

MICCAI 2010 Tutorial

# Intensity-based Deformable Registration

## Similarity Measures

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# Quantify the similarity between images

- Do images contain the same object?



Image retrieval

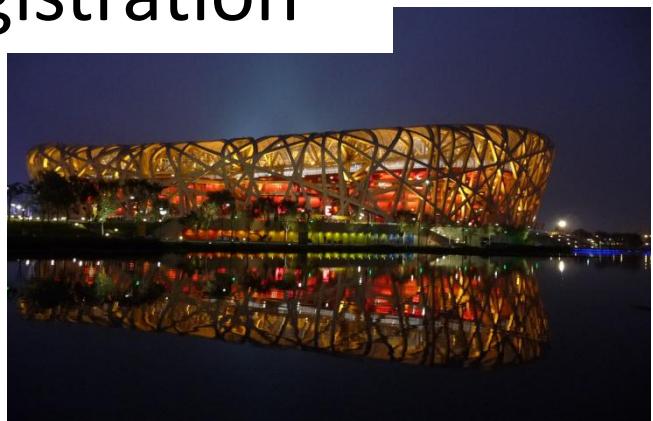
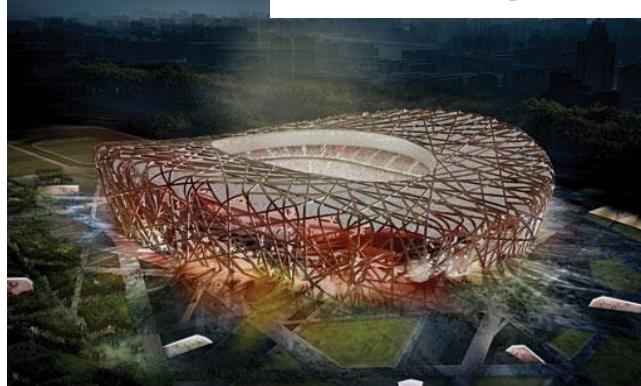


# Quantify the similarity between images

- How are images correctly aligned?

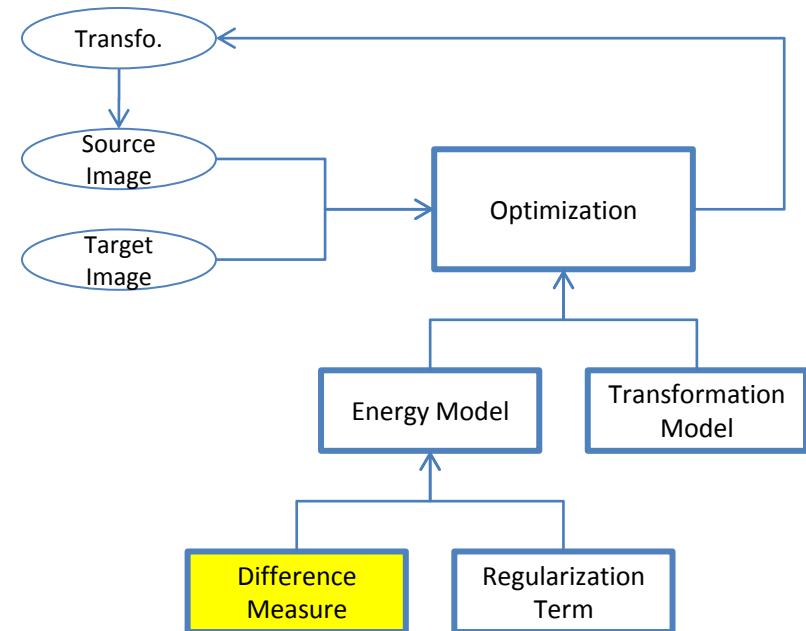


Image registration



# Requirements on similarity measure

- Extremum for correctly aligned images
- Smooth, best convex
- Fast computation
- Differentiable



# Overview

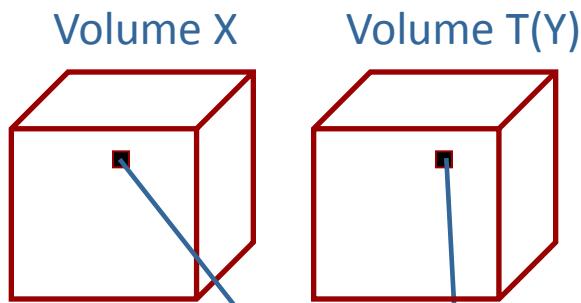
1. Standard similarity measures
2. Probabilistic framework
3. Pre-processing steps
4. Recent approaches
5. Linear vs. non-linear registration

# PART I

## Standard Similarity Measures

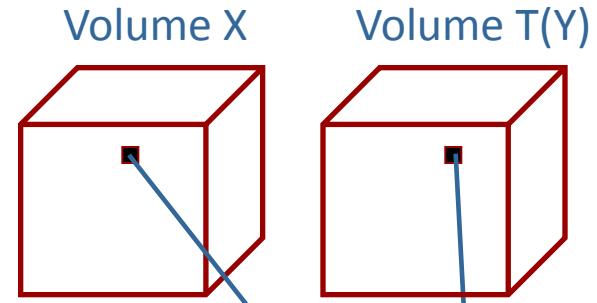
- SSD-SAD
- Cross Correlation
- Mutual information
- Derivatives

# Difference Measures



$$SSD = \frac{1}{N} \sum_i (x_i - y_i)^2$$

**Sum of Squared Differences**

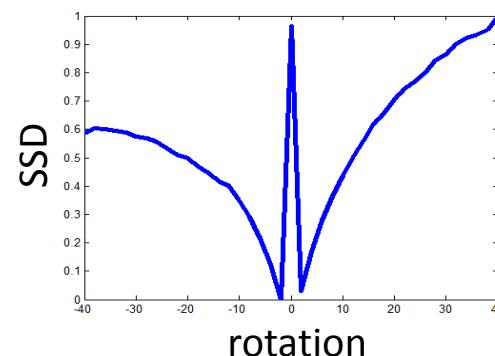


$$SAD = \frac{1}{N} \sum_i |x_i - y_i|$$

**Sum of Absolute Differences:**  
Less sensitive on large intensity  
differences than SSD

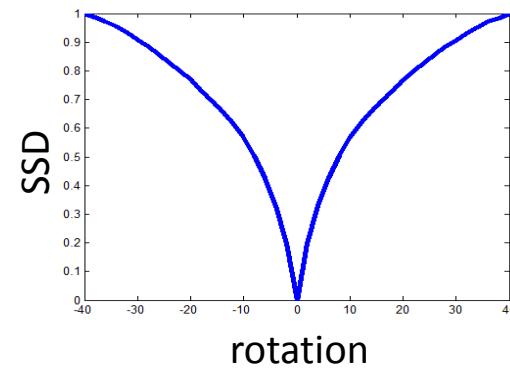
# Limitations of SSD

- Illumination change affects similarity function



- Idea: normalization of images

$$\tilde{X} = \frac{X - \mathbb{E}[X]}{\sigma(X)}$$



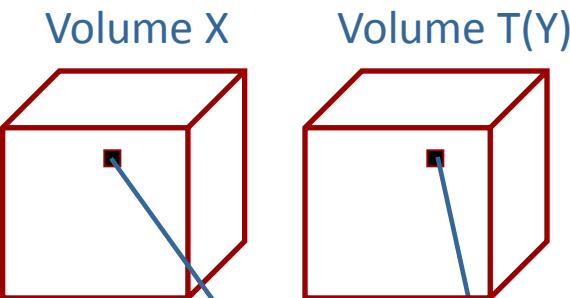
# SSD on Normalized Images

$$\mathbb{E}[(\tilde{X} - \tilde{Y})^2]$$

$$\tilde{X} = \frac{X - \mathbb{E}[X]}{\sigma(X)}$$

$$\tilde{Y} = \frac{Y - \mathbb{E}[Y]}{\sigma(Y)}$$

# Normalized Cross Correlation (NCC)



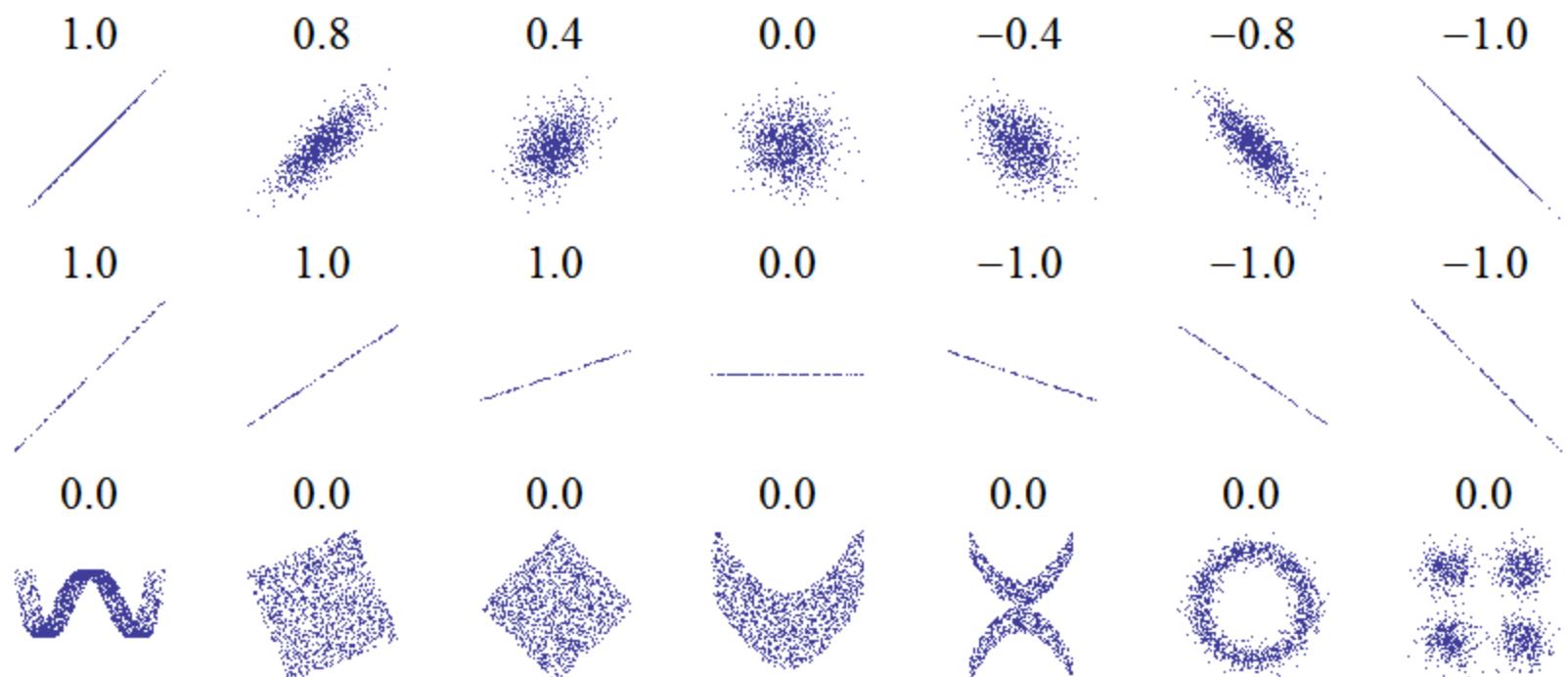
$\bar{x}$  : Mean  
 $\sigma_x$  : Standard deviation  
 $N$  : Number of pixels

$$NCC = \frac{1}{N} \sum_i \frac{(x_i - \bar{x})(y_i - \bar{y})}{\sigma_x \sigma_y}$$

**Normalized Cross Correlation:**

Expresses the linear relationship between voxel intensities in the two volumes

# NCC - example

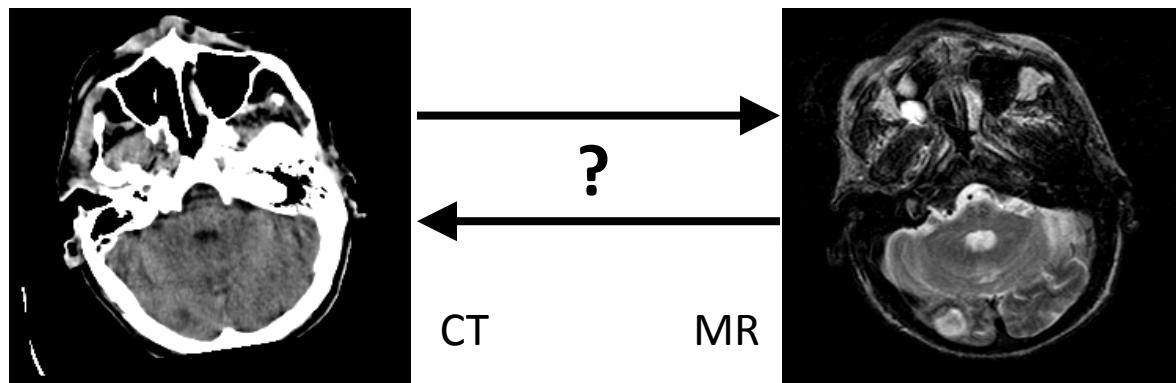


Source: wikipedia

Sets of (x,y) points, with the NCC of x and y for each set.

