## Numerical Methods for Image Registration

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### **Motivation**

#### **Image Registration**

Given a reference image  $\mathcal{R}$  and a template image  $\mathcal{T}$ , find a reasonable transformation *y*, such that

the transformed image  $\mathcal{T}[y]$  is similar to  $\mathcal{R}$ 





#### transformed template $\mathcal{T}[y]$

template T

Titlepag

Motivation

Outline

### **Motivation**

#### Image Registration

Given a reference image  ${\mathcal R}$  and a template image  ${\mathcal T},$ 

find a reasonable transformation y, such that

the transformed image  $\mathcal{T}[y]$  is similar to  $\mathcal{R}$ 

#### Questions:

- What is a transformed image T[y]?
- What is similarity of  $\mathcal{T}[y]$  and  $\mathcal{R}$ ?
- What is reasonability of y?

#### Image Registration: Variational Problem

$$\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min,$$

$$y_{\rm reg}(x) = x$$

### Outline

- Applications
- ▶ Variational formulation  $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \mathcal{S}[y] \xrightarrow{y} \min$ 
  - image models T[y]
  - distance measures  $\mathcal{D}[T[y], R]$
  - regularizer  $\mathcal{S}[y]$
- Numerical methods
- Constrained image registration
- Conclusions





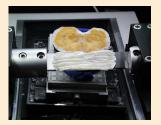
## **Applications**





### **HNSP: Sectioning**

with Oliver Schmitt, Institute of Anatomy, University Rostock, Germany



- sliced
- flattened
- stained
- mounted
- ▶ ...
- digitized



#### large scale digital images, up to $10.000 \times 20.000$ pixel



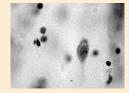


## **HNSP: Microscopy**







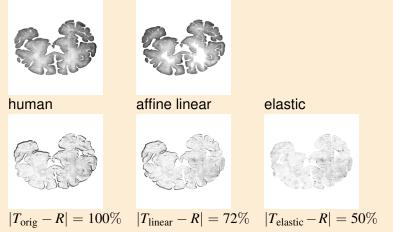






### HNSP: Deformed Images

#### sections 3.799 and 3.800 out of about 5.000

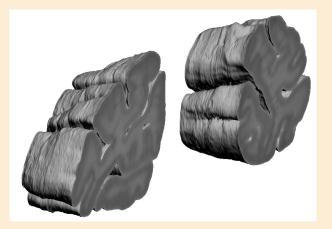






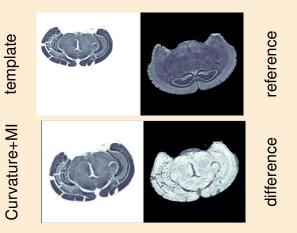
### **HNSP: Results**

3D elastic registration of a part of the visual cortex (two hemispheres; 100 sections á  $512 \times 512$  pixel)





#### Multi-Modal Registration with Stefan Heldmann, SAFIR, University of Lübeck





#### Registration of *Sprague-Dawley* Brain with Stefan Wirtz, SAFIR, University of Lübeck





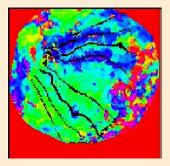


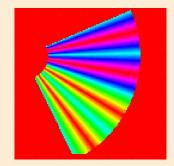


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## Neuroimaging (fMRI)

with Brian A. Wandell, Department of Psychology, Stanford Vision Science and Neuroimaging Group



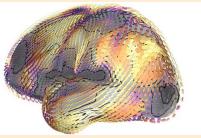


"flattened visual cortex"



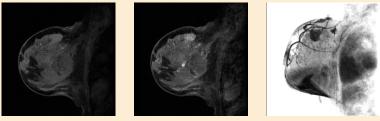
#### DTI: Diffusion Tensor Imaging with Brian A. Wandell, Department of Psychology, Stanford Vision Science and Neuroimaging Group







#### MR-mammography, biopsy (open MR) with Bruce L. Daniel, Department of Radiology, Stanford University



pre contrast

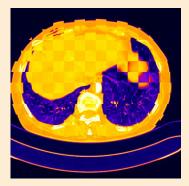
post contrast

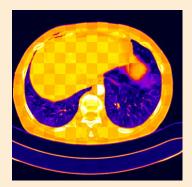
3D





#### Liver Registration with Stefan Wirtz, SAFIR, University of Lübeck & Siemens Erlangen, Germany







### Virtual Surgery Planning

S. Bommersheim & N. Papenberg, SAFIR, BMBF/FUSION Future Environment for Gentle Liver Surgery Using Image-Guided Planning and Intra-Operative Navigation



