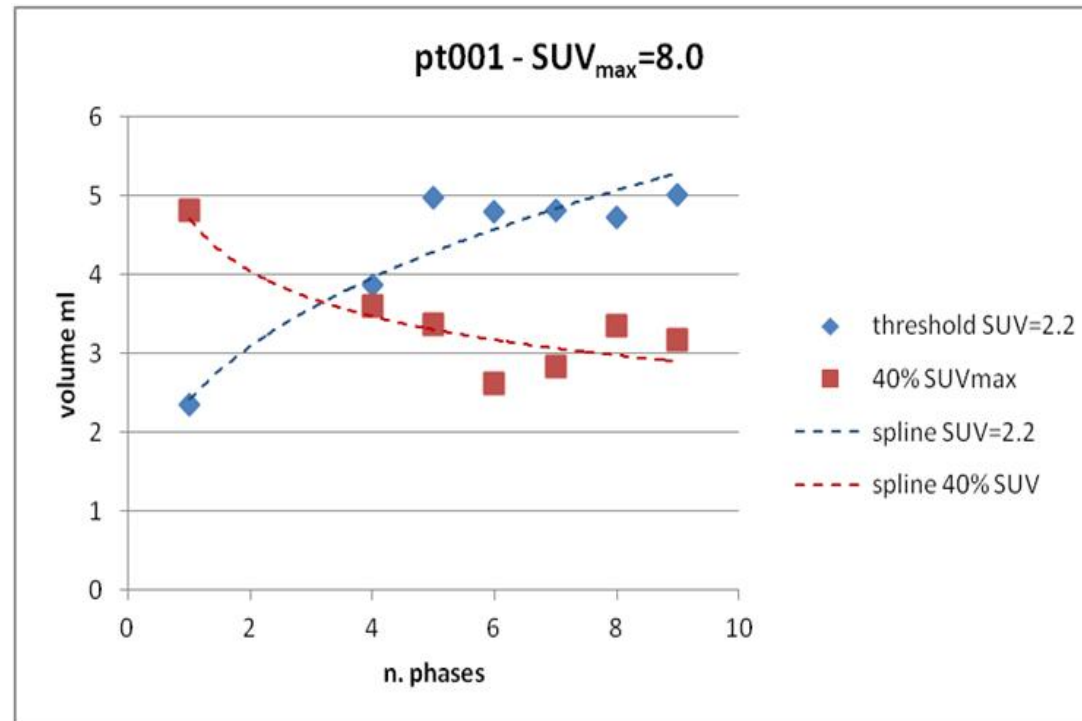
THE EFFECT OF MOTION ON SUV-BASED VOLUMES



CASE 1

- threshold-based algorithms => underestimation of volume
- %SUVmax algorithms => overestimation of volume

THE EFFECT OF MOTION ON SUV-BASED VOLUMES



CASE 2

- threshold-based algorithms => ?
- %SUVmax algorithms => ?
- more complex algorithms needed for accuracy

PET-CT REGISTRATION TO CT

- PET-CT has an **inherent CT dataset** that might be used for treatment planning if the required parameters and conditions are used
- PET-CT can be **registered to a different (setup) CT** – usually through **CT-CT (intra-modality)** registration whose transformation is then applied to the PET dataset
- Multi-modality PET-to-CT registration is **feasible but should be avoided** (poor “common information”)

IMAGE REGISTRATION - METHODS

- **Spatial coherence** between different imaging modalities used for treatment planning is (thought to be) a key factor for treatment success
- **Manual registration** methods must be **avoided** when co-registering 3D datasets
- **Automatic methods** are implemented on modern treatment planning systems for **rigid registration**
- **Deformable registration** is seldom implemented and requires careful evaluation of results

