# graphics and vision gravis



### Registration – Deformation models

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#### Registration as analysis by synthesis



#### Priors



Define the Gaussian process  $u \sim GP(\mu, k)$ with mean function  $\mu: \Omega \to \mathbb{R}^2$ and covariance function

 $k: \Omega \times \Omega \to \mathbb{R}^{2 \times 2}$ .

Characteristics of deformation fields

Zero mean:

$$\mu(x) = \begin{pmatrix} 0 \\ 0 \end{pmatrix}$$

Squared exponential covariance function (Gaussian kernel)

$$k(x, x') = \begin{pmatrix} s_1 \exp\left(-\frac{\|x - x'\|^2}{\sigma_1^2}\right) & 0\\ 0 & s_2 \exp\left(-\frac{\|x - x'\|^2}{\sigma_2^2}\right) \end{pmatrix}$$



$$s_1 = s_2$$
 small,  $\sigma_1 = \sigma_2$  large



$$s_1=s_2$$
 small,  $\sigma_1=\sigma_2$  small



$$s_1=s_2$$
 large,  $\sigma_1=\sigma_2$  large

#### Why are priors interesting?

 $\theta^* = \arg \max_{\theta} p(\varphi[\theta]) p(I_T | I_R, \varphi[\theta])$ 



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## Intermezzo – The space of samples

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## Gaussian processes - Deeper Insights

#### Scalar-valued Gaussian processes

#### Vector-valued (this course)

• Samples u are deformation fields:

 $u: \mathbb{R}^n \to \mathbb{R}^d$ 

#### Scalar-valued (more common)

• Samples f are real-valued functions  $f: \mathbb{R}^n \to \mathbb{R}$ 





$$u \sim \mu + \sum_{i} \alpha_{i} \sqrt{\lambda_{i}} \phi_{i} = \sum_{i} \beta_{i} k(x_{i}, \cdot)$$
 for some  $\beta$ 

Argument:

- Covariance function k is symmetric and positive definite
- For any finite sample it holds that:
   => the covariance matrix is symmetric
   => rowspace = columnspace = eigenspace

Samples are linear combinations of the "rows" of k

#### Example: Gaussian kernel

- Click to edit Master  $(\underline{telkt stylles})$  Second level
  - - Third level
      - Fourth level





#### Example: Gaussian kernel

$$k(x, x') = \exp\left(-\frac{\|x - x'\|^2}{\sigma^2}\right)$$



 $\sigma = 3$ 



• 
$$k(x, x') = \exp\left(-\left\|x - \frac{x'}{1}\right\|^2\right) + 0.1 \exp\left(-\left\|x - \frac{x'}{0.1}\right\|^2\right)$$





#### Periodic kernels

• Define 
$$u(x) = \begin{pmatrix} \cos(x) \\ \sin(x) \end{pmatrix}$$

• 
$$k(x, x') = \exp(-\|(u(x) - u(x')\|^2) = \exp(-4\sin^2\left(\frac{\|x - x'\|}{\sigma^2}\right))$$





#### Symmetric kernels

- Enforce that f(x) = f(-x)
- k(x, x') = k(-x, x') + k(x, x')





#### Changepoint kernels

• 
$$k(x, x') = s(x)k_1(x, x')s(x') + (1 - s(x))k_2(x, x')(1 - s(x'))$$
  
•  $s(x) = \frac{1}{1 + \exp(-x)}$ 





#### Combining existing functions

k(x, x') = f(x)f(x')



#### Combining existing functions

k(x, x') = f(x)f(x')





#### Combining existing functions

$$k(x, x') = \sum_{i} f_i(x) f_i(x')$$





#### Statistical models



$$\mu(x) = \overline{u}(x) = \frac{1}{n} \sum_{i=1}^{n} u^{i}(x)$$
$$k_{SM}(x, x') = \frac{1}{n-1} \sum_{i=1}^{n} (u^{i}(x) - \overline{u}(x)) (u^{i}(x') - \overline{u}(x'))^{T}$$

Statistical shape models are linear combinations of example deformations  $u^1, \dots u^n$ .

