

# Is Learning Based Image Registration Really the Future?

Jamie McClelland

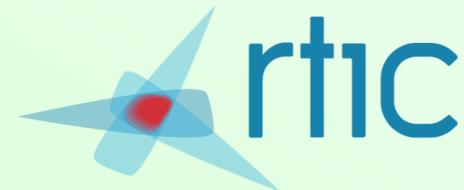
Radiotherapy Image Computing (RTIC) group

Centre for Medical Image Computing (CMIC)

Wellcome/EPSRC centre Interventional and Surgical Science (WEISS)

Dept. Medical Physics and Biomedical Engineering

University College London



# Medical Image Registration: Past, Present, and Future...

Jamie McClelland

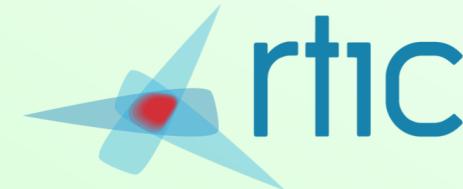
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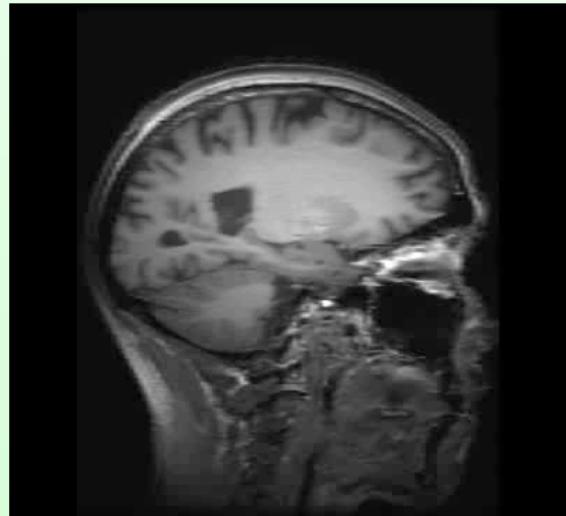


# Overview

- Past: Classical image registration
- Present: Learning-based image registration
- Future: ???

# What is Image Registration?

- Process of finding a spatial transformation that optimally aligns image A to image B
  - Image A called Source (or floating or moving) image
  - Image B called Target (or reference or fixed) image



MRI Target image (B)  
PET Source image (A)

# What is Image Registration?

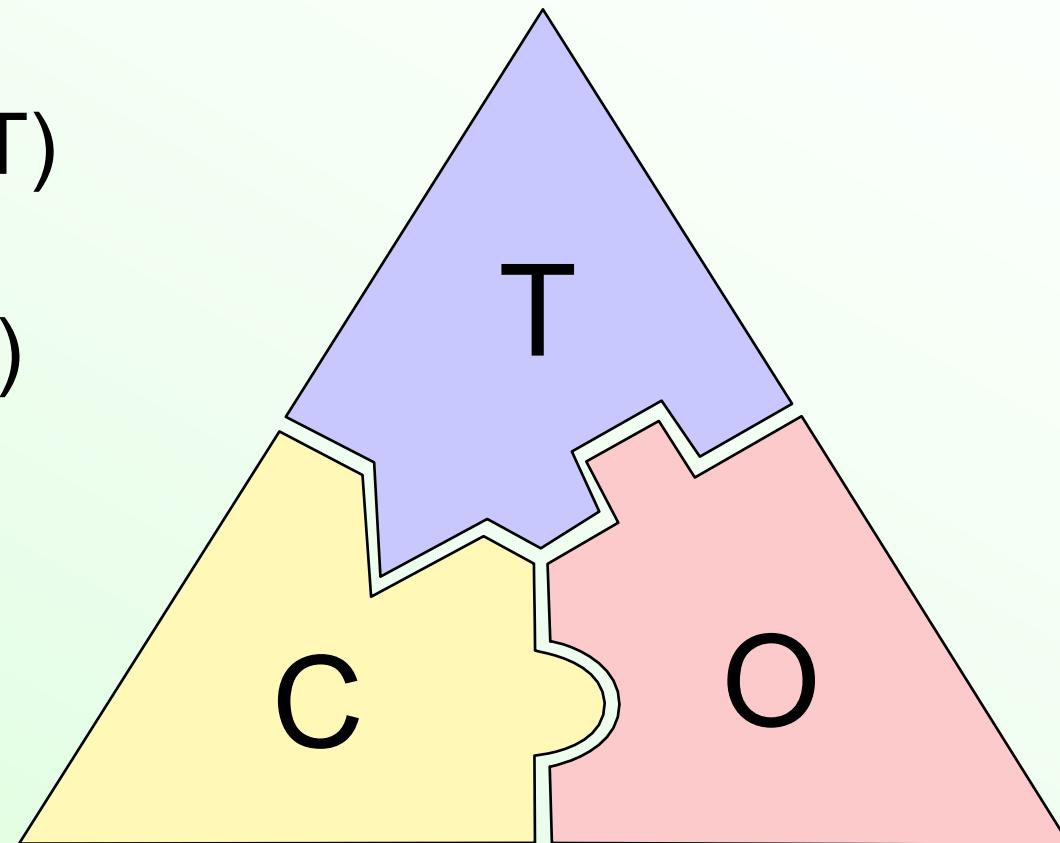
- What do we mean by 'aligned'?
  - Establishing correspondence
  - Not always obvious what 'correct' correspondence is
    - Images contain different information,
      - e.g. MRI and PET
    - Anatomy has changed between images,
      - e.g. before and after surgery
    - Images are from different subjects

