

# Lecture 11: Medical Image Registration Mutual Information, Elastic Regularisation, Landmarks

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## Medical Image Registration (1)

# Medical Image Registration

## What is the Medical Image Registration Problem?

**Given:** Two medical images (e.g. CT or MRT scans).

**Wanted:** Realistic deformation field that allows to compare/combine information.

## Example for a Typical Medical Image Registration Task

- ◆ Hand Matching with CT images  
(*Martin-Fernández/Munoz-Moreno/Martin-Fernández/Alberola-Lopez 2003*)



(a) Left: Reference Image. (b) Middle: Template image. (c) Right: Matched template image.

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### Fields of Application

- ◆ Radiotherapy (e.g. tumor therapy, improved diagnosis)
- ◆ Pathology/Histology (e.g. slice registration)

### Challenges in Medical Image Data

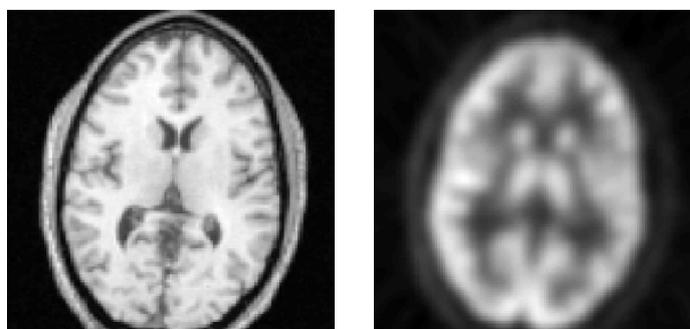
- ◆ Large displacements
- ◆ Deformations constrained to elastic motion
- ◆ Intensity differences
- ◆ Images may be created by different image acquisition methods
- ◆ Heavy noise (e.g. speckle noise in ultrasound images)
- ◆ Manually set displacements (landmarks) have to be respected

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### Mutual Information

#### Combining Information from Different Image Acquisition Methods

- ◆ Registration of MRT and PET Scans



Images of a human brain. **(a) Left:** MRT scan. **(b) Right:** PET scan. *Author:* W.-H. Liao (2003).

- ◆ Problem: How to register images that visualise completely different properties of the same object. Is there some common information that still allows registration?

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## Mutual Information (2)

MI  
A

### Mutual Information

- ◆ *Idea:* Use distance measure in the data term that specifies how independent the grey value distributions in both images are (probabilistic measure).
- ◆ *Known:* Defining  $I_1 = f(x, y, t)$  and  $I_2 = f(x + u, y + v, t + 1)$  we usually consider data terms that comply with the following distance measure

$$d_{\text{OF}}(I_1, I_2) = \Psi\left((I_1 - I_2)^2\right).$$

- ◆ *New:* Mutual information is defined using **probabilistic density functions** (Viola 1995, Viola/Wells 1997)

$$d_{\text{MI}}(I_1, I_2) = \int_{[0,255] \times [0,255]} p^{[I_1, I_2]}(a, b) \log \frac{p^{[I_1, I_2]}(a, b)}{p^{[I_1]}(a) p^{[I_2]}(b)} da db.$$

- $p^{[I_1]}(a)$ ,  $p^{[I_2]}(b)$  are the **marginal** probability density functions of  $I_1$  and  $I_2$
- $p^{[I_1, I_2]}(a, b)$  is the **joint** probability density function of  $I_1$  and  $I_2$

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## Mutual Information (3)

MI  
A

### What Is Actually Measured by Mutual Information?

- ◆ *Explanation:* Mutual information measures the distance between the joint probability density function of  $I_1$  and  $I_2$  in the independent case given by

$$p_{\text{independent}}^{[I_1, I_2]}(a, b) = p^{[I_1]}(a) p^{[I_2]}(b)$$

and the actually measured joint probability density function of  $I_1$  and  $I_2$

$$p_{\text{measured}}^{[I_1, I_2]}(a, b) = p^{[I_1, I_2]}(a, b).$$

Given a grey value  $a$  in  $I_1$  and a grey value  $b$  in  $I_2$ , we can thus measure if the relation between both grey values exceeds the random probability by

$$\frac{p_{\text{measured}}^{[I_1, I_2]}(a, b)}{p_{\text{independent}}^{[I_1, I_2]}(a, b)} = \frac{p^{[I_1, I_2]}(a, b)}{p^{[I_1]}(a) p^{[I_2]}(b)}.$$

Here, larger ratios than 1 indicate a dependency of the grey values  $a$  and  $b$ .