IMAGE REGISTRATION – transformation types

- **Rigid registration** – described by 6 parameters
 - three translations and three rotations corresponding to the principal axes in 3D
- **Deformable registration – affine** – 12 parameters
 - 3 translations + 3 rotations + 3 scaling f. + 3 shear factors
- **Deformable registration – local**
 - locally rigid registration – free to deform on a large scale
 - B-splines (B-cubic-splines)
 - locally affine
 - biomechanical models (finite elements method - FEM)
 - elastic or visco-elastic models

STRUCTURE OF A (DEFORMABLE) REGISTRATION ALGORITHM

$$T = \arg_T \max(sim(I_{Ref}, I_{fl} \ T) + \lambda Reg(T))$$



- *similarity measure*



- *regularization term
(deformable only)*

- similarity measurements vary as a function of the nature of co-registration (intramodality, multimodality ...)
- the regularization term charges a penalty on improbable transformations

SIMILARITY MEASURES

- Least-squares distance (set of **fiducial points**)
- Least-squares distance (**surfaces**)
- Intra-modality problem (e.g. CT-to-CT): **cross-correlation** (or mutual information, see below)
- Multimodality problem (e.g. MR-to-CT): maximization of the **mutual information** index/ **normalized mutual information (NMI)**
- ...

Multimodality image registration: joint histogram-based co-registration

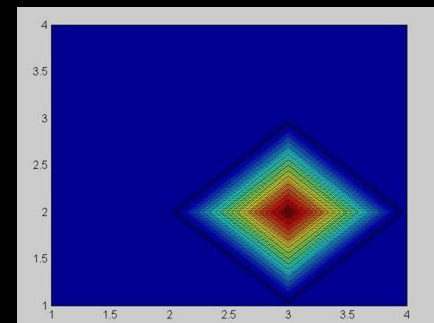
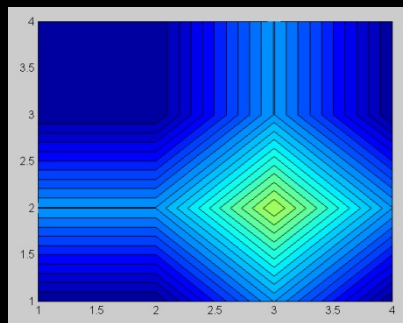
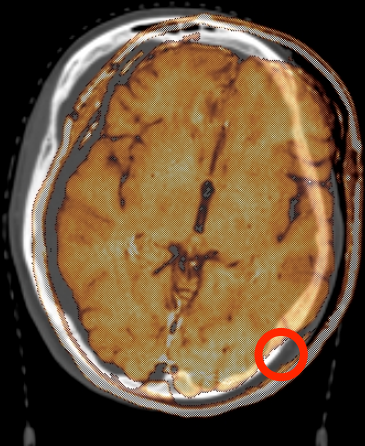
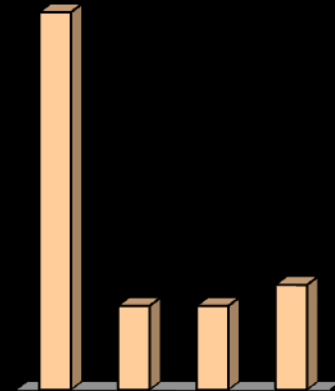
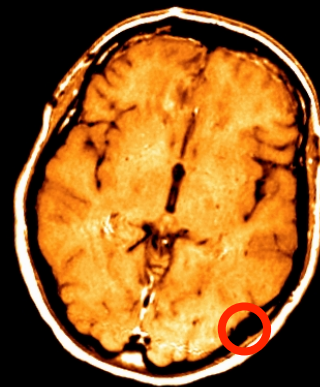
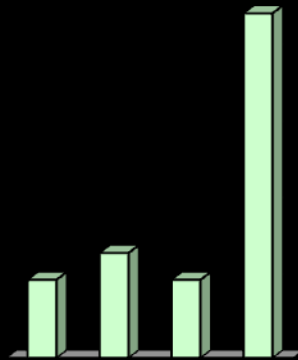
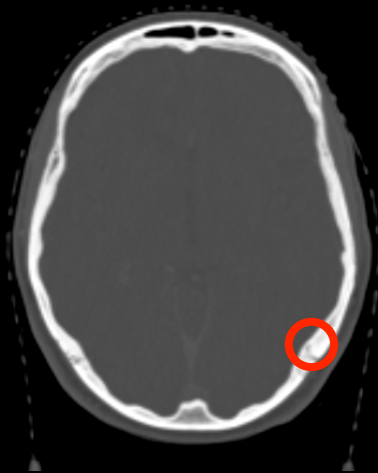