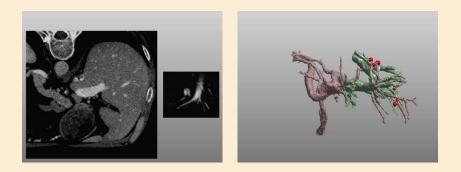








### **Results for 3D US/CT**



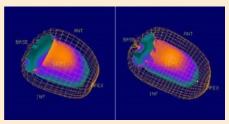


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### Cardiac Images

#### with Tracy Faber, Department of Radiology, Emory University, Atlanta,USA

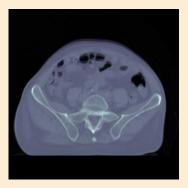


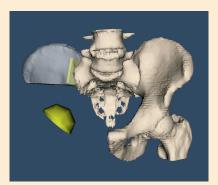




### Virtual Surgery Planning

#### from Jan Ehrhardt, Medical Computer Science, University of Hamburg







#### Laser Surgery from Joachim Noack, Medical Laser Center Lübeck



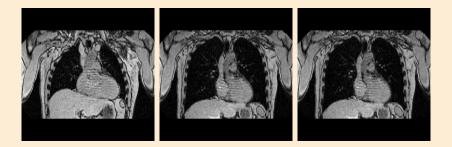


#### (Macula pucker, warped retina)



## **Motion Correction**

from Thomas Netsch, Philips Research, Hamburg, Germany





## Human Knee, 3D

from Astrid Franz, Philips Research, Hamburg, Germany

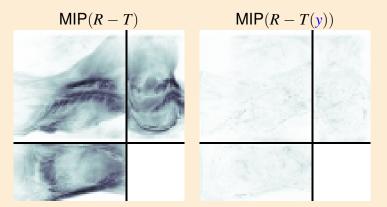
reference template  $T(\mathbf{y})$ B 2D slice



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### Human Knee, 3D

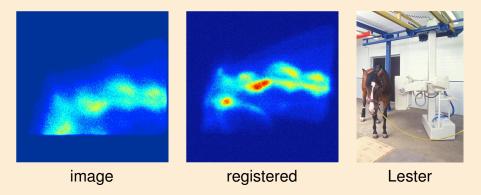
#### Maximum Intensity Projections





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#### SPECT: Single Photon Emissions CT with Oliver Mahnke, SAFIR, University of Lübeck & MiE GmbH, Seth, Germany





### **Registration in Medical Imaging**

- Comparing/merging/integrating images from different
  - ► times, e.g., pre-/post surgery
  - devices, e.g., CT-images/MRI
  - perspectives, e.g., panorama imaging
  - objects, e.g., atlas/patient mapping
- Template matching, e.g., catheter in blood vessel
- Atlas mapping, e.g., find 2D view in 3D data
- Serial sectioning, e.g., HNSP

Registration is not restricted to medical applications



▶ ...

### **Classification of Registration Techniques**

- feature space
- search space
- search strategy
- distance measure
- dimensionality of images (d = 2, 3, 4, ...)
- modality (binary, gray, color, ...)
- mono-/multimodal images
- acquisition (photography, FBS, CT, MRI, ...)
- inter/intra patient





## **Image Registration**

## **Transforming Images**

# $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min$





Variational Approach for Image Registration

 $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min$ 

 $\blacktriangleright$  Continuous models  $\mathcal{R}, \mathcal{T}$  for reference and template:

discrete data X, T  $\rightsquigarrow$   $\mathcal{T}(x) = interpolation(X, T, x)$ 

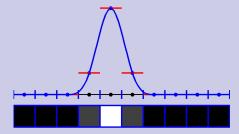
• Transformation  $y : \mathbb{R}^d \to \mathbb{R}^d$ 

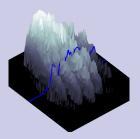
 $\mathcal{T}[y](x) = \mathcal{T}(y(x)) = \text{interpolation}(X, T, y(x))$ 