#### Gaussian process regression

- Given: observations  $\{(x_1, y_1), ..., (x_n, y_n)\}$
- Model:  $y_i = f(x_i) + \epsilon$ ,  $f \sim GP(\mu, k)$
- Goal: compute  $p(y_*|x_*, x_1, ..., x_n, y_1, ..., y_n)$



#### Gaussian process regression

• Solution given by posterior process  $GP(\mu_p,k_p)$  with

$$\mu_p(x_*) = K(x_*, X) [K(X, X) + \sigma^2 I]^{-1} y$$

$$k_p(x_*, x_*') = k(x_*, x_*') - K(x_*, X)[K(X, X) + \sigma^2 I]^{-1}K(X, x_*')$$

- The covariance is independent of the value at the training points
  - Structure of posterior GP determined solely by kernel.
- The most likely solution is a linear combination of kernels evaluated at the training points
  - This is known as the **Representer Theorem** in machine learning.
  - Structure of solution determined solely by kernel.

### Illustration: Representer theorem





• Gaussian kernel ( $\sigma = 1$ )



• Gaussian kernel ( $\sigma = 5$ )



Examples

• Periodic kernel



• Changepoint kernel



Examples

• Symmetric kernel



Examples

• Linear kernel



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## Deformation models for registration

#### Basic assumption: Deformation fields are smooth

- Typical assumption:
  - Deformation field is smooth
- GP approach
  - Choose smooth kernel functions  $k(x, x') = s \exp(-\frac{\|x - x'\|^2}{\sigma^2})$
- Regularization operators
  - Penalize large derivatives

$$\mathcal{R}[u] = \|Ru\|^2 = \sum_{i=0}^n \alpha_i \|D_i u\|^2$$



#### Green's functions and covariance functions

$$\mathcal{R}[u] = ||Ru||^2 = \sum_{i=0}^n \alpha_i ||D_iu||^2$$

Corresponding covariance function for GP is the Greens function G:

$$R^*RG(x,y) = \delta(x-y)$$

• We can define Gaussian processes, which mimic typical regularization operators.

T. Poggio and F. Girosi; Networks for Approximation and Learning, Proceedings of the IEEE, 1990

Example: Gaussian kernel

$$k(x, x') = \exp(-\frac{\|x - x'\|^2}{\sigma^2})$$

$$\mathcal{R}[u] = \|Ru\|^2 = \sum_{i=0}^{\infty} \frac{\sigma^{2i}}{i! \, 2^i} \|D_i u\|^2$$

- Non-zero functions are penalized
  - pushes functions to zero away from data

Yuille, A. and Grzywacz M. A mathematical analysis of the motion coherence theory. International Journal of Computer vision

#### Example: Exponential kernel (1D case)

$$k(x, x') = \frac{1}{2\alpha} \exp(-\alpha ||x - x'||)$$

$$\mathcal{R}[u] = \|Ru\|^2 = \alpha^2 u + \|D_1 u\|^2$$

Rasmussen, Carl Edward, and Christopher KI Williams. *Gaussian processes for machine learning*. Vol. 1. Cambridge: MIT press, 2006.

#### Matérn class of kernels

$$k(x, x') = s \frac{2^{1-\nu}}{\Gamma(\nu)} \left( 2\sqrt{2\nu} \frac{\|x - x'\|}{\rho} \right)^{\nu} K_{\nu}(\sqrt{2\nu} \frac{\|x - x'\|}{\rho})$$

- $\Gamma$  is the  $\Gamma$  function,  $k_{
  u}$  the modified Bessel function and u, 
  ho are parameters
- Process  $u \sim GP(0, k)$  is v 1 times m.s. differentiable
- Special cases:

• 
$$\nu = \frac{1}{2}$$
:  $k(x, x') = s \exp(-\frac{\|x - x'\|}{\rho})$   
•  $\nu = \frac{3}{2}$ :  $k(x, x') = s(1 + \frac{\sqrt{3}\|x - x'\|}{\rho}) \exp(-\frac{\sqrt{3}\|x - x'\|}{\rho})$ 

