



Analyse d'Images Médicales et Modélisation: de l'acquisition à la quantification

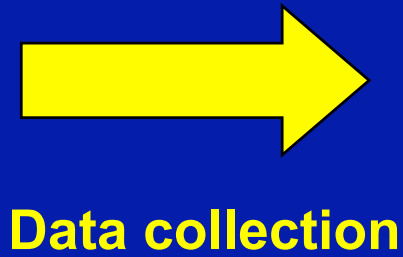
Patrick CLARYSSE

CREATIS-LRMN, CNRS UMR5220,
Inserm U630, Lyon, France

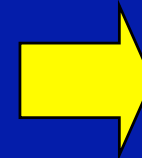
Patrick.clarysse@creatis.insa-lyon.fr
<http://www.creatis.insa-lyon.fr/~clarysse>



Patient



clinician

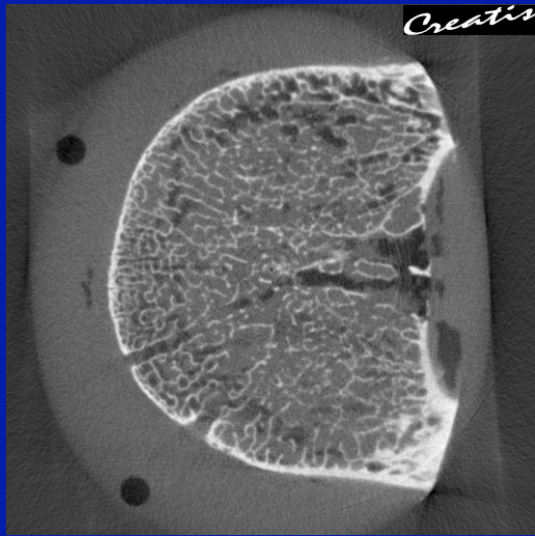


Decision

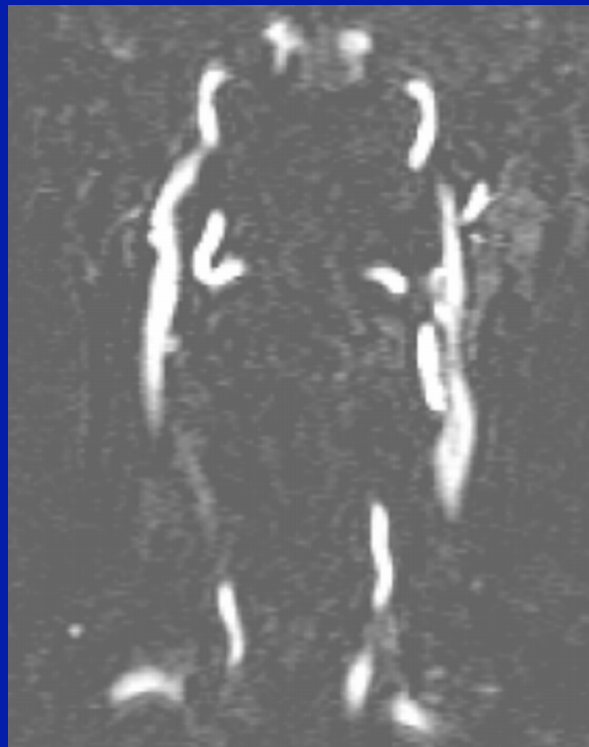
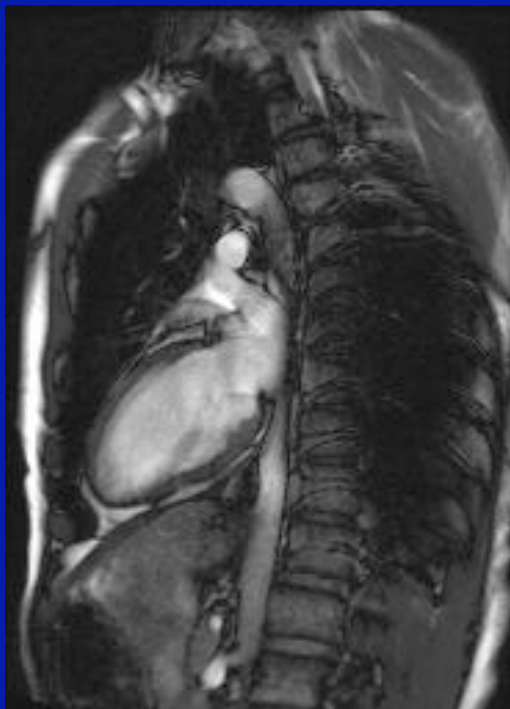


action

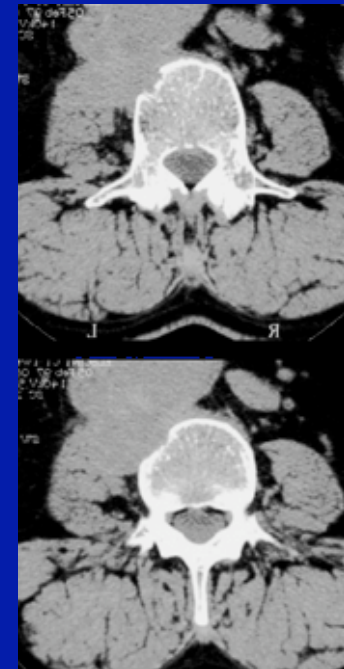


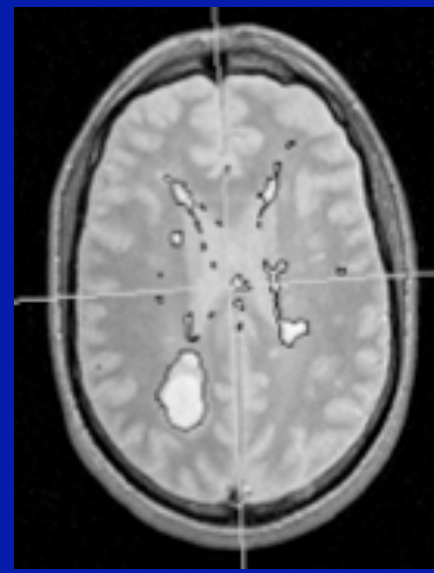
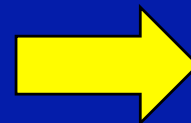
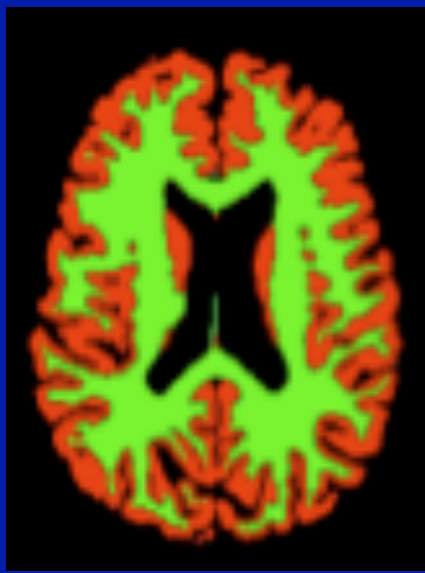
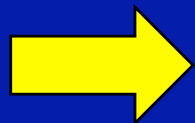


Bone image HRX



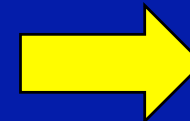
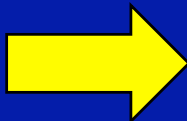
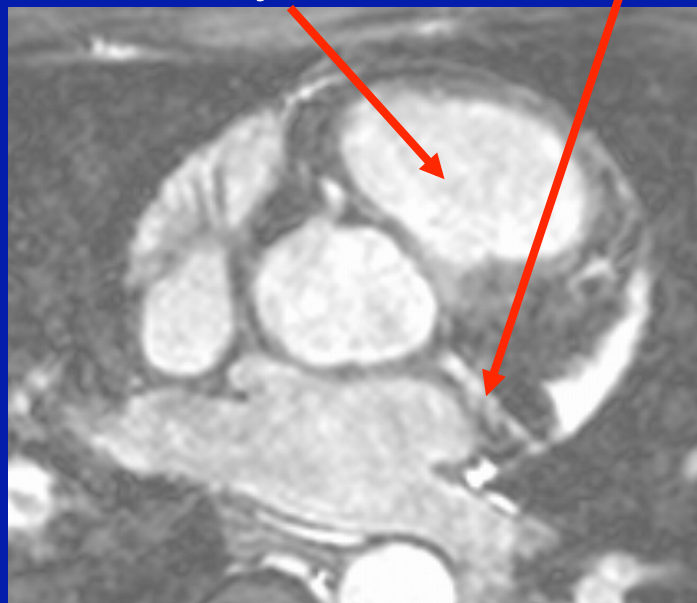
MRA Carotid





Cavity

Vessel



Stenosis
detection

MRA coronary

Difficulties

- Images = Partial measurements
- Objectives: Recover the truth
 - shapes, functions...
 - normal and abnormal patterns

➔ Inverse problem:

- Not a unique solution
- Stability upon the data

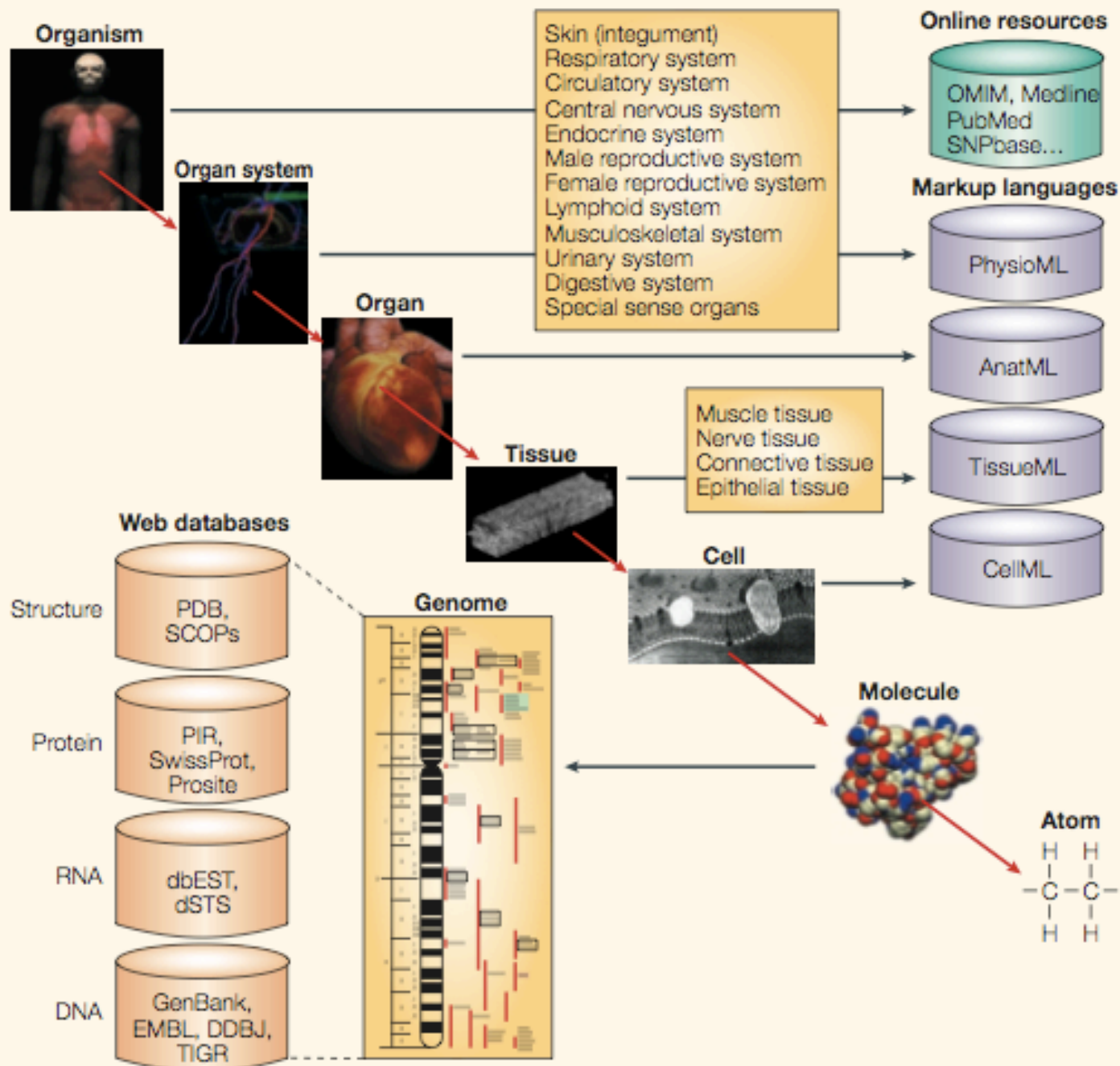
From ill-posed to well-posed

The image segmentation example

- Regularization (smoothness constraint)
- Adding prior information:
 - Shape constraints: average shape, topology, shape statistics...
 - Static and dynamic properties
 - Other functional properties
 - Appearance model (image)

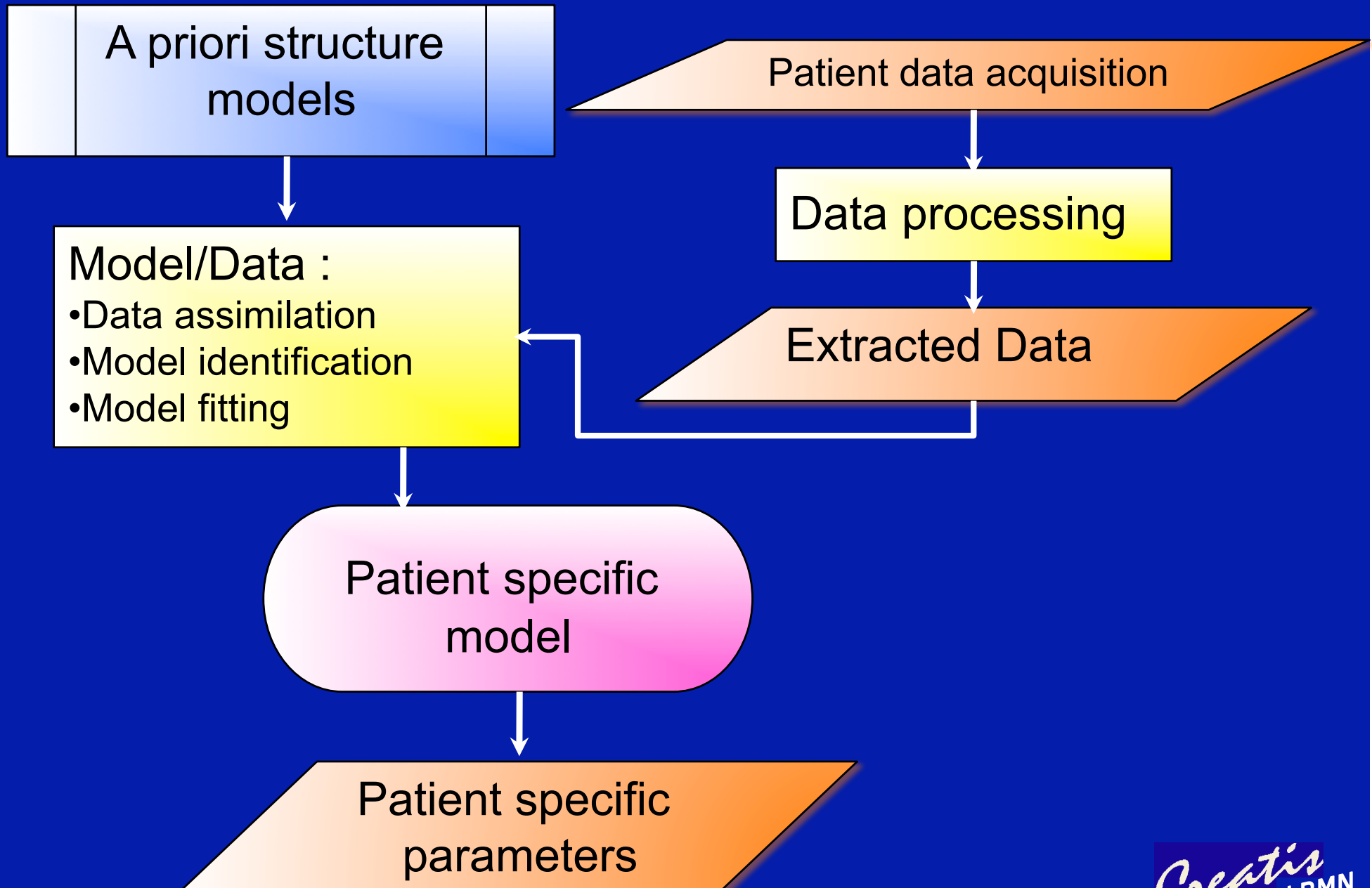
➤ ***a priori* structure models**

The Modeling perspective

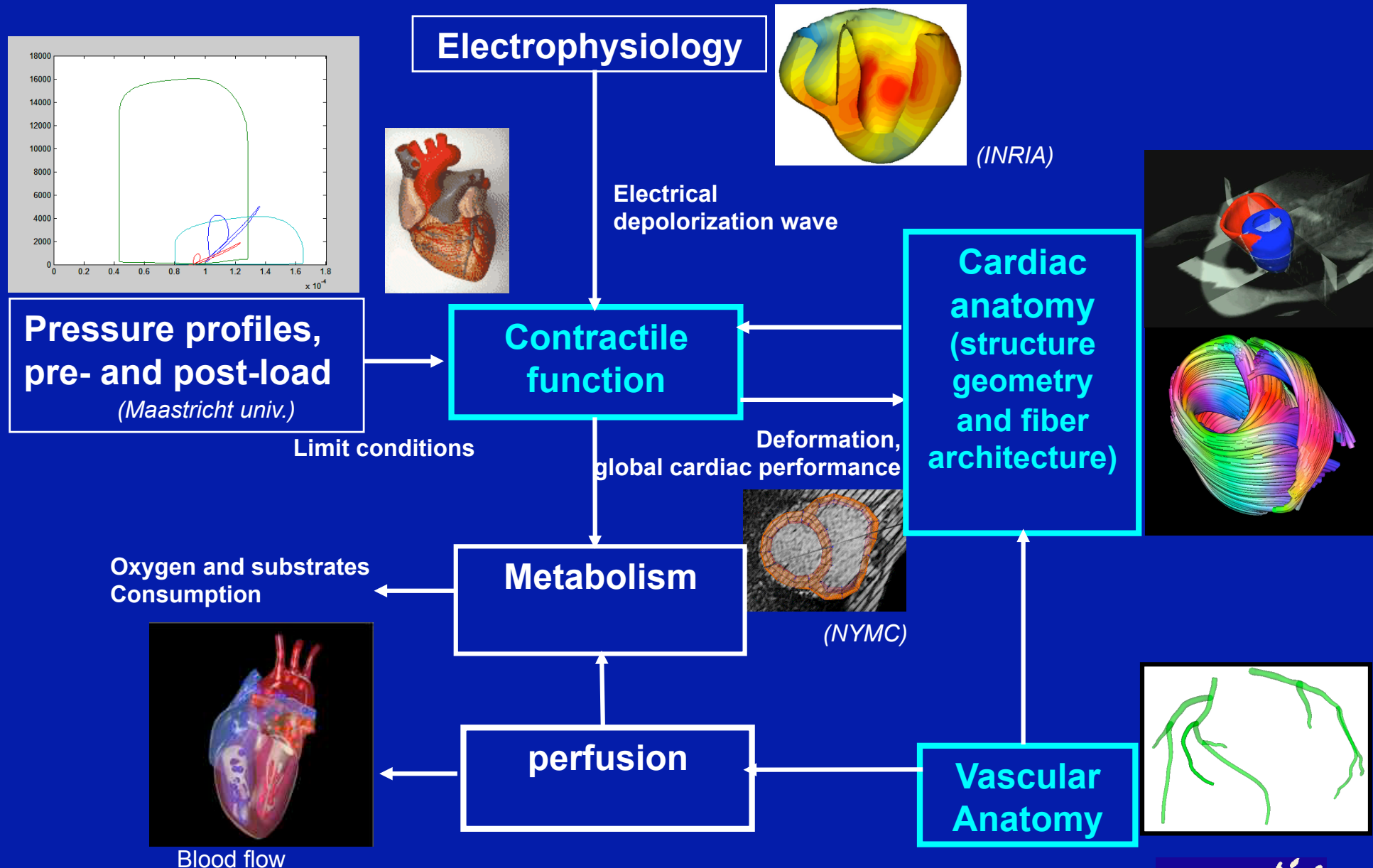


From Hunter & Borg
Nature Rev, 2003
(Physiome & VPH
projects)