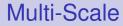


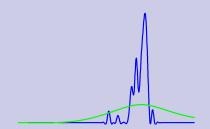
### Interpolation









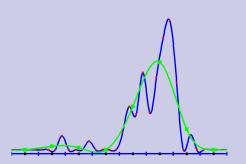






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### **Multilevel**







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### **Transforming Images**

$$\mathcal{T}[\mathbf{y}](\mathbf{x}) = \mathcal{T}(\mathbf{y}(\mathbf{x})) = \text{interpolation}(\mathbf{X}, \mathbf{T}, \mathbf{y}(\mathbf{x}))$$

non-linear







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## **Distance Measures**

# $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min$





#### **Distance Measures**

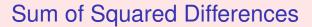
Feature Based

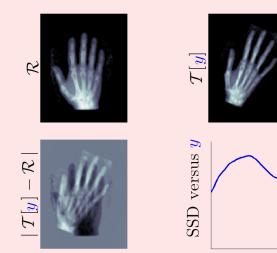
(Markers / Landmarks / Moments / Localizer)

- ► *L*<sub>2</sub>-norm, *Sum of Squared Differences (SSD)*  $\mathcal{D}^{\text{SSD}}[\mathcal{T}[y], \mathcal{R}] = \frac{1}{2} \int_{\Omega} [\mathcal{T}(y(x)) - \mathcal{R}(x)]^2 dx,$
- correlation
- Mutual Information (multi-modal images)
- Normalized Gradient Fields







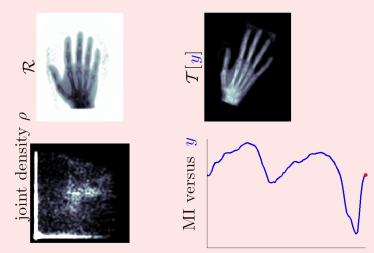




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## **Mutual Information**





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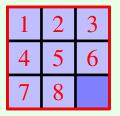
# Regularization

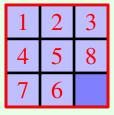
# $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min$



FIELDS

## Transformation y







- Registration is severely ill-posed
- Restrictions onto the transformation y needed
- Goal: implicit physical restrictions



### Implicit versus Explicit Regularization ...

Registration is ill-posed ~ requires regularization

- Parametric Registration
  - restriction to (low-dimensional) space (rigid, affine linear, spline,...)
  - regularized by properties of the space (implicit)
  - not physical or model based
- Non-parametric Registration
  - regularization by adding penalty or likelihood (explicit)
  - allows for a physical model
  - ► ~→ y is no longer parameterizable



#### ... implicit versus explicit regularization

registration is ill-posed  $\rightsquigarrow$  requires regularization

parametric registration

parametric registration

 $\mathcal{D}[R,T;y] \stackrel{y}{=} \min$  s.t.  $y \in \mathcal{Q} = \{x + \sum w_j q_j, w \in \mathbb{R}^m\}$ 

non-parametric registration

non-parametric registration

$$\mathcal{D}[R,T;y] + \alpha \mathcal{S}[y-y_{\text{reg}}] \stackrel{y}{=} \min$$



#### **References for Well-Posedness**

- M. Droske and M. Rumpf.
   A variational approach to non-rigid morphological registration. SIAM Appl. Math., 64(2):668–687, 2004.
- B. Fischer and J. Modersitzki.

A unified approach to fast image registration and a new curvature based registration technique.

Linear Algebra and its Applications, 380:107–124, 2004.

J. Weickert and C. Schnörr.

A theoretical framework for convex regularizers in PDE-based computation of image motion.

Int. J. Computer Vision, 45(3):245-264, 2001.



## Regularizer S

y(x) = x + u(x), displacement  $u : \mathbb{R}^d \to \mathbb{R}^d$ 

- "elastic registration"  $S^{elas}[u] = elastic potential of u$
- "fluid registration"  $S^{\text{fluid}}[u] = \text{elastic potential of } \partial_t u$
- "diffusion registration"  $S^{\text{diff}}[u] = \frac{1}{2} \sum_{\ell=1}^{d} \int_{\Omega} \|\nabla u_{\ell}\|_{\mathbb{R}^2}^2 dx$
- "curvature registration"  $S^{\text{curv}}[u] = \frac{1}{2} \sum_{\ell=1}^{d} \int_{\Omega} (\Delta u_{\ell})^2 dx$



**>** . . .

## **Elastic Registration**

Transformation/displacement, y(x) = x + u(x)

$$S^{\text{elas}}[u] = \text{elastic potential of } u$$
$$= \int_{\Omega} \frac{\lambda + \mu}{2} \|\nabla \cdot u\|^2 + \frac{\mu}{2} \sum_{i=1}^d \|\nabla u_i\|^2 dx$$

#### image painted on a rubber sheet



C. Broit. *Optimal Registration of Deformed Images.* PhD thesis, University of Pensylvania, 1981.

Bajcsy & Kovačič 1986, Christensen 1994, Bro-Nielsen 1996, Gee et al. 1997, Fischer & M. 1999, Rumpf et al. 2002, ...