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## Mutual Information (4)

MI  
A

### Properties of Mutual Information

- ◆ *Positivity:* Mutual information is always nonnegative (without proof), i.e. it holds

$$d_{\text{MI}}(I_1, I_2) = \int_{[0,255] \times [0,255]} p^{[I_1, I_2]}(a, b) \log \frac{p^{[I_1, I_2]}(a, b)}{p^{[I_1]}(a) p^{[I_2]}(b)} da db \geq 0 .$$

- ◆ *Symmetry:* Since  $p^{[I_1, I_2]}(a, b) = p^{[I_2, I_1]}(a, b)$  the following holds for  $I_1$  and  $I_2$

$$d_{\text{MI}}(I_1, I_2) = d_{\text{MI}}(I_2, I_1) .$$

- ◆ *Minimum:* If  $I_1$  and  $I_2$  are independent,  $p^{[I_1, I_2]}(a, b)$  can be factorised as follows

$$p^{[I_1, I_2]}(a, b) = p^{[I_1]}(a) p^{[I_2]}(b) \rightarrow \log \frac{p^{[I_1, I_2]}(a, b)}{p^{[I_1]}(a) p^{[I_2]}(b)} = \log 1 = 0$$

Then the mutual information attains its minimum given by  $d_{\text{MI}}(I_1, I_2) = 0$ .

## Mutual Information (5)

MI  
A

### Using Mutual Information in Practice

- ◆ *Attention:* Since we are actually interested in maximising the mutual information (the dependency of both distributions), we have to consider the **negative version**

$$\overline{d_{\text{MI}}}(I_1, I_2) = - d_{\text{MI}}(I_1, I_2) = - \int_{[0,255] \times [0,255]} p^{[I_1, I_2]}(a, b) \log \frac{p^{[I_1, I_2]}(a, b)}{p^{[I_1]}(a) p^{[I_2]}(b)} da db .$$

- ◆ *Computation of the Probabilistic Densities:* The probabilistic densities are computed via the **Parzen estimators**

$$p^{[I_1]}(a) = K_\sigma * \frac{1}{|\Omega|} \int_{\Omega} \mathbf{1}_{(I_1(x,y)=a)} dx dy ,$$

$$p^{[I_1, I_2]}(a, b) = K_\sigma * \frac{1}{|\Omega|} \int_{\Omega} \mathbf{1}_{((I_1(x,y)=a) \wedge (I_2(x+u, y+v)=b))} dx dy ,$$

which can be considered as **Gaussian smoothed normalised histograms**.

## Mutual Information (6)

MI  
A

### Mutual Information for Block Matching Techniques

- ◆ *Idea:* Use mutual information as matching cost for block matching
- ◆ *Example:* Using a 2-D search space  $\mathcal{S}_d$  with displacements  $u, v \in \{-d, \dots, d\}$  yields the following block matching approach

$$\mathbf{u}_{i,j} = \operatorname{argmin}_{u,v \in \mathcal{S}_d} \overline{d_{\text{MI}}} \left( \mathcal{N}_m(f_{i,j}), \mathcal{N}_m(g_{i+u,j+v}) \right).$$

where  $\mathcal{N}_m(f, i, j)$  is a patch of size  $(2m+1) \times (2m+1)$  around the pixel  $f_{i,j}$

- ◆ *Attention:* The marginal and joint probability density functions are only computed for the **local patches**  $\mathcal{N}_m(f_{i,j})$  and  $\mathcal{N}_m(g_{i+u,j+v})$  (and not for the whole image).
- ◆ *Realisation:* The required cost map is derived in three steps
  - compute the normalised histograms for the separate and the joint case
  - smooth histograms with Gaussian to obtain probability density functions
  - compute cost map using the negative mutual information, i.e.  $\overline{d_{\text{MI}}}$

## Mutual Information (7)

MI  
A

### Mutual Information for Variational Methods

- ◆ *Idea:* Use mutual information as data term for variational methods
- ◆ *Example:* Replacing the data term of the method of Horn and Schunck with the expression for mutual information yields the functional (Hermosillo 2001)

$$E(\mathbf{w}) = - \int_{[0,255] \times [0,255]} p^{[I_1, I_2^{\mathbf{w}}]}(a, b) \log \frac{p^{[I_1, I_2^{\mathbf{w}}]}(a, b)}{p^{[I_1]}(a) p^{[I_2^{\mathbf{w}}]}(b)} da db + \alpha \int_{\Omega} (|\nabla u|^2 + |\nabla v|^2) dx dy$$

with  $p^{[I_1]}(a)$ ,  $p^{[I_2^{\mathbf{w}}]}(b)$  and  $p^{[I_1, I_2^{\mathbf{w}}]}(a, b)$  defined as before.