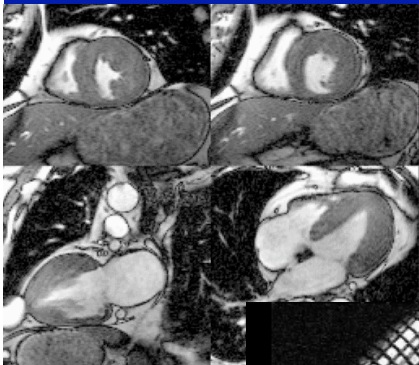
Patient specific Modeling



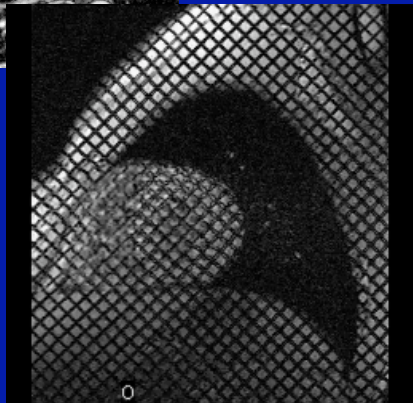
Patient specific cardiac modeling



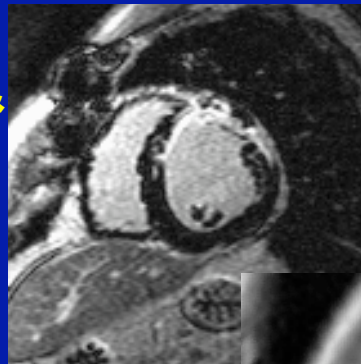
Available data



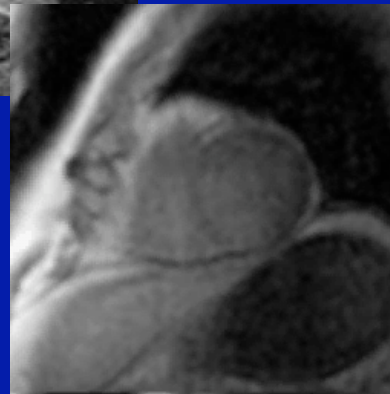
MRI:
Anatomy &
contractile
function



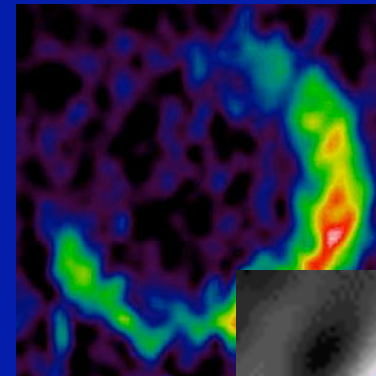
→ cm.s^{-1} , cm.s^{-2}



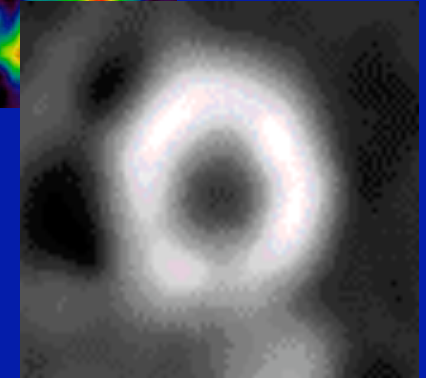
MRI:
perfusion



→ $\text{ml.g}^{-1}.\text{min}^{-1}$

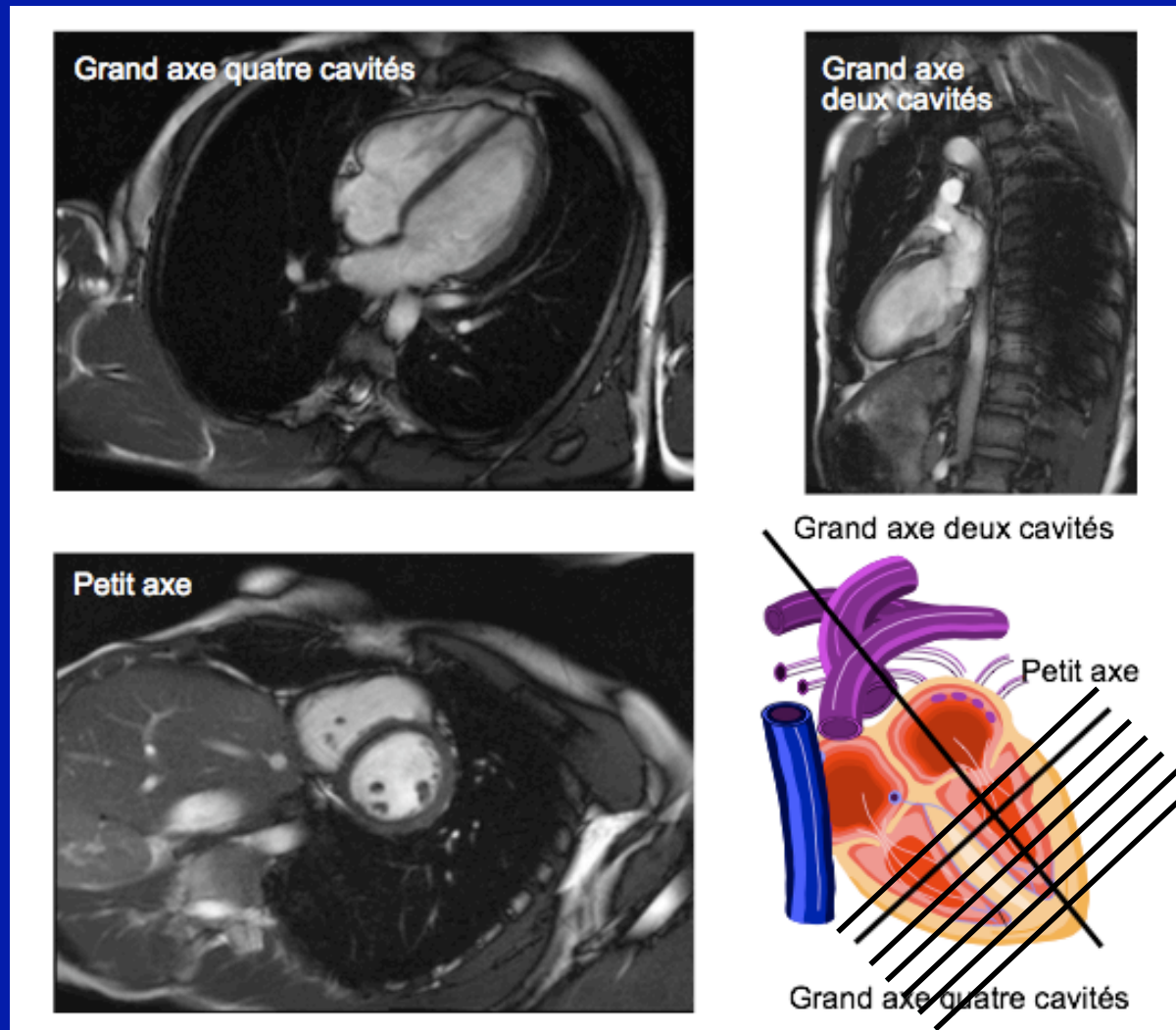


PET:
metabolism



→ $\text{mmol.g}^{-1}.\text{min}^{-1}$

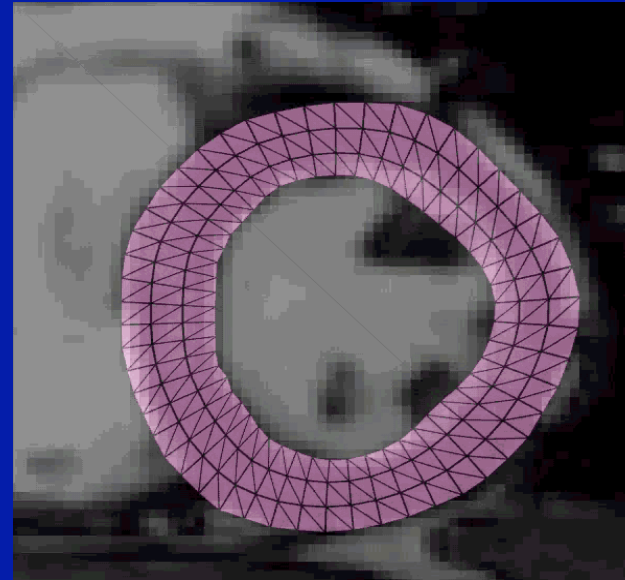
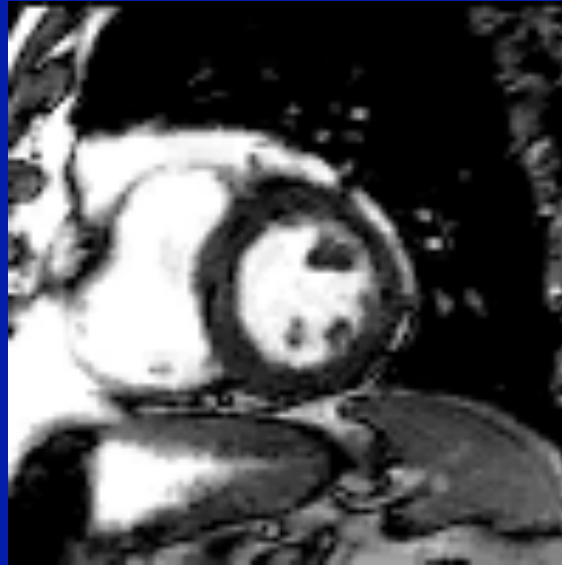
MR imaging



Standard multi-slice cine images

➔ Human & small animal

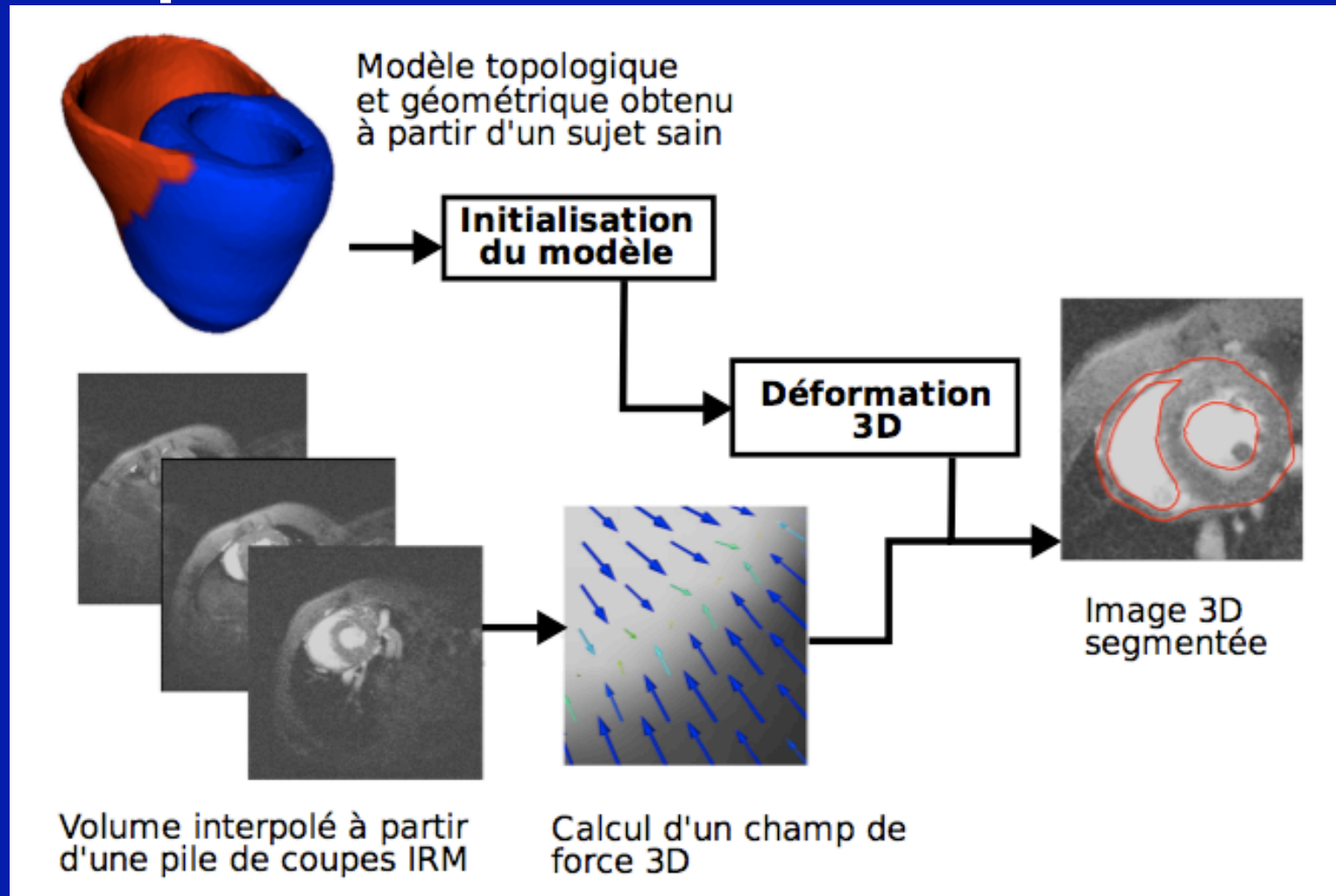
Dynamic cardiac segmentation



- **Objective:**
 - segmentation and tracking of the heart in temporal image sequences (MRI)
 - Characterizing the shape and dynamics of normal and pathological hearts

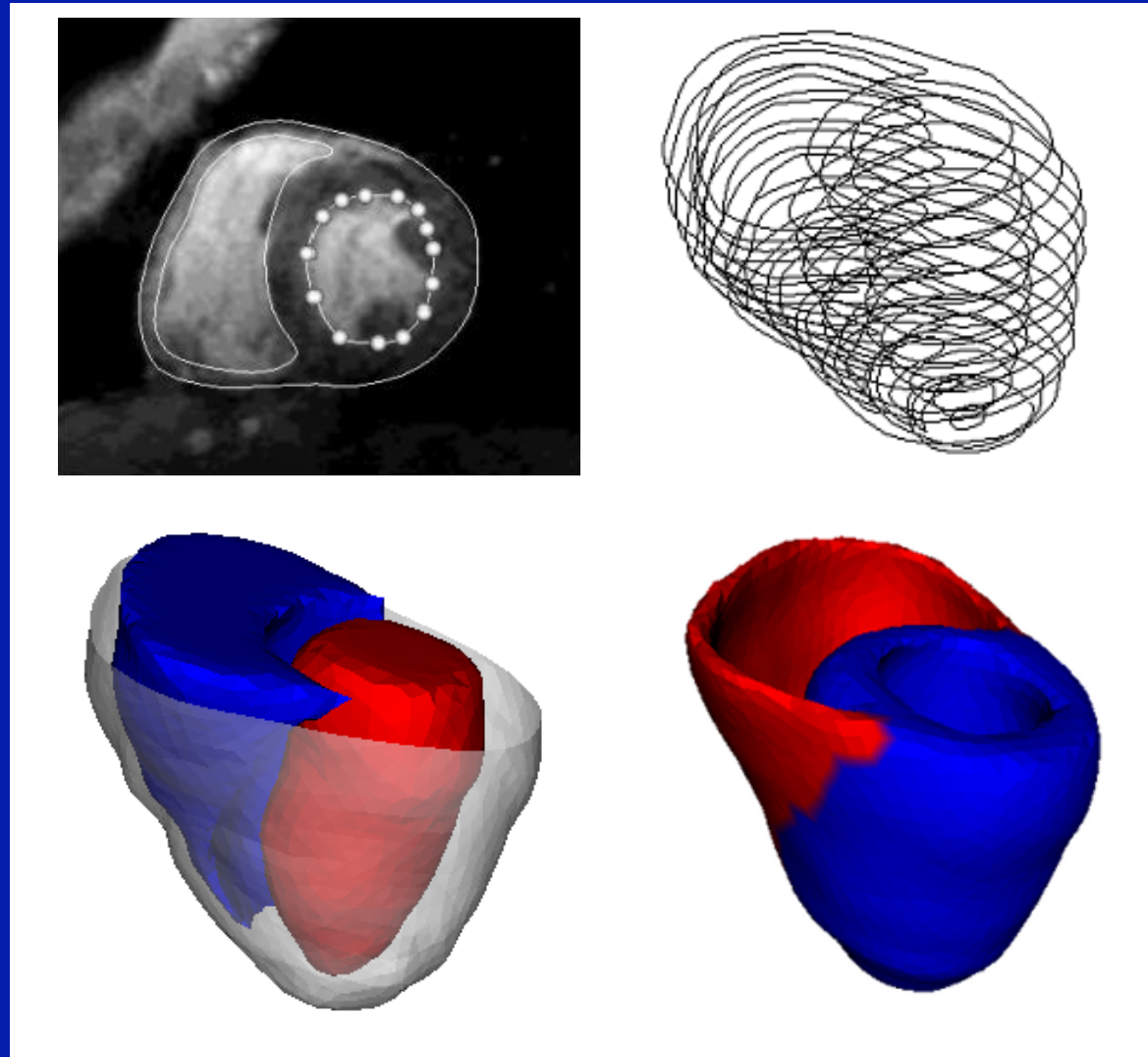
The Deformable Elastic Template (DET)