# Multi-Modal Registration

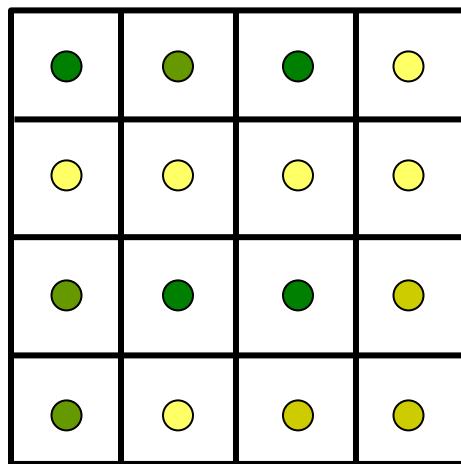
- More complex intensity relationship



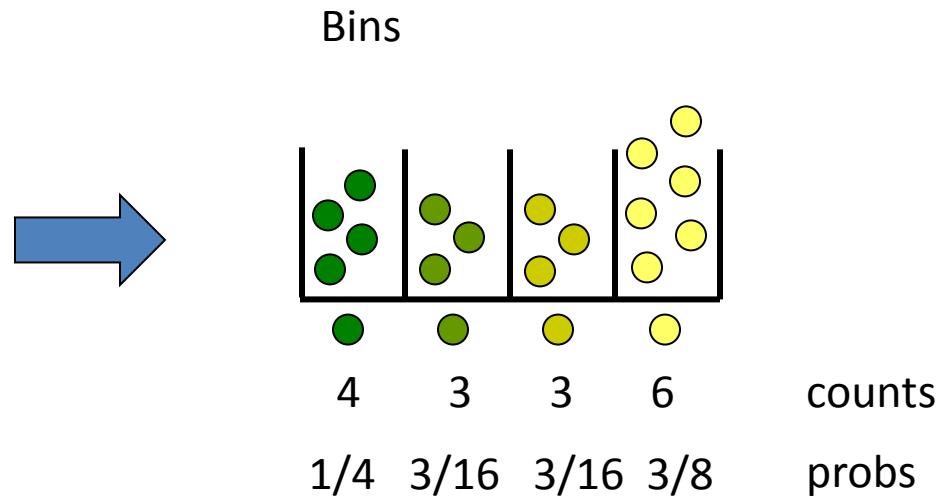
- Approaches:
  - Simulate one modality from the other one
  - Apply sophisticated similarity measures

# Information Theoretic Approach

# Histogram calculation



Image



Histogram

# Joint histogram calculation

Image X

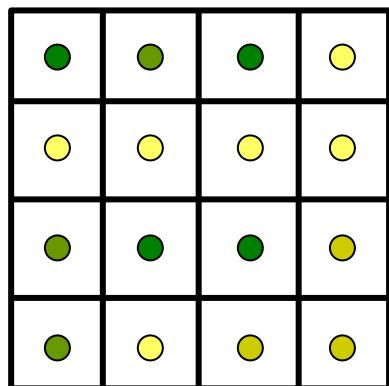
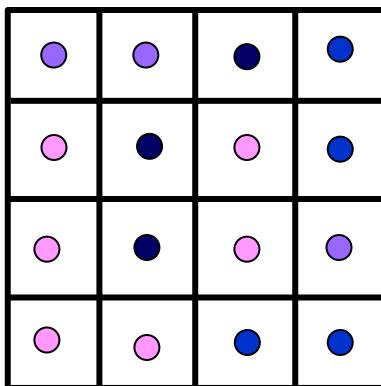
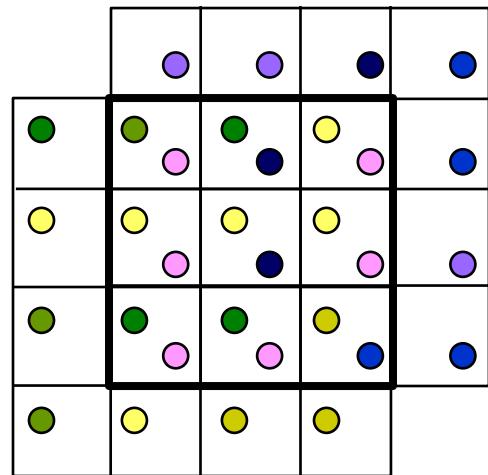


Image Y

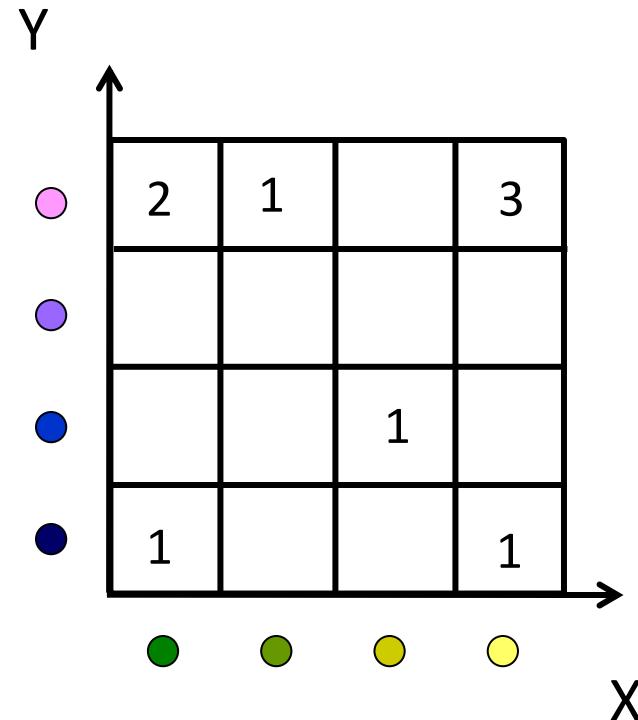


Overlap



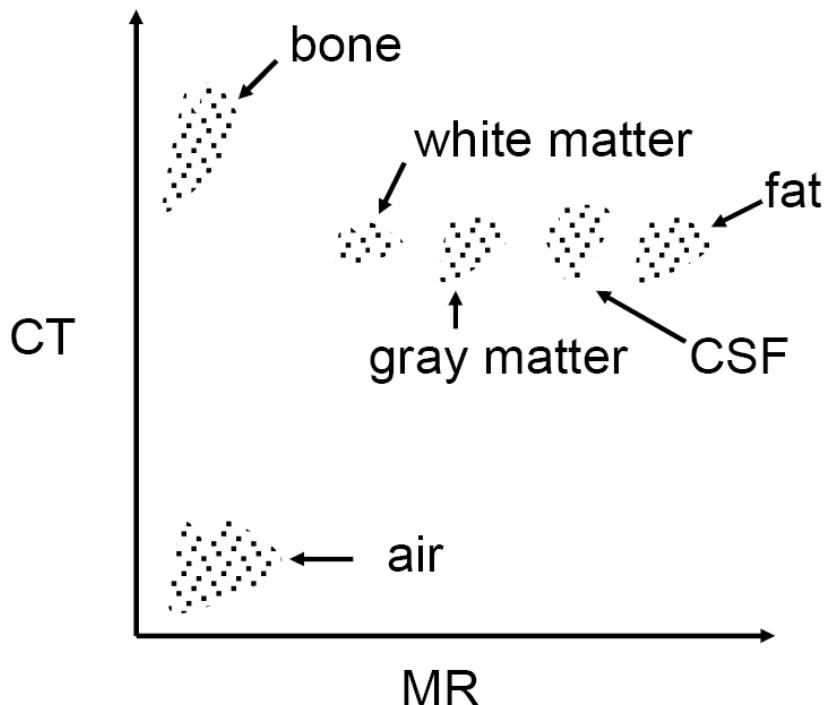
$$\begin{pmatrix} t_x = 1 \\ t_y = 1 \end{pmatrix}$$

Joint Histogram

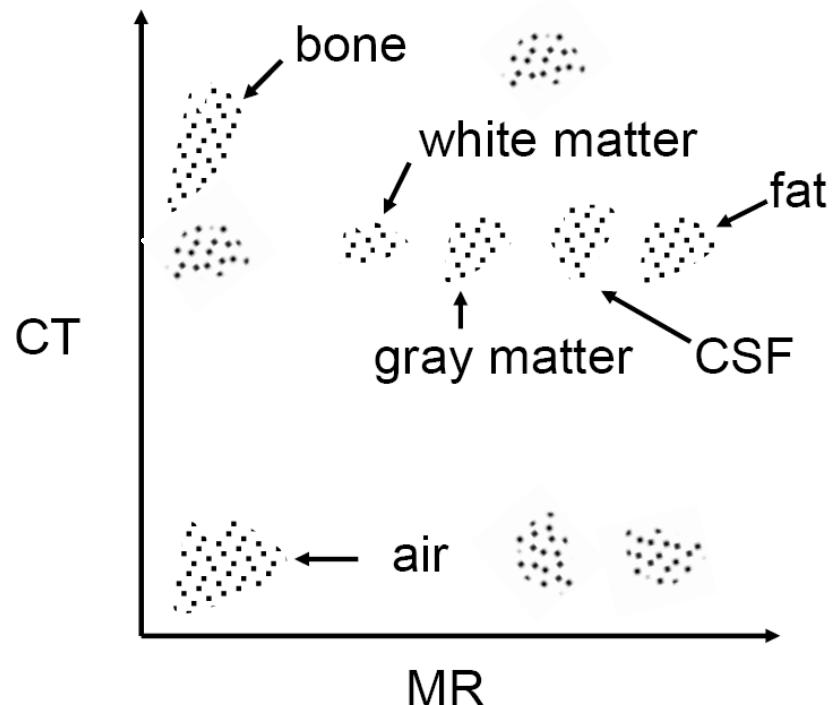


# Information Theoretic Approach

Registered



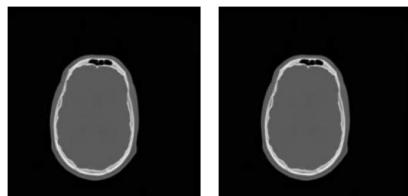
Not Registered



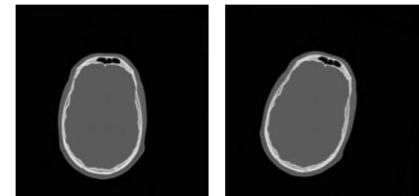
Source: W. Wells, MICCAI 2009

# Joint Histogram

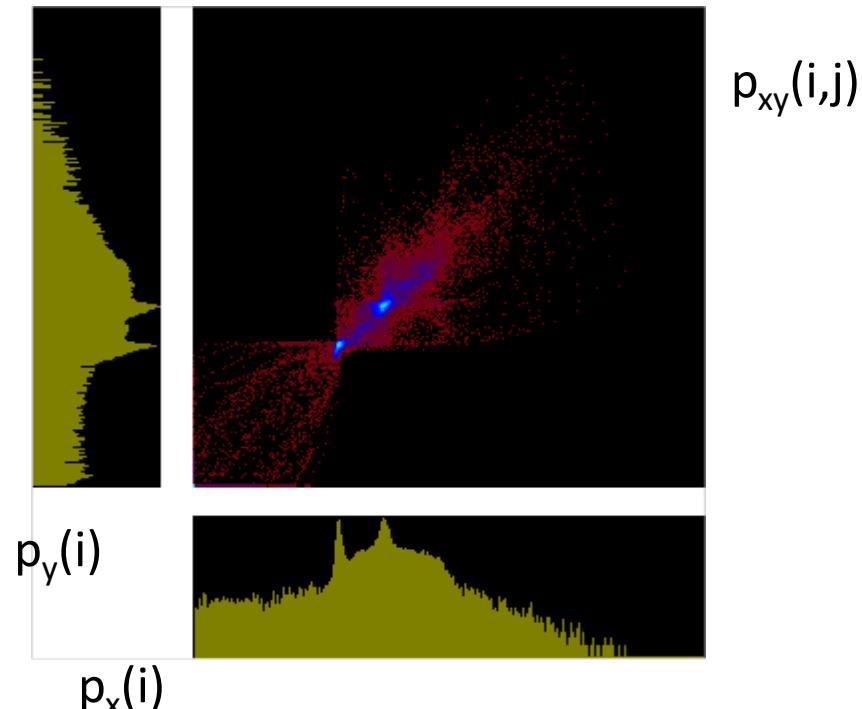
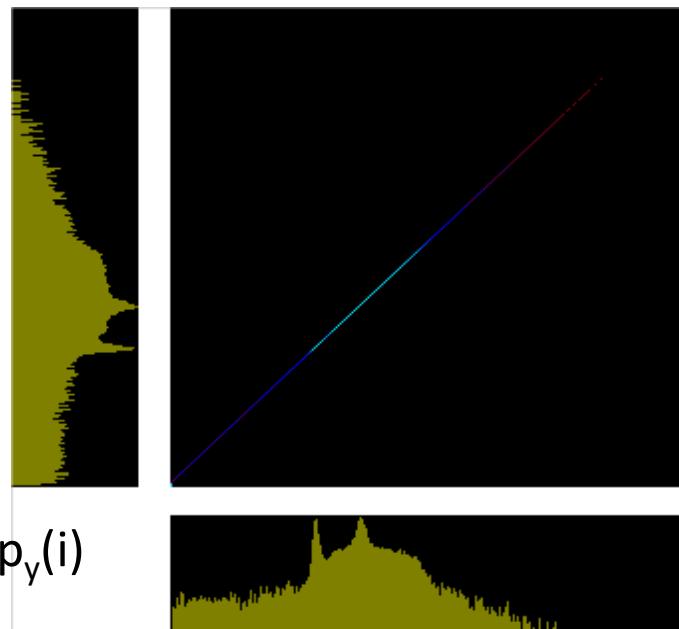
X and Y identical