### Fluid Registration

Transformation/displacement, y(x, t) = x + u(x, t)

 $S^{\text{fluid}}[u] = \text{elastic potential of } \partial_t u$ 

image painted on honey

#### GE. Christensen.

Deformable Shape Models for Anatomy. PhD thesis, Sever Institute of Technology, Washington University,

1994.

Bro-Nielsen 1996, Henn & Witsch 2002, ...



## **Diffusion Registration**

Transformation/displacement, y(x) = x + u(x)

 $\mathcal{S}^{\text{diff}}[u] = \text{oszillations of } u$  $= \frac{1}{2} \sum_{\ell=1}^{d} \int_{\Omega} \|\nabla u_{\ell}\|_{\mathbb{R}^{2}}^{2} dx$ 

heat equation

2003....

 B. Fischer and J. Modersitzki. Fast diffusion registration. AMS Contemporary Mathematics, Inverse Problems, Image Analysis, and Medical Imaging, 313:117–129, 2002.
 Horn & Schunck 1981, Thirion 1996, Droske, Rumpf & Schaller



## **Curvature Registration**

Transformation/displacement, y(x) = x + u(x)

$$\mathcal{S}^{\text{curv}}[\boldsymbol{u}] = \text{oscillations of } \boldsymbol{u} \\ = \frac{1}{2} \sum_{\ell=1}^{d} \int_{\Omega} \|\Delta \boldsymbol{u}_{\ell}\|_{\mathbb{R}^{2}}^{2} dx$$

#### bi-harmonic operator

B. Fischer and J. Modersitzki.
 Curvature based image registration.
 J. of Mathematical Imaging and Vision, 18(1):81–85, 2003.

#### Stefan Henn.

A multigrid method for a fourth-order diffusion equation with application to image processing. *SIAM J. Sci. Comput.*, 2005.



#### Registration of a

#### Curvature Registration

- ▶ Goal: do not penalize affine linear transformations  $S[Cx + b] \stackrel{!}{=} 0$  for all  $C \in \mathbb{R}^{d \times d}$  and  $b \in \mathbb{R}^{d}$
- But:  $S^{\text{diff},\text{elas},\text{fluid},\dots}[Cx+b] \neq 0$  !
- ► Idea:  $S^{\text{curv}}[y] = \sum_{\ell} \int_{\Omega} (\Delta y_{\ell})^2 dx \Rightarrow S^{\text{curv}}[Cx + b] = 0$





## Summary Regularization

- ► Registration is ill-posed ~> requires regularization
- Regularizer controls reasonability of transformation
- Application conform regularization
- Enabling physical models (linear elasticity, fluid flow, ...)
- ► ~ high dimensional optimization problems



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# Numerical Methods for Image Registration





### Optimize \leftrightarrow Discretize

**Image Registration** 

$$\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min, \quad y_{\text{reg}}(x) = x$$

Numerical Approaches:

- ► Optimize → Discretize
- ► Discretize → Optimize



relatively large problems:
 2.000.000 – 500.000.000 unknowns



## Optimize → Discretize: ELE

Image Registration

$$\mathcal{J}[y] = \mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \min, \quad y_{\text{reg}}(x) = x$$

- Euler-Lagrange eqs. (ELE) give necessary condition:  $\mathcal{D}_y + \alpha \mathcal{S}_y = 0 \iff f[y] + \alpha \mathcal{A}y = 0$ system of non-linear partial differential eqs. (PDE)
- outer forces *f*, drive registration
- inner forces Ay, tissue properties
- ► ELE ~→ PDE: balance of forces



## Optimize → Discretize: Summary

Continuous Euler-Lagrange equations

 $f[y] + \alpha \mathcal{A} y = 0, \quad f[y^k] + \alpha \mathcal{A} y^{k+1} = 0, \quad f[y] + \alpha \mathcal{A} y = y_t$ 

all difficulties dumped into right hand side *f* spatial discretization straightforward
 efficient solvers for linear systems
 small controllable steps (~> movies)

- moderate assumptions on f and A (smoothness)
- -
- no optimization problem behind
- non-linearity only via f
- small steps