•  $\nu \rightarrow \infty$  Gaussian kernel

• Minimizes the bending energy of a metal sheet

$$R[u] = \left\| \left( \nabla^T \nabla u \right) \right\|^2$$

• Corresponding covariance function

$$k(x, x') = \frac{1}{12} \left( 2\|x - x'\|^3 - 3R(\|x - x'\|^2 + R^3) \right)$$
  
where  $R = \max_{x, x' \in \Omega} \|x - x'\|$ 

Rohr, Karl, et al. "Landmark-based elastic registration using approximating thin-plate splines." *IEEE Transactions on medical imaging* 20.6 (2001): 526-534. Williams, Oliver and Fitzgibbon Andrew, "Gaussian process implicit surfaces"

• We can build a covariance function from B-Spline basis functions  $\beta$  (*s* is a scaling constant)

$$k(x,y) = \sum_{k \in \mathbb{Z}^d} \beta(sx - k)\beta(sy - k)$$

• Corresponding deformation model often called "free form deformations"

- Rueckert, Daniel, et al. "Nonrigid registration using free-form deformations: application to breast MR images." *IEEE transactions on medical imaging* 18.8 (1999): 712-721.
- Klein, Stefan, et al. "Elastix: a toolbox for intensity-based medical image registration." *IEEE transactions on medical imaging* 29.1 (2010): 196-205.

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Many standard models for registration can be formulated using Gaussian processes

- Yields probabilistic interpretation
- We can sample and visualize deformation fields
- Can use them as building blocks for more complicated priors

Constructing s.p.d. kernels



1. 
$$k(x, x') = f(x) f(x')^T$$
,  $f: X \to \mathbb{R}^d$   
2.  $k(x, x') = \alpha k_1(x, x'), \alpha \in \mathbb{R}_+$  (scaling)  
3.  $k(x, x') = B^T k_1(x, x')B$ ,  $B \in \mathbb{R}^{r \times d}$  (lifting)  
4.  $k(x, x') = k_1(x, x') + k_2(x, x')$  (or relationship)  
5.  $k(x, x') = k_1(x, x') \cdot k_2(x, x')$  (and relationship)

Add kernels that act on different scales:

$$k(x,x') = \sum_{i=0}^{n} \sum_{k \in \mathbb{Z}^d} \beta \left( 2^{-i}x - k \right) \beta \left( 2^{-i}y - k \right)$$

• Wavelet like multiscale representation

Opfer, Roland. "Multiscale kernels." *Advances in computational mathematics* 25.4 (2006): 357-380.

#### Multi-scale kernel



Scale deformations differently in each direction

$$k(x,x') = R^T \begin{pmatrix} \sqrt{s_1} & 0\\ 0 & \sqrt{s_2} \end{pmatrix} k(x,x') \begin{pmatrix} \sqrt{s_1} & 0\\ 0 & \sqrt{s_2} \end{pmatrix} R$$

- R is a rotation matrix
- k is scalar valued
- $s_1$ ,  $s_2$  scaling factors

#### Anisotropic priors



Spatially-varying priors

Use different models for different regions

 $k(x, x') = \chi(x)\chi(x')k_1(x, x') + (1 - \chi(x))(1 - \chi(x'))k_2(x, x')$ 

 $\chi(x) = \begin{cases} 1 & \text{if } x \in \text{thumb region} \\ 0 & \text{otherwise} \end{cases}$ 



Freiman, Moti, Stephan D. Voss, and Simon K. Warfield. "Demons registration with local affine adaptive regularization: application to registration of abdominal structures." *Biomedical Imaging: From Nano to Macro, 2011 IEEE International Symposium on*. IEEE, 2011.

Spatially-varying priors



#### Statistical deformation models

#### Estimate mean and covariance function from data:



#### Example 5: Statistical deformation models



#### Landmark registration using GP Regression



Given:

- Gaussian process:  $u \sim GP(\mu, k)$
- Observations:  $\{(l_i^R, \tilde{u}_i), i = 1, ..., n\}$

Assume:  $\tilde{u}_i = u(l_i) + \epsilon$  with  $\epsilon \sim N(0, \sigma^2 I_{2\times 2})$ .