X and Y misaligned



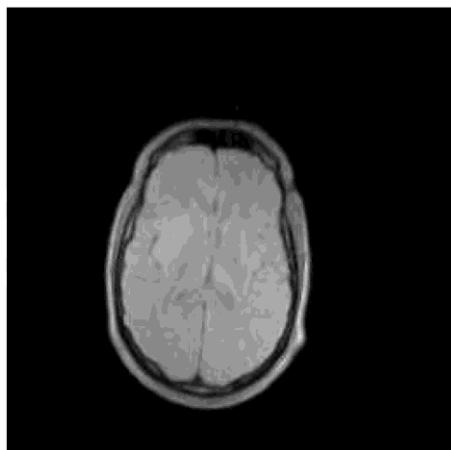
$$p_{xy}(i,j)$$



# Joint Histogram

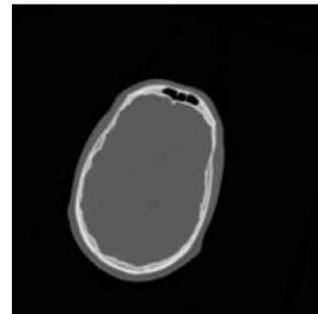
- Histogram for images from different Modalities

Source Image

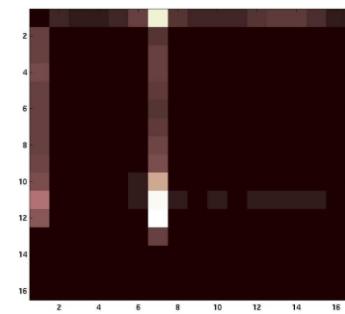


Not Aligned

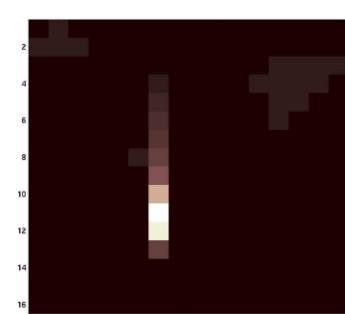
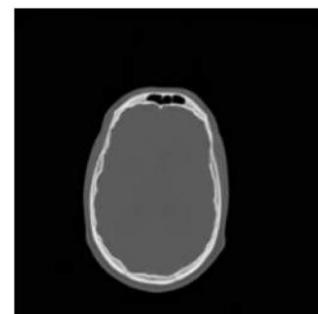
Target Image



Joint Histogram

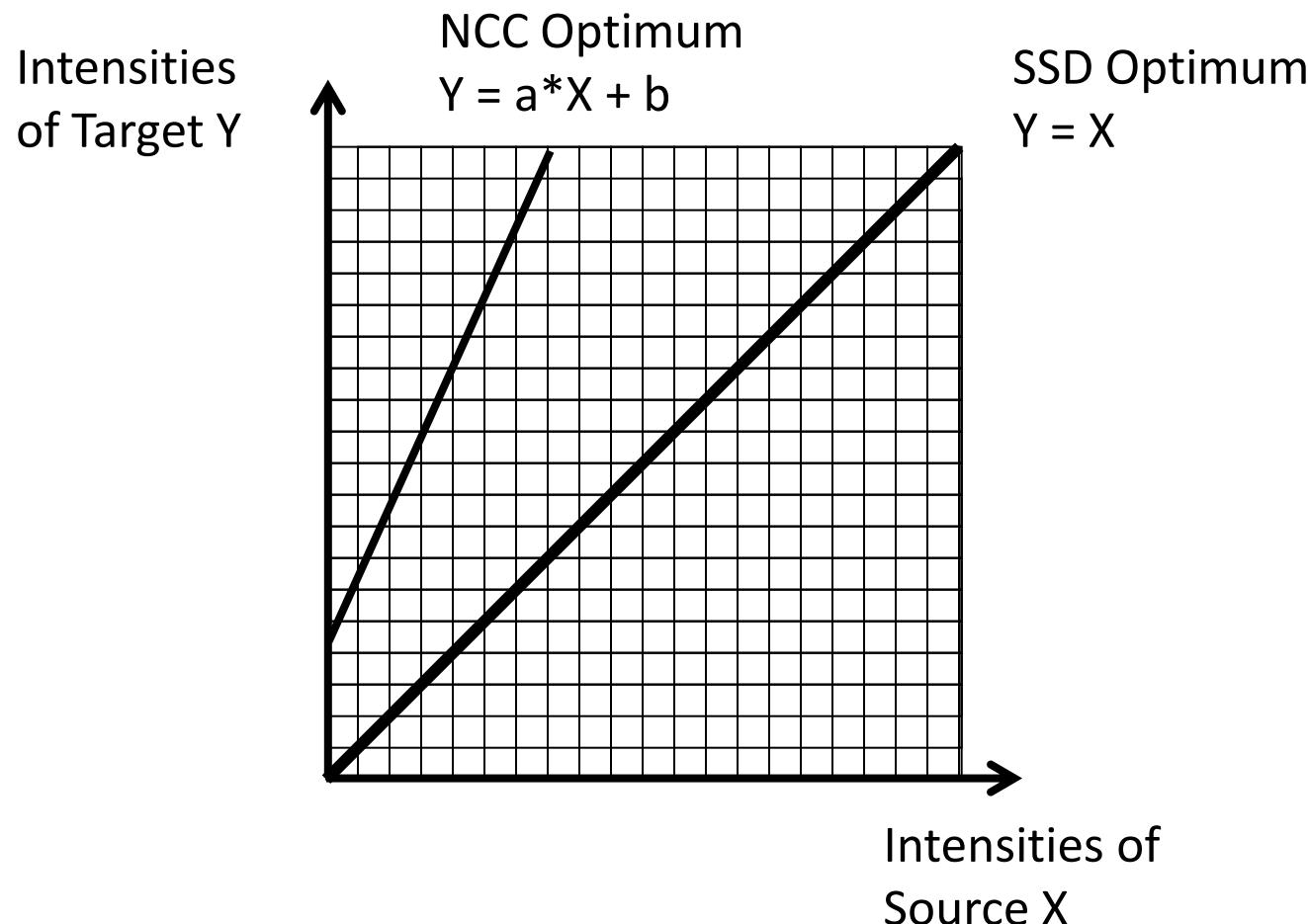


Aligned

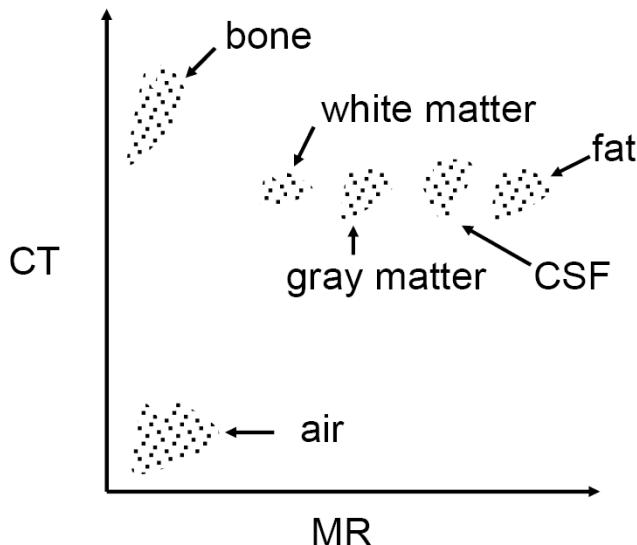


Source: PhD Thesis, L. Zöllei

# Joint Histogram



# Information Theoretic Approach



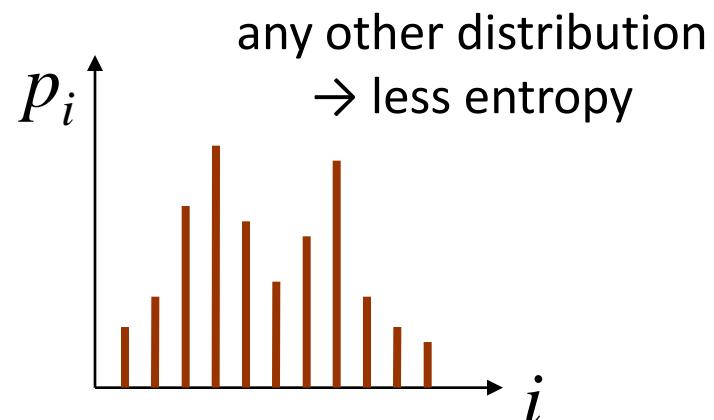
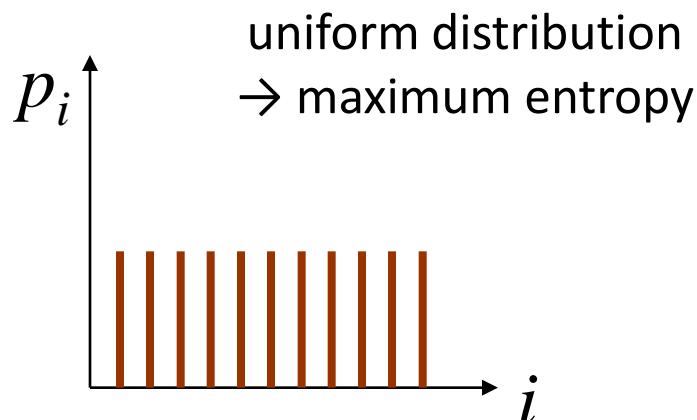
- How to quantify the quality of alignment between MR and CT?
- ✓ Measure the *structure* of the joint distribution
- How to measure the structure?
- ✓ Shannon Entropy

Source: W. Wells, MICCAI 2009

# Entropy

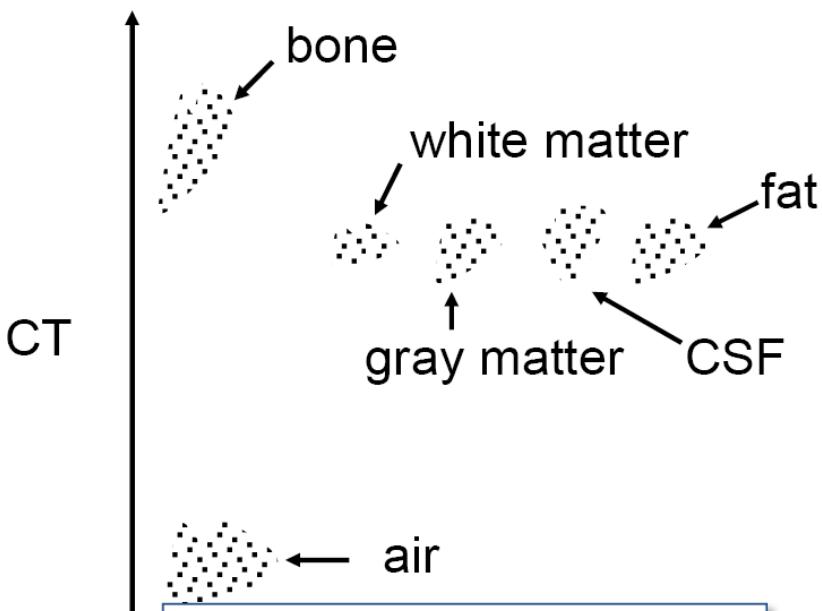
Shannon Entropy, developed in the 1940s  
(communication theory)

$$H = - \sum_i p_i \log p_i$$

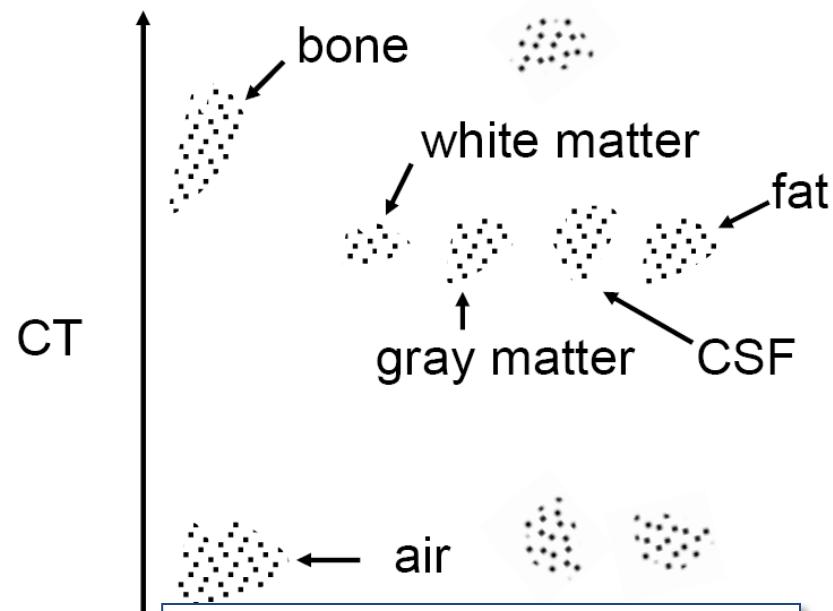


# Maximize or minimize entropy?

Registered



Not Registered



# Mutual Information (MI)

$$MI(X, Y) = H(X) + H(Y) - H(X, Y)$$

$$= \sum_i \sum_j p_{xy}(i, j) \log \frac{p_{xy}(i, j)}{p_x(i)p_y(j)}$$

- Maximized if X and Y are perfectly aligned
- $H(X)$  and  $H(Y)$  help to make the measure more robust
- Maximization of mutual information leads to minimization of joint entropy

# Historical Note

- Minimum Entropy Registration
  - Collignon A., Vandermeulen, D., Suetens, P., and Marchal, G. *3D multi-modality medical image registration using feature space clustering*. CVRMED April 1995.
- Maximum Mutual Information Registration
  - Viola, P. and Wells, W.. *Alignment by maximization of mutual information*. In Proceedings of the 5th International Conference of Computer Vision, June 20 – 23, 1995.
  - Collignon A, Maes F, Delaere D, Vandermeulen D, Suetens P, Marchal G, *Automated multi-modality image registration based on information theory*. IPMI June 26, 1995.
  - Viola, P. *Alignment by maximization of Mutual Information*. MIT PhD Thesis, June 1995.