**Software:** http://www.math.uni-luebeck.de/SAFIR



#### Discretize → Optimize: Summary

Discretization  $\rightsquigarrow$  finite dimensional problem:  $y^h \approx y(x^h)$ 

$$D(y^h) + \alpha S(y^h) \xrightarrow{y^h} \min, \quad y^h \in \mathbb{R}^n, \qquad h \longrightarrow 0$$

efficient optimization schemes (Newton-type) linear systems of type  $H \delta_y = -rhs$ ,

 $H = M + \alpha B^{\top} B$ ,  $M \approx D_{yy}$ ,  $\text{rhs} = D_y + \alpha (B^{\top} B) y^h$ 



efficient multigrid solver for linear systems large steps



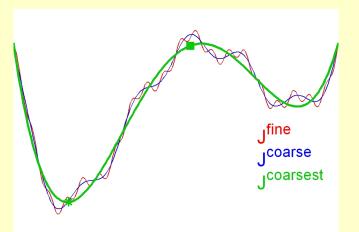
discretization not straightforward (multigrid)

all parts have to be differentiable (data model)



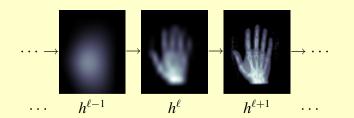
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### **Multilevel**





#### **Multilevel**



for  $\ell = 1 : \ell_{max}$  do

transfer images to level  $\ell$ approximately solve problem for yprolongating y to finer level  $\rightsquigarrow$  perfect starting point end for



#### Advantages of Multilevel Strategy

#### Regularization

Focusses on essential minima

Creates extraordinary starting value

Reduces computation time

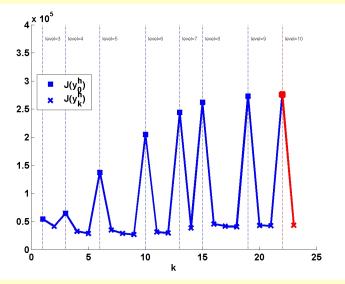








#### Example: Multilevel Iteration History





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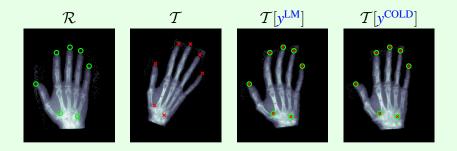
# **Constrained Image Registration**





## Example: COLD

#### Combining Landmarks and Distance Measures



Patent AZ 10253 784.4; Fischer & M., 2003



## Adding Constraints

Constrained Image Registration

$$\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] + \beta \int_{\Omega} \psi \left( \mathcal{C}^{\text{soft}}[y] \right) dx \xrightarrow{y} \min$$

subject to  $\mathcal{C}^{hard}[y](x) = 0$  for all  $x \in \Omega_{\mathcal{C}}$ 

#### Example: landmarks/volume preservation

- soft constraints (penalty)
- hard constraints
- both constraints



# **Rigidity Constraints**





# Soft Rigidity Constraints

FAIR with Soft Rigidity

$$\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] + \beta \mathcal{C}[y] \xrightarrow{y} \min$$

C soft constraints / penalty:

$$C[\mathbf{y}] = \frac{1}{2} \| \underbrace{\mathbf{r}^{\text{linear}}(\mathbf{y})}_{\text{linear}} \|_{\mathcal{Q}}^{2} + \frac{1}{2} \| \underbrace{\mathbf{r}^{\text{orth}}(\mathbf{y})}_{\mathcal{Q}} \|_{\mathcal{Q}}^{2} + \frac{1}{2} \| \underbrace{\mathbf{r}^{\text{det}}(\mathbf{y})}_{\mathcal{Q}} \|_{\mathcal{Q}}^{2}$$
orientation
$$r^{\text{linear}}(\mathbf{y}) = [\partial_{1,1}y_{1}, \dots, \partial_{d,d}y_{1}, \partial_{1,1}y_{2}, \dots]$$

$$r^{\text{orth}}(\mathbf{y}) = \nabla \mathbf{y}^{\top} \nabla \mathbf{y} - I_{d}$$

$$r^{\text{det}}(\mathbf{y}) = \det(\nabla \mathbf{y}) - 1$$

$$\mathbf{y} \text{ rigid } \iff [ \mathbf{r}^{\text{linear}} = 0 \land \mathbf{r}^{\text{orth}} = 0 \land \mathbf{r}^{\text{det}} = 0 ]$$