IMAGE ENTROPY (INFORMATION)

$$H = \sum_i p_i \log \frac{1}{p_i}$$

3 3 3 3 3

$p(3)=1$
 $\Rightarrow \mathbf{H = 0}$ “PREDICTABLE” MESSAGE – no
information added at each step

1 5 4 3 2

$p(1)=0.2$ $p(2)=0.2$ $p(3)=0.2$ $p(4)=0.2$ $p(5)=0.2$
 $\Rightarrow \mathbf{H = 1.61}$
“UNPREDICTABLE” MESSAGE – new
information added at each step

1 3 3 3 5

$p(1)=0.2$ $p(3)=0.6$ $p(5)=0.2$
 $\Rightarrow \mathbf{H = 0.95}$
INTERMEDIATE CASE

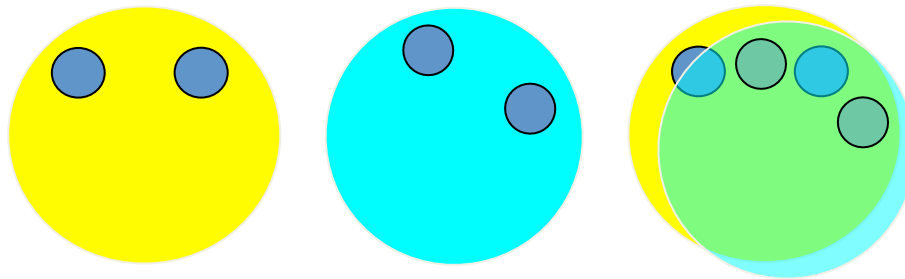
The MUTUAL INFORMATION index

Subtraction of the “joint entropy” (“false” information)

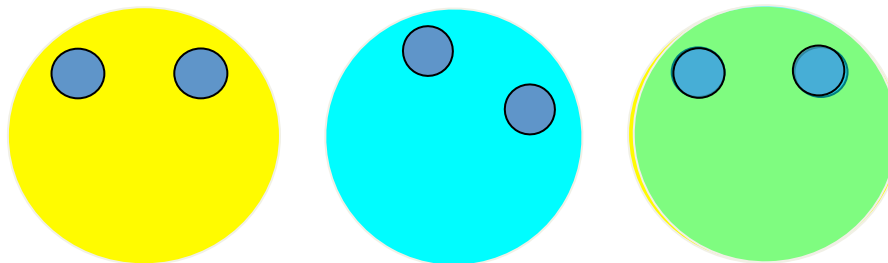
=> maximization of the mutual information index