- Principle



→ A bio-inspired model

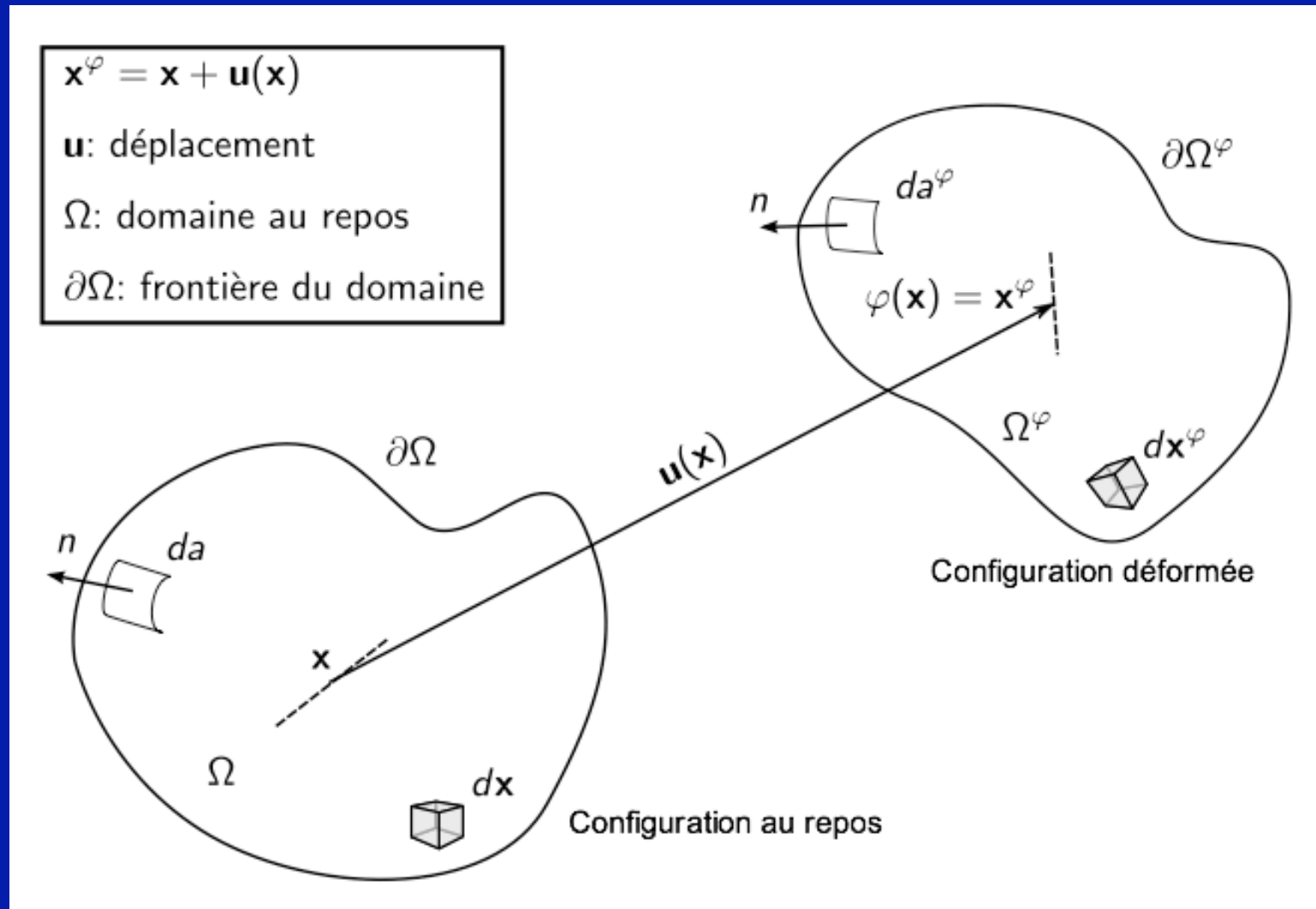
Reference *a priori* model



Mailleur GHS3D, Gamma Project, INRIA

DET Patient specific model adaptation

- Elastic deformation



- **Elastic solid under constraints :**

- **Equilibrium equation**

$$\begin{cases} \operatorname{div} \boldsymbol{\sigma} = \mathbf{0} & \text{in } \Omega_0 \\ \boldsymbol{\sigma}_n = f & \text{on } \partial\Omega_0 \end{cases}$$

- **Potential energy**

$$E(u) = \underbrace{\frac{1}{2} \int_{\Omega_0} \boldsymbol{\varsigma}^T(u) \boldsymbol{\epsilon}(u) dx}_{E_{elastic}} - \underbrace{\int_{\delta\Omega_0} f^T \cdot u ds}_{E_{image}}$$

- **Elastic Energy**

$$E_{elastic} = \frac{1}{2} \int_{\Omega} \mu \nabla u^T \nabla u + (\lambda + \mu) \operatorname{div}(u) \operatorname{div}(u) d\Omega$$

With λ, μ the Lamé Coefficients

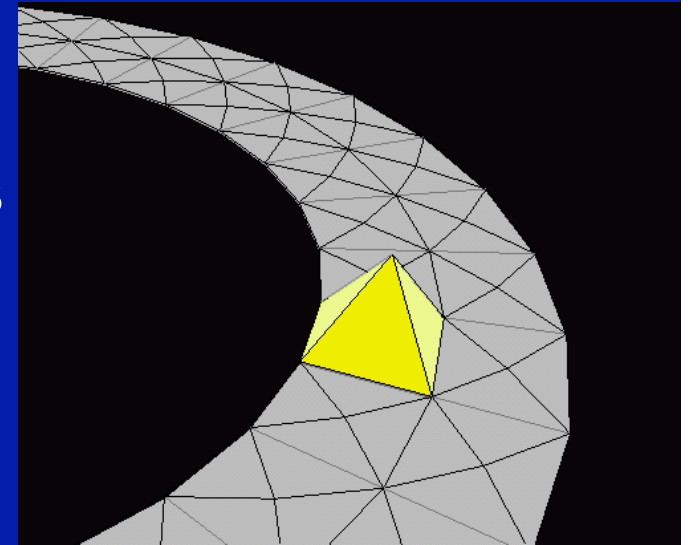
- Optimization: FEM

- Partitioning the domain in elements
- Linear basis functions

- Solving the system

- Algorithm M1: incremental load

$$\text{M1} \quad \begin{cases} u^{k+1} - \Delta t \operatorname{div} \sigma(u^{k+1}) = u^k & \text{dans } \Omega_0 \\ \sigma_n(u^{k+1}) = \underline{f(\mathbf{I} + u^k)} & \text{sur } \partial\Omega_0 \end{cases}$$



$$\mathbf{KU} = \mathbf{F}$$

- FEM solution: evolution equation

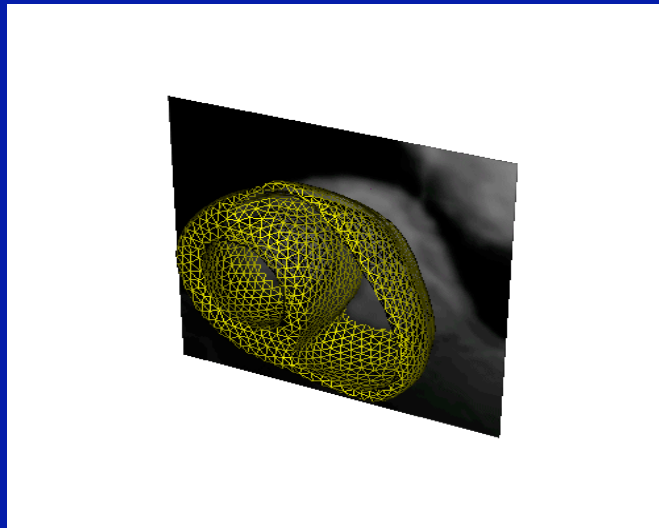
$$\frac{d\mathbf{U}}{dt} + \mathbf{KU}(t) = \underline{\mathbf{F}(\mathbf{U}(t))}$$

$$\mathbf{U} = (u_1, u_2, u_3, \dots, u_N)$$

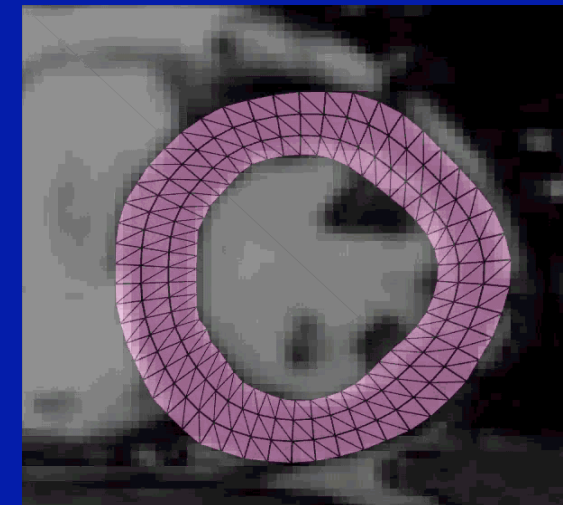
K: stiffness matrix

$$(\mathbf{I} + \Delta t \mathbf{K})\mathbf{U}^{k+1} = \mathbf{U}^k + \Delta t \mathbf{F}(\mathbf{U}^k)$$

DET story



- 2D linear static DET :
[PhD thesis F. Vincent, 2001]
- 3D linear static DET, bi-ventricular model :
[PhD thesis Q-C. Pham, 2002]
- Non-linear static DET,
[PhD thesis Y. Rouchdy, 2005]
(Collab. ICJ, J. Pousin)
- Dynamique DET,
[PhD thesis J. Schaerer, 2008]
(Collab. ESIEE, L. Najman, J. Cousty)



- **Non linear algorithm [Rouchdy, 2005]**
 - To avoid the linearization of the def tensor

$$\begin{cases} \operatorname{div} \sigma(u(\lambda)) = 0 & \text{in } \Omega \\ \sigma(u(\lambda)) \cdot n = \lambda f(u(\lambda)) & \text{on } \partial\Omega \end{cases}$$

- Where $\lambda [0,1]$ is an evolution variable**
- **With an explicit Euler scheme:**

$$\mathbf{A}^k \mathbf{U}_{k+1} = \mathbf{A}^k \mathbf{U}_k + (\lambda_{k+1} - \lambda_k) \mathbf{F}(\mathbf{U}_k)$$

with $\mathbf{A}^k = \mathbf{K}^k - \lambda_k \mathbf{D}^k$

*Collab. Inst. C. Jordan, INSA Lyon
MATH-STIC CNRS project*

- **Border constraint:** $f(\mathbf{I} + u) = 0$
 - Singular perturbation approach
[Schaerer, 2008]

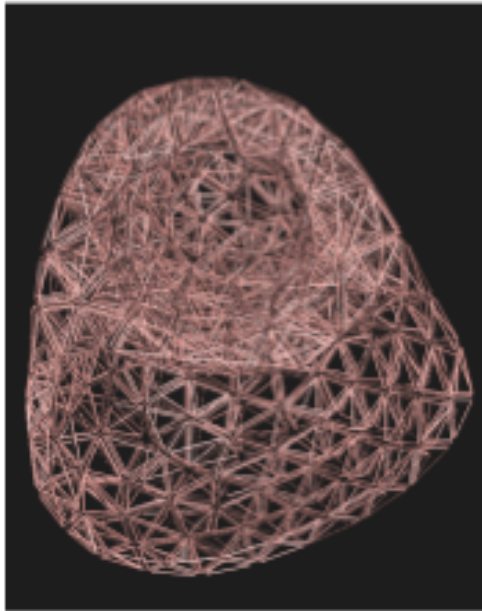
$$\left(\Delta t e^{-\beta k} \mathbf{K} + \mathbf{I}\right) \mathbf{U}^k = \Delta t F\left(\mathbf{U}^{k-1}\right) + \mathbf{U}^{k-1}$$

With Δt the time step and β a parameter controlling the decrease

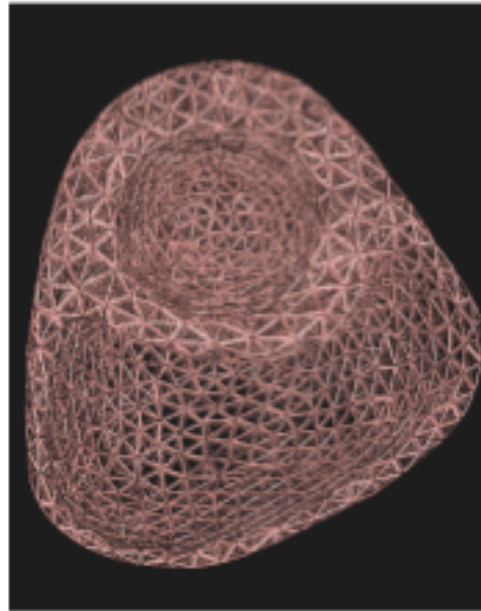
- *If the force field derive from a potential and is Lipschitz then the algo converges*

*Collab. Inst. C. Jordan, INSA Lyon
MATH-STIC CNRS project*

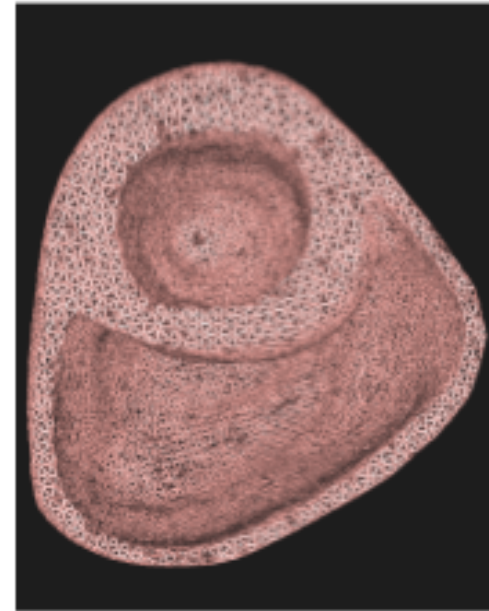
- **Multiresolution meshes**



149 points

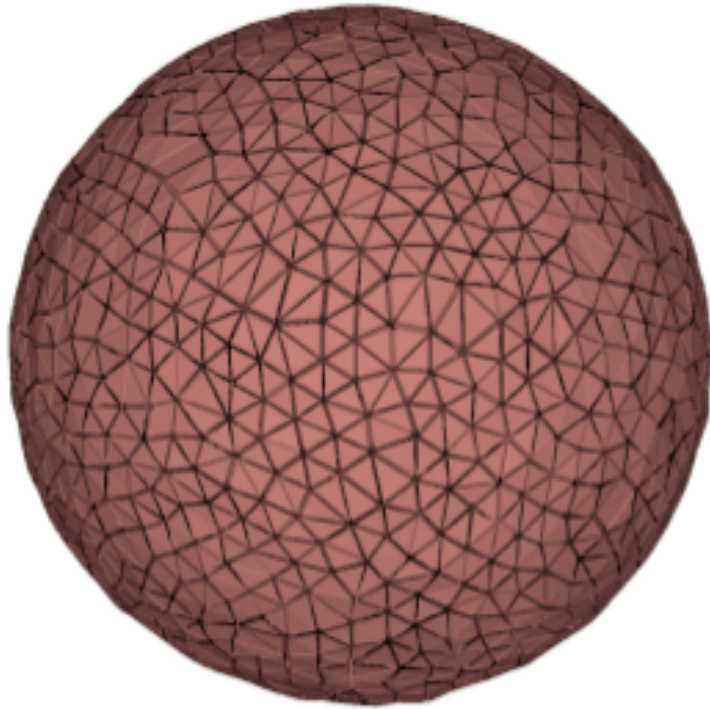


362 points

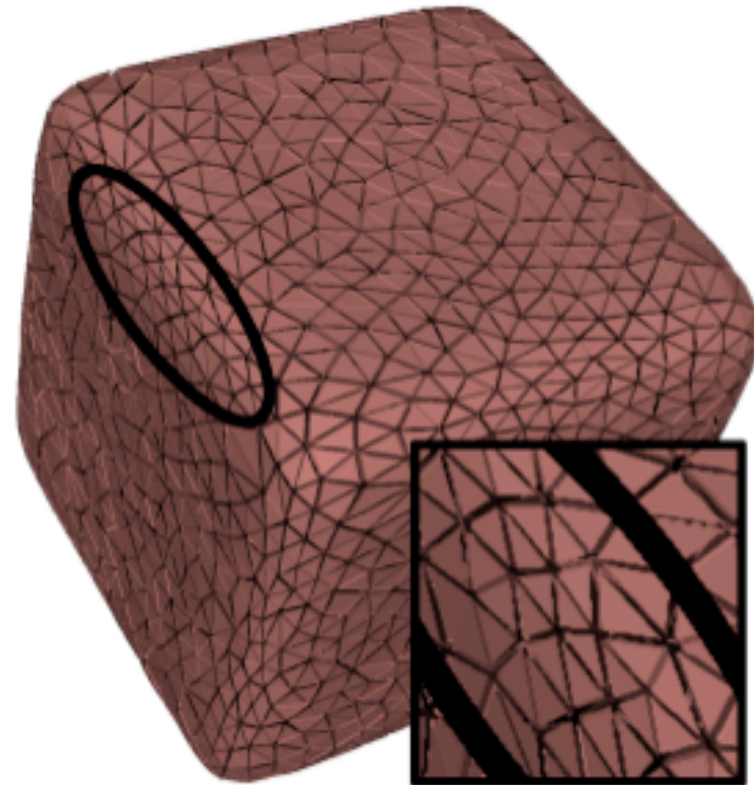


3438 points

- **Impact of the non linear DET**



Maillage initial de boule
(≈ 6000 points)