- ◆ *Remark:* In order to make explicit that we consider the motion compensated second frame  $f(\mathbf{u} + \mathbf{w})$  in our data term, we replaced  $I_2$  by  $I_2^{\mathbf{w}}$ .
- ◆ *Attention:* This energy functional is highly non-convex and its optimisation is non-trivial. However, the Euler-Lagrange equations can be stated explicitly.

## Mutual Information (8)

### Mutual Information for Variational Methods

- ◆ *Minimisation:* The associated Euler-Lagrange equations are given by the PDEs

$$0 = -\left(\frac{\partial}{\partial b} L^{[I_1, I_2^w]}\right)(f(\mathbf{x}), f(\mathbf{x} + \mathbf{w})) f_x(\mathbf{x} + \mathbf{w}) - \alpha \Delta u$$

$$0 = -\left(\frac{\partial}{\partial b} L^{[I_1, I_2^w]}\right)(f(\mathbf{x}), f(\mathbf{x} + \mathbf{w})) f_y(\mathbf{x} + \mathbf{w}) - \alpha \Delta v$$

with (reflecting) Neumann boundary conditions  $\mathbf{n}^\top \nabla u = 0$  and  $\mathbf{n}^\top \nabla v = 0$  where the non-linear operator  $L^{[I_1, I_2^w]}$  is defined as follows

$$L^{[I_1, I_2^w]}(a, b) = 1 + p^{[I_1, I_2^w]}(a, b) \log \frac{p^{[I_1, I_2^w]}(a, b)}{p^{[I_1]}(a) p^{[I_2^w]}(b)}.$$

- ◆ *Attention:* Since these equations are difficult to linearise, one often performs a steepest descent/gradient descent approach. While such an approach works in most cases, it easily gets trapped in the next local minimum.

## Mutual Information (9)

### The Steepest Descent/Gradient Descent Approach

- ◆ *Known:* The first variation of a typical 2-D energy functional is given by

$$\mathbf{v}(u, v) = \begin{pmatrix} v_1(u, v) \\ v_2(u, v) \end{pmatrix} = \begin{pmatrix} F_u - \frac{\partial}{\partial x} F_{u_x} - \frac{\partial}{\partial y} F_{u_y} \\ F_v - \frac{\partial}{\partial x} F_{v_x} - \frac{\partial}{\partial y} F_{v_y} \end{pmatrix}$$

known from the Euler-Lagrange equations, where we seek  $\mathbf{v} = 0$ .

- ◆ *New:* In order to minimize an energy functional it makes sense to walk from a given solution in the direction of the steepest descent

$$u^{k+1} = u^k - \gamma v_1(u^k, v^k)$$

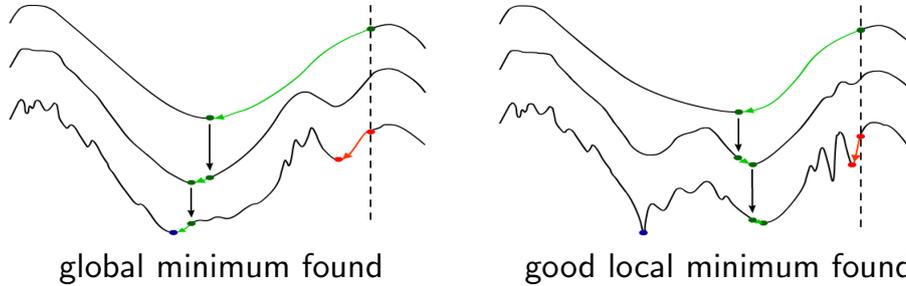
$$v^{k+1} = v^k - \gamma v_2(u^k, v^k)$$

where  $\gamma$  is the speed factor (step size) and  $-\mathbf{v} = (-v_1, -v_2)^\top$  is the direction of the steepest descent (since  $\mathbf{v}$  is the direction of the steepest ascent).

- ◆ *Remark:* The Euler-Lagrange equations represent the steady-state of the gradient descent approach, since nothing changes for  $\mathbf{v} = 0$ .

The Steepest Descent/Gradient Descent Approach

- ◆ *Visualisation:* Since a gradient descent (red) gets trapped in the next local minima, one often uses a coarse-to-fine initialisation strategy (green)



- ◆ *Application to Our Problem:* In the case of the energy functional for mutual information the gradient descent equations are given by