$$I(A, B) = H(A) + H(B) - H(A, B)$$

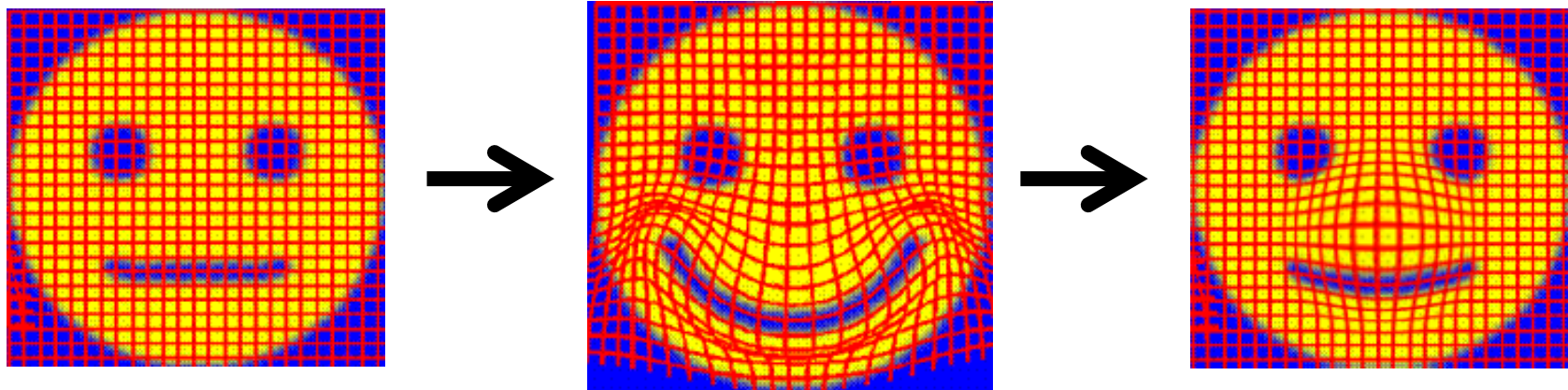
NON-REGISTERED
IMAGES:



REGISTERED
IMAGES:



STRUCTURE OF A (DEFORMABLE) REGISTRATION ALGORITHM

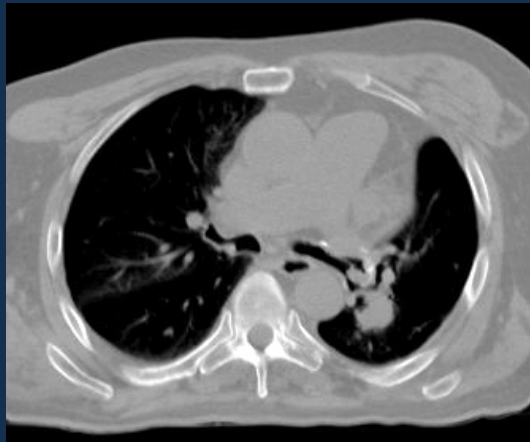


$$F_c = C_{sim} + \omega_{pen} C_{pen}$$

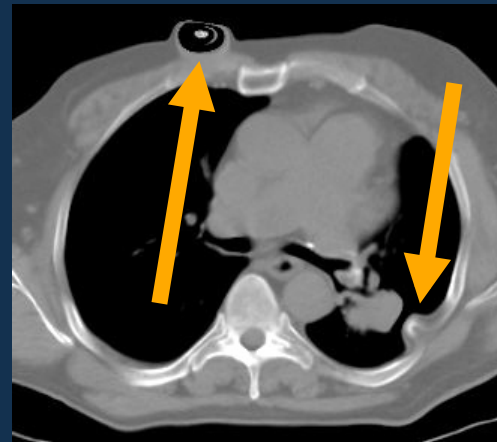
Regularization term:

$$1 + J_\tau J_\tau^T; \quad 1 + \det(J_\tau); \quad K$$

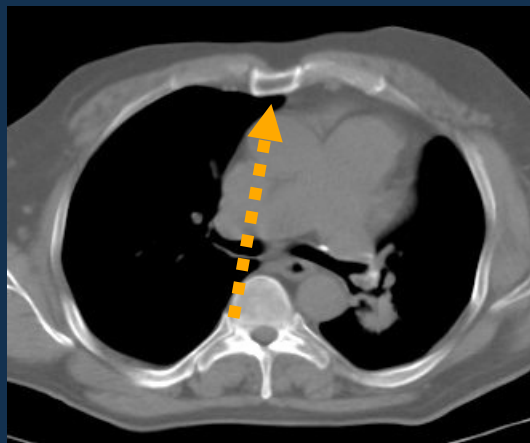
ROLE OF THE REGULARIZATION TERM



ORIGINAL IMAGE
(INSPIRATION)