Maillage déformé

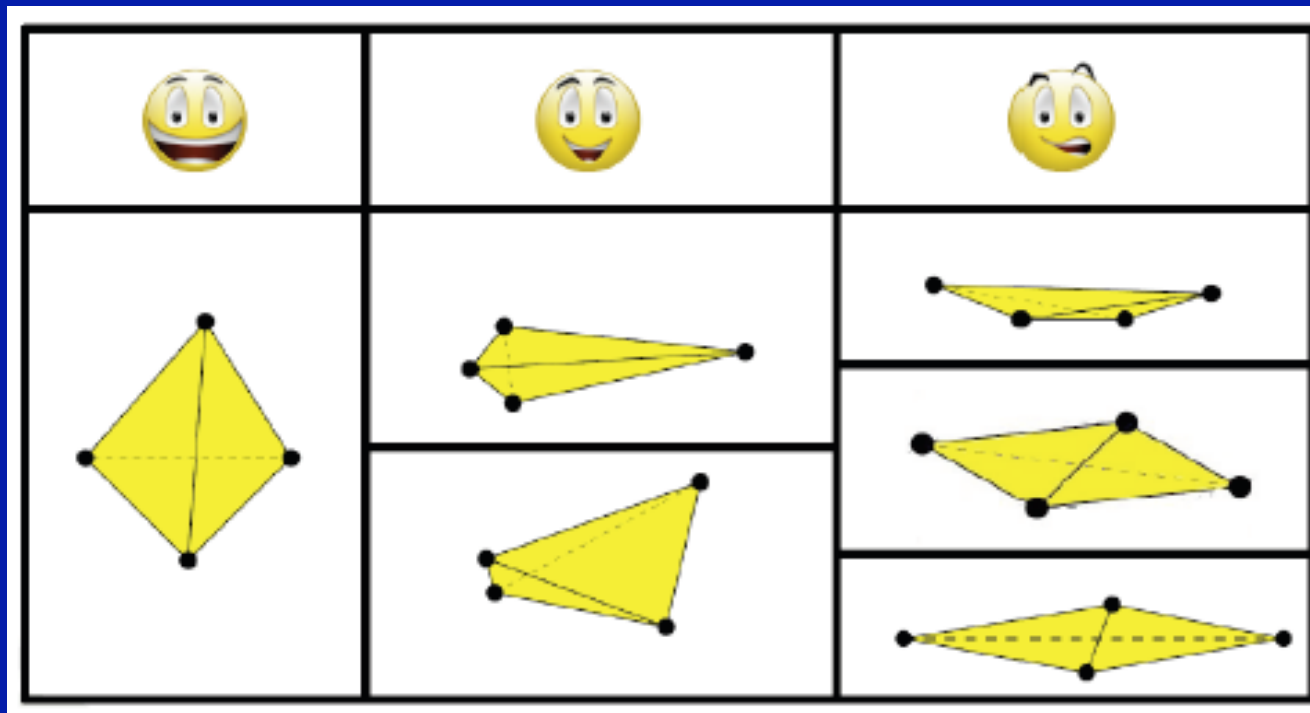
- **Critère de qualité:** rayon de la sphère inscrite au tétraèdre sur la longueur de sa plus grande arête, compris entre 0 et 1

	Cube linéaire	Cube non-linéaire
Sans multirésolution		
Temps de calcul	54s	1m25s
Qualité moyenne	0.58	0.61
Qualité minimale	0.10	0.17
Multirésolution à 3 niveaux		
Temps de calcul	26s	46s
Qualité moyenne	0.58	0.61
Qualité minimale	0.13	0.17

⇒ L'algorithme non-linéaire permet une meilleure stabilité numérique

Mesh Quality for FE methods

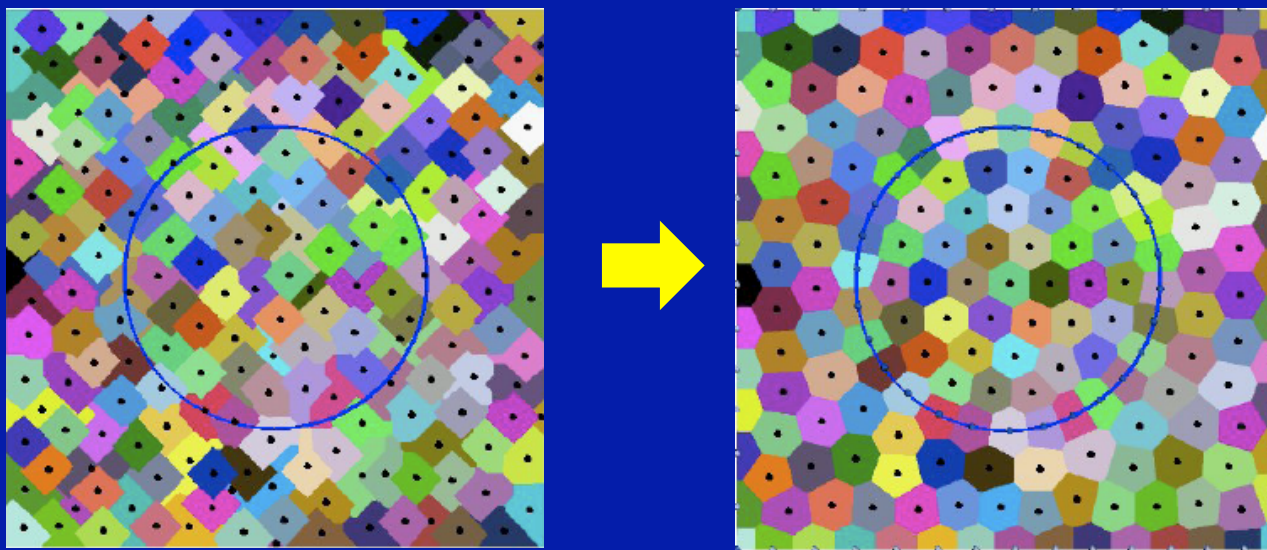
- Discretization error
- Interpolation error
- Matrix conditionning



Reconstructing 3D meshes from voxels

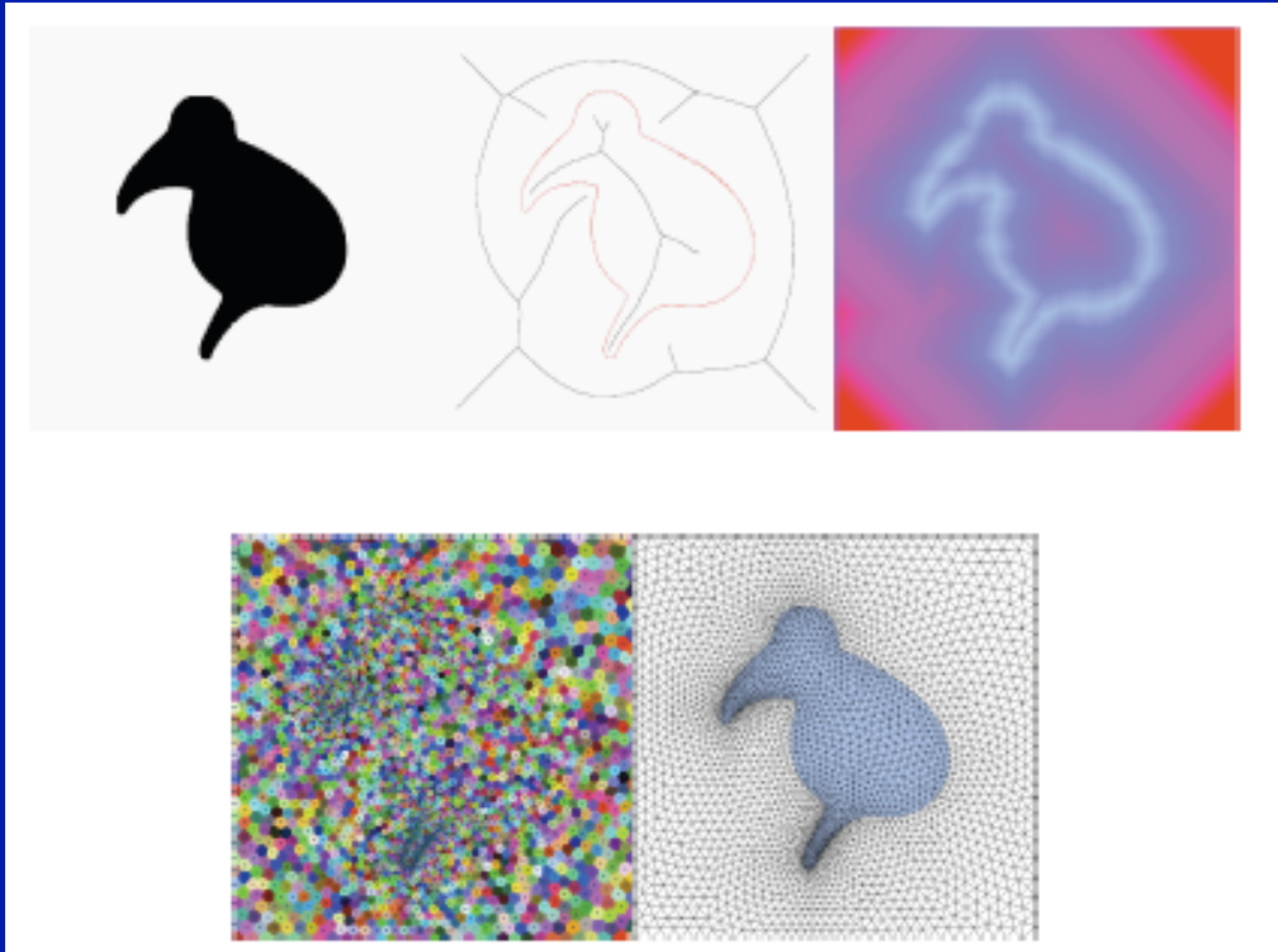
J. Dardenne thesis

- Domain partitioning with frontier preservation (energy minimization approach)



J. Dardenne, S. Valette et al., Variational tetraedral mesh generation from discrete volume data. *The Visual Computer*, pp (in-press), 2009.

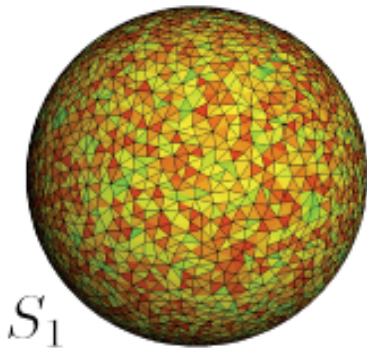
- Adaptive mesh density through a density map



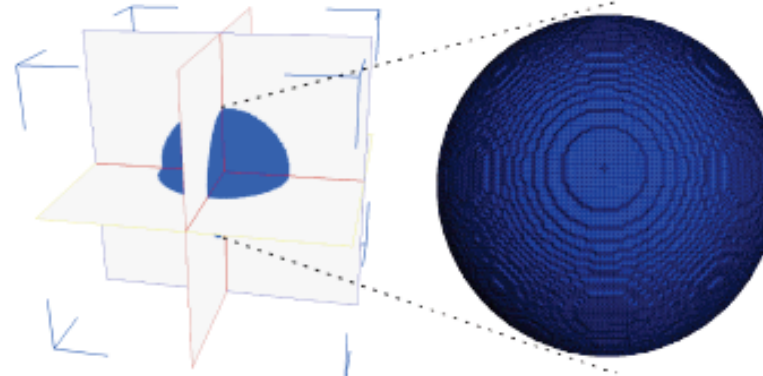
Iso-Surface



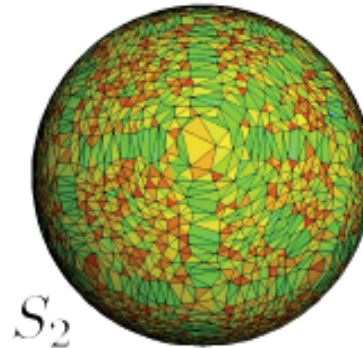
Amira



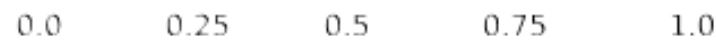
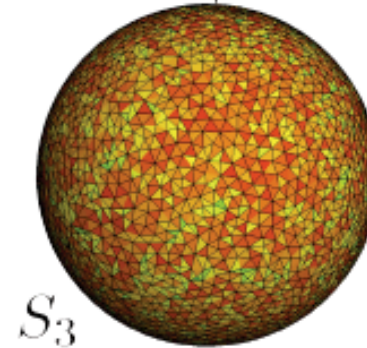
Ensemble de Voxels



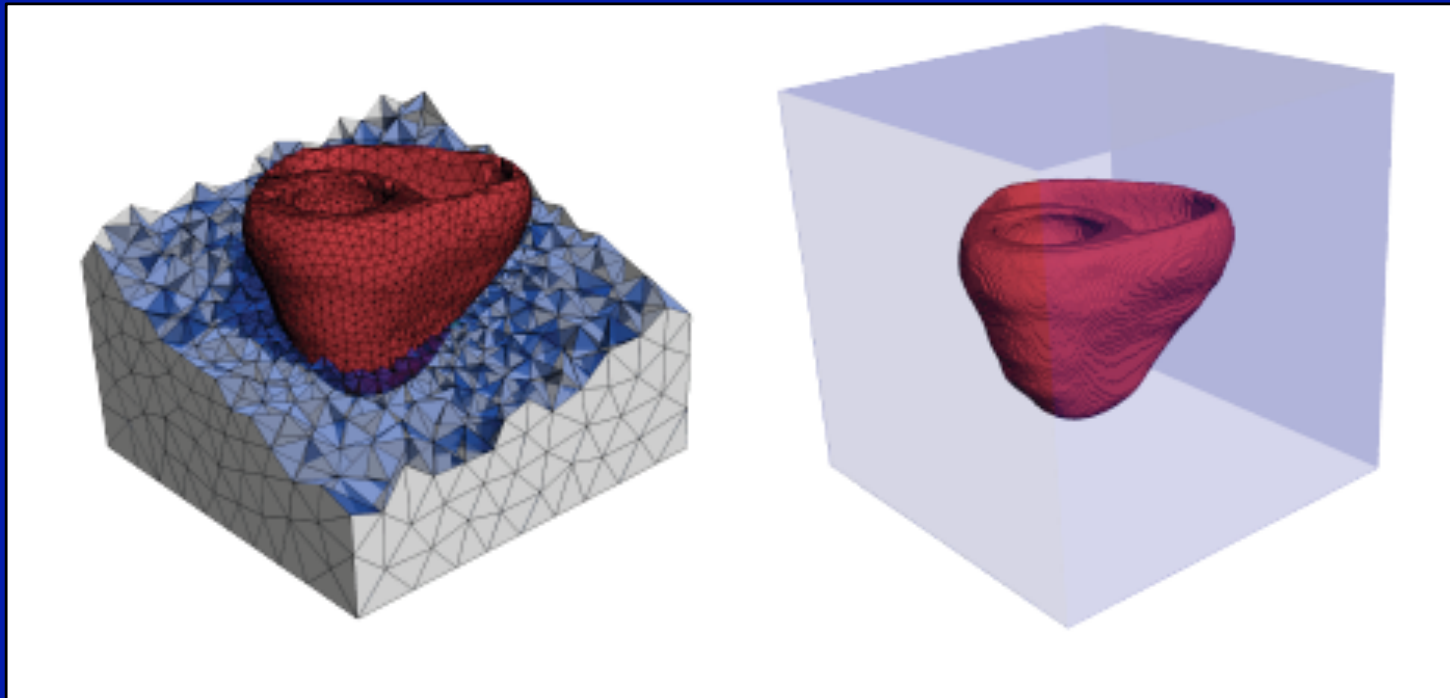
Amira



Notre
approche



J. Dardenne, S. Valette et al., Variational tetrahedral mesh generation from discrete volume data. *The Visual Computer*, pp (in-press), 2009.



DET automatic initialization

- Looking for a 3D affine transform \mathbf{T} which minimizes:

$$J(p) = \sum_{i=1}^{N_c} d_n^2(\mathbf{T}(p, \mathbf{x}_i)) + \sum_{i=1}^{N_{nod}} (I(\mathbf{T}(p, \mathbf{x}_i)) - I_{nod}(\mathbf{x}_i))^2$$

Where

$$p = (t_x, t_y, t_z, r_x, r_y, r_z, s_x, s_y, s_z, sh_{xy}, sh_{xz}, sh_{yz})$$

And

$$\mathbf{T} = \mathbf{T}_r \circ \mathbf{S} \circ \mathbf{R}$$

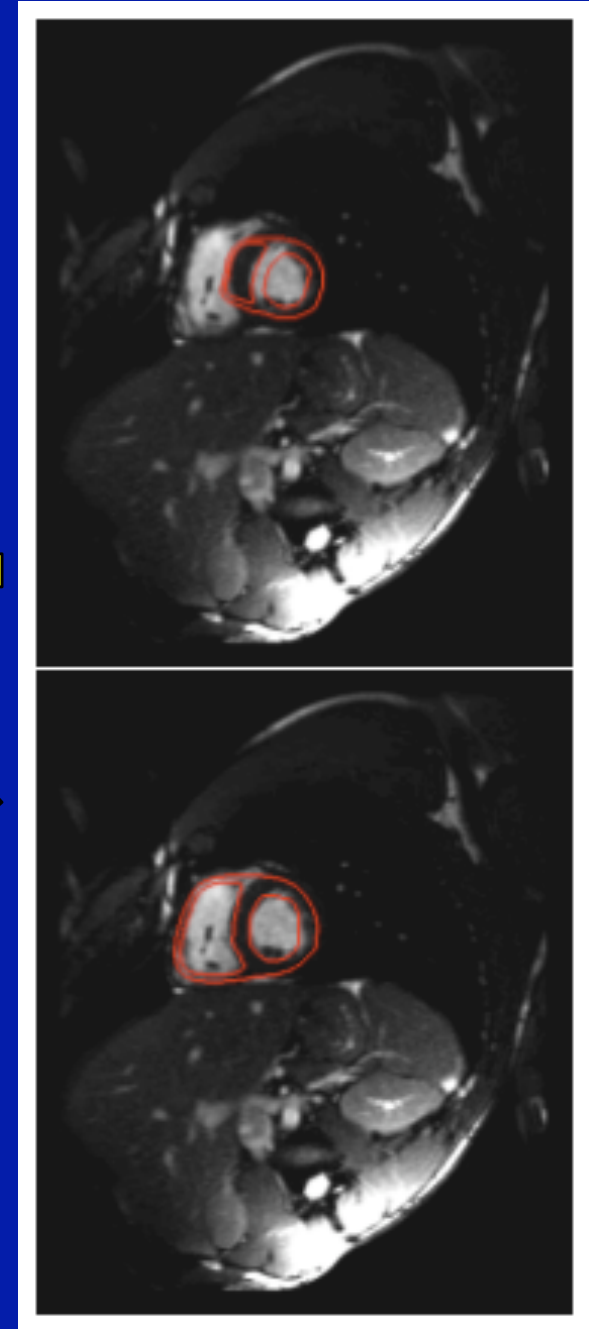
- Based on endocardial contours
- Multi-random-start simplex optimization method