# Normalization of MI

- Problem: changing overlap affects MI
- Entropy Correlation Coefficient (ECC)

$$\text{ECC}(X, Y) = \sqrt{2 - \frac{2 \cdot H(X, Y)}{H(X) + H(Y)}}$$

Maes F, Collignon A, Vandermeulen D, Marchal G, Suetens P. *Multimodality image registration by maximization of mutual information*, TMI, 1997

- Normalized MI (NMI)

$$\text{NMI}(X, Y) = \frac{H(X) + H(Y)}{H(X, Y)}$$

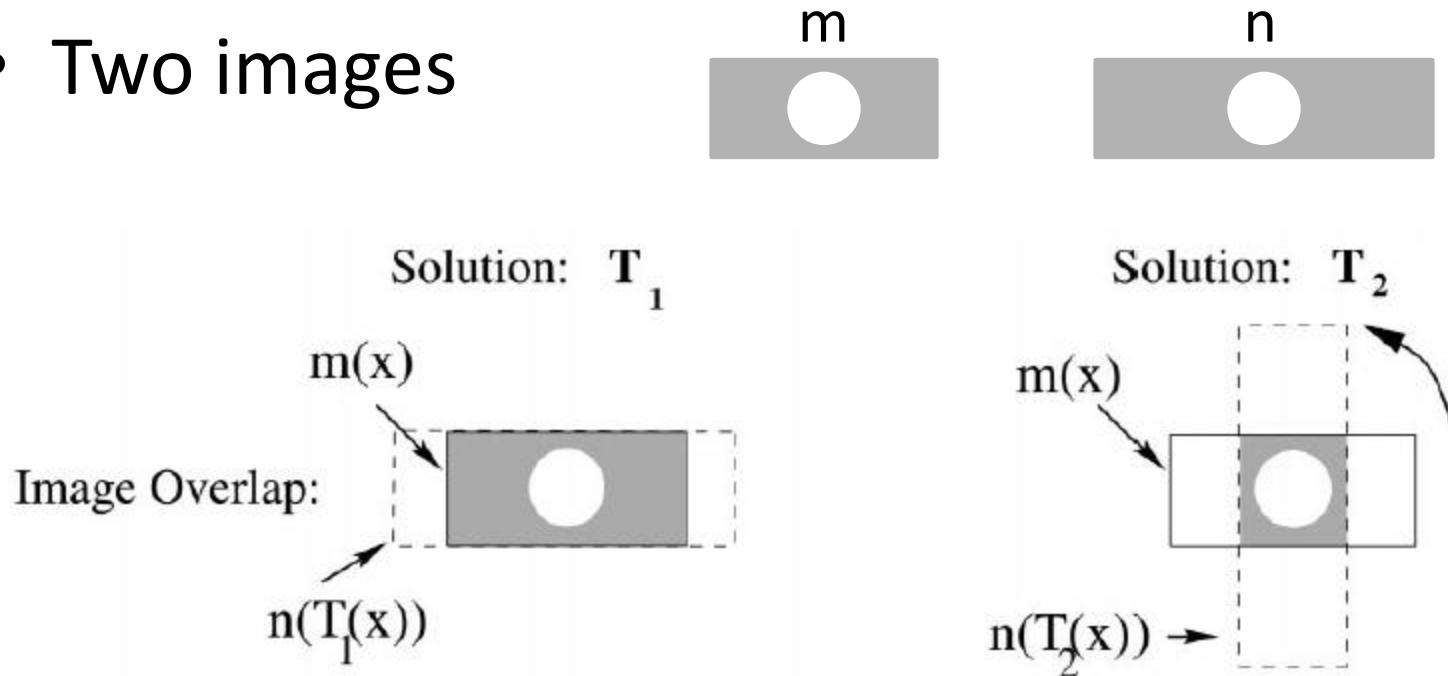
C. Studholme, D.L.G.Hill, D.J. Hawkes, *An Overlap Invariant Entropy Measure of 3D Medical Image Alignment*, Pattern Recognition, Vol. 32(1), Jan 1999, pp 71-86.

- Revisiting overlap invariance

Nathan D. Cahill, Julia A. Schnabel, J. Alison Noble, David J. Hawkes, *"Revisiting overlap invariance in medical image alignment,"* MMBIA, 2008

# NMI – change of overlap

- Two images



- $T_2$  has higher information content
- Overlap changes for deformable registration?

C. Studholme, D.L.G.Hill, D.J. Hawkes, *An Overlap Invariant Entropy Measure of 3D Medical Image Alignment*, *Pattern Recognition*, Vol. 32(1), Jan 1999, pp 71-86.

# Improvements to MI

- Density estimation
  - Parzen window
  - Partial volume distribution
  - Uniform volume histogram
  - NP windows
- Spatial information
- Alternative entropy measures
- Tutorial at MICCAI 2009:  
*Information theoretic similarity measures for image registration and segmentation:* Maes, Wells, Pluim  
<http://ubimon.doc.ic.ac.uk/MICCAI09/a1882.html>

# Derivatives of Similarity Measures

- General form of derivative of similarity metrics

$$\frac{\partial \text{SM}(X, Y(T_p))}{\partial p} = \boxed{\frac{\partial \text{SM}(X, Y)}{\partial Y}} \quad \frac{\partial Y}{\partial T_p} \quad \frac{\partial T_p}{\partial p}$$

– SSD:

$$\nabla \text{SSD}(X, Y) = -2 \cdot (X - Y)$$

– MI

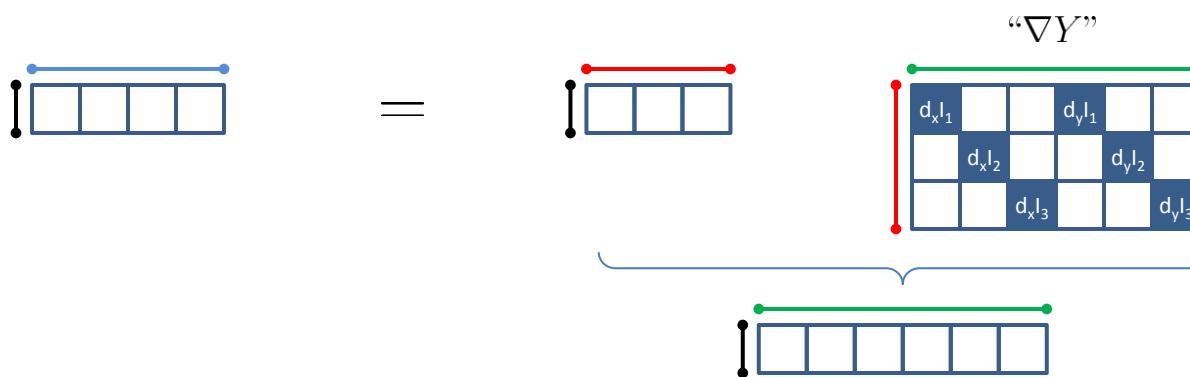
$$\nabla \text{MI}(X, Y) = G_\Psi * \frac{1}{|\Omega|} \left( \frac{\partial_2 p(X, Y)}{p(X, Y)} - \frac{p'(Y)}{p(Y)} \right)$$

Hermosillo, G., Chefd'Hotel, C., Faugeras, O.: *Variational Methods for Multimodal Image Matching*, International Journal of Computer Vision 50(3) (2002)

# Derivatives of Similarity Measures

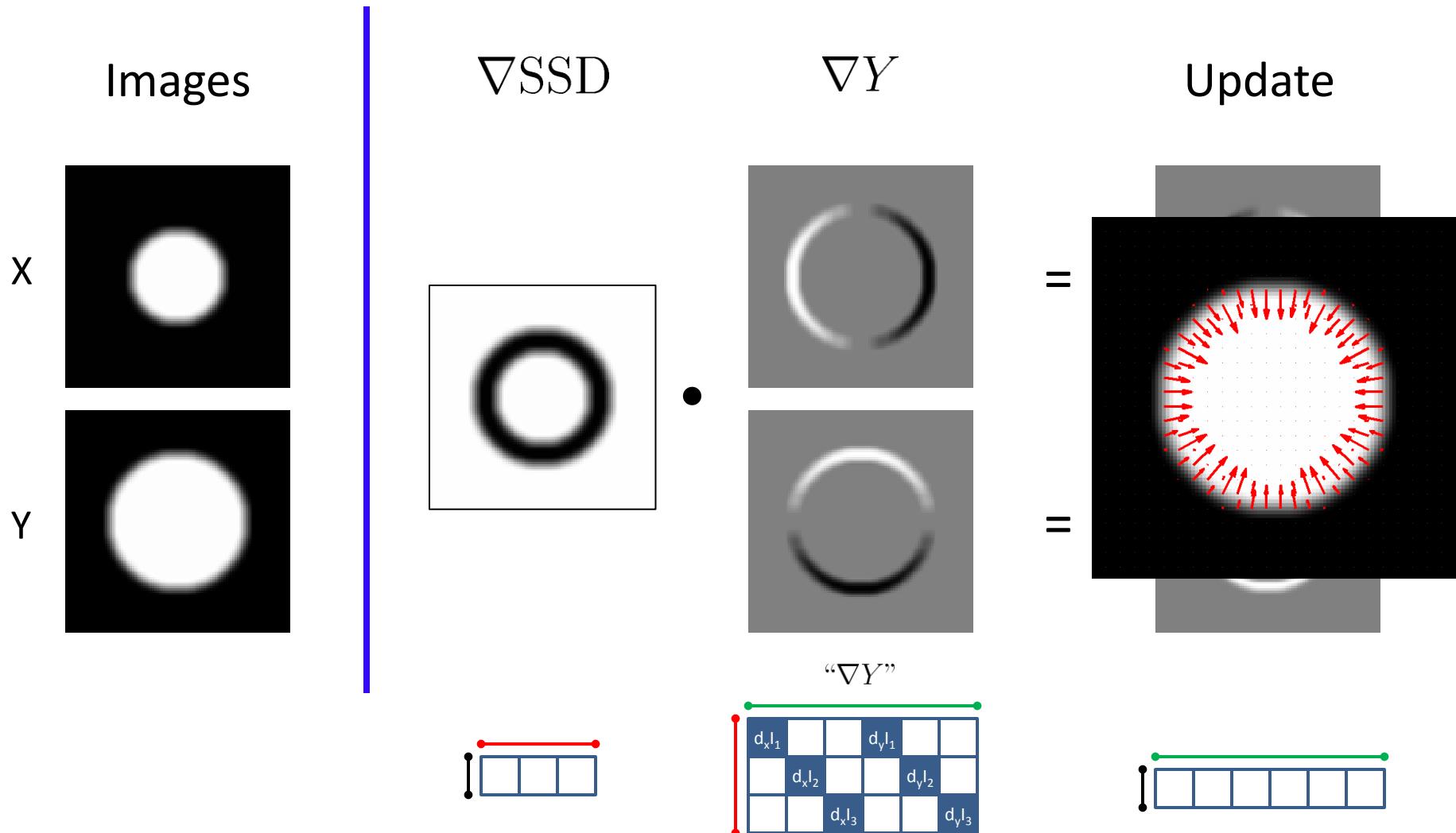
- General form of derivative of similarity metrics

$$\frac{\partial \text{SM}(X, Y(T_p))}{\partial p} = \frac{\partial \text{SM}(X, Y)}{\partial Y} \frac{\partial Y}{\partial T_p}$$

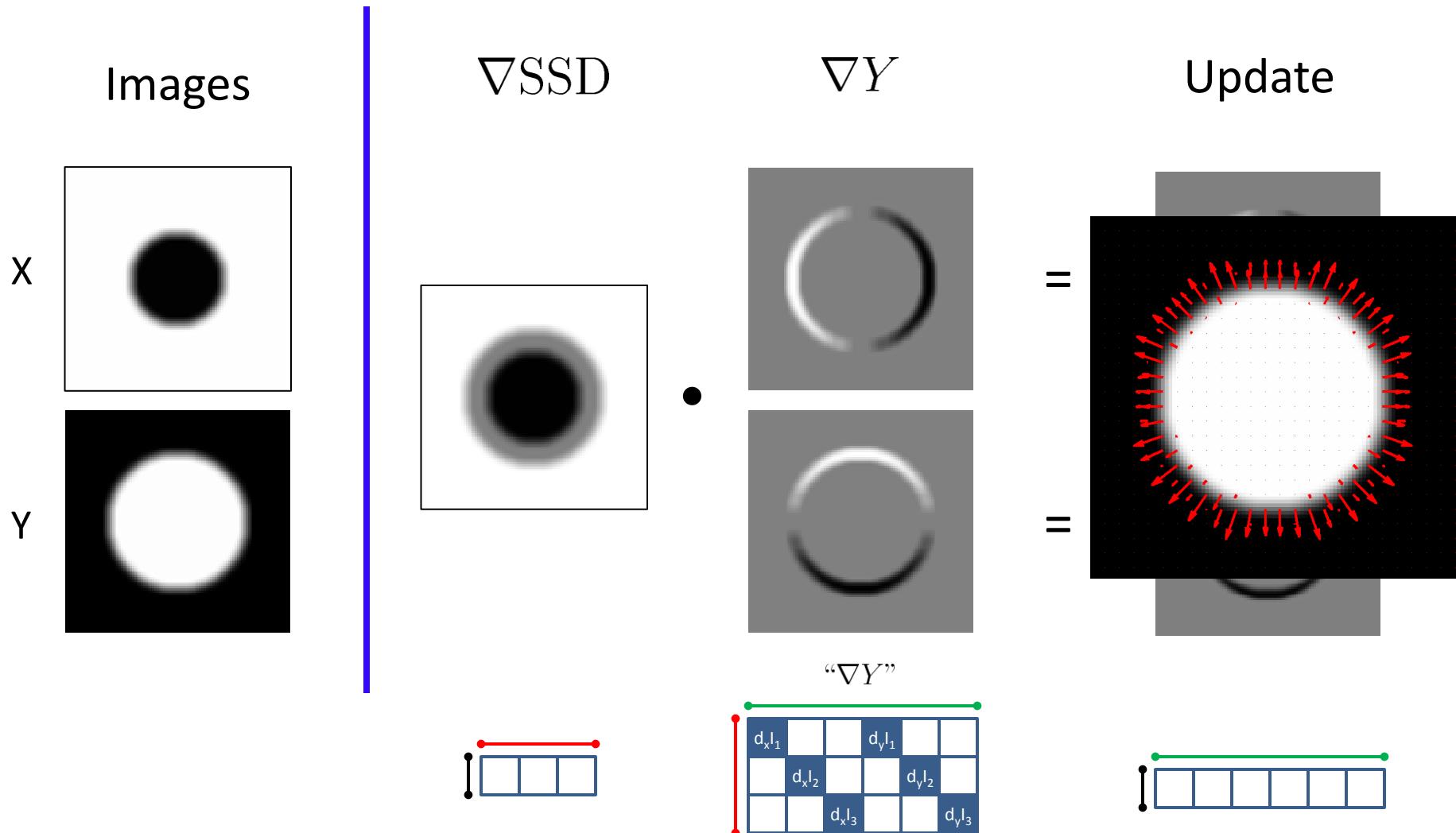


The diagram shows a 5x5 grid of squares. A large red 'X' is drawn across the entire grid, indicating that the current configuration is invalid. A green vertical line is positioned to the left of the first column, and a blue horizontal line is positioned above the first row. These lines highlight specific rows and columns that are being analyzed or modified.