# Components of a registration algorithm

Transformation (T)

Cost Function (C)

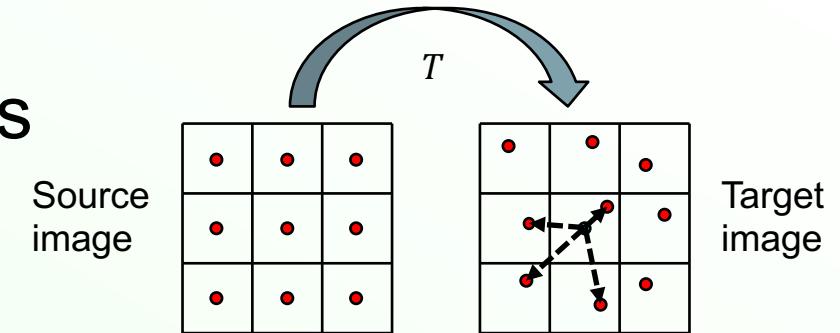
Optimiser (O)



# What is a transformation?

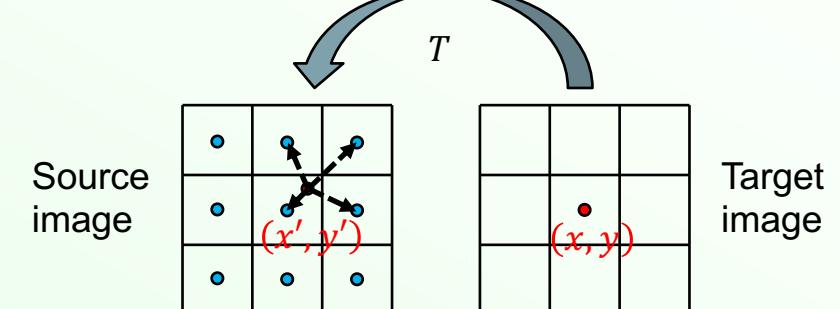
- Spatial mapping between coordinate systems

$$(x', y', z') = T(x, y, z)$$
$$(\vec{x}') = T(\vec{x})$$



- Used to resample (warp) the source image into the space of the target image

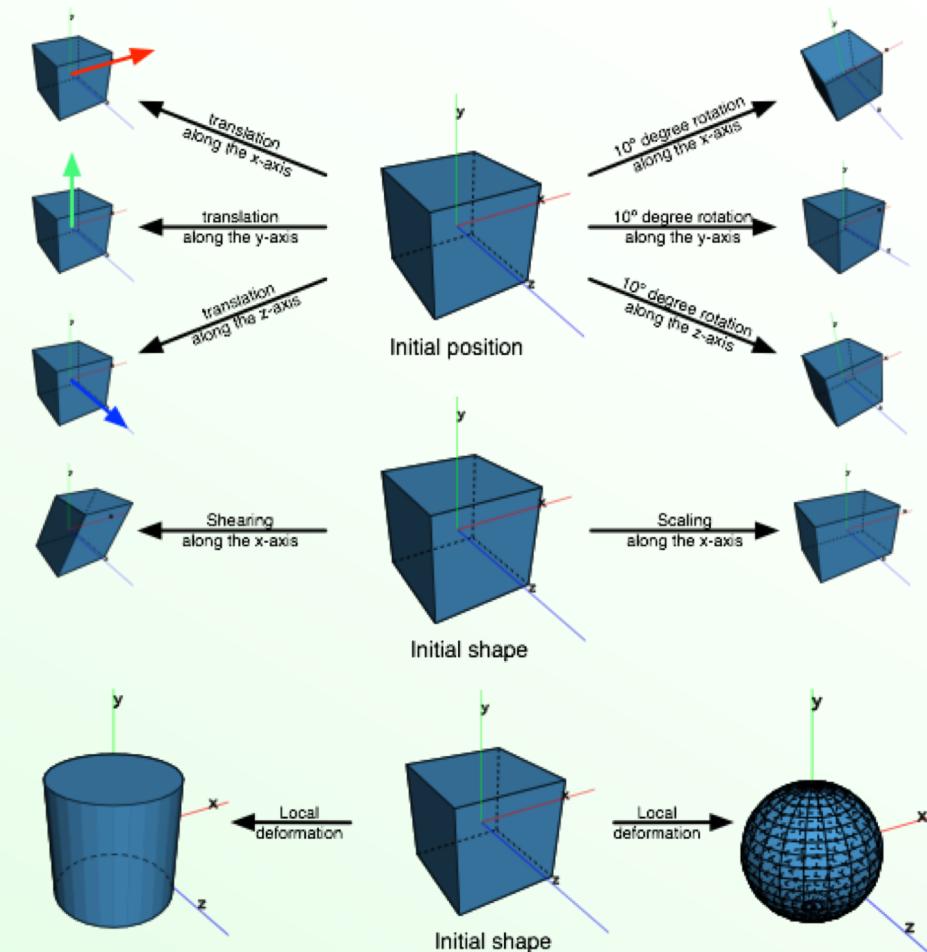
- Requires interpolation
- Transformation usually maps from target to source image