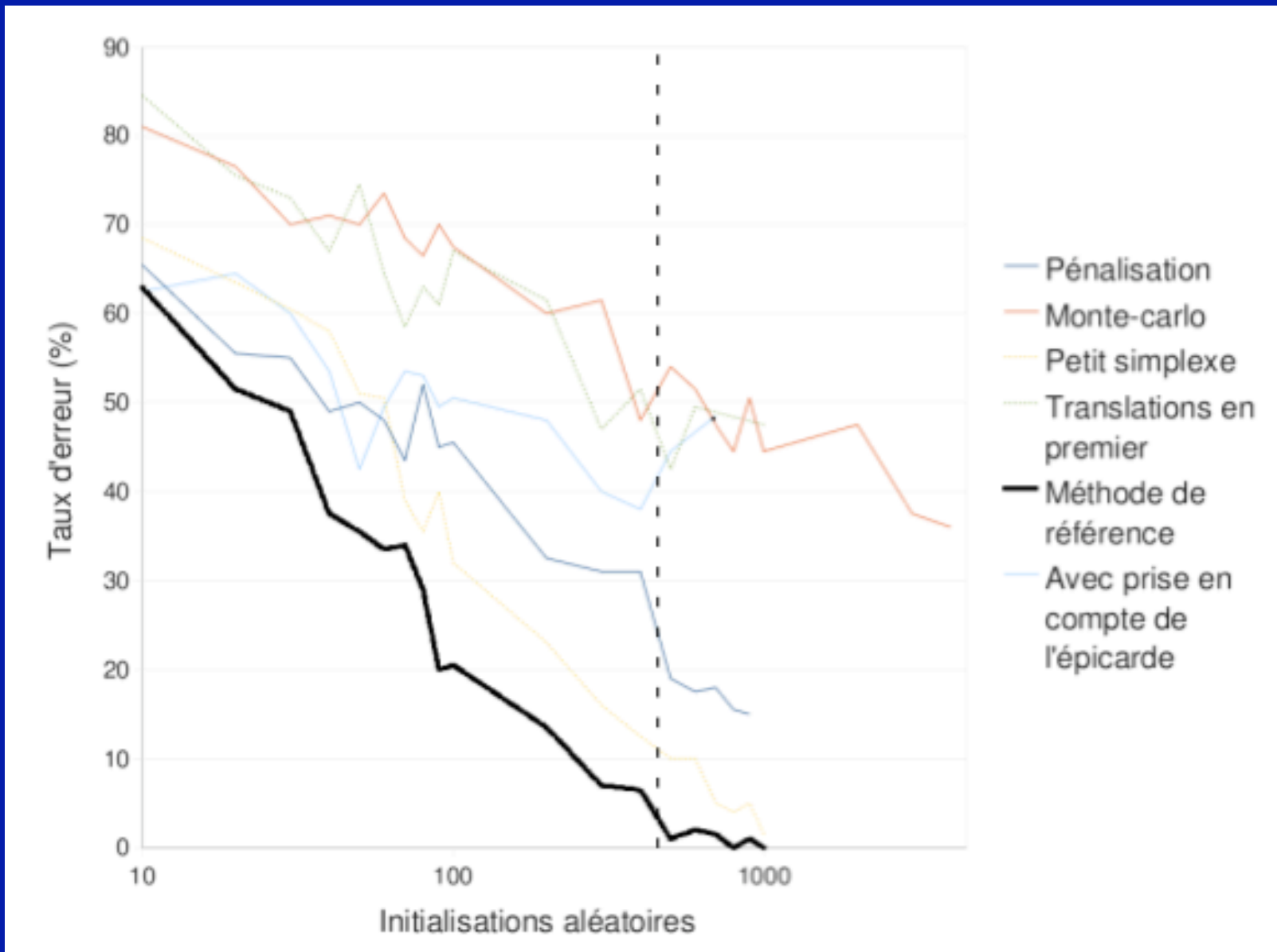
- **Initialization evaluation**

- One ED 3D image
- Comparison to a manual established reference

$$\|p_{ideal} - p\|^2 < \Delta_p$$

- 10-4000 random starts
- 200 registrations to estimate the error rate for each configuration

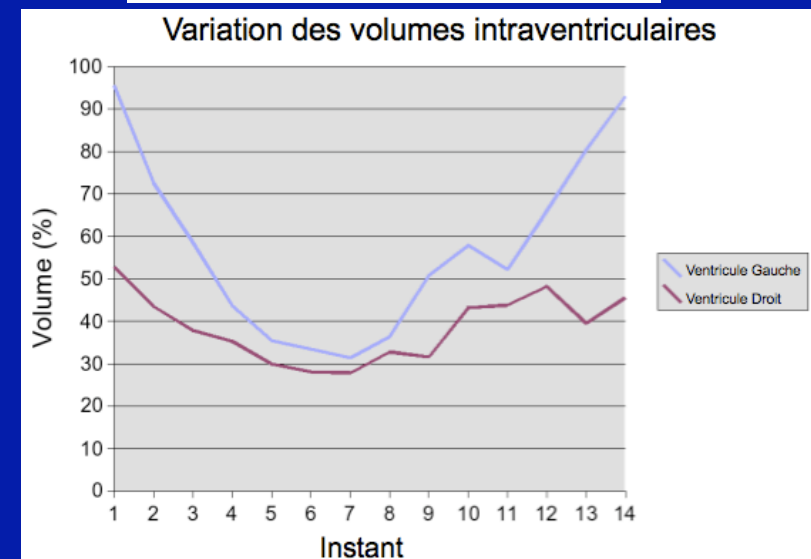
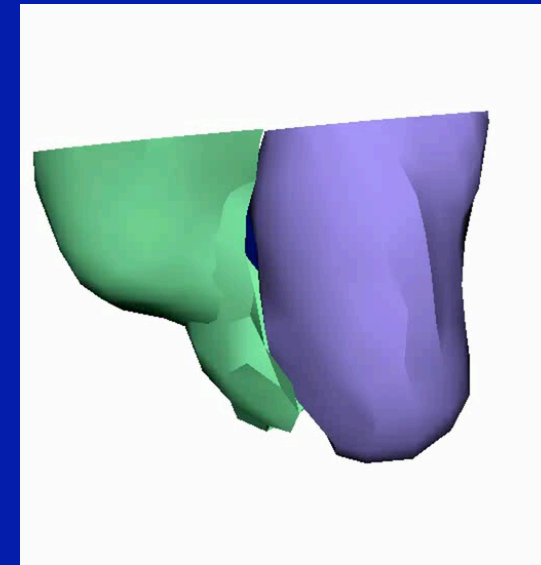
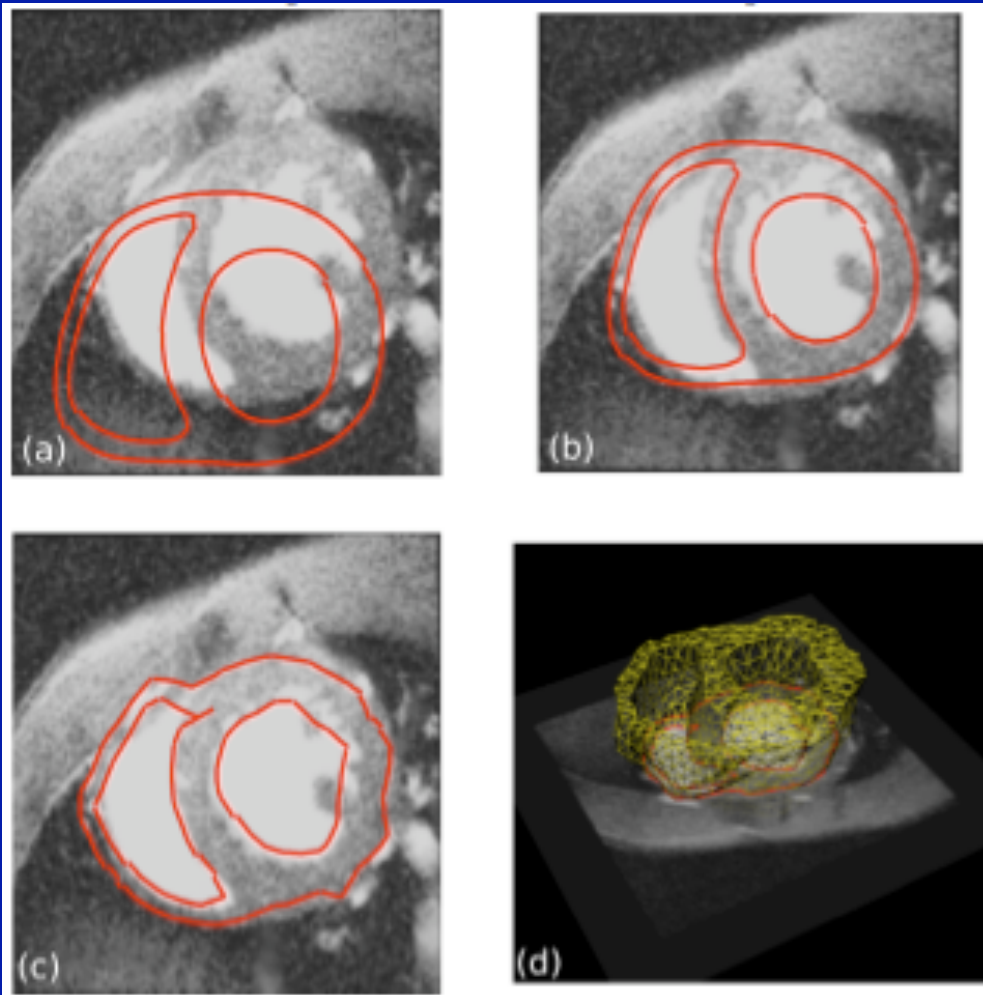




➔ The proposed approach offers the best compromise

Some results

- Mouse Images 7T MRI(FLASH sequence FOV=25 mm² , 256 ×256 pixels, slice thickness=1 mm), cardiac freq. = 450 bpm.



DET Spatio-temporal Model

- Assumptions
 - Continuous & periodic motion
- Simplified fundamental equation of dynamics :

$$\mathbf{D}\dot{\mathbf{U}}(t) + \mathbf{K}\mathbf{U}(t) = \mathbf{F}(\mathbf{U}(t), t)$$

- Trajectory representation (Fourier)

$$\mathbf{U}(t) = \sum_{n=0}^N \mathbf{A}_n \cos\left(\frac{2\pi n}{N}t\right) + \mathbf{B}_n \sin\left(\frac{2\pi n}{N}t\right)$$

Implementation of the non-linear problem

- Pseudo instationnary scheme:
a series of linear problems

$$\begin{cases} \frac{d\mathbf{U}}{d\tau} = \mathbf{F}(\mathbf{U}) - \mathbf{A}\mathbf{U} \\ \mathbf{U}(0) = 0. \end{cases}$$

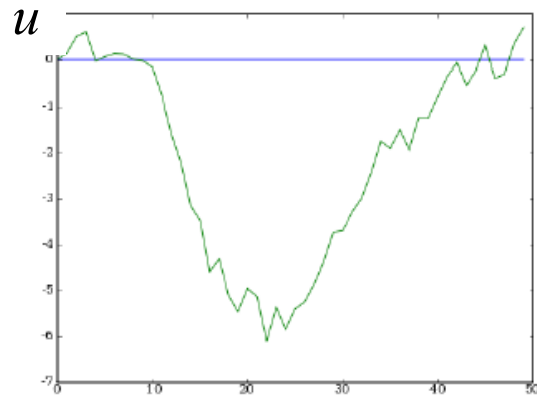
$$\left(\frac{1}{\Delta\tau} + \mathbf{A}\right)\mathbf{U}^\tau = \mathbf{F}(\mathbf{U}^{\tau-1}) + \frac{1}{\Delta\tau}\mathbf{U}^{\tau-1}$$

→ Existence, uniqueness and asymptotic behavior with respect to τ can be demonstrated

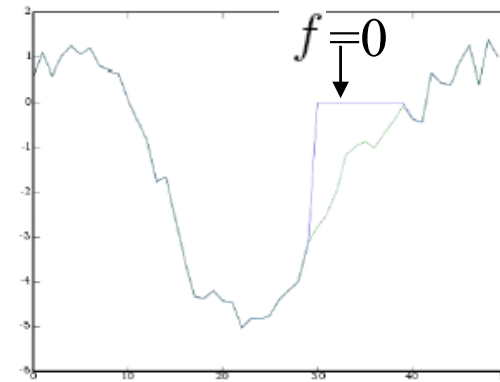
[Schaerer et al., in revision]

1D+time illustration

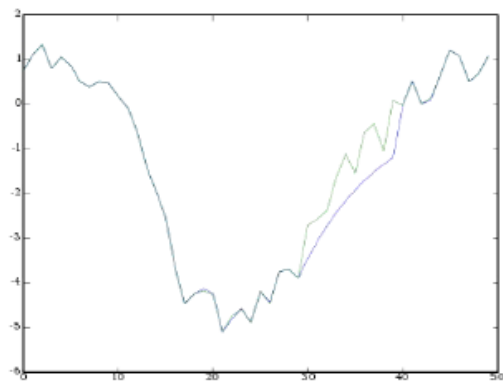
$$\alpha u'(t) + ku(t) = f(t)$$



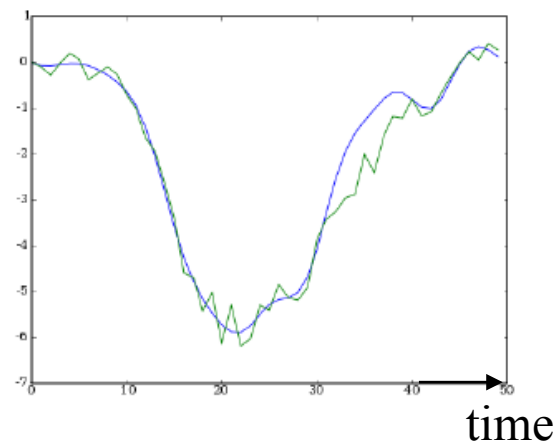
(a)



(b)



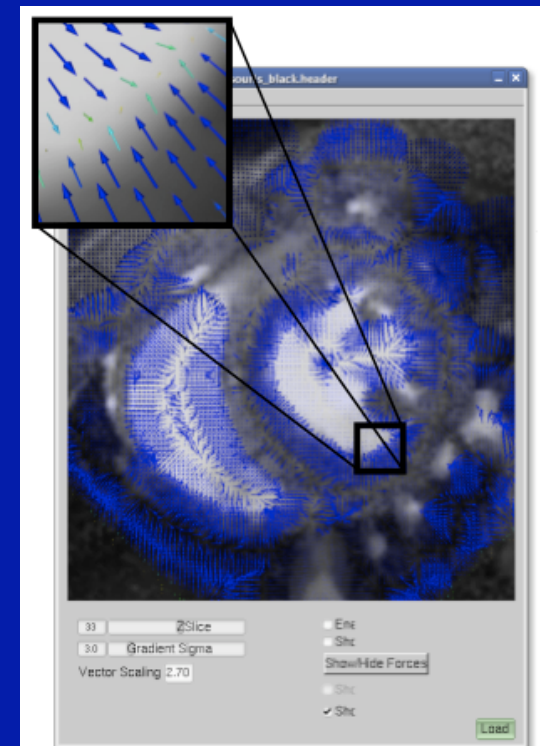
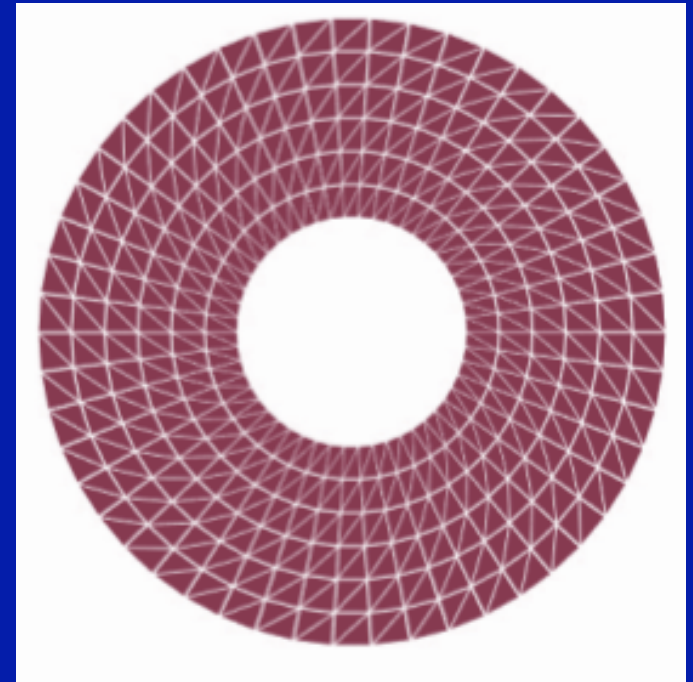
(c)



(d)

Fig. 1. Sample 1D results: (a) Target signal (green curve) and initialization ($U = 0$), (b) Results without filtering or damping, (c) Results with damping but no filtering ($d=0.2, k=1$), (d) Results with damping and filtering ($d=0.2, k=1$ and 8 harmonics)

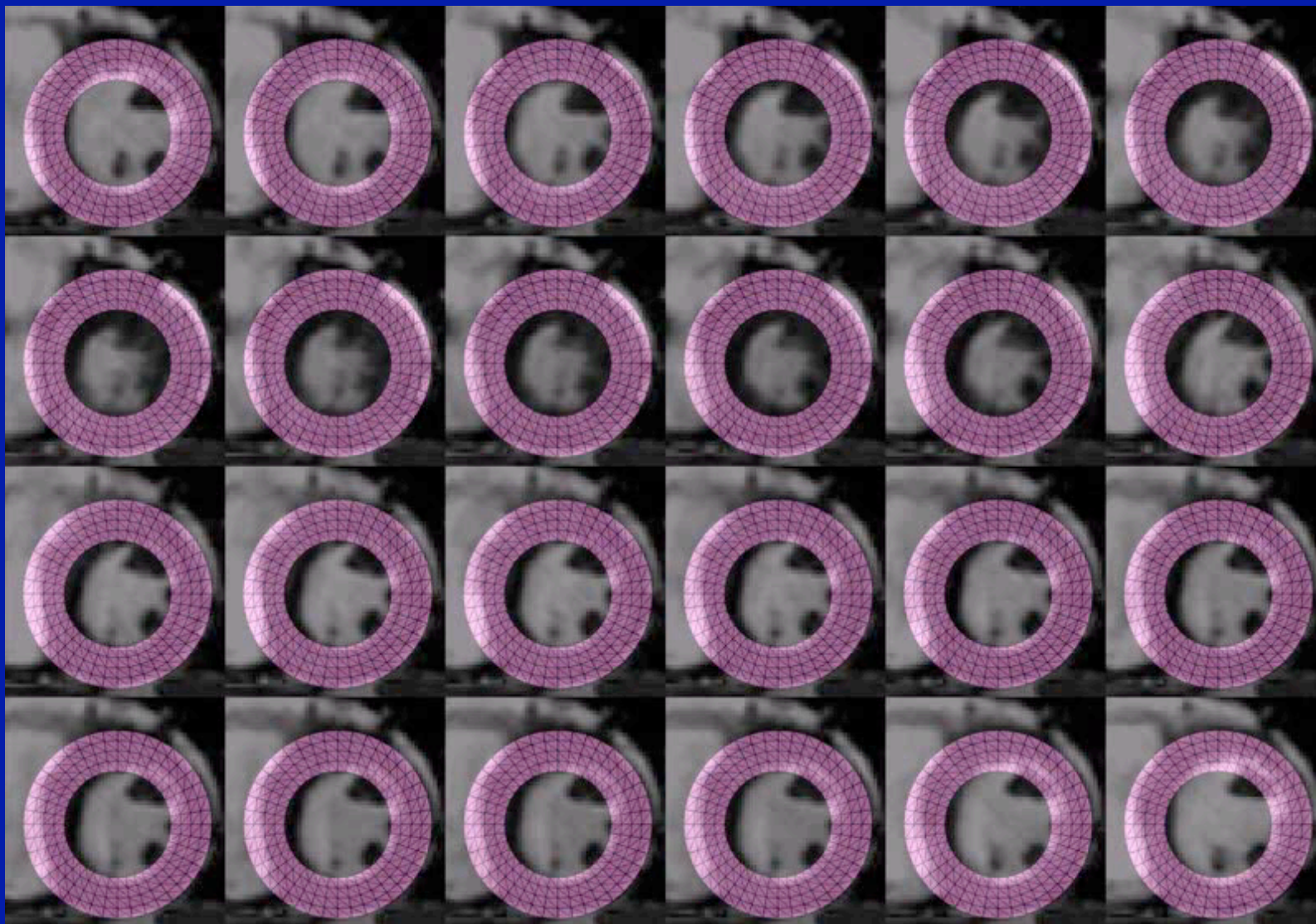
- **2D Implementation**
 - Initial myocardial model: annulus
 - Constraint field from images
 - Gradient + GVF, but
 - Temporal jittering
 - Local lack of data
 - Morphological pre-segmentation as a constraint (PhD thesis J. Cousty, ESIEE) + GVF