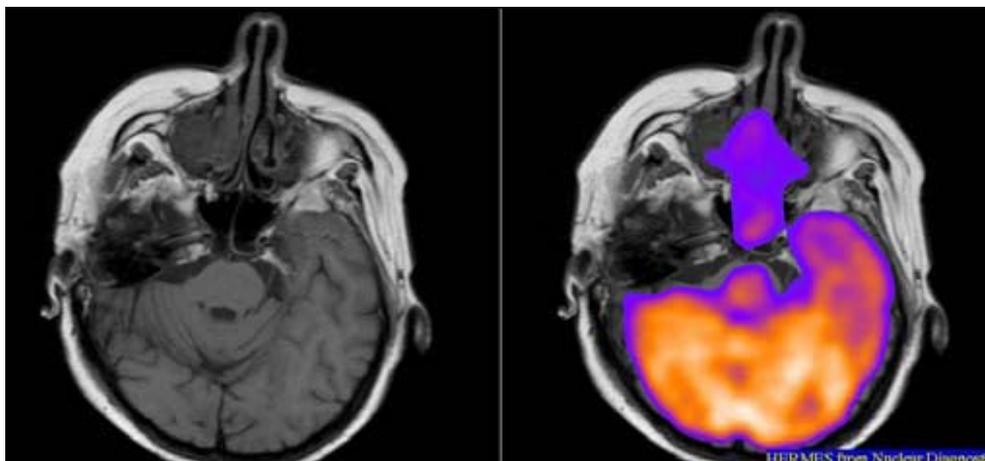
$$u^{k+1} = u^k - \gamma \left( - \left( \frac{\partial}{\partial b} L [I_1, I_2^{w^k}] \right) (f(\mathbf{x}), f(\mathbf{x} + \mathbf{w}^k)) f_x(\mathbf{x} + \mathbf{w}^k) - \alpha \Delta u^k \right),$$

$$v^{k+1} = v^k - \gamma \left( - \left( \frac{\partial}{\partial b} L [I_1, I_2^{w^k}] \right) (f(\mathbf{x}), f(\mathbf{x} + \mathbf{w}^k)) f_y(\mathbf{x} + \mathbf{w}^k) - \alpha \Delta v^k \right).$$

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Results for Mutual Information

- ◆ Mapping of Magneto Resonance Tomography (MRT) and Positron Emission Tomography (PET) Images via Mutual Information  
(Willdendrup/Svarer 2004)

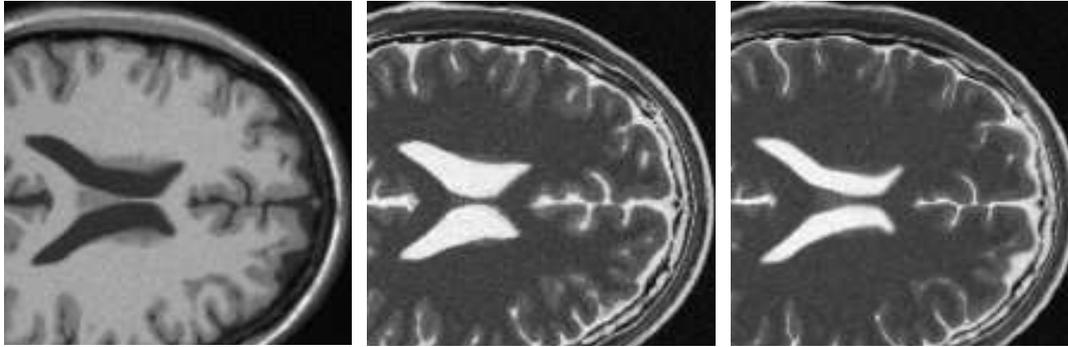


(a) Left: MRT image. (b) Right: Registered PET image blended over MRT image.

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## Results for Mutual Information

- ◆ Mapping of T1/T2 MRT Brain Images via Mutual Information  
(Heidmann/Mahnke/Potts/Modersitzki/Fischer 2004)



Mapping of T1/T2 brain images. **(a) Left:** T1-MRT image (reference) **(b) Center:** T2-MRT image (template). **(c) Right:** Registered T2-MRT image (matched template).

Information from different image acquisition methods can be combined!

## Normalised Gradient Fields (1)

## Normalised Gradient Fields

## Can We Find A Simpler Useful Distance Measure Than Mutual Information?

- ◆ *Idea:* Use normalised gradient fields to construct data term
- ◆ *Example:* Given an image  $f(\mathbf{x})$  the corresponding normalised gradient field is given by the directional vector

$$\mathbf{n}(f, \mathbf{x}) = \begin{cases} \frac{\nabla f(\mathbf{x})}{|\nabla f(\mathbf{x})|} & \text{if } \nabla f(\mathbf{x}) \neq 0, \\ 0 & \text{else} \end{cases}$$