REGISTERED TO EXP – no
regularization

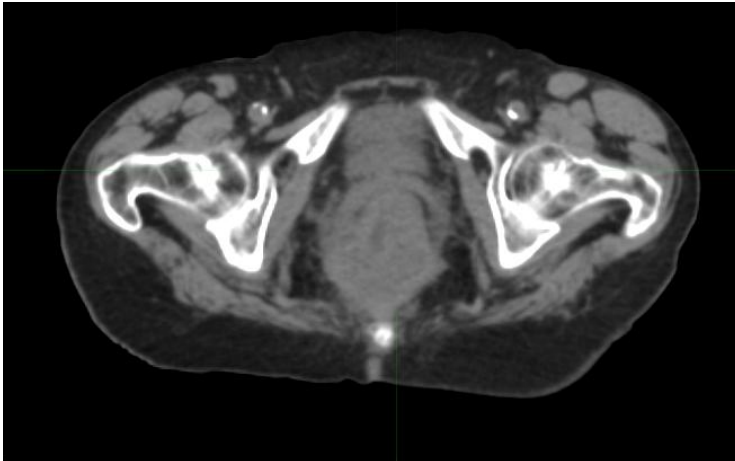


REGISTERED TO EXP – light
regularization

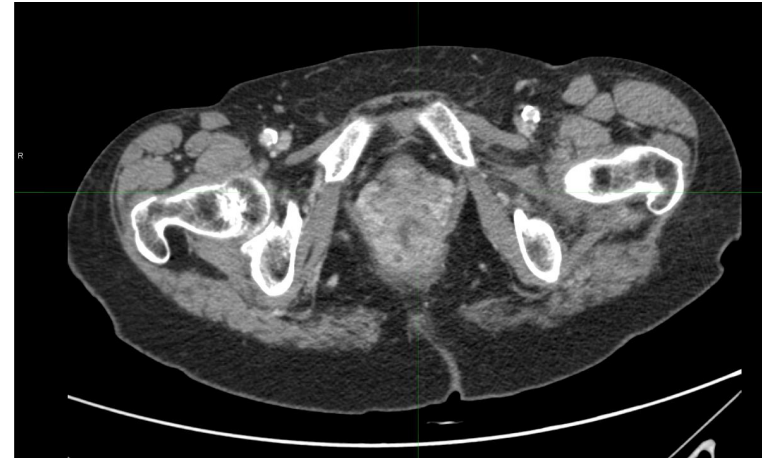


REGISTERED TO EXP –
strong regularization