- Does NOT necessarily represent physical changes between images

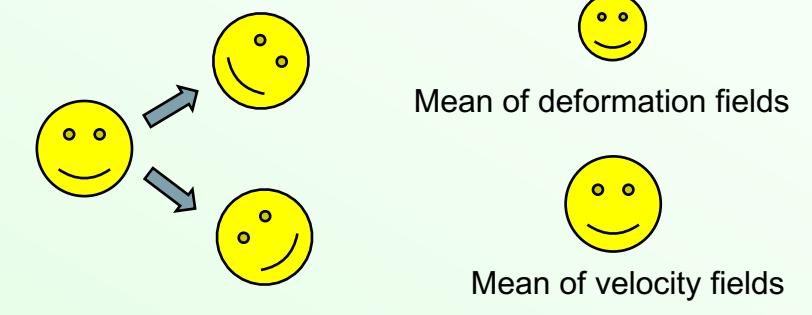
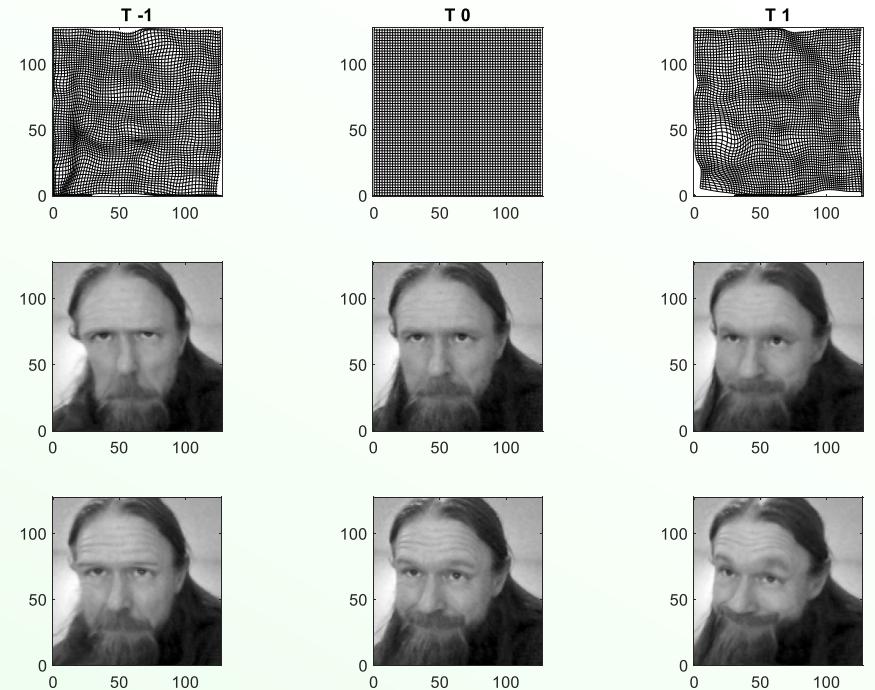
# Types of transformation

- Rigid
  - Translation, rotation
- Affine
  - Rigid, shearing, scaling
- Deformable (non-rigid, non-linear)
  - Free-form deformations
  - ‘Non-parametric’ (voxel-based)
  - Velocity fields
    - Often used for diffeomorphic transformations
- 2D-2D, 3D-3D, 4D-4D, 3D-4D, 2D-3D,...