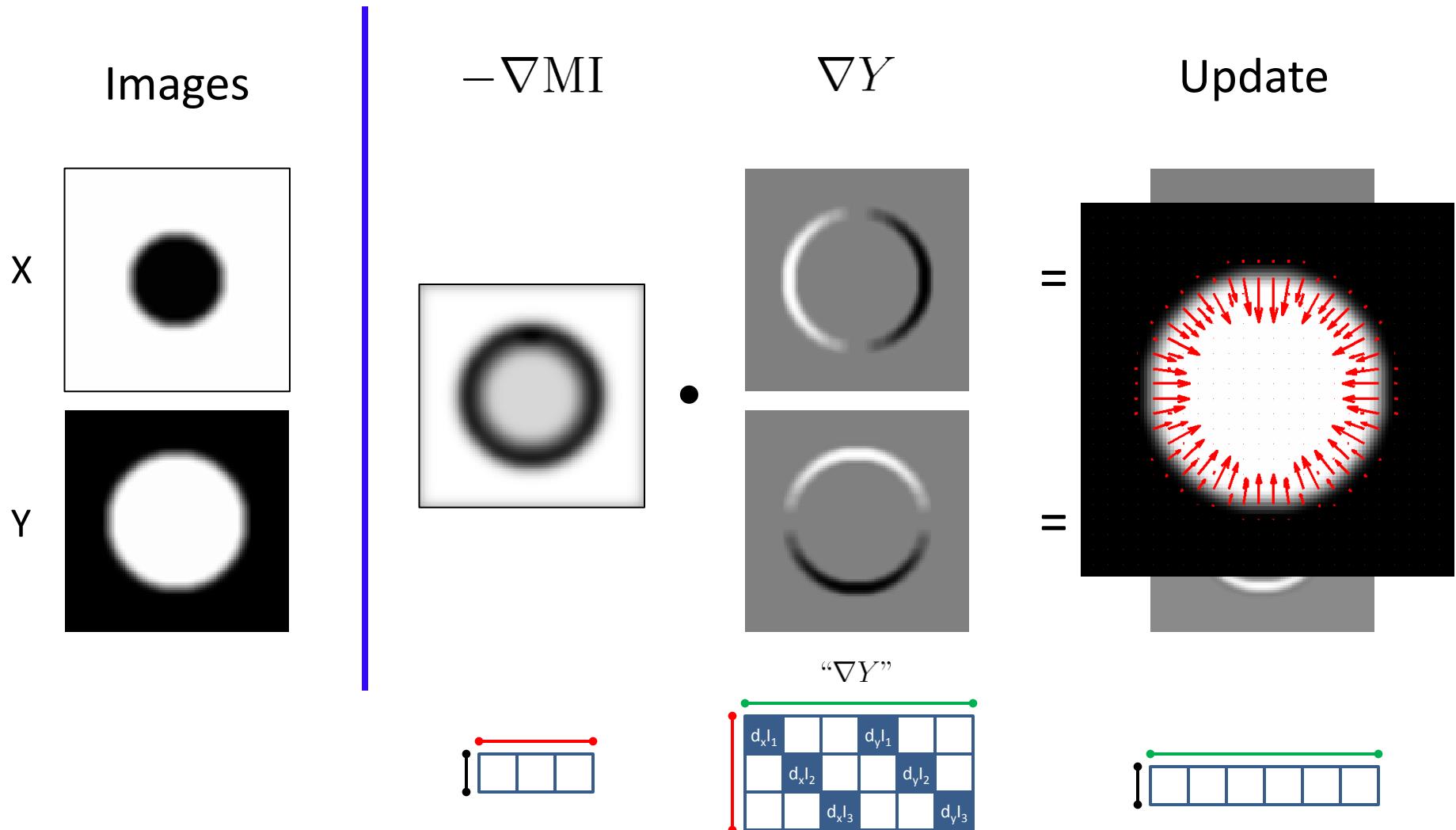
# Derivatives of Similarity Measures



# Derivatives of Similarity Measures



# Derivatives of Similarity Measures



# Overview

1. Standard similarity measures
2. Probabilistic framework
3. Pre-processing steps
4. Recent approaches
5. Linear vs. non-linear registration

# PART II

## Probabilistic Framework for Image Registration

- Viola, PhD thesis, 1995
- Roche et al., 2000
- Zöllei, PhD thesis, 2006

# Probabilistic Framework for registration

- Maximum Likelihood Estimation (MLE)
  - Probability for the model  $m$  having the observations  $a$

$$\arg \max_m P(a = \{a_1, \dots, a_n\} | m) = \arg \max_m \prod_{i=1}^n P(a_i | m)$$

- Log-likelihood function

$$\log \mathcal{L}(m|a) = \sum_{i=1}^n \log P(a_i | m)$$

- Formulate registration as likelihood maximization

# MLE framework for registration

- Model

$$X = f(Y(T)) + \varepsilon$$

$X, Y$  : Images  
 $\varepsilon$  : Noise  
 $f$  : Intensity map

- Probability function

$$P(X|Y, T, \varepsilon, f)$$

- Log-likelihood function

$$\begin{aligned} \log \mathcal{L}(T, \varepsilon, f) &= \log P(X|Y, T, \varepsilon, f) \\ &= \frac{1}{N} \sum_{v \in \Omega} \log P(X(v)|Y(T(v)), \varepsilon, f) \end{aligned}$$

$$\log \mathcal{L}(T, \varepsilon, f) = \frac{1}{N} \sum_{v \in \Omega} \log P(X(v)|Y(T(v)), \varepsilon, f)$$

- Stationary white Gaussian noise

$$P(X|Y, T, f, \sigma) = \frac{1}{\sqrt{2\pi}\sigma} \cdot e^{-\frac{1}{2} \frac{[X - f(Y(T))]^2}{\sigma^2}}$$

- Log-likelihood with Gaussian noise

$$\log \mathcal{L}(T, \sigma, f) = -N \log \sqrt{2\pi}\sigma - \frac{1}{2\sigma^2} \frac{1}{N} \sum_{v \in \Omega} (X(v) - f(Y(T(v))))^2$$

Assuming  $f$  to be the identity

$$\log \mathcal{L}(T, \varepsilon, f) = \frac{1}{N} \sum_{v \in \Omega} \log P(X(v)|Y(T(v)), \varepsilon, f)$$

- Stationary white Gaussian noise

$$P(X|Y, T, f, \sigma) = \frac{1}{\sqrt{2\pi}\sigma} \cdot e^{-\frac{1}{2} \frac{[X-f(Y(T))]^2}{\sigma^2}}$$

- Log-likelihood with Gaussian noise

$$\log \mathcal{L}(T, \sigma) = -N \log \sqrt{2\pi}\sigma - \frac{1}{2\sigma^2} \frac{1}{N} \sum_{v \in \Omega} (X(v) - Y(T(v)))^2$$

SSD

# Intensity relationships

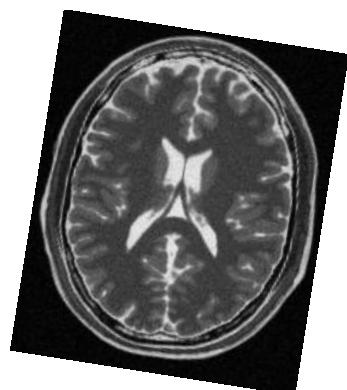
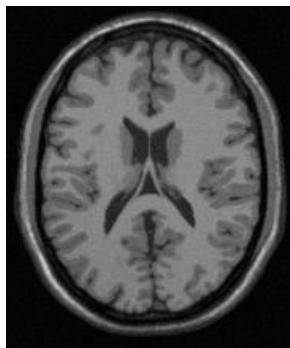
- Identity : SSD
- Affine : NCC
- Functional : Correlation Ratio
- Statistical : Mutual Information

Viola, P. *Alignment by maximization of Mutual Information*. MIT PhD Thesis, June 1995.  
Roche et al., *Unifying maximum likelihood approaches in medical image registration*, 2000

# PART III

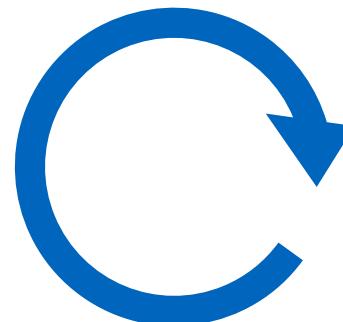
## Pre-Processing Steps

Images



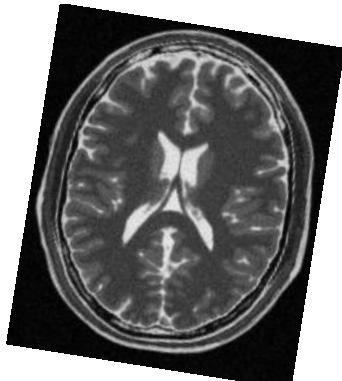
Registration Framework

Similarity Measure



Optimization

## Images

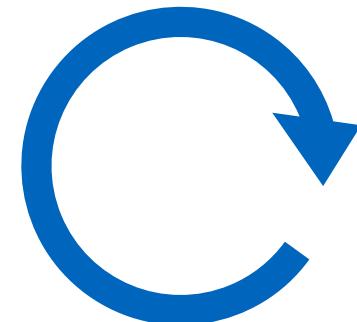


## Pre-processing

1. Image gradients
2. Entropy images
3. Phase
4. Multi-resolution
5. Attribute vectors

## Registration Framework

Similarity Measure



Optimization

# Image Gradients

- Normalized gradient fields

$$n(X) := \begin{cases} \frac{\nabla X}{\|\nabla X\|}, & \text{if } \nabla X \neq 0 \\ 0, & \text{otherwise} \end{cases}$$

*“two images are considered similar, if intensity changes occur at the same locations”*

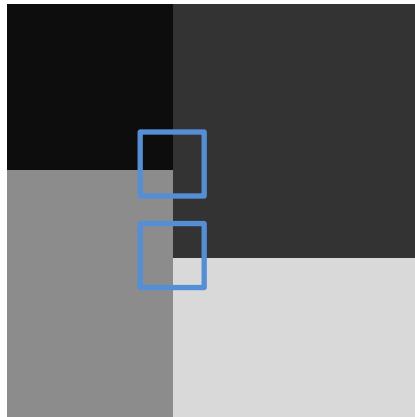
- Maximize square of cosine

$$d(X, Y) = \langle n(X), n(Y) \rangle^2$$

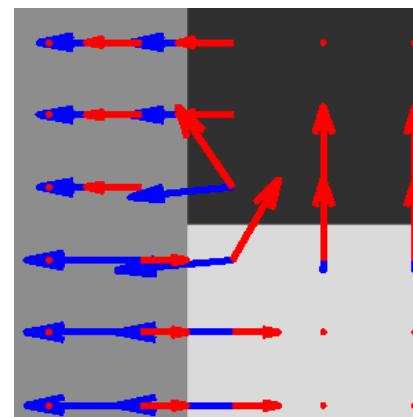
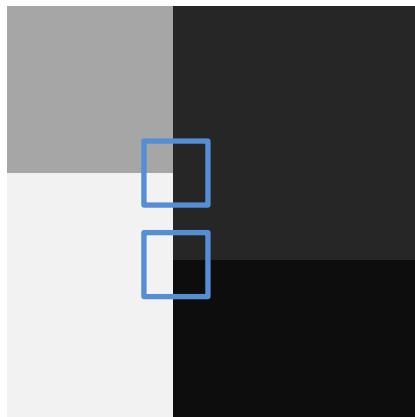
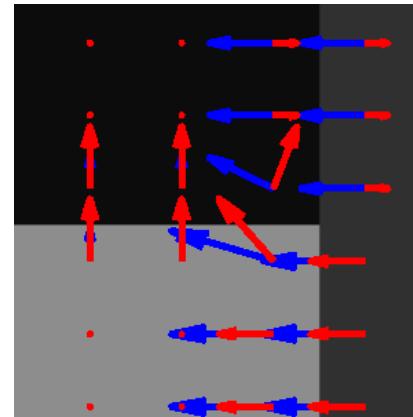
Haber, E. and Modersitzki, J., *Intensity gradient based registration and fusion of multi-modal images*, MICCAI 2006