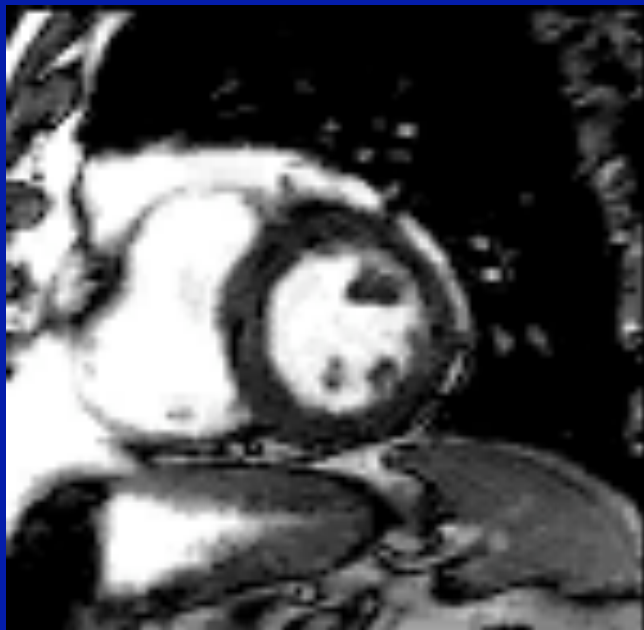


Dynamic DET segmentation Results

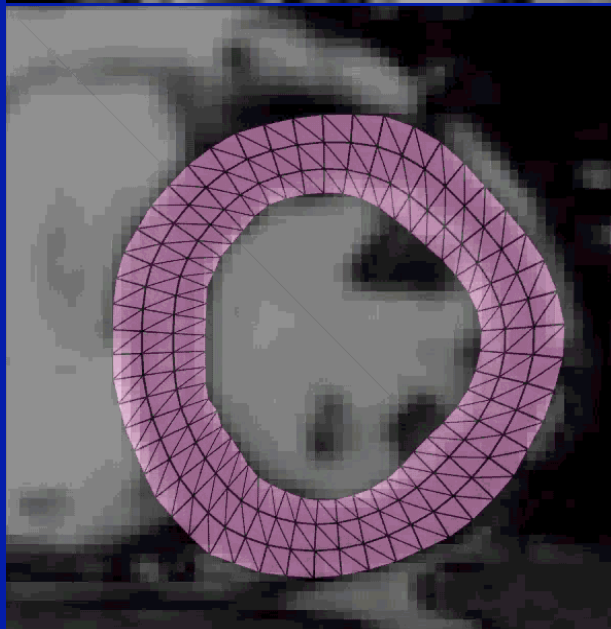
- **MRI in vivo Sequences**
 - Human cardiac MRI database at <http://laurentnajman.org/heart>
 - 18 cases
 - multi-phase, multi-slice cine-MRI
 - 5-10 slice levels 256x256
 - ~1mm in plane, 5-10mm slice thickness
 - 15-30 phases
 - **IMPEIC project: 'Initiative Multicentrique pour une Plateforme d'Evaluation en Imagerie Cardiaque'**
(supported by GDR STIC-Santé)

Time →

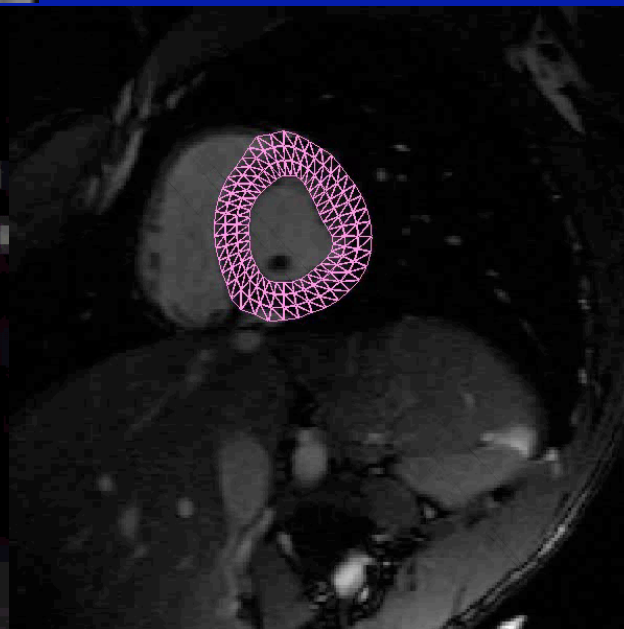




**in vivo MRI Sequence
from the database**

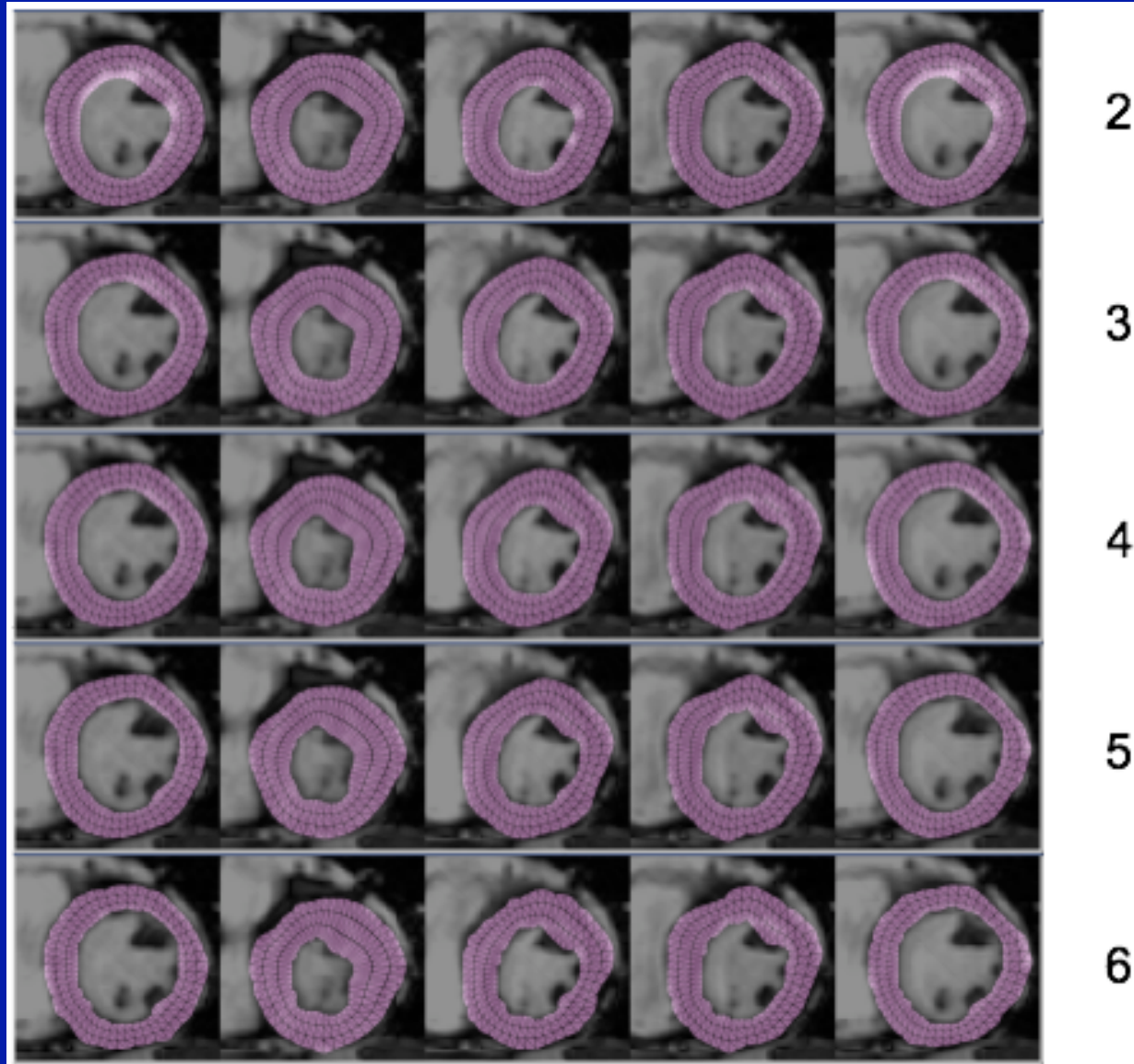


**Segmentation with
pre-segmentation**



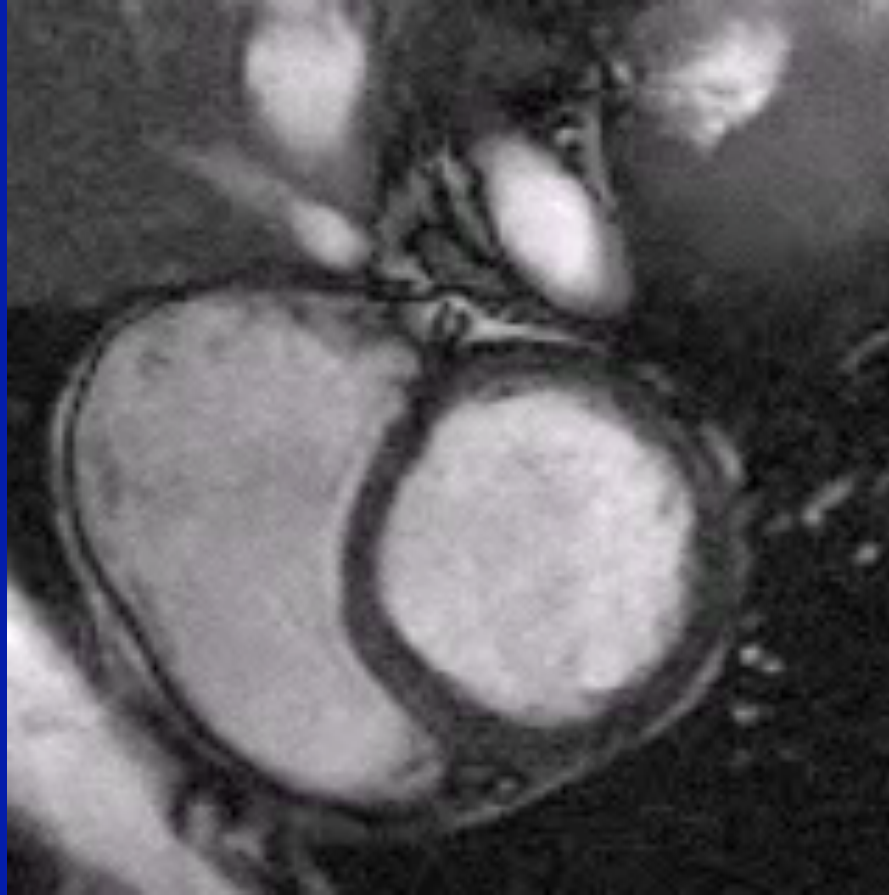
**GVF based
Segmentation**

- **Harmonic order impact**

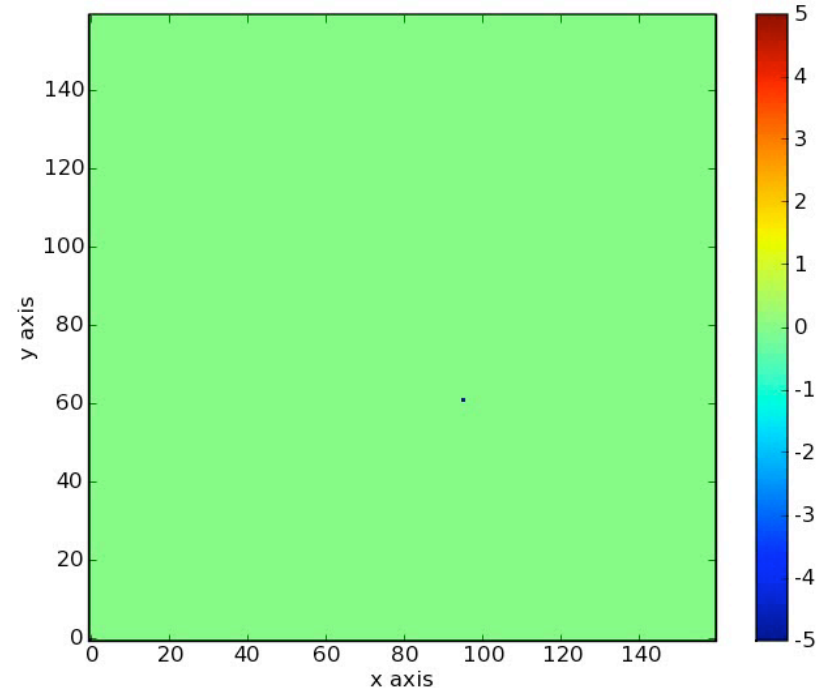
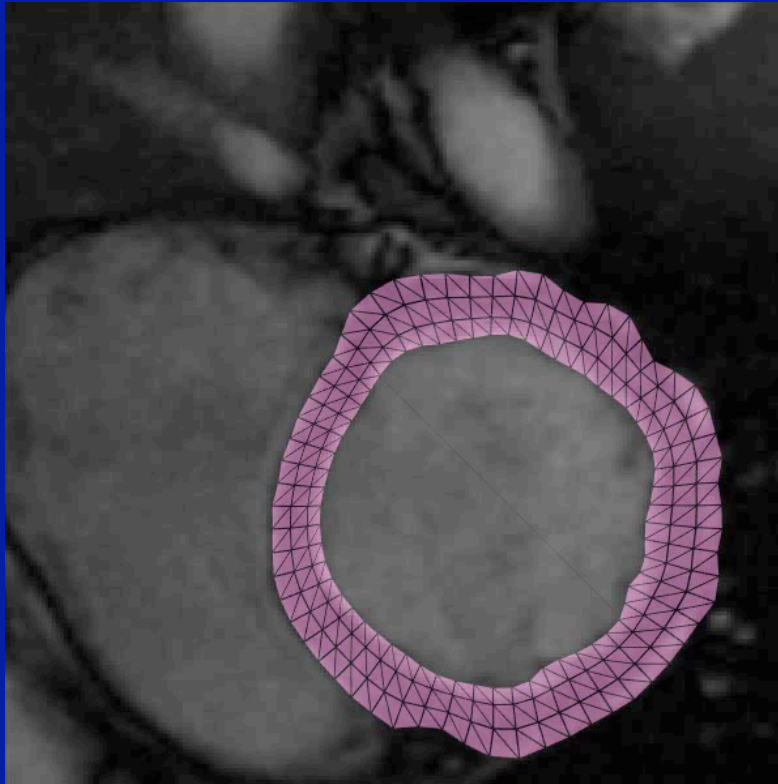


Dynamic DET motion tracking Results

- **Synthetic 2D MRI Sequences**
 - Spatio-temporal reference heart model
[Clarysse, MEDIA 2000]
 - Simulated MRI Sequence
[Delhay, submitted]
 - Parameters
 - Model: normal/patho
 - Image: variable noise level



Simulated Sequence 'normal subject',
~ basal level



Radial displacement
error (mm)

Conclusions on DET model

- **Generic concept**
 - Adaptable to various geometries : Tested on 3D US imaging
 - Dynamic DET: motion tracking in Cine MRI → tagged MRI
- **Still...**
 - Parameter tuning
 - Template initialization
- **Remarks**
 - Computing a convenient force field (modality dependent)
 - Simplifying image information
 - low level processing
 - multi-resolution approaches

A Priori Statistical dynamical model

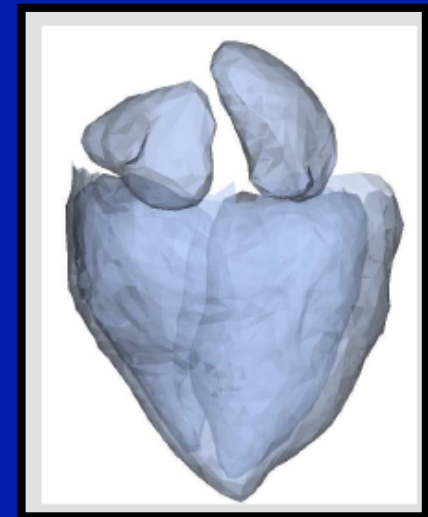
Learning based static model (4 chambers)

- 15 healthy subjects [Lötjönen et al., 2004]
- Short Axis + Long Axis slices (End- Diastole)