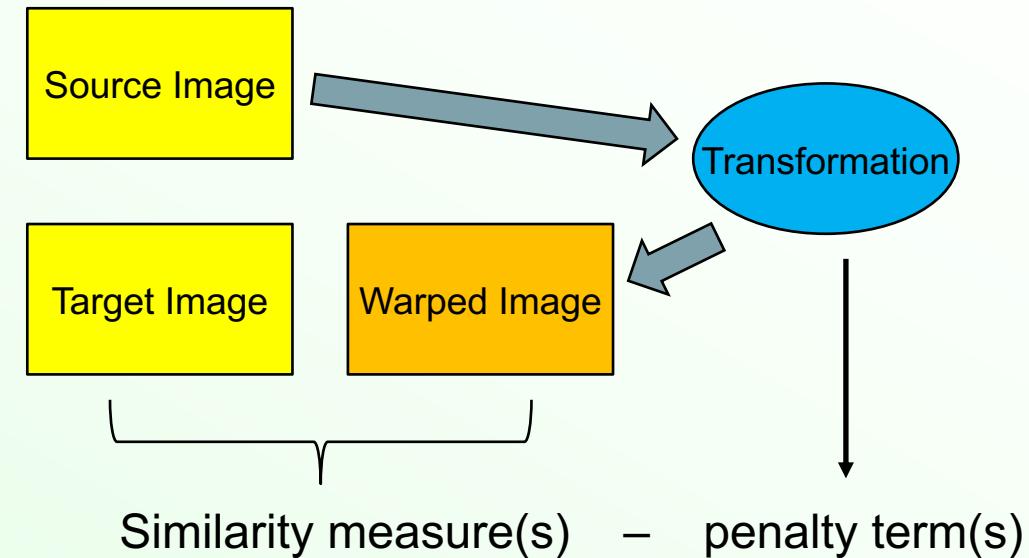
# What is a diffeomorphism?

- One-to-one mapping (bijective)
  - Transformation has an inverse
  - No folding
- Transformation and inverse are differentiable
  - No discontinuities
- Nice mathematical properties
  - Topology is preserved
    - Structures preserved
  - Composing diffeomorphisms gives a diffeomorphism
  - Statistics on velocity fields (Log-Euclidean framework)



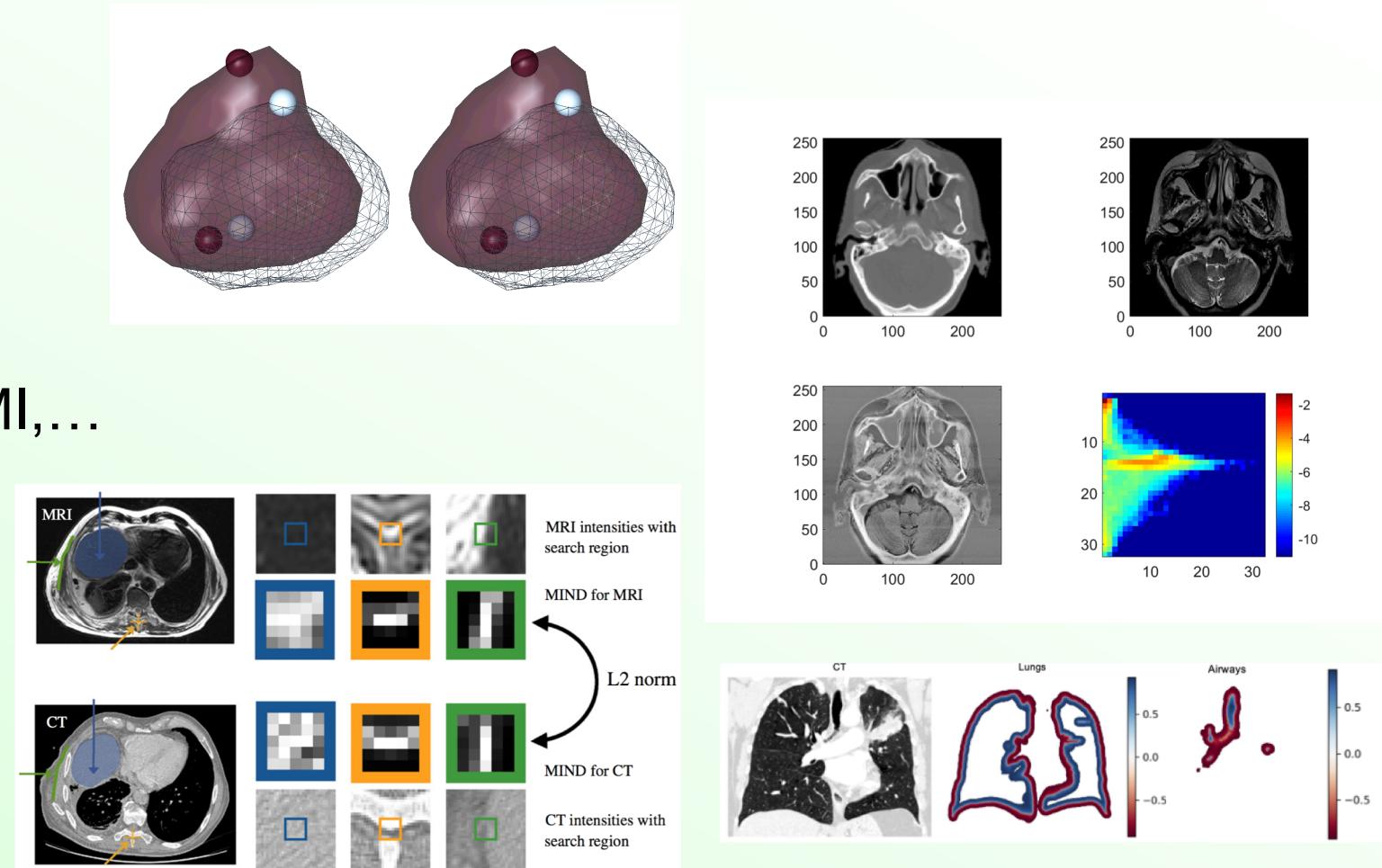
# What is a cost function?