## The Weight $\mathcal{Q}$

- only locally rigid
- use weight function Q
- regions to be kept rigid move with y



$$||f||_{\mathcal{Q}}^2 = \int_{\Omega} f(x) \ \mathcal{Q}(\mathbf{y}(x))^2 \ dx$$



#### Numerical Scheme

- $\blacktriangleright \quad Q(y^h) \approx \mathcal{Q}(y(x^h))$
- ►  $r(y^h) = [\operatorname{diag}(Q(y^h)) r_1(y^h), \dots, \operatorname{diag}(Q(y^h)) r_{\operatorname{end}}(y^h)]$
- $C(y^h) = \frac{1}{2}r(y^h)^\top r(y^h)$
- $C_y(y^h) =$  lengthy formula
- $D(y^h) + \alpha S(y^h) + \beta C(y^h) \xrightarrow{y^h} \min$
- Optimizer: Gauß-Newton type approach,

$$H \approx "\nabla^2 \mathcal{D}" + \alpha B^\top B + \beta r_y^\top r_y$$

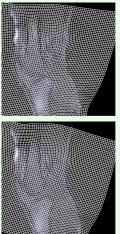


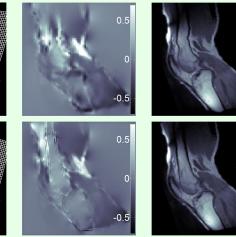


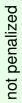
# Example: Knee

 $\det(\nabla y) - 1$ 







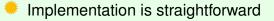


penalized





#### Results are OK





Constraints are not fulfilled



How to pick penalty  $(\beta, \psi)$ ?



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# Hard Rigidity Constraints

FAIR with Hard Rigidity

 $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{\text{reg}}] \xrightarrow{y} \text{min subject to } y \text{ rigid on } \mathcal{Q}$ 

Eulerian  $\rightarrow$  Lagrangian

computations of  $\mathcal{D}$  and  $\mathcal{S}$ involve det $(\nabla y)$ 

rigidity in  $\mathcal{T}$  domain  $\rightsquigarrow$  linear constraints

$$y(x) = D_k x + t_k, \quad k = 1 : \#$$
segments





## Lagrangian Model of Rigidity (2D)

rigid on segment i

$$\mathbf{y}(\mathbf{x}) = \mathbf{Q}(\mathbf{x})\mathbf{w}^{i} = \begin{pmatrix} \cos w_{1}^{i} & -\sin w_{1}^{i} \\ \sin w_{1}^{i} & \cos w_{1}^{i} \end{pmatrix} \begin{pmatrix} x_{1} \\ x_{2} \end{pmatrix} + \begin{pmatrix} w_{2}^{i} \\ w_{3}^{i} \end{pmatrix}$$

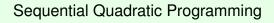
• 
$$w = (w^1, \dots, w^m), \quad \mathcal{C} = (\mathcal{C}^1, \dots, \mathcal{C}^m), \quad m = \#$$
segments

$$\mathcal{C}^{i}[y,w] = y(x) - Q(x)w^{i}, \qquad i = 1, \dots, m$$

Lagrangian:

$$L(\mathbf{y}, \mathbf{w}, p) = \mathcal{D}[\mathbf{y}] + \alpha \mathcal{S}[\mathbf{y}] + p^{\top} \mathcal{C}[\mathbf{y}, \mathbf{w}]$$

Numerical Scheme:

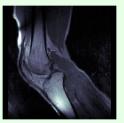


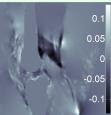




## Rigidity as a Hard Constraint

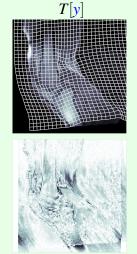
R













## Summary of Hard Rigidity Constraints

#### Results are OK

#### Implementation is interesting



Constraints are fulfilled



No additional Parameters



FIELDS



# Summary

FIELDS

#### Summary

- Introduction to image registration: important, challenging, interdisciplinary
- ► General framework based on a variational approach:  $\mathcal{D}[\mathcal{T}[y], \mathcal{R}] + \alpha \mathcal{S}[y - y_{reg}] \xrightarrow{y} min$
- Discussion of various building blocks:
  - ▶ image model T[y]
  - distance measures D
  - regularizer S
- ► Numerical methods: multilevel, optimize ↔ discretize
- ► Constraints *C*:

landmarks, local rigidity, intensity correction, ...

#### Solutions and Algorithms For Image Registration