Set of n geometrical models
whose points are spatially coherent

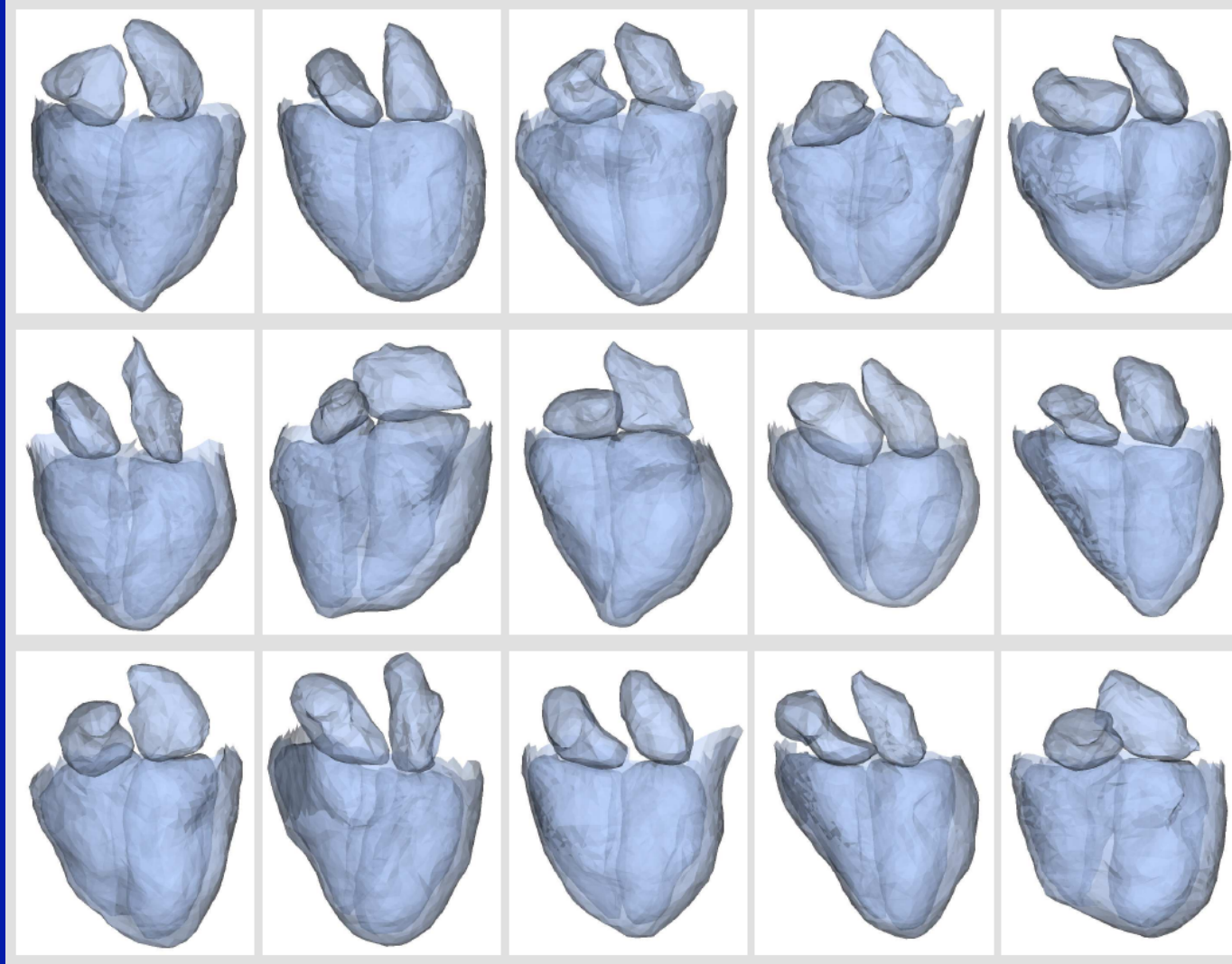
$$\bar{\mathcal{S}} = \frac{1}{n} \sum_{p=1}^n \mathcal{S}(p) = \frac{1}{n} \sum_{p=1}^n (\mathbf{x}^{i,p})_{1 \leq i \leq m}$$

Semi-landmarks number = 2086



- Inter-individual variability

Subject 1



Subject 15

■ Learning based dynamic model

[Delhay *et al.* Cinc 05]

➔ Atlas transport through the sequence

For each subjects:
sequential non-rigid
registrations

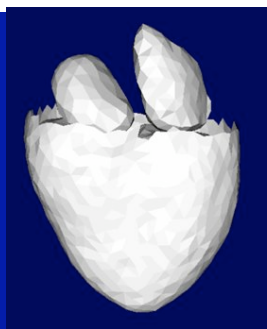
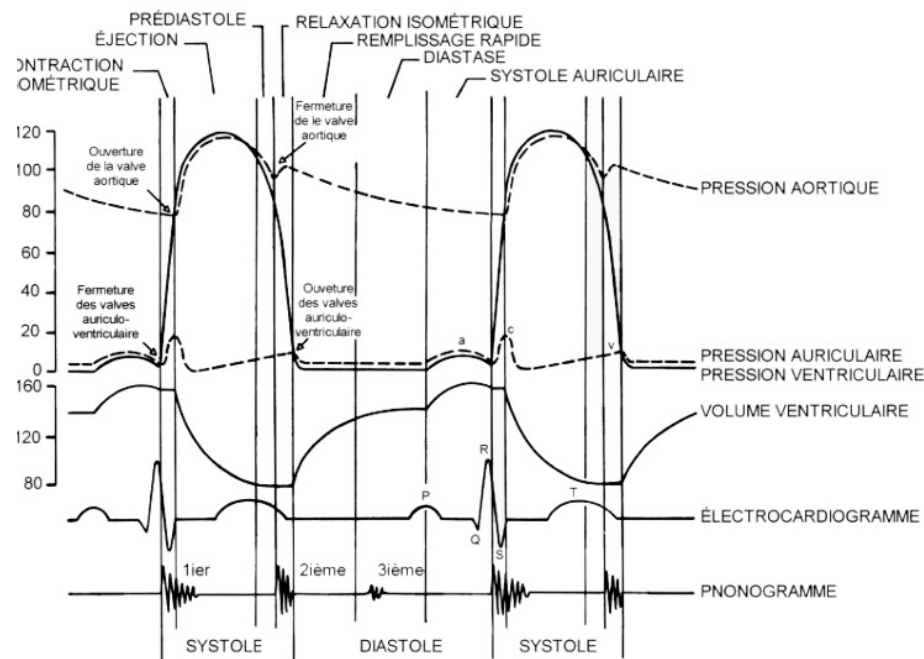
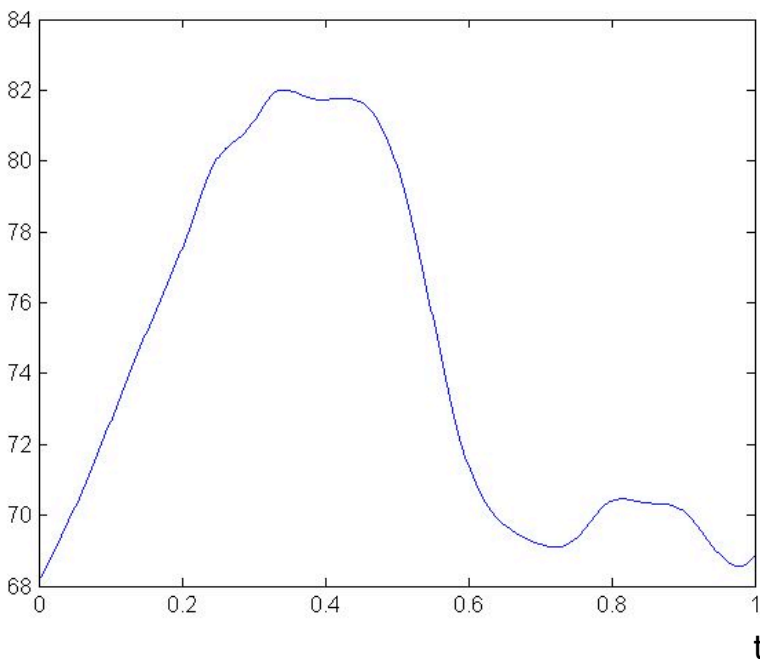
[Lötjönen and Mäkelä, 2001]

+

Manual corrections

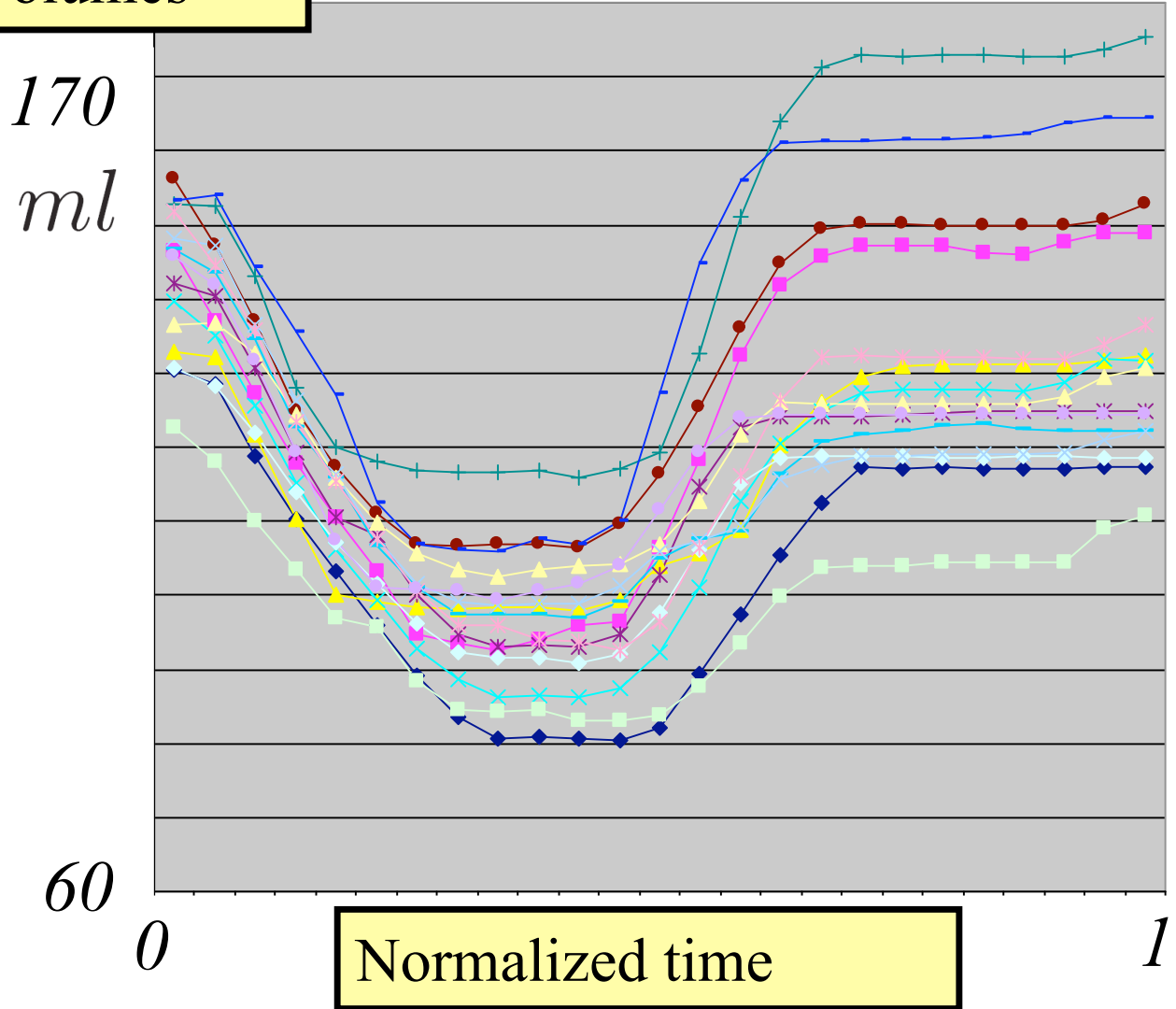
Trajectory normalisation for 15 healthy volunteers, ~2100 pseudo-landmarks over the whole heart

y (cm)

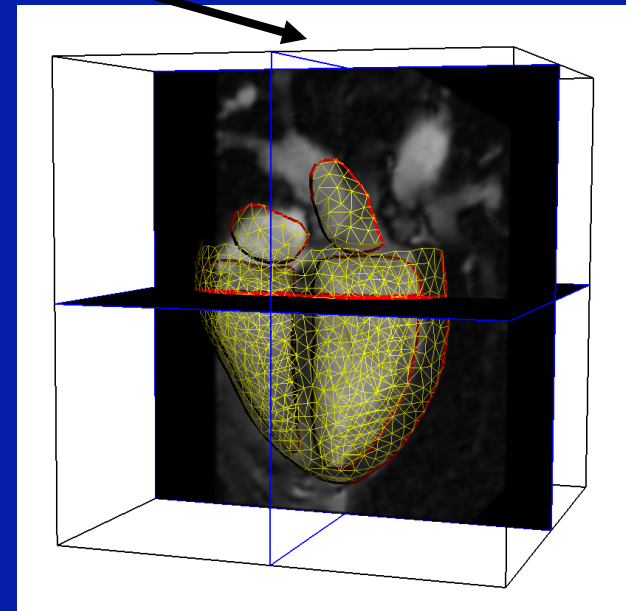
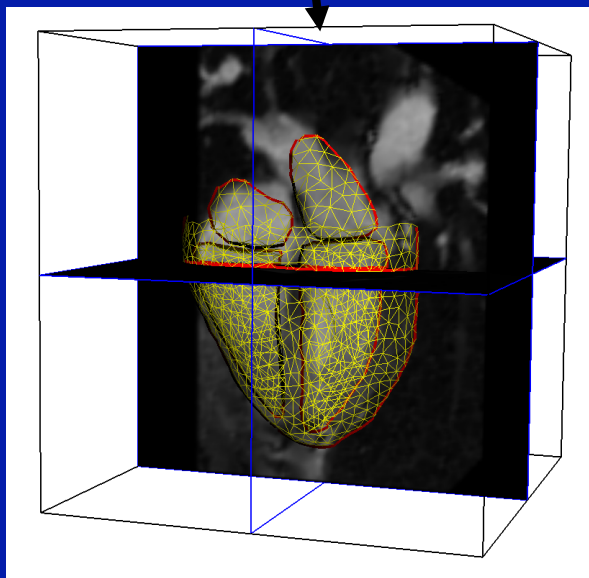
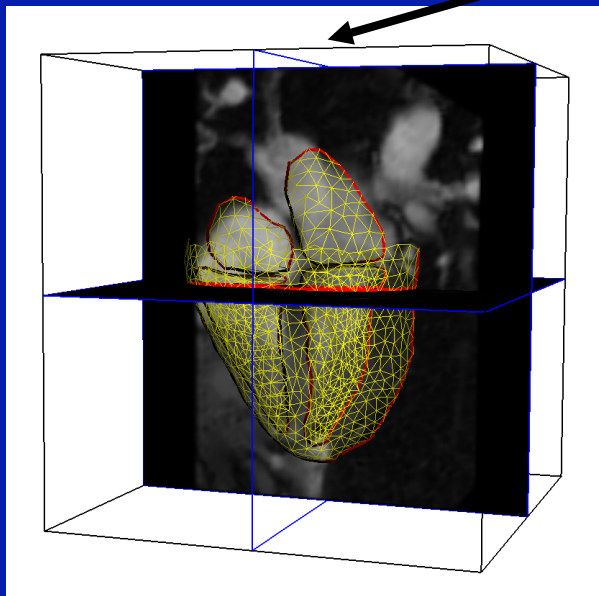
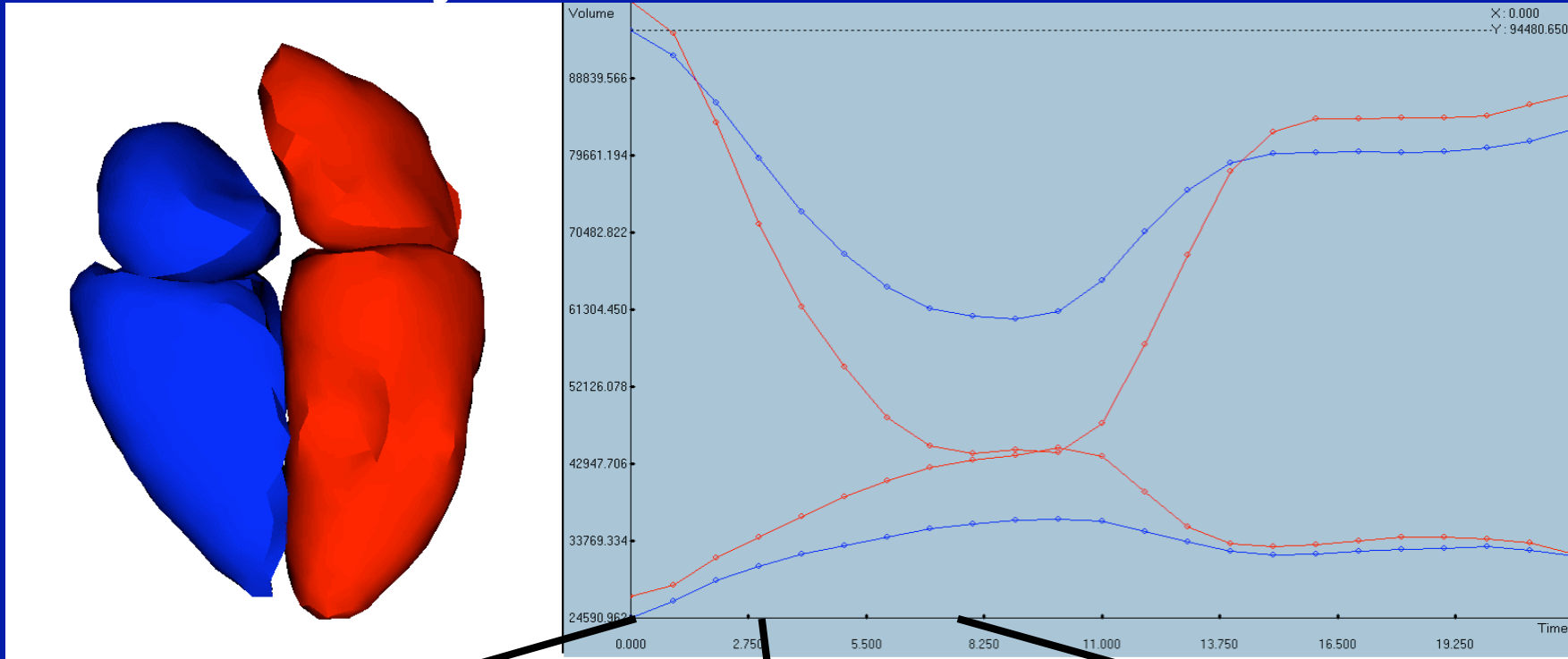


0
D S D
1

LV
Volumes



Mean dynamical model



Landmark variability modeling

→ Parzen Windowing

- Only one parameter
- Continuous probability density function (*PDF*)

$$p(\mathbf{x}^i(t)) = \frac{1}{nh} \sum_{p=1}^n K \left(\frac{\|\mathbf{x}^i(t) - \mathbf{x}^{i,p}(t)\|^2}{h} \right)$$

n : number of subjects in the data set

$\mathbf{x}^{i,p}(t)$: Coordinates of the landmark i at time t

K : Gaussian kernel

h : Kernel size

■ A priori driven segmentation

A priori regularizing term C

$$\mathcal{J}(I_c, I_s, \varphi) = \mathcal{A}(I_c, I_s, \varphi) + \alpha \mathcal{R}(\varphi) + \gamma \mathcal{C}(\varphi)$$

$$\mathcal{C}(\varphi, h) = \frac{1}{2} \sum_{i=1}^m (1 - p(\mathbf{x}^i(t_j), h))^2$$