# Image Gradients

Synthetic images

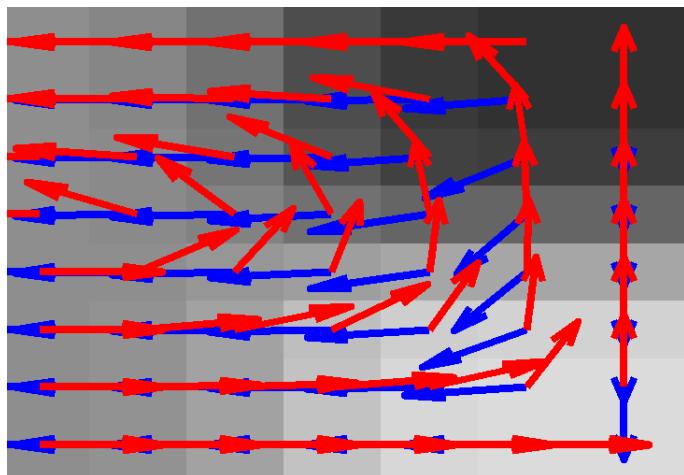
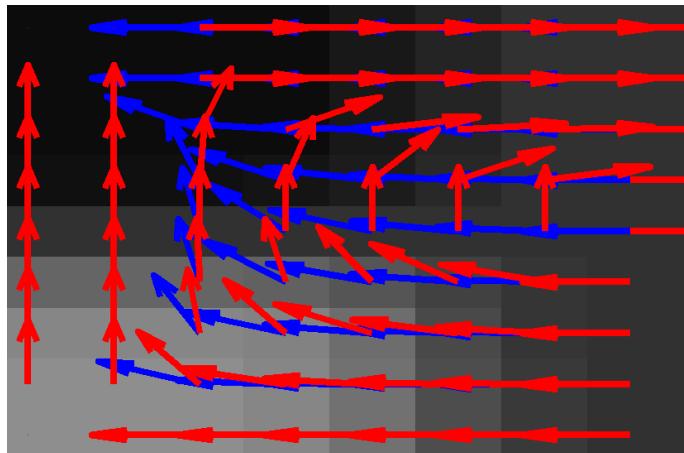


Gradient images

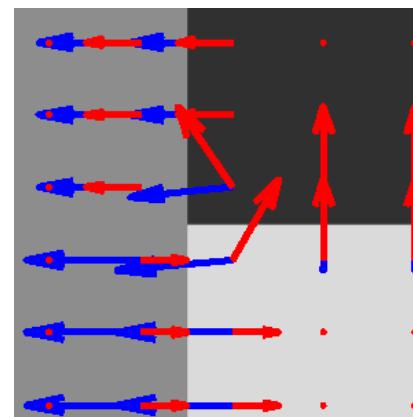
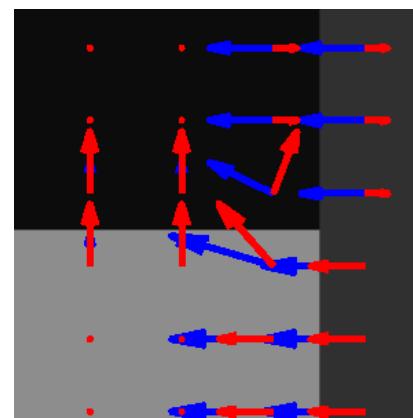


# Image Gradients

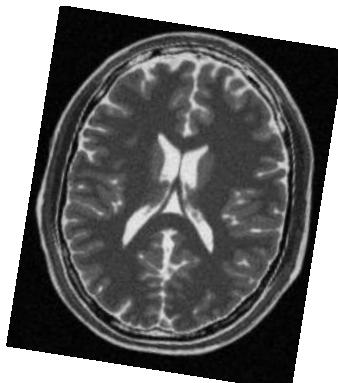
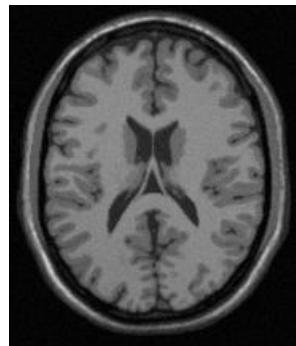
Gradient of smoothed images



Gradient images



## Images

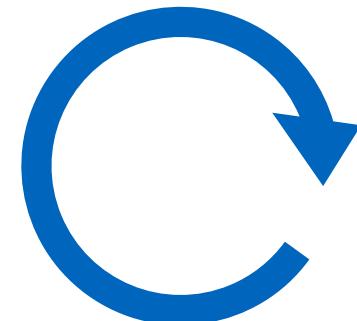


## Pre-processing

1. Image gradients
2. Entropy images
3. Phase
4. Multi-resolution
5. Attribute vectors

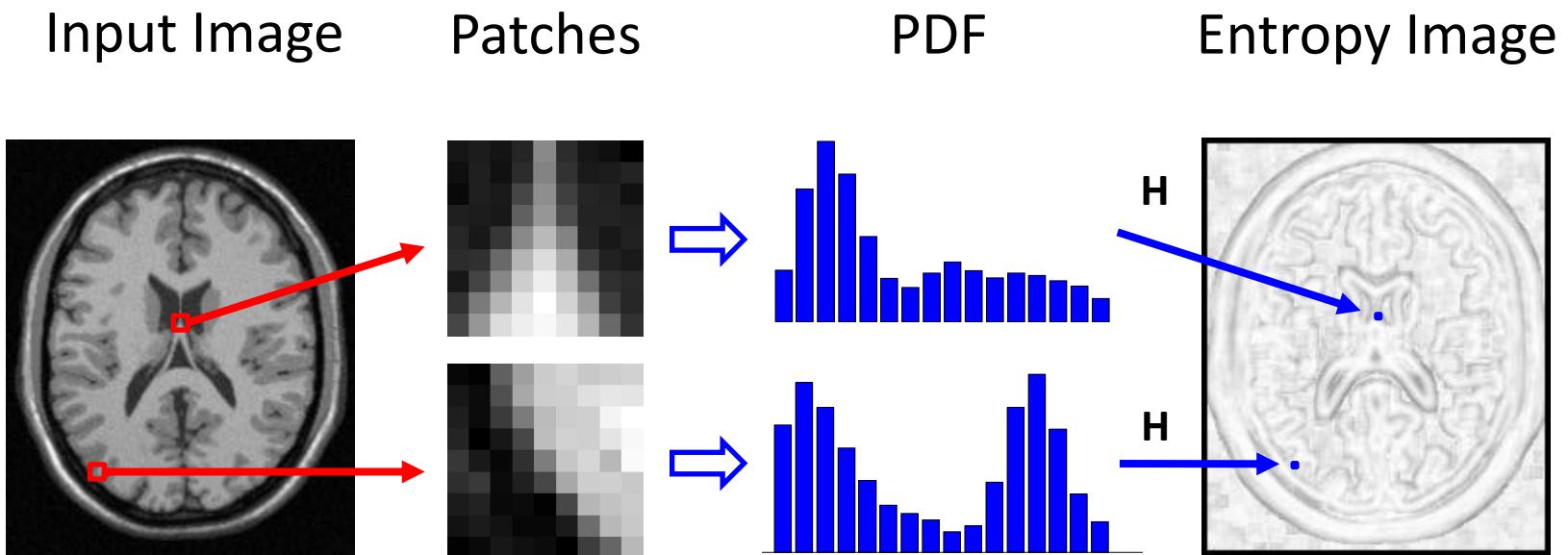
## Registration Framework

Similarity Measure



Optimization

# Entropy Images

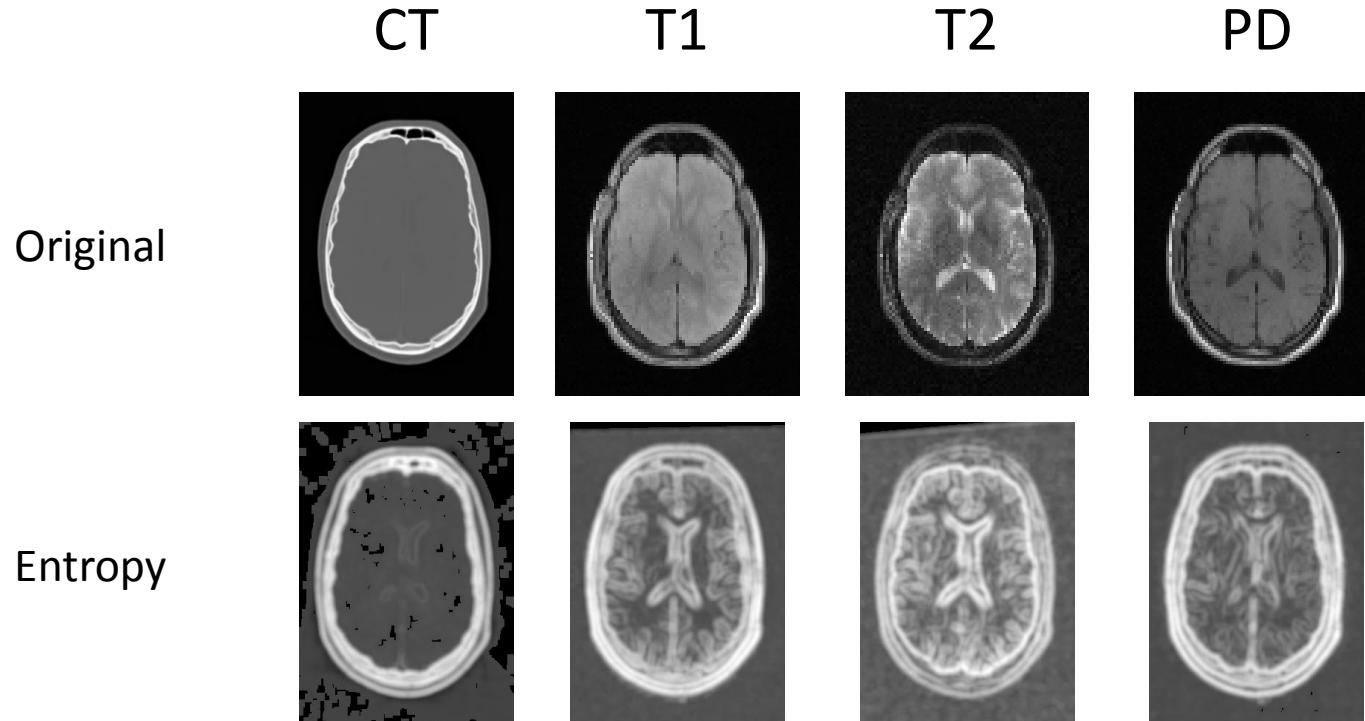


Wachinger, Navab, *Structural Images for Image Registration*, MMBIA, 2010.

# Entropy Images

- Examples from the RIRE dataset

<http://www.insight-journal.org/rire/>



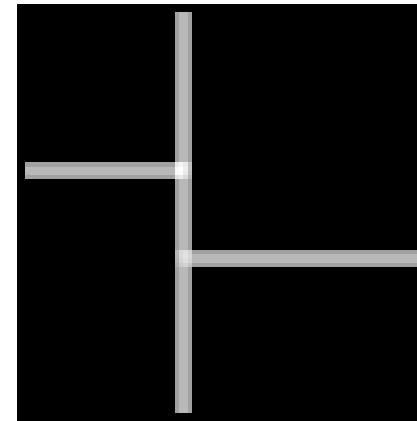
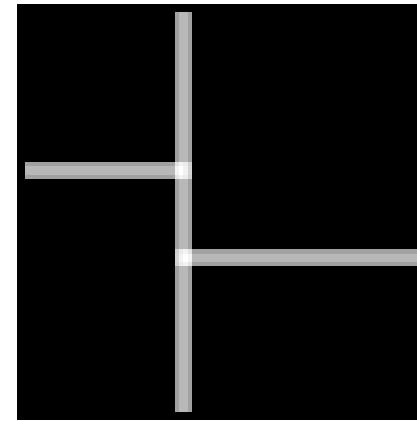
Wachinger, Navab, *Structural Images for Image Registration*, MMBIA, 2010.

# Entropy Images

Synthetic images



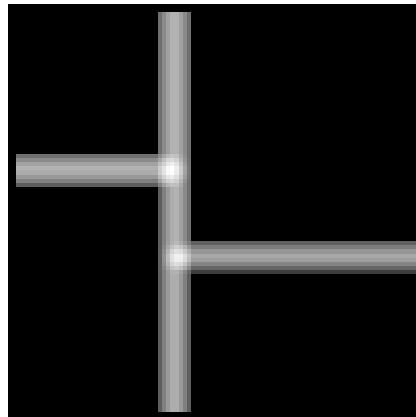
Entropy images



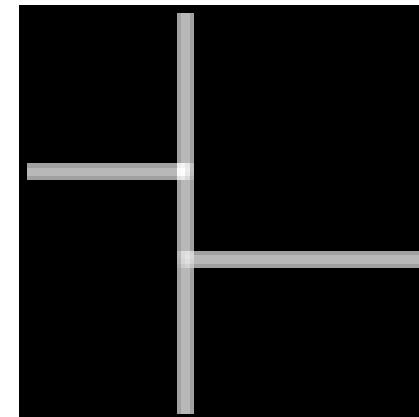
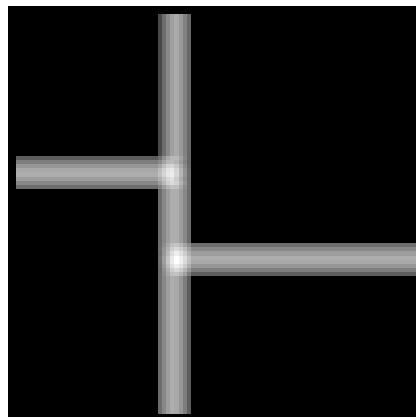
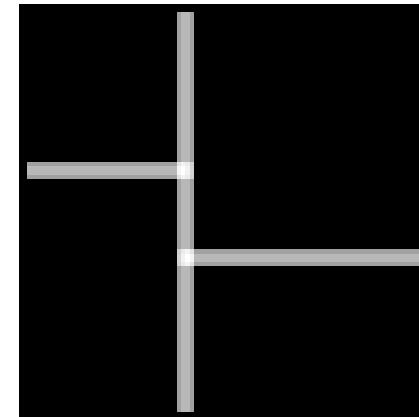
Wachinger, Navab, *Structural Images for Image Registration*, MMBIA, 2010.