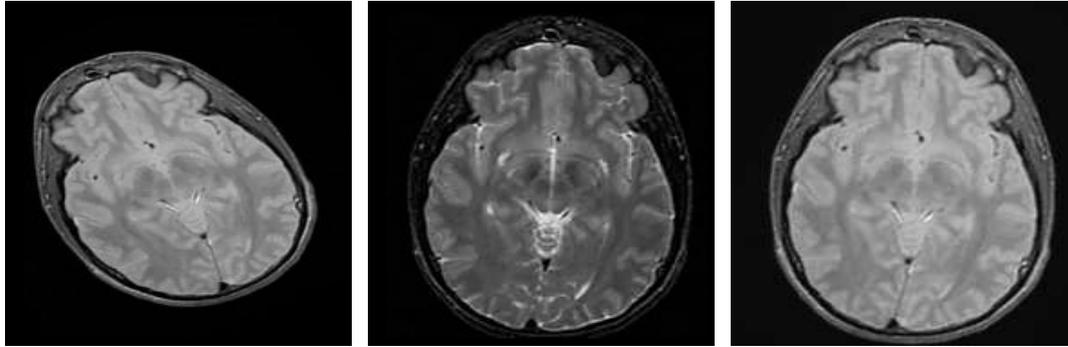
A suitable distance that favours small angles between the gradient field of both images is thus given by the negative squared scalar product

(Haber/Modersitzki 2006)

$$d_{\text{NGF}}(I_1, I_2) = -\left(\mathbf{n}(I_1, \mathbf{x})^\top \mathbf{n}(I_2, \mathbf{x})\right)^2.$$

Results for Normalised Gradient Fields

- ◆ Mapping of T1/T2 MRT Brain Images via Normalised Gradient Fields (Haber/Modersitzki 2004)



Mapping of T1/T2 brain images. (a) Left: T1-MRT image (reference) (b) Center: T2-MRT image (template). (c) Right: Registered T2-MRT image (matched template).

Normalised gradient fields are a simple but good alternative

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Elastic Regularisation

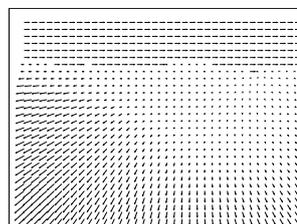
Can We Model the Elastic Deformations of a Tissue in a Suitable Way?

- ◆ Idea: Use smoothness term that can be derived from elasticity theory
- ◆ Example: If the tissue is elastic, no new tissue should appear or vanish during the registration. This can be encouraged via the penaliser

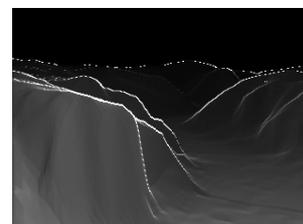
$$\left(\operatorname{div} \begin{pmatrix} u \\ v \end{pmatrix}\right)^2 = \left(\nabla \cdot \mathbf{u}\right)^2 = (u_x + v_y)^2$$

which models a **divergence-free flow field** (containing neither sinks nor sources).

- ◆ Visualisation: By the example of the Yosemite sequence we can see that large divergence expressions occur at locations where information appears or vanishes (e.g. at discontinuities).



Flow field



Magnitude of divergence

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## Elastic Regularisation (2)

MI  
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### Elastic Regularisation in Variational Approaches

- ◆ *Idea:* Assume both a smooth and elastically deformed flow field
- ◆ *Example:* The energy functional for the Horn and Schunck method with elastic deformation term is given by

$$E(u, v) = \int_{\Omega} \underbrace{(f_x u + f_y v + f_t)^2}_{\text{data term}} + \underbrace{\mu (|\nabla u|^2 + |\nabla v|^2)}_{\text{smoothness term}} + (\lambda + \mu) \underbrace{((u_x + v_y)^2)}_{\text{elasticity term}} dx dy,$$

where the weights  $\mu$  and  $\lambda$  are the so-called **Lamé constants** that are related to parameters that describe the elasticity of the deformed material.

- ◆ *Minimisation:* The corresponding Euler-Lagrange equations are given by the following **linear** equations which are known as **Navier-Lamé (NLE)** equations

$$\begin{aligned} 0 &= J_{11} u + J_{12} v + J_{13} - \mu \Delta u - (\lambda + \mu)(u_{xx} + v_{xy}), \\ 0 &= J_{12} u + J_{22} v + J_{23} - \mu \Delta v - (\lambda + \mu)(u_{xy} + v_{yy}) \end{aligned}$$

with associated **boundary conditions** that read

$$\mathbf{n}^T \begin{pmatrix} (\lambda + 2\mu)u_x + (\lambda + \mu)v_y \\ \mu u_y \end{pmatrix} = 0 \quad \text{and} \quad \mathbf{n}^T \begin{pmatrix} \mu v_x \\ (\lambda + 2\mu)v_y + (\lambda + \mu)u_x \end{pmatrix} = 0.$$

## Elastic Regularisation (3)

MI  
A

### Elastic Regularisation in Variational Approaches

- ◆ *Generic Form:* These Euler-Lagrange equations can be rewritten in the form

$$\begin{aligned} 0 &= J_{11} u + J_{12} v + J_{13} - \operatorname{div}(D_1 \nabla u) - \operatorname{div}(D_2 \nabla v), \\ 0 &= J_{12} u + J_{22} v + J_{23} - \operatorname{div}(D_2 \nabla u) - \operatorname{div}(D_3 \nabla v), \end{aligned}$$

with (reflecting) Neumann boundary conditions  $\mathbf{n}^T \nabla u = 0$  and  $\mathbf{n}^T \nabla v = 0$ .