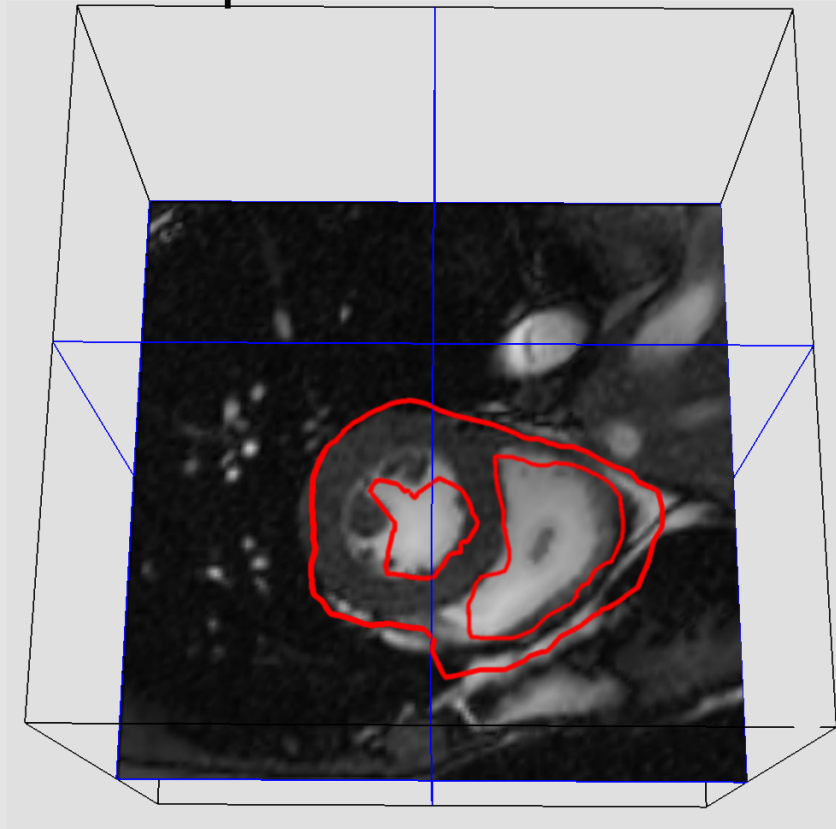
Jacobian of the transformation

$$\frac{\partial \mathcal{C}_i(\boldsymbol{\xi}, h)}{\partial \boldsymbol{\xi}} = \left(1 - p(\varphi_{t_0; t_j}(\mathbf{x}^i(t_0), \boldsymbol{\xi}), h) \right) \times J \times \frac{\partial p(\varphi_{t_0; t_j}(\mathbf{x}^i(t_0), \boldsymbol{\xi}), h)}{\partial \mathbf{x}}$$

Derivative of the Parzen function

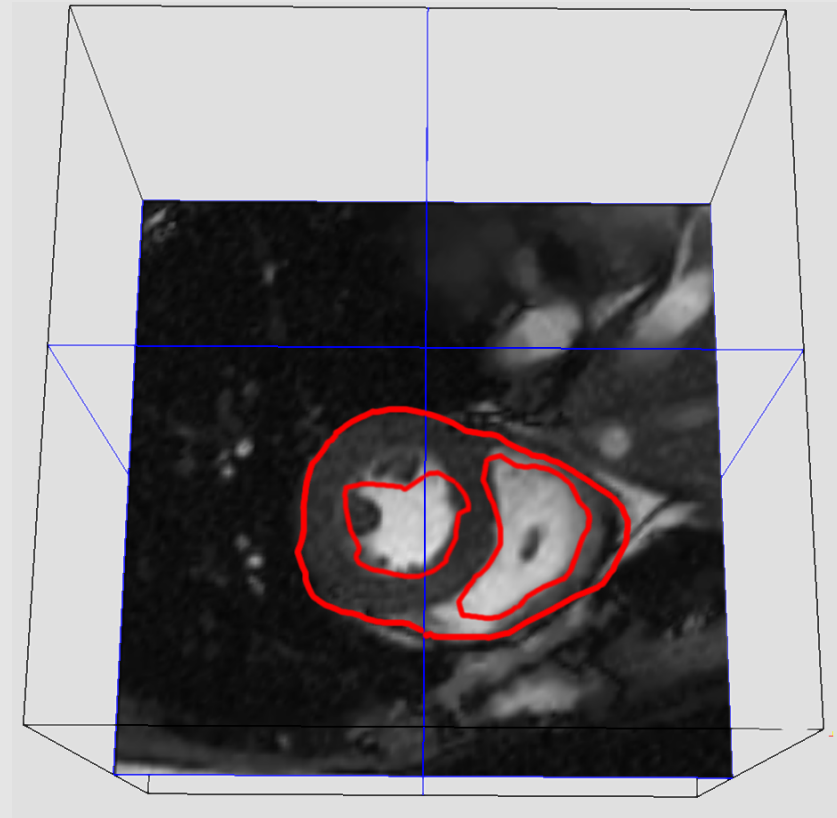
- Preliminary results : on 3D real cardiac data
(Structures tracking)

Sequential estimations



instant: 5/25

Sequential estimations + model



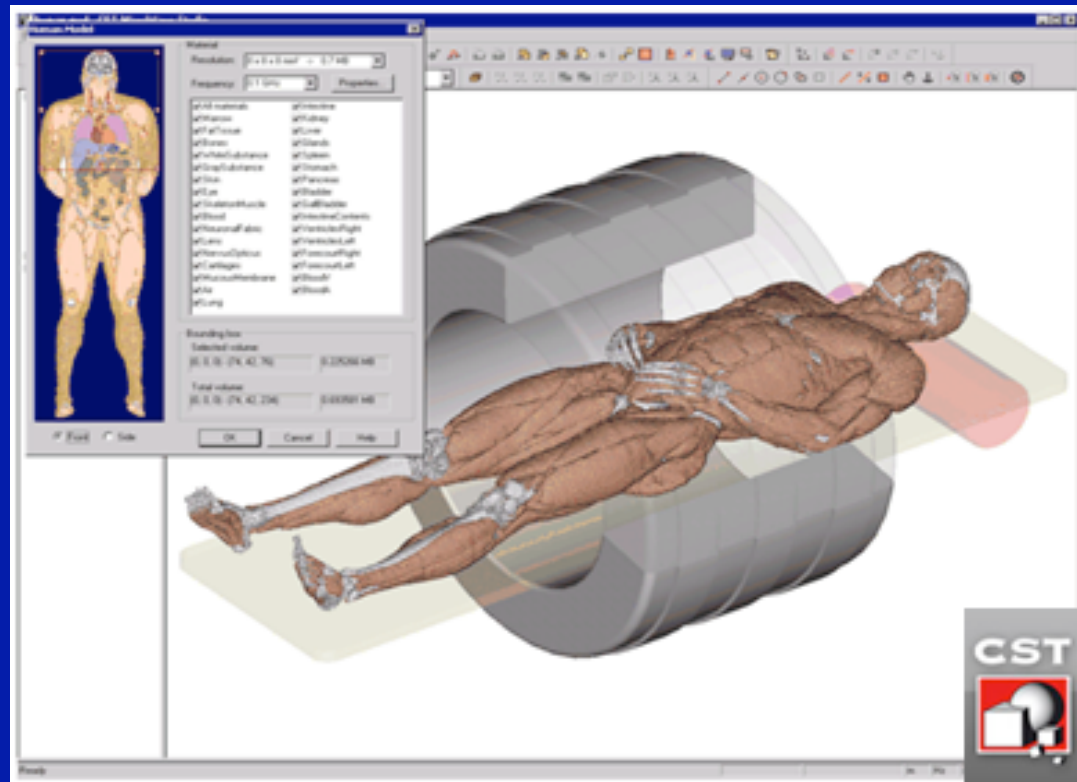
$\alpha = 0.1$ $\gamma = 0.4$
 $h = 5.0mm$

Conclusions on Learning based models

- **Generic concept**
 - Adaptable to various anatomical structures
 - Adaptable to deformable structures
- **But**
 - Construction of the a priori model is laborious
 - Difficulties to construct large sample sets
 - ➔ limitation in shape variations (normal & abnormal)
- **Artificial enlargement [Koikalainen, IEEEETMI-2008]**
 - *Partitioning the structure into separate parts*
 - *Non-rigid local perturbations*
- **Artificial smoothness**
 - Covariance matrix to induce smoothness to def. modes
 - PCA + FEM

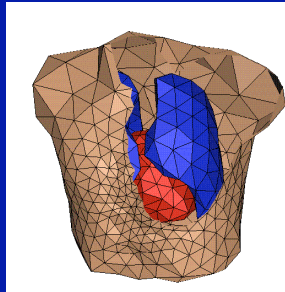
Method's evaluation : a whole numeric approach

→ 3D virtual model and imaging



<http://www.cst.com/>

- **Components**



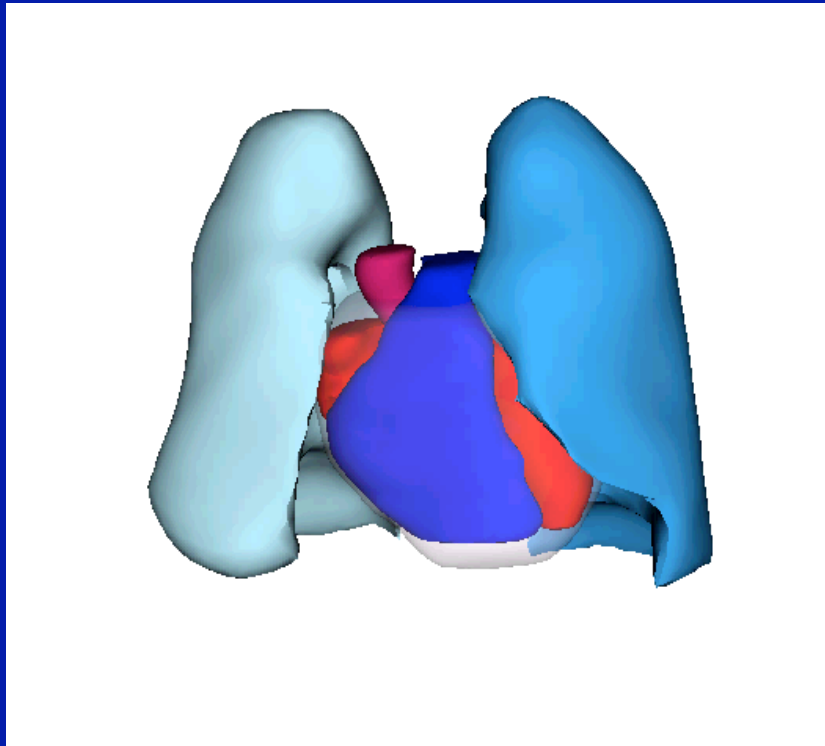
Models

Medical
imaging
modality
simulators

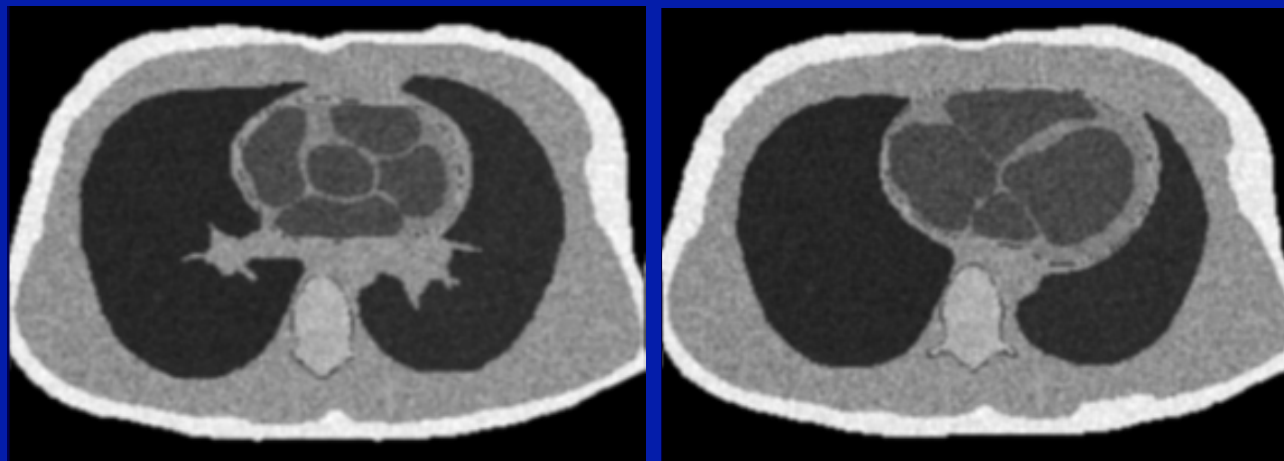
Simulated
images



Breathing thorax & beating heart model



*[Haddad, PhD thesis 2007],
[Haddad et al, IEEE EMBS,
2007]*



Simulated images,
MRI Simulator
SIMRI, CREATIS-
LRMN
*[Benoit-Cattin et al,
JMR,05]*

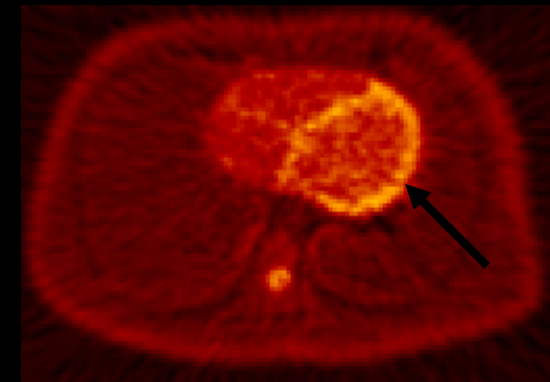
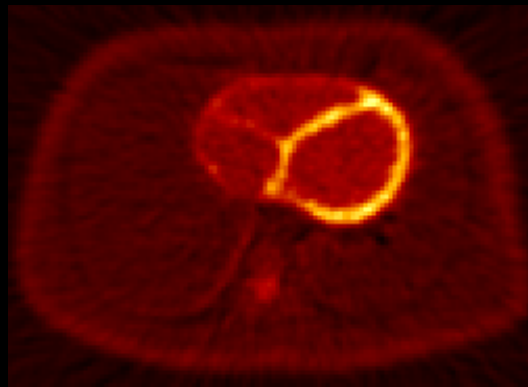
Creatis
LRMN

PET simulation with SORTEO (Cermep, Lyon): Simulated emission PET images for healthy and ischemic models (hypocaptation of glucosis)

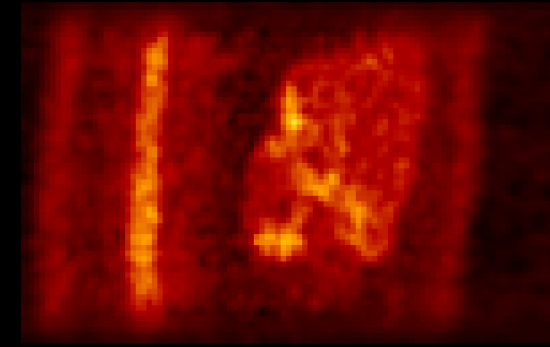
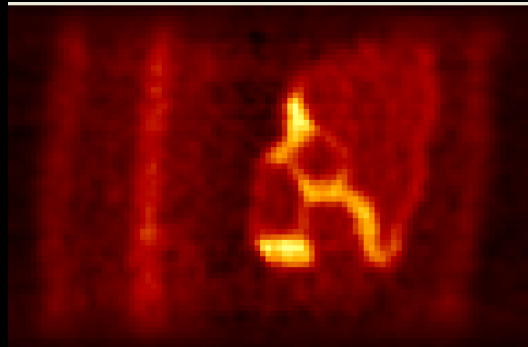
Healthy model

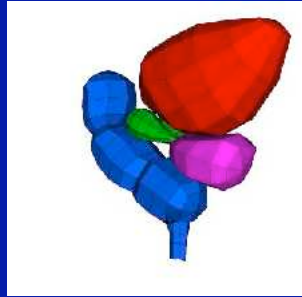
Ischemic model (F-18
collecting in the myocardium
with 40% of its initial value)

Axial slice



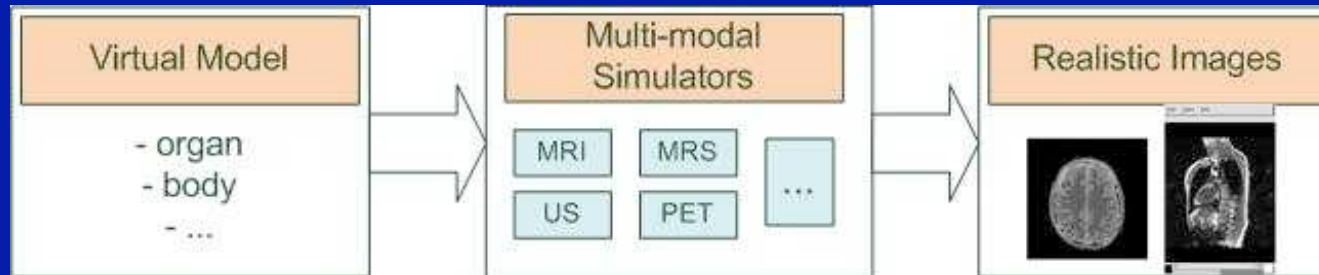
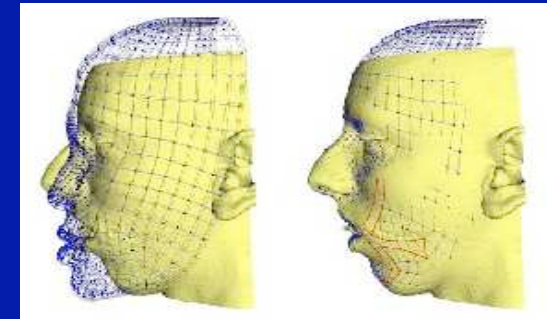
Sagittal slice





→ **Simed** project
(*P. Clarysse – E. Promayon*)

Simulation en Imagerie Médicale
pour le Diagnostic et la
Thérapie



→ Virtual Physiological Human European NOE



Collaborations / Support



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- N. Pauna
- R. Haddad
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- J. Vandemeulebroucke
- L. Grezes-Besset
- T. Schilling

Post-docs:

- J. Schaerer

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- J-M Morvan, IGD, Lyon
- J. Pousin, Inst. C. Jordan, Lyon
- C. Ginestet, CLB, Lyon
- D. Clot, LASS, Lyon

Supports

Région Rhône-Alpes: Cluster ISLE, ADemo, Santé & HPC, Cancéropole

CNRS: PICS 1932, AS ICOMIM

GDR STIC-Santé: Projet IMPEIC

Ministère de la Recherche: ACI AGIR

ANR: GWENDIA

Europe: EGEE, VPH

HUT, Finland

Prof. T. Katila
J. Nenonen



Technical Research
Centre of Finland (VTT)

J. Lötjönen



- Thank you!