deformable registration - regularization



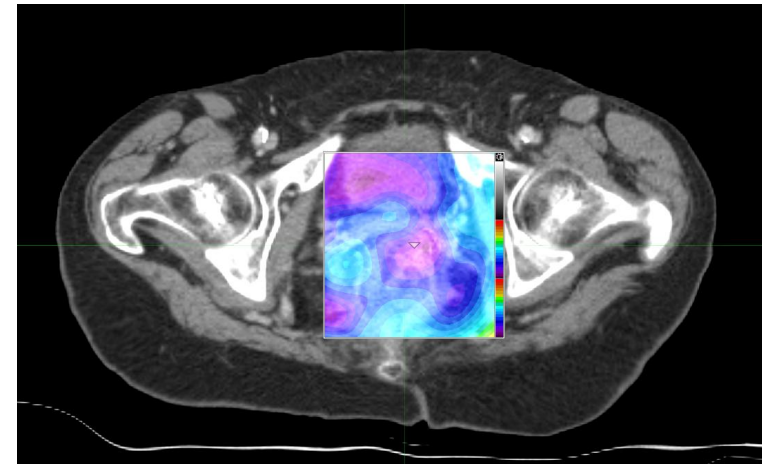
target



source



deformed



deformation map

deformable registration - regularization



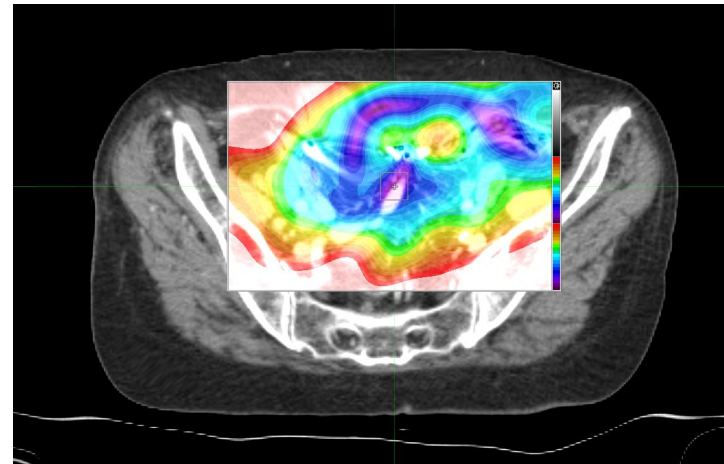
target



source

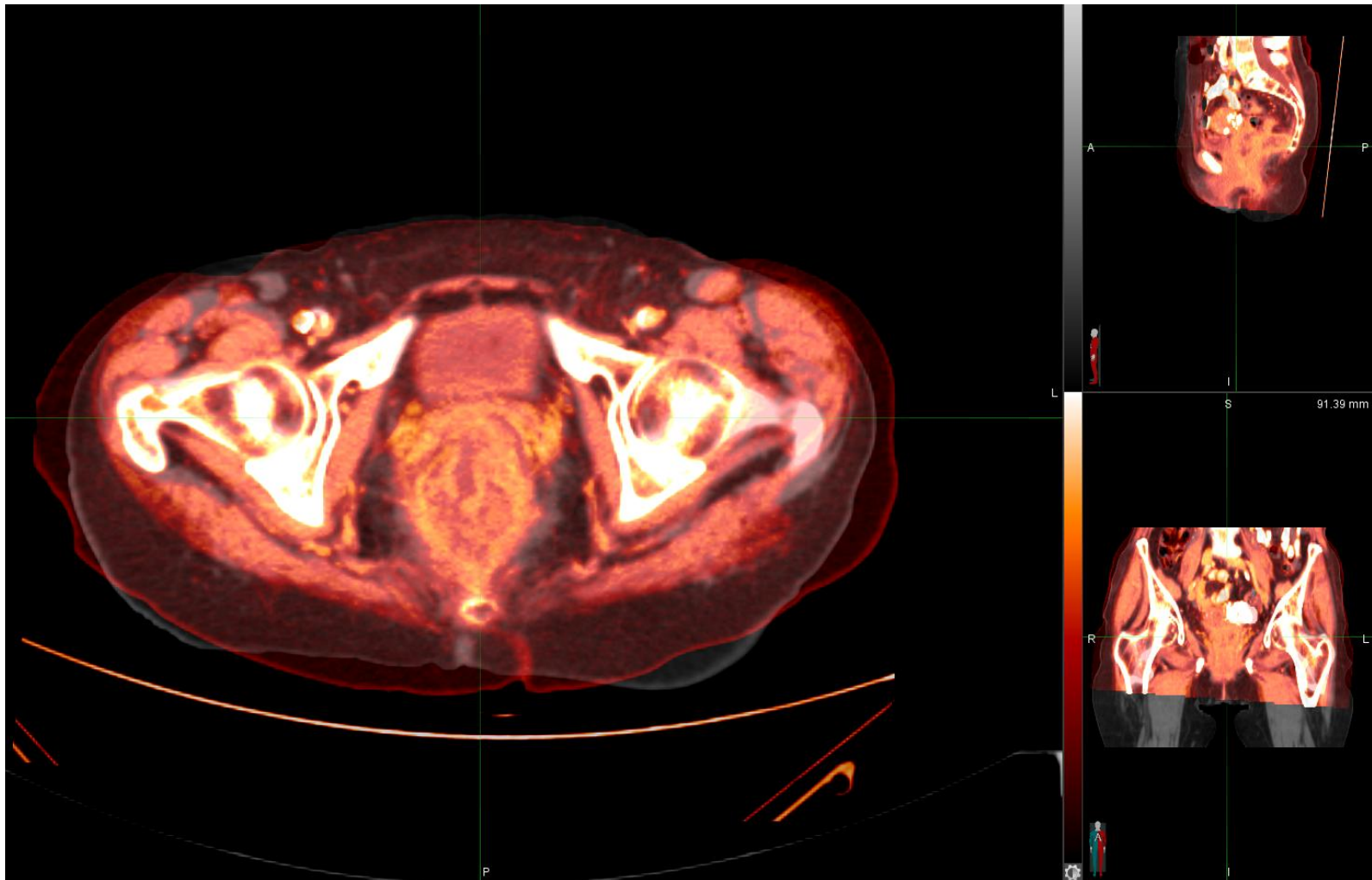


deformed



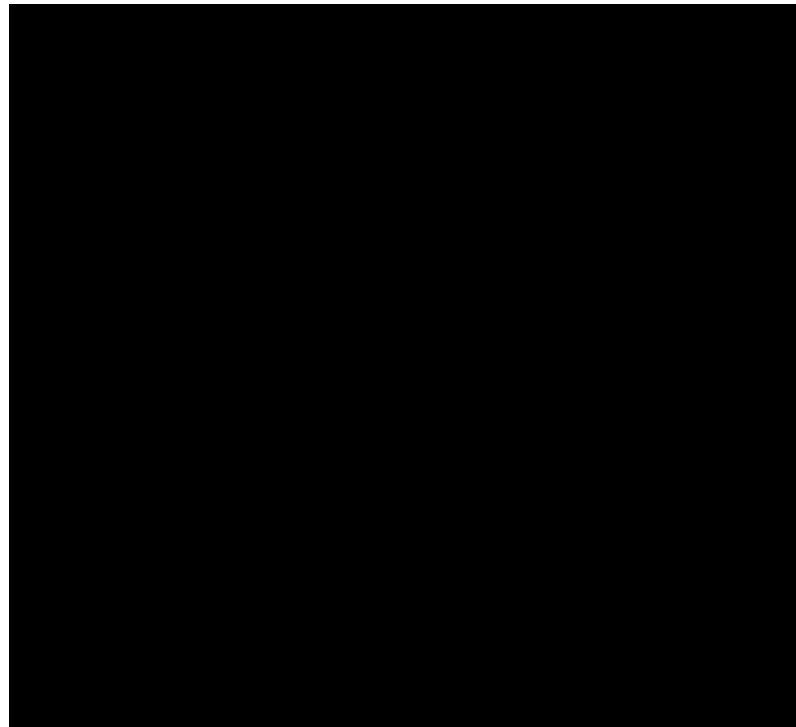
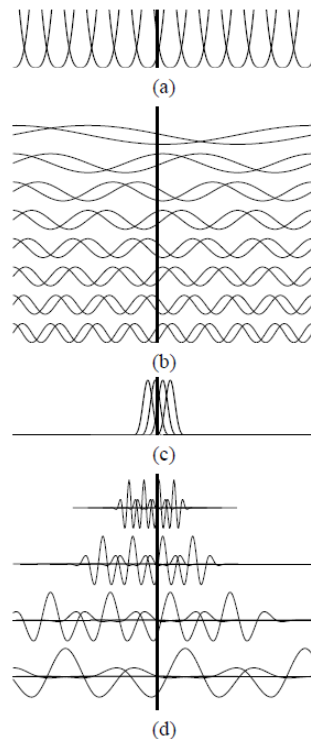
deformation map