- ◆ *New:* So far we have obtained only Euler-Lagrange equations that contain either divergence expressions for  $u$  or for  $v$ . Now expressions for **both unknowns** occur.
- ◆ *Notation:* The **anisotropic** diffusion tensors  $D_1$ ,  $D_2$  and  $D_3$  are given by

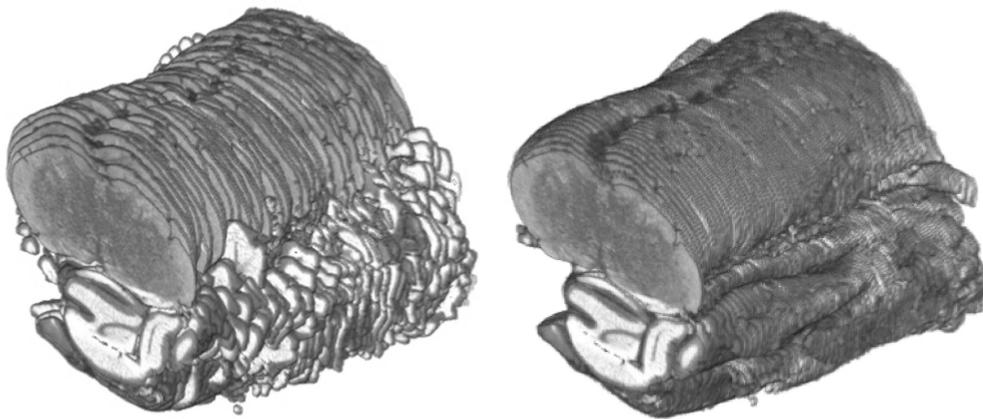
$$D_1 = \begin{pmatrix} (\lambda + 2\mu) & 0 \\ 0 & \mu \end{pmatrix}, \quad D_2 = \begin{pmatrix} 0 & \frac{(\lambda + \mu)}{2} \\ \frac{(\lambda + \mu)}{2} & 0 \end{pmatrix}, \quad D_3 = \begin{pmatrix} \mu & 0 \\ 0 & (\lambda + 2\mu) \end{pmatrix}.$$

- ◆ *Attention:* The Euler-Lagrange equations for elastic regularisation are **linear**, since the associated diffusion tensors do not depend on the unknowns  $u$  and  $v$ .

## Elastic Regularisation (4)

### Results for Elastic Regularisation (Variational Approach with Warping)

- ◆ Registering Histologic Slices of a Rat Brain Acquired by a Flat Bed Scanner  
(Wirtz/Fischer/Modersitzki/Schmitt 2004)



(a) Left: Original slices (unregistered). (b) Right: Registered slices.

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## Elastic Regularisation (5)

### Results for Elastic Regularisation (Non-Variational Approach)

- ◆ Hand Matching with CT Images  
(Martin-Fernández/Munoz-Moreno/Martin-Fernández/Alberola-Lopez 2003)



(a) Left: Reference Image. (b) Middle: Template image. (c) Right: Matched template image.

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## Curvature-Based Regularisation

### Are There Any Alternatives for the Smoothness Term?

- ◆ *Idea:* Assume locally affine transformations (instead of locally constant ones)
- ◆ *Example:* Instead of penalising the first derivatives and thus assuming the displacement field to be locally constant, one penalises the second derivatives

$$(\Delta u)^2 + (\Delta v)^2 = u_{xx}^2 + 2u_{xx}u_{yy} + u_{yy}^2 + v_{xx}^2 + 2v_{xx}u_{yy} + v_{yy}^2 .$$

- ◆ *Explanation:* Affine transformations are a **subset** (for  $s_1 = s_2 = 0$ ) of the following functions that are all mapped to zero by the curvature regulariser

$$\begin{aligned} u(x, y) &= s_1 xy + ax + by + c , \\ v(x, y) &= s_2 xy + dx + ey + f . \end{aligned}$$

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### Curvature-Based Regularisation in Variational Approaches

- ◆ *Example:* The energy functional for the Horn and Schunck method with curvature-based regularisation is given by

$$E(u, v) = \int_{\Omega} \underbrace{(f_x u + f_y v + f_t)^2}_{\text{data term}} + \alpha \underbrace{((\Delta u)^2 + (\Delta v)^2)}_{\text{smoothness term}} dx dy .$$