- Used to measure how good the registration result is
- Composed of two parts:
  - One or more similarity measures
    - Function of target image and warped source image
    - Choice will depend on type of images being registered
  - Zero or more penalty (constraint) terms
    - Function of transformation parameters
    - Penalises undesirable transformations
      - Encourages plausible/~~realistic~~ transformations.
    - Different terms used depending on what is considered desirable behaviour
  - Often only one similarity measure and zero-to-two penalty terms are used



# Similarity measures

- Point/surface based
- Voxel-wise
  - SSD, MSD, NCC, MI, NMI,...
  - Locally normalised
    - LNCC, LNMI
  - Feature/structure based
    - MIND, distance maps
- Hybrid



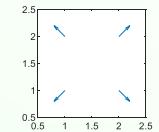
# Penalty terms

- Using 1<sup>st</sup> order derivative
  - Diffusion (L2 norm), Linear energy,
  - **L1 norm**
  - Log of Jacobian Determinant

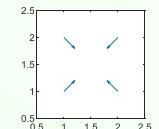
$$\text{Jac}(T(\vec{x}_n)) = \begin{bmatrix} \frac{\partial T_1(\vec{x}_n)}{\partial x_1} & \dots & \frac{\partial T_1(\vec{x}_n)}{\partial x_{N_{dim}}} \\ \vdots & \ddots & \vdots \\ \frac{\partial T_{N_{dim}}(\vec{x}_n)}{\partial x_1} & \dots & \frac{\partial T_{N_{dim}}(\vec{x}_n)}{\partial x_{N_{dim}}} \end{bmatrix}$$

- Using 2<sup>nd</sup> order derivative
  - Bending energy
- Encourage transformations to be:
  - Smooth / **piece-wise smooth** / volume preserving

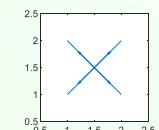
$$|\text{Jac}(T(\vec{x}_n))| > 1$$



$$1 > |\text{Jac}(T(\vec{x}_n))| > 0$$



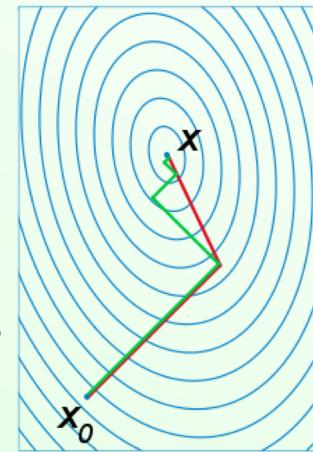
$$|\text{Jac}(T(\vec{x}_n))| \leq 0$$



# Optimisation

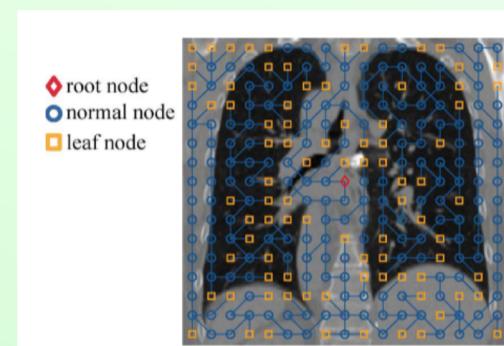
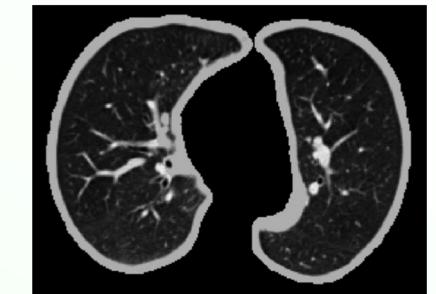
- Optimisation methods:

- Gradient based methods
  - Gradient descent/ascent
  - Conjugate gradient
  - Other gradient based methods
- Variational calculus
  - Partial differential equations
- Discrete methods
  - Graph based



- Approaches/strategies

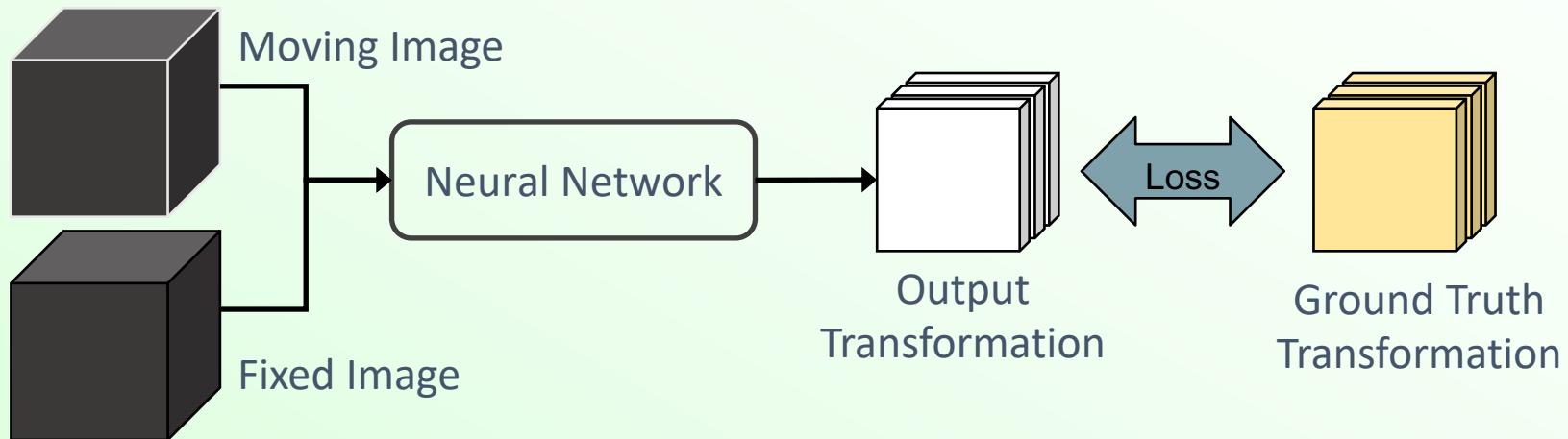
- Multi-resolution
- Cropping and masking images
- Symmetric registration
- Inverse-consistent registrations
- Group-wise registrations



# Learning based registrations

- Almost all current research on learning-based registrations
- Learn relationship between images and transformations from training data
  - Can calculate transformation very quickly for new images
  - Requires lots of images and computing resources for training
- Training can be:
  - Supervised, self-supervised, weakly supervised