deformable registration - regularization



DEFORMABLE REGISTRATION - LUNG

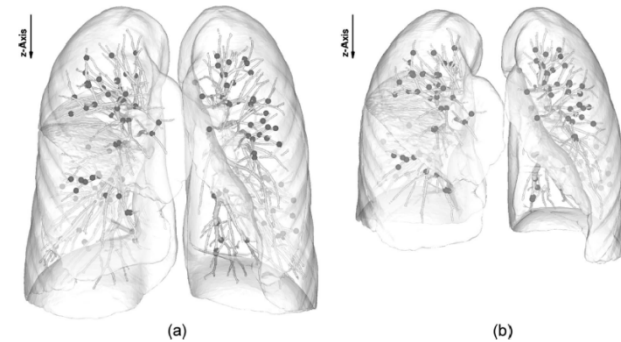
- B-spline-based** deformable registration
- continuous and differentiable functions
- simple implementation – calculation speed
- critical aspects in “**anatomic discontinuities**”



DEFORMABLE REGISTRATION - LUNG

- regularization**: conditions on the transf. Jacobian
- for example $D \cdot D^T = I$ or $J+1 = 0$ etc.

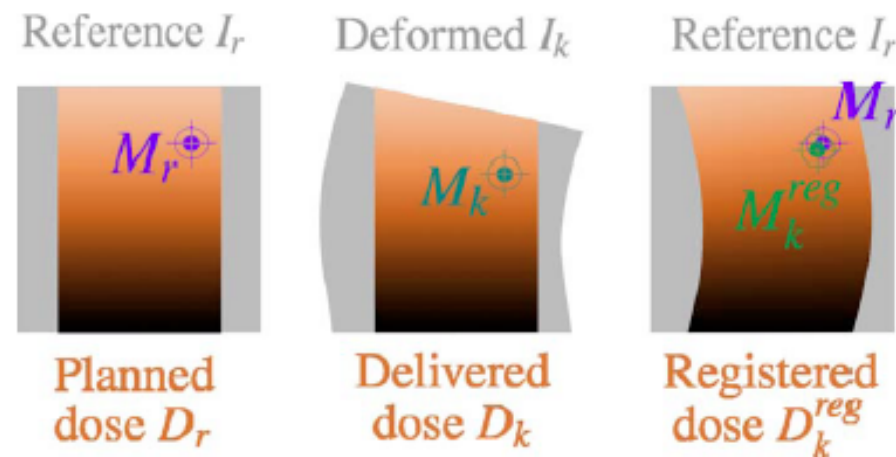
$$J(\mathbf{x}; \phi) = \det(\mathbf{D}) \quad \text{with } \mathbf{D} = \begin{pmatrix} \frac{\partial T_x^c}{\partial x} & \frac{\partial T_x^c}{\partial y} & \frac{\partial T_x^c}{\partial z} \\ \frac{\partial T_y^c}{\partial x} & \frac{\partial T_y^c}{\partial y} & \frac{\partial T_y^c}{\partial z} \\ \frac{\partial T_z^c}{\partial x} & \frac{\partial T_z^c}{\partial y} & \frac{\partial T_z^c}{\partial z} \end{pmatrix}.$$



- corresponds to **volume preservation**
- false** in general in the lung =>
alternative condition **mass preservation**

IMAGE REGISTRATION – beyond multimodality image integration for treatment planning

-Dose tracking – dose accumulation in **Adaptive Radiation Therapy**



G Janssens, J Orban de Xivry, S Fekkes, A Dekker, B Macq, P Lambin, W van Elmpt, "Evaluation of nonrigid registration models for interfraction dose accumulation in radiotherapy". Med. Phys. 36(9), 4268-4276 (2009)

TAKE HOME MESSAGES

1. Image registration is the process that makes two or more image sets **spatially coherent to each other**

2. Applications to Radiation Oncology include **treatment planning and treatment verification/adaptation**

3. **Rigid transformation** is to be preferred, **if possible**, but deformations shall be considered as potential sources of error

4. **Deformable registration** is powerful but difficult to control – expert judgment needed!

5.... see following module for other considerations on image registration applied to motion management ...