- ◆ *Minimisation:* The corresponding Euler-Lagrange equations are given by the following **linear** equations

$$\begin{aligned} 0 &= J_{11} u + J_{12} v + J_{13} + \alpha \overbrace{(u_{xxxx} + 2u_{xxyy} + u_{yyyy})}^{\Delta^2 u} , \\ 0 &= J_{12} u + J_{22} v + J_{23} + \alpha \overbrace{(v_{xxxx} + 2v_{xxyy} + v_{yyyy})}^{\Delta^2 v} , \end{aligned}$$

with Neumann (reflecting) boundary conditions

$$\begin{aligned} \mathbf{n}^\top \nabla \Delta u &= 0 , & \mathbf{n}^\top \begin{pmatrix} \Delta u \\ 0 \end{pmatrix} &= 0 , & \mathbf{n}^\top \begin{pmatrix} 0 \\ \Delta u \end{pmatrix} &= 0 , \\ \mathbf{n}^\top \nabla \Delta v &= 0 , & \mathbf{n}^\top \begin{pmatrix} \Delta v \\ 0 \end{pmatrix} &= 0 , & \mathbf{n}^\top \begin{pmatrix} 0 \\ \Delta v \end{pmatrix} &= 0 . \end{aligned}$$

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### Curvature-Based Regularisation in Variational Approaches

- ◆ *New:* Since the smoothness term contains derivatives of order 2, we used an extended version of our general formulation of the Euler-Lagrange equations

$$0 \stackrel{!}{=} F_u - \frac{\partial}{\partial x} F_{u_x} - \frac{\partial}{\partial y} F_{u_y} + \frac{\partial}{\partial x x} F_{u_{xx}} + \frac{\partial}{\partial x y} F_{u_{xy}} + \frac{\partial}{\partial y x} F_{u_{yx}} + \frac{\partial}{\partial y y} F_{u_{yy}},$$

$$0 \stackrel{!}{=} F_v - \frac{\partial}{\partial x} F_{v_x} - \frac{\partial}{\partial y} F_{v_y} + \frac{\partial}{\partial x x} F_{v_{xx}} + \frac{\partial}{\partial x y} F_{v_{xy}} + \frac{\partial}{\partial y x} F_{v_{yx}} + \frac{\partial}{\partial y y} F_{v_{yy}}.$$

with associated boundary conditions that read

$$\mathbf{n}^\top \begin{pmatrix} F_{u_x} - \partial_x F_{u_{xx}} - \partial_y F_{u_{xy}} \\ F_{u_y} - \partial_x F_{u_{yx}} - \partial_y F_{u_{yy}} \end{pmatrix} = 0, \quad \mathbf{n}^\top \begin{pmatrix} F_{u_{xx}} \\ F_{u_{xy}} \end{pmatrix} = 0, \quad \mathbf{n}^\top \begin{pmatrix} F_{u_{yx}} \\ F_{u_{yy}} \end{pmatrix} = 0,$$

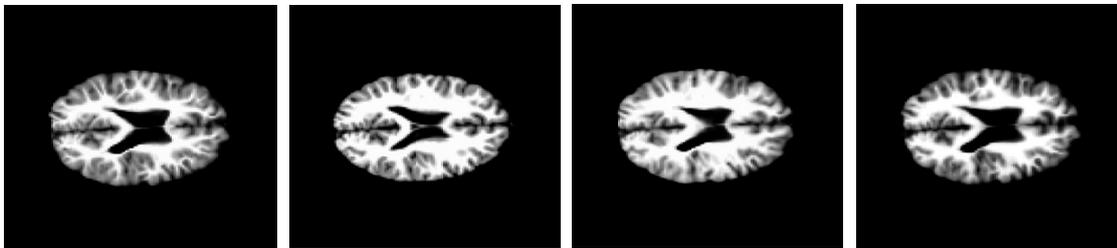
$$\mathbf{n}^\top \begin{pmatrix} F_{v_x} - \partial_x F_{v_{xx}} - \partial_y F_{v_{xy}} \\ F_{v_y} - \partial_x F_{v_{yx}} - \partial_y F_{v_{yy}} \end{pmatrix} = 0, \quad \mathbf{n}^\top \begin{pmatrix} F_{v_{xx}} \\ F_{v_{xy}} \end{pmatrix} = 0, \quad \mathbf{n}^\top \begin{pmatrix} F_{v_{yx}} \\ F_{v_{yy}} \end{pmatrix} = 0.$$

- ◆ *Attention:* The actual implementation of the boundary conditions for higher order regularisers is highly non-trivial.