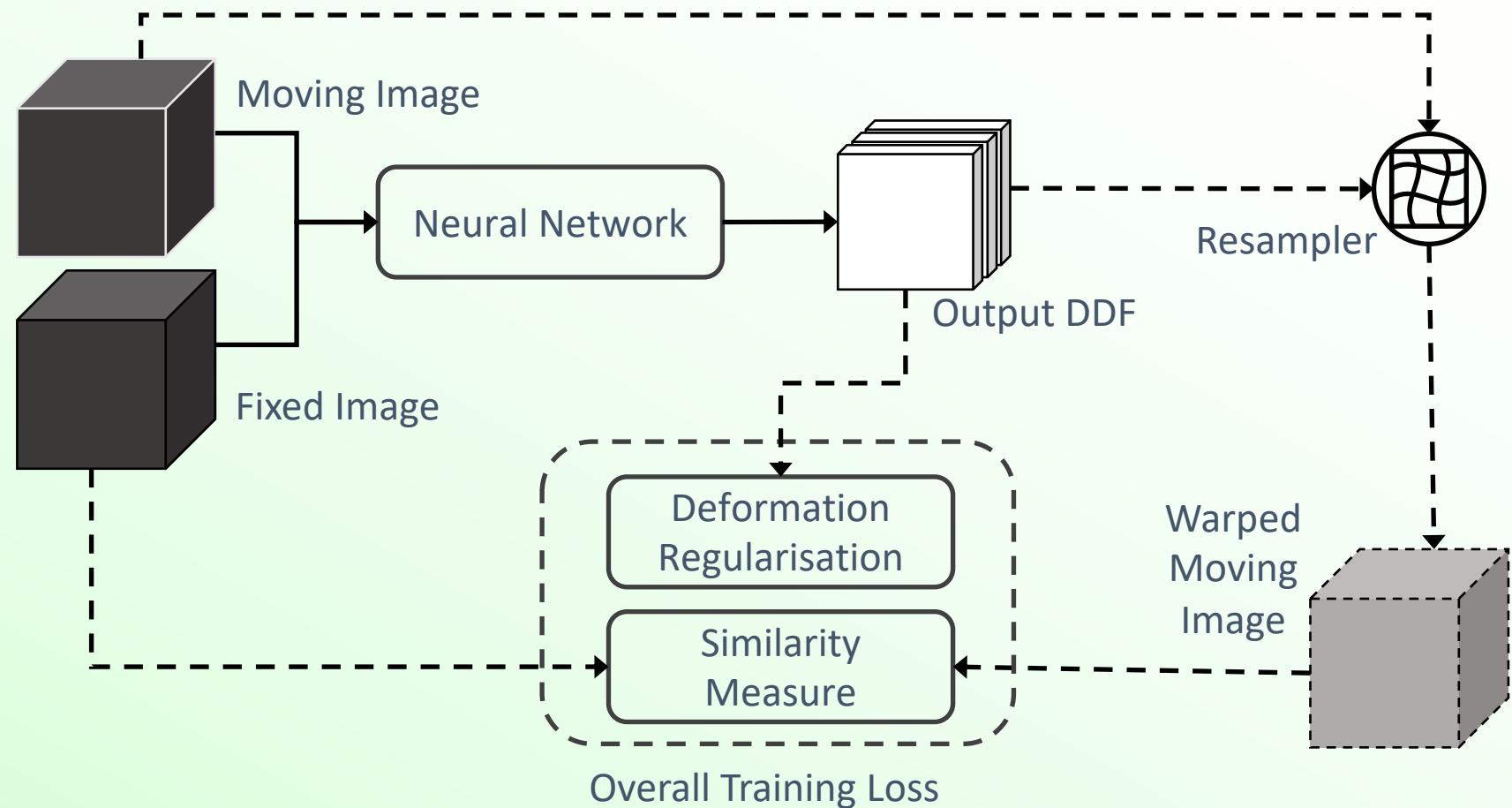
# Supervised training



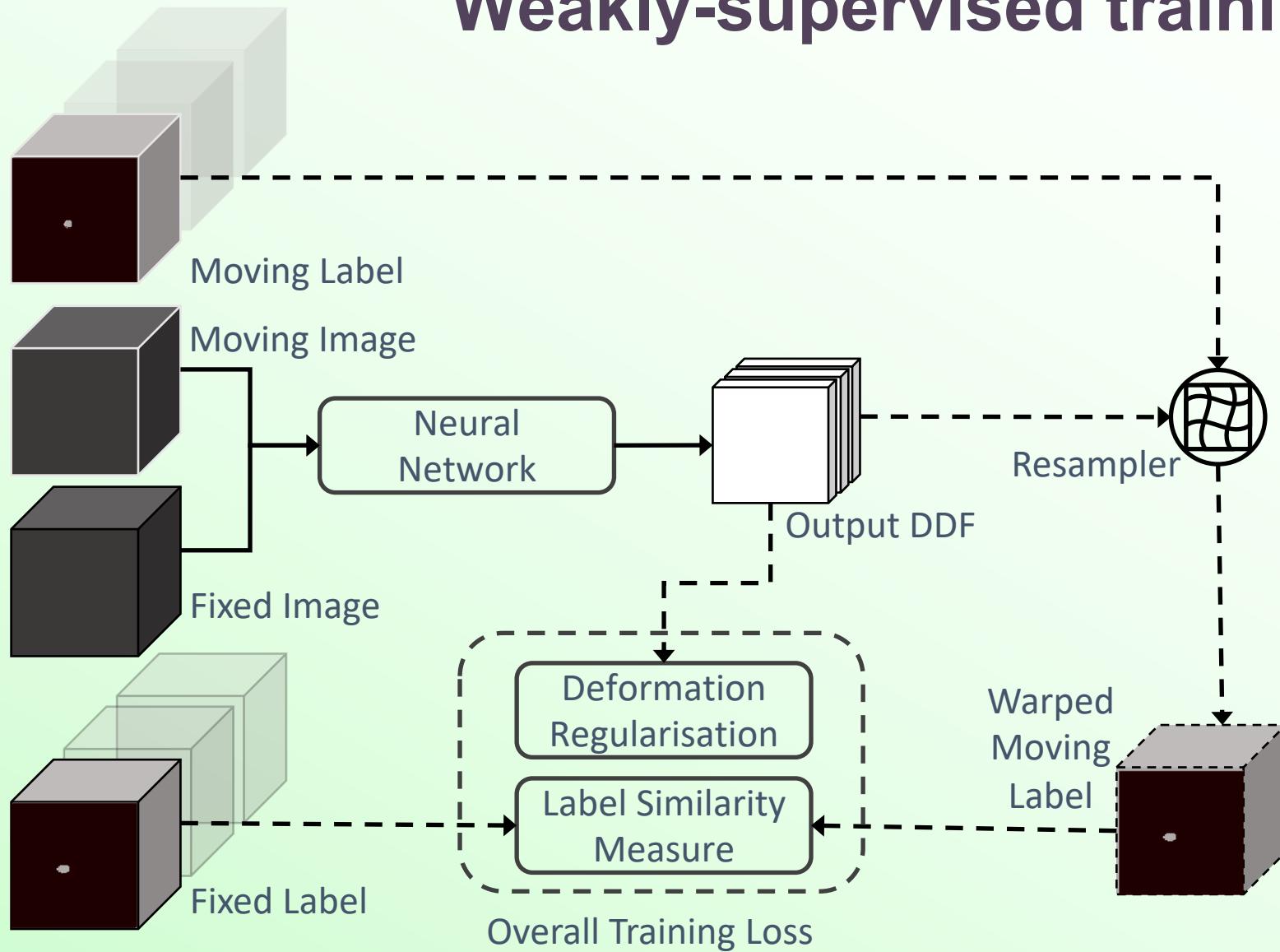
- Required ground truth transformation
  - Simulation, manual alignment, classical image registration