Goal:

• Find posterior distribution

$$u \mid l_1^R$$
, ...,  $l_n^R$ ,  $\tilde{u}_1$ , ...,  $\tilde{u}_n$ 

#### Gaussian process regression



The posterior  $u \mid l_1^R, ..., l_n^R, \tilde{u}_1, ..., n$ is a Gaussian process  $GP(\mu_p, k_p)$ Its parameters are known analytically.

 $\mu_p(x) = \mu(x) + K(x, Y)(K(Y, Y) + \sigma^2 I_{2n \times 2n})^{-1}(\tilde{u} - \mu(Y))$  $k_p(x, x') = k(x, x') - K(x, Y)(K(Y, Y) + \sigma^2 I_{2n \times 2n})^{-1}K(Y, x')$ 

#### Landmark registration using GP Regression





#### Hybrid registration

- We can now combine landmark registration with intensity:
  - 1. Use Gaussian process regression to obtain posterior from  $GP(\mu, k)$  from landmarks
  - 2. Use  $GP(\mu_p, k_p)$  as new prior model for registration

Wörz, Stefan, and Karl Rohr. "Hybrid spline-based elastic image registration using analytic solutions of the navier equation." *Bildverarbeitung für die Medizin 2007*. Springer Berlin Heidelberg, 2007. 151-155.

Lu, Huanxiang, Philippe C. Cattin, and Mauricio Reyes. "A hybrid multimodal non-rigid registration of MR images based on diffeomorphic demons." *Engineering in Medicine and Biology Society (EMBC), 2010 Annual International Conference of the IEEE*. IEEE, 2010.

#### Demo: Priors and interactive registration



## **Skull Segmentation in MRI** Lab-meeting

Slides by Patrick Kahr



## **Skull Segmentation in MRI**

**Problem**: Bone and air have similar intensities in MRI

 $\rightarrow$  unlike CT, no threshold segmentation possible







 $\operatorname{Cube}(C_1)$  3D  $\operatorname{Cross}(R_1)$ 

 $(O_1) = 3D \operatorname{Octagon}(O_2)$ 

#### Mathematical Morphology (Dogdas et al. 2005)



Multi-level model fitting (Lerch, Lüthi 2008)





Multi-Atlas matching (Torrado-Carvajal et al. 2015)

#### Modeling deformations with SSMs



deformation field defined on reference mesh

Model deformations of reference shape using an SSM  $GP(\mu_{ssm}, k_{ssm})$ 

Deformations are only defined on surface of reference shape:

 $\rightarrow\,$  deformation field needs to be interpolated for the rest of the image.

### Modeling deformations with SSMs



deformation field defined on complete image domain

Hybrid kernel: mix SSM with smooth Gaussian kernel

$$(1 - w(x))k_{gaussian}(x, x')(1 - w(x')) + w(x)k_{SSM}(x, x')w(x')$$

where

x', y': closest points to x,y on the surface,

w = 1 if x, y on the surface, w $\rightarrow$ 0 for x, y far away from surface.

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#### Registration accuracy: KGaussian vs KHybrid

Register 14 CT templates to 12 MR targets (168 registrations).

Measure average distance to a set of 10 anatomical landmarks.



Registration accuracy (K\_Gaussian vs K\_Hybrid)





#### Segmentation accuracy

#### Multi-atlas matching:

Target segmentation is mean shape obtained from 14 CT registrations.







### Segmentation example







#### Segmentation accuracy

Segmentation accuracy (K\_Gaussian vs K\_Hybrid)



#### Variance

Lower variance between registrations for each of the 12 targets with  $K_{\mbox{hybrid.}}$ 



Total Reg. Variance (K\_Gaussian vs K\_Hybrid)







### Summary

- GPs provide probabilistic interpretation to classic registration models
  - But, can visualize assumptions
  - New ways to combine priors to individual applications.
- Modelling and model fitting are separated
  - Change in prior does not lead to change in algorithm
  - No increase in complexity
  - Can tailor model to application without increase in complexity
- Can use SSMs of individual organs to guide image registration