# Entropy Images

Entropy on smoothed images

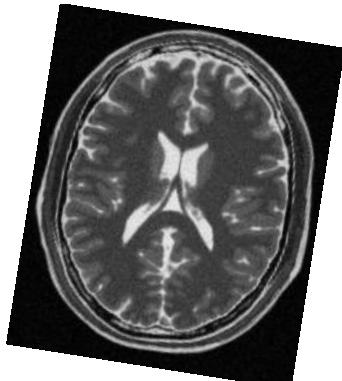


Entropy images



Wachinger, Navab, *Structural Images for Image Registration*, MMBIA, 2010.

## Images

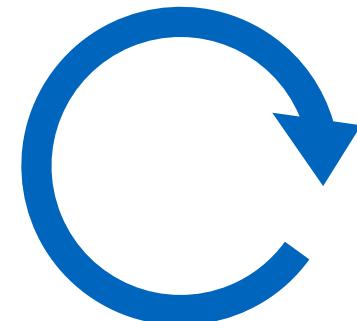


## Pre-processing

1. Image gradients
2. Entropy images
3. Phase
4. Multi-resolution
5. Attribute vectors

## Registration Framework

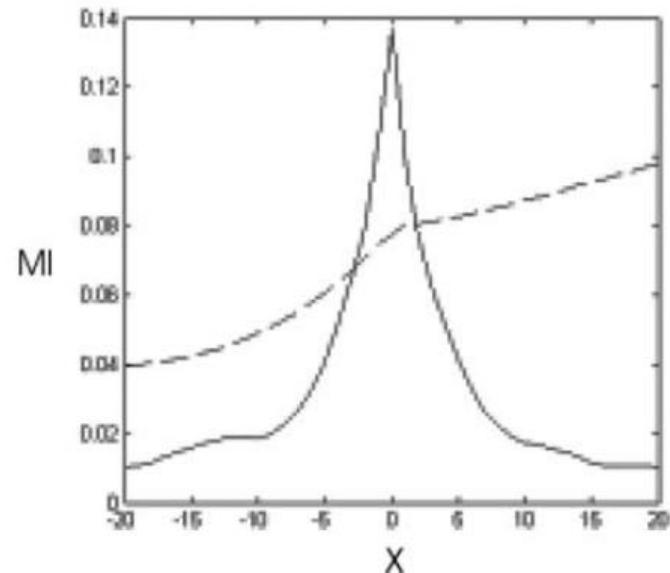
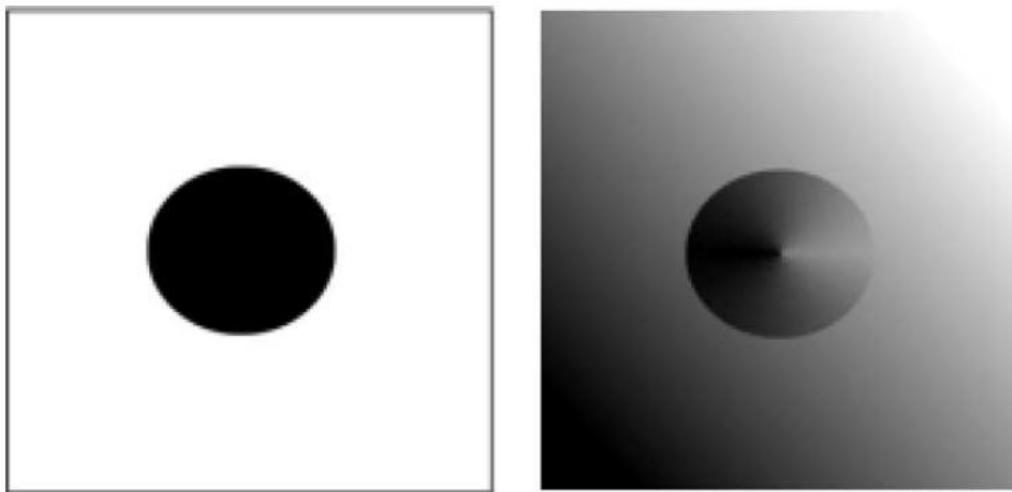
Similarity Measure



Optimization

# Phase-base registration

Dashed: MI  
Solid: Phase MI



- Fourier representation

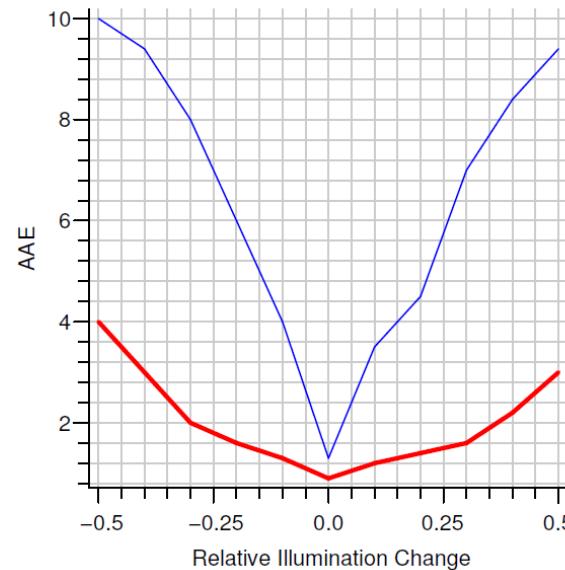
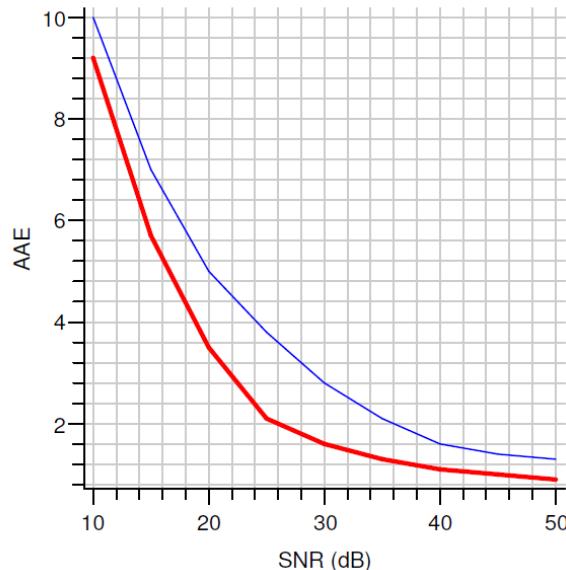
$$f(x) = \int_0^{\infty} A(\omega) \cdot \sin(\omega x + \phi(\omega)) d\omega$$

$A(\omega)$  : Amplitude  
 $\phi(\omega)$  : Phase

M. Mellor and M. Brady. *Phase mutual information as a similarity measure for registration*. Medical Image Analysis, 2005.

# Optical Flow with Phase

- Idea: replace the assumption of brightness constancy with phase constancy
- Error for chaning noise and illumination



blue: gradient  
red: phase

L. Wietzke, G. Sommer, O. Fleischmann, *The Geometry of 2D Image Signals*, CVPR, 2009

## Images

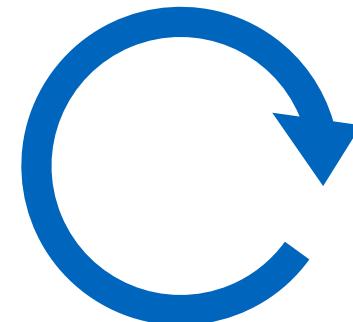


## Pre-processing

1. Image gradients
2. Entropy images
3. Phase
4. Multi-resolution
5. Attribute vectors

## Registration Framework

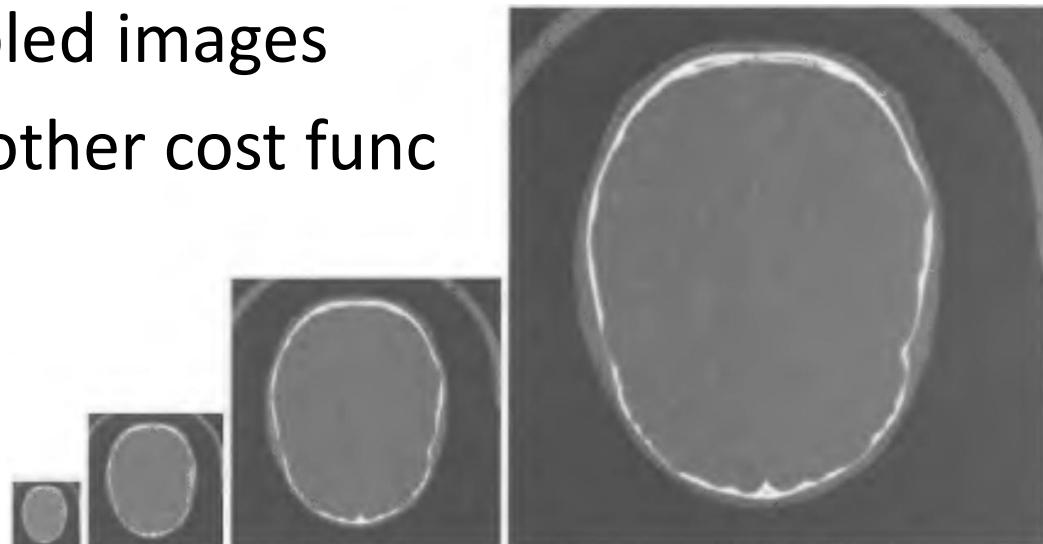
Similarity Measure



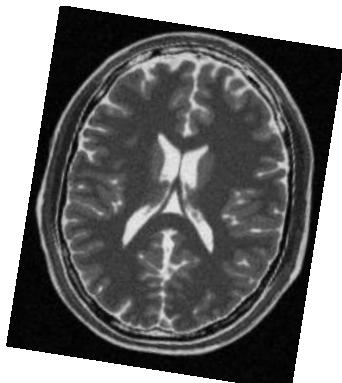
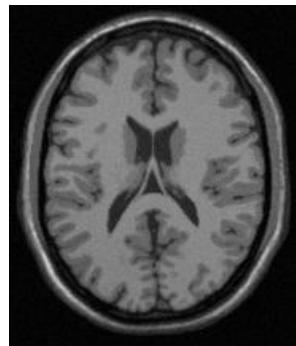
Optimization

# Multi-Resolution Registration

- Perform registration on multiple resolutions
  1. Smooth
  2. Downsample
- Advantages:
  - Speed: down-sampled images
  - Convergence: smoother cost func



## Images

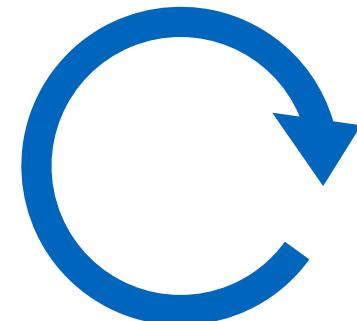


## Pre-processing

1. Image gradients
2. Entropy images
3. Phase
4. Multi-resolution
5. Attribute vectors

## Registration Framework

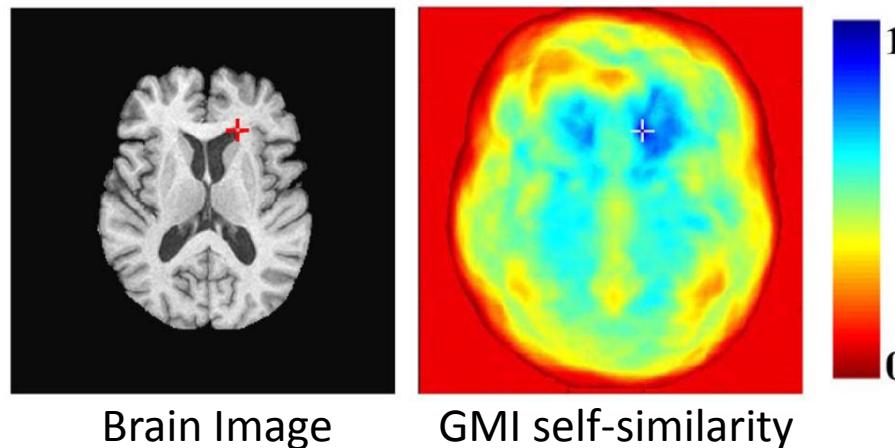
Similarity Measure



Optimization

# Attribute Vectors

- Registration with attribute vectors  
 $a = [ \text{gradient}, \text{intensity}, \text{GMI} ]$
- GMI = geometric moment invariants



Shen, D. and Davatzikos, C., *HAMMER: hierarchical attribute matching mechanism for elastic registration*, IEEE Transactions on Medical Imaging, 2002