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### Results for Curvature-Based Regularisation (Variational Approach)

- ◆ Matching of T1 MRT Brain Images with Curvature-Based and Elastic Regulariser (*Agostino/Modersitzki/Maes/Vandermeulen/Fischer/Suetens 2003*)



(a) **Outer Left:** T-1 Reference image. (b) **Middle Left:** T-1 Template image. (c) **Middle Right:** Matched template image curvature-based regularisation. (d) **Outer Right:** Matched template image with elastic regularisation.

**Curvature-based regularisation yields reasonable results, but is not as good elastic regularisation**

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## Landmarks

## How Can We Integrate Correspondences that are Pre-Given by the User?

- ◆ *Idea:* Set displacements at certain points fixed to pre-given values.
- ◆ *Example:* By introducing a binary confidence function  $c(x, y)$  that is defined as

$$c(x, y) = \begin{cases} 1 & \text{if the displacement } u, v \text{ is known at location } (x, y) \\ 0 & \text{else} \end{cases}$$

we can modify the Euler-Lagrange equations of the method of Horn and Schunck in such a way that they become

(Fischer/Modersitzki 2003)

$$0 = (1 - c) (J_{11} u + J_{12} v + J_{13} - \alpha \Delta u) + c (u - u_{\text{given}})$$

$$0 = (1 - c) (J_{12} u + J_{22} v + J_{23} - \alpha \Delta v) + c (v - v_{\text{given}})$$

with (reflecting) Neumann boundary conditions  $\mathbf{n}^T \nabla u = 0$  and  $\mathbf{n}^T \nabla v = 0$ .

## Landmarks

- ◆ *Explanation:* For a better understanding let us consider the following two cases
- ◆ *Case 1:* If  $c(x, y) = 0$ , i.e. the displacement for the point is **not known** we obtain the original Euler-Lagrange equations given by

$$0 = J_{11} u + J_{12} v + J_{13} - \alpha \Delta u ,$$

$$0 = J_{12} u + J_{22} v + J_{23} - \alpha \Delta v .$$

- ◆ *Case 2:* If  $c(x, y) = 1$ , i.e. the displacement for the point is **known** we obtain the following equations

$$\left. \begin{array}{l} 0 = u - u_{\text{given}} \\ 0 = v - v_{\text{given}} \end{array} \right\} \rightarrow v = v_{\text{given}} , \quad u = u_{\text{given}} .$$

- ◆ *Attention:* The given information propagates via the smoothness term also to all other points of the solution (**global effect**).

## Why Can't We Use a Variational Model instead of Modifying the PDEs?

- ◆ *Straightforward Idea:* Let us consider the following approach to integrate landmarks explicitly in the model

$$E(u, v) = \int_{\Omega} (1-c) \left( \underbrace{(f_x u + f_y v + f_t)^2}_{\text{data term}} + \alpha \underbrace{(|\nabla u|^2 + |\nabla v|^2)}_{\text{smoothness term}} \right) + c \underbrace{\left( (u - u_{\text{given}})^2 + (v - v_{\text{given}})^2 \right)}_{\text{landmark term}} dx dy .$$

The corresponding Euler-Lagrange equations are then given by

$$\begin{aligned} 0 &= (1-c) \left( J_{11} u + J_{12} v + J_{13} \right) - \alpha \operatorname{div} \left( (1-c) u \right) + c \left( u - u_{\text{given}} \right) , \\ 0 &= (1-c) \left( J_{12} u + J_{22} v + J_{23} \right) - \alpha \operatorname{div} \left( (1-c) v \right) + c \left( v - v_{\text{given}} \right) . \end{aligned}$$

- ◆ *Attention:* Unfortunately we do not obtain exactly the desired result. Since  $c$  depends on  $x, y$  it remains within the divergence expression of the smoothness term as a **binary diffusivity** (similar to image-driven methods).

## Results for Landmarks (PDE-Based Approach)

- ◆ Hand Matching of CT Images with and without Landmarks  
(Fischer/Modersitzki 2003)



(a) **Outer Left:** Reference image. (b) **Middle Left:** Template image. (c) **Middle Right:** Matched template image without landmarks. (d) **Outer Right:** Matched template image with landmarks.

## Summary

- ◆ The concept of mutual information from information theory allows the registration of images obtained by different image acquisition devices.
- ◆ A simpler alternative is given by normalised gradient fields which are much more convenient to implement.
- ◆ Elastic regularisers from elasticity theory allow to address the problem of non-rigid deformations of tissues in medical images.
- ◆ Curvature based regularisers can estimate of locally affine transformations.
- ◆ Landmarks allow interaction with user by integrating pre-given displacements. This yields semi-automatic registration approaches.

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