# Self-supervised (unsupervised) training

- Training loss utilises same terms as classical image registration
- Much current research
- Well known example:
  - Voxelmorph

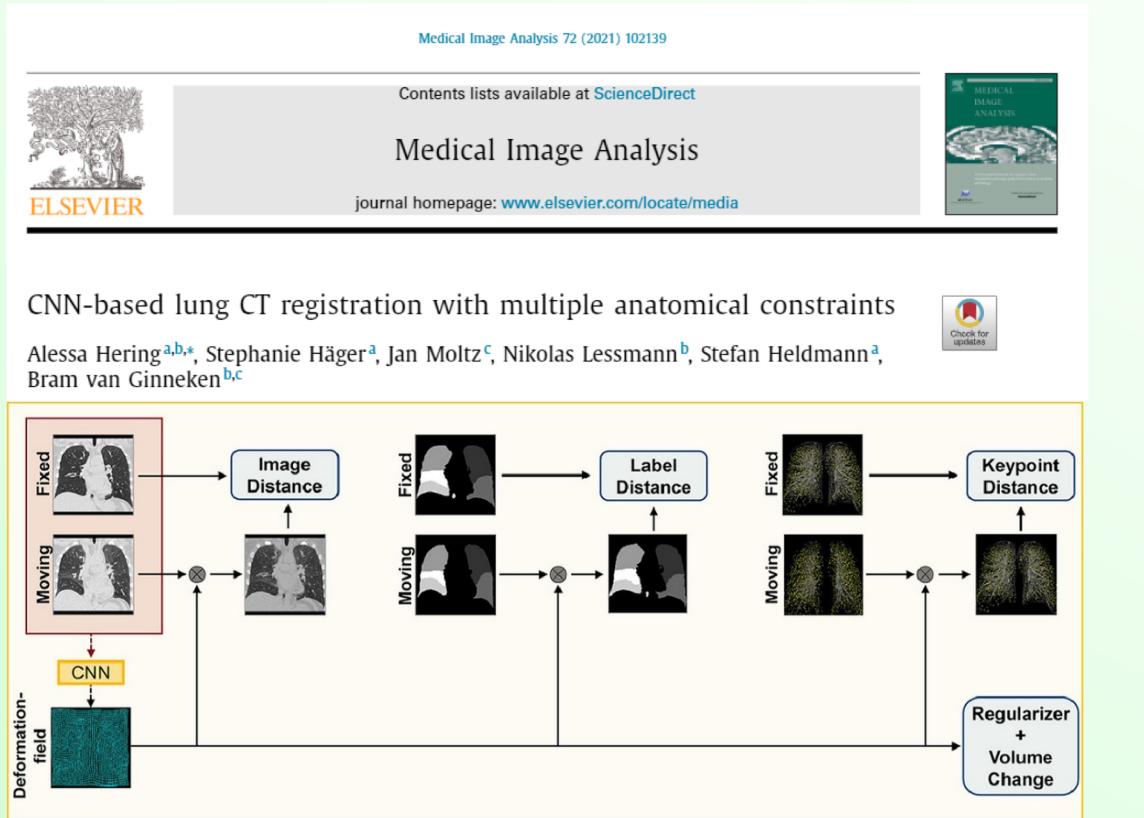


# Weakly-supervised training