M. Wacker and F. Deinzer. *Automatic robust medical image registration using a new democratic vector optimization approach with multiple Measures*, MICCAI, 2009

# **PART IV**

## **Recent Approaches**

# Recent examples...

- mainly compare to MI as state of the art
- address the following problems of MI
  - intensity does not represent tissue (e.g. US-CT)



CT



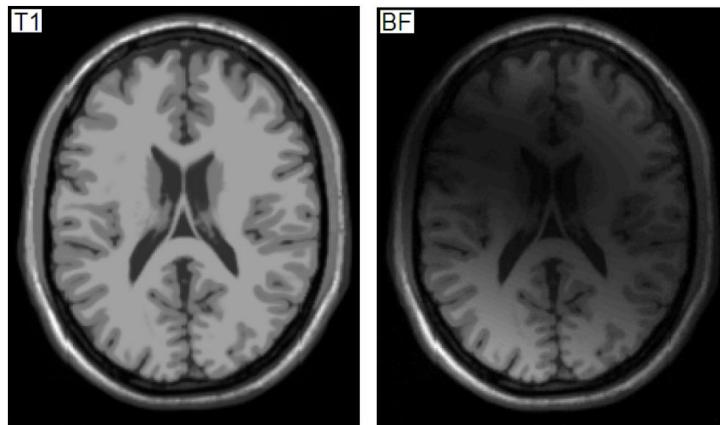
US



US-CT

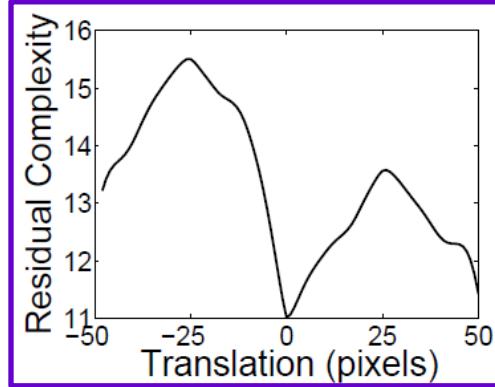
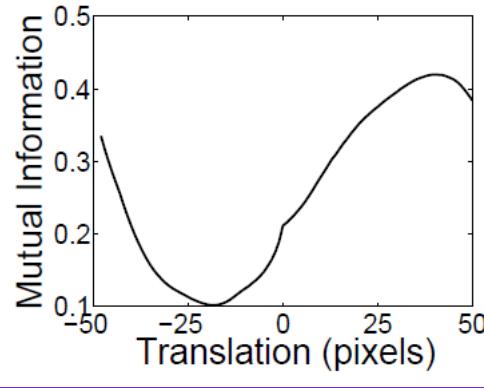
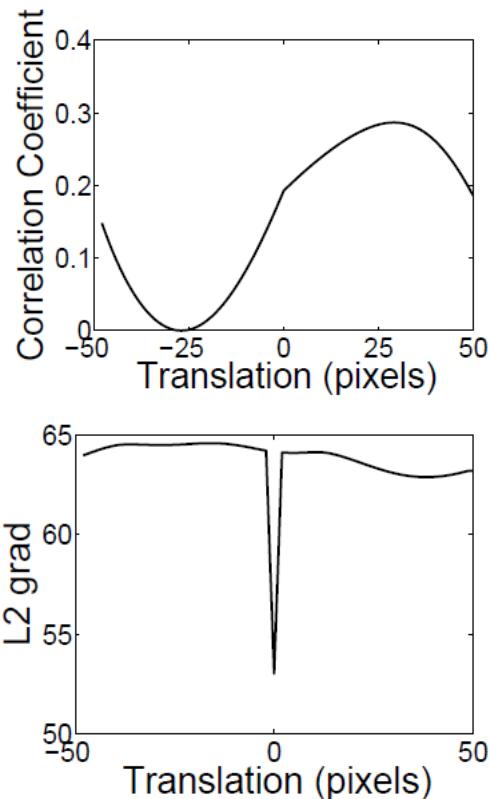
# Recent examples...

- mainly compare to MI as state of the art
- address the following problems of MI
  - intensity does not represent tissue (e.g. US-CT)
  - intensity non-uniformity (bias)



# Minimizing Residual Complexity

- Motivation: MI fails for these images



Model

$$X = Y(T) + S + \varepsilon$$

Myronenko A., Song X.: "Intensity-based Image Registration by Minimizing Residual Complexity", IEEE Trans. on Medical Imaging, 2010

# Minimizing Residual Complexity

- Compress the difference image

$$\sum_{i=1}^N \mathbf{q}_i \cdot (X - Y) \quad \mathbf{q}_i : i\text{-th DCT basis function}$$

- Similarity measure

$$RC(X, Y) = \sum_{i=1}^N \log \left( \frac{(\mathbf{q}_i(X - Y))^2}{\alpha} + 1 \right)$$

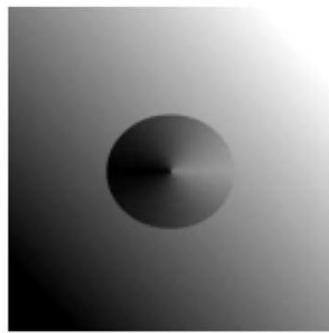
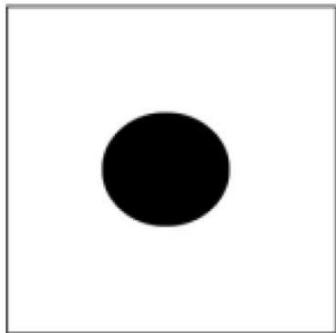
- Related to entropy of difference image

$$H(X - Y)$$

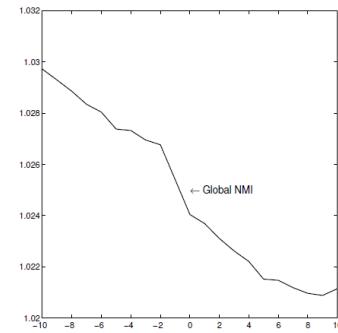
Buzug et al.: *Image registration: convex weighting functions*, CVRMed, 1997

Myronenko A., Song X.: "Intensity-based Image Registration by Minimizing Residual Complexity", IEEE Trans. on Medical Imaging, 2010

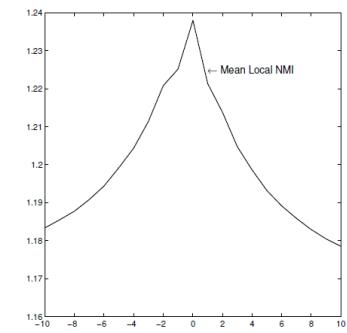
# Local and Global Statistics



global



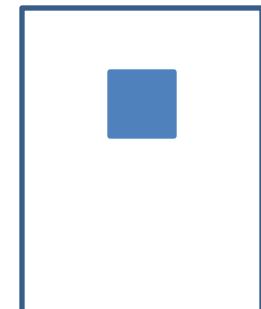
local



$\text{NMI}_g$

$\text{NMI}_l(x)$

$$\text{NMI}_{l+g} = \frac{1}{|\Omega|} \sum_{x \in \Omega} (1 - \alpha_x) \cdot \text{NMI}_g + \alpha_x \cdot \text{NMI}_l(x)$$



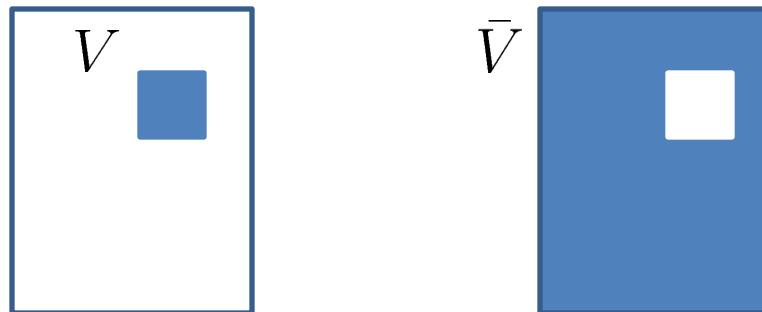
$$\alpha_x = \text{NMI}_l(x) - 1$$

Z. Yi and S. Soatto. *Nonrigid Registration Combining Global and Local Statistics*, CVPR, 2009.

# Local and Global Statistics

- Combining local and global densities

$$P(X, Y) = \alpha \cdot P_V(X, Y) + (1 - \alpha) \cdot P_{\bar{V}}(X, Y)$$



- $\alpha$ : manually, emphasize on local or global influence

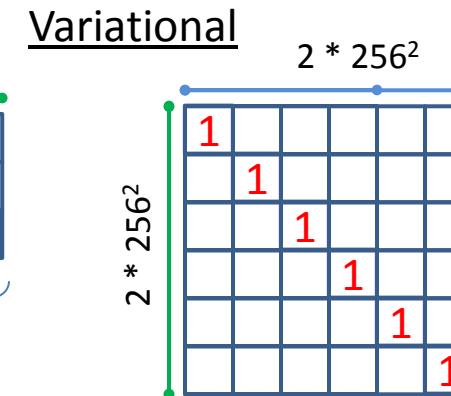
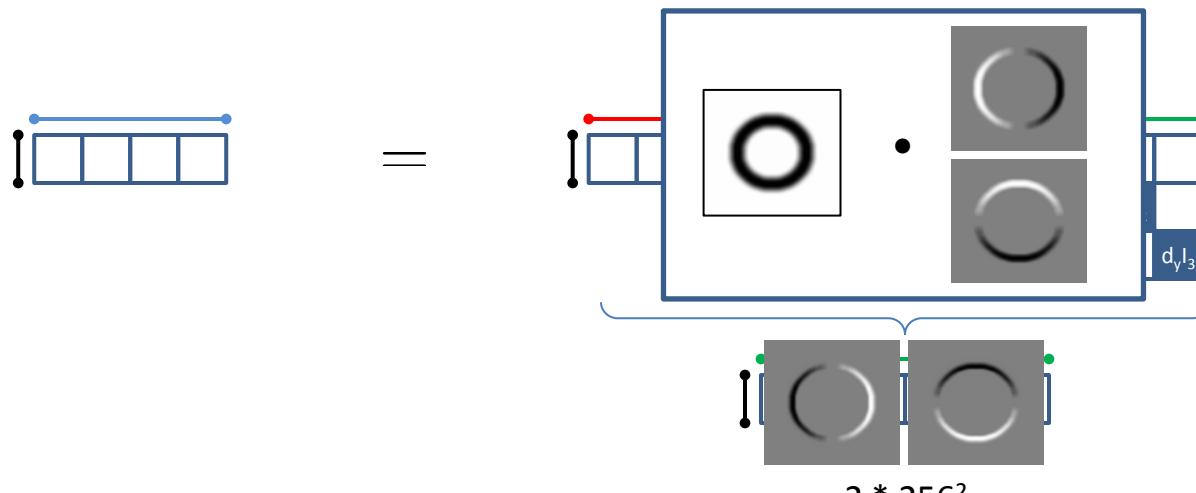
Zhuang, X. and Hawkes, D. and Ourselin, *Unifying Encoding of Spatial Information in Mutual Information for Nonrigid Registration*, IPMI, 2009  
Loeckx, Slagmolen, Maes, Vandermeulen, Suetens, *Nonrigid image registration using conditional mutual information*, IPMI, 2007

# PART V

## Linear vs. Non-Linear Registration

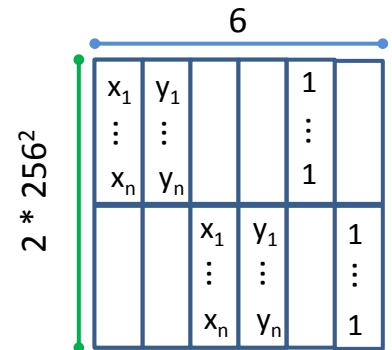
# Linear vs. Non-Linear Registration

$$\frac{\partial \text{SM}(X, Y(T_p))}{\partial p} = \frac{\partial \text{SM}(X, Y)}{\partial Y} \frac{\partial Y}{\partial T_p} \frac{\partial T_p}{\partial p}$$



Affine

$$\begin{pmatrix} a_1 & a_2 \\ a_3 & a_4 \end{pmatrix} \begin{pmatrix} x \\ y \end{pmatrix} + \begin{pmatrix} t_x \\ t_y \end{pmatrix}$$



- Deformable registration is more sensitive to errors from poor modeling
- Averaged out in linear registration

# Linear vs. Non-Linear Registration

- Choice of similarity measure has influence on the optimization strategy

EXAMPLE: Gauß-Newton Optimization with SSD

$$\left( \begin{array}{ccc|ccc} \nabla I_S^x & \nabla I_S^x & & \nabla I_S^x & \nabla I_S^y & \\ & \nabla I_S^y & \nabla I_S^x & & \nabla I_S^y & \nabla I_S^z \\ & & \nabla I_S^z & \nabla I_S^x & & \nabla I_S^z & \nabla I_S^z \end{array} \right) - \lambda \left( \begin{array}{ccc} \Delta & & \\ & \Delta & \\ & & \Delta \end{array} \right) = \left( \begin{array}{c} h_x \\ h_y \\ h_z \end{array} \right) = \left( \begin{array}{c} f_x \\ f_y \\ f_z \end{array} \right)$$

$$J_e^\top J_e$$

$$h = f$$

# Linear vs. Non-Linear Registration

- Overlap invariance only for linear ?
  - MI vs. NMI
  - Joint Entropy vs. MI
- Transition between linear and non-linear
  - Parameterization
  - Regularization

# Summary

- A large variety of concepts for measuring the similarity is available in the literature
- Choose similarity measure that best fits the application
- Appropriate choice is more important for deformable registration

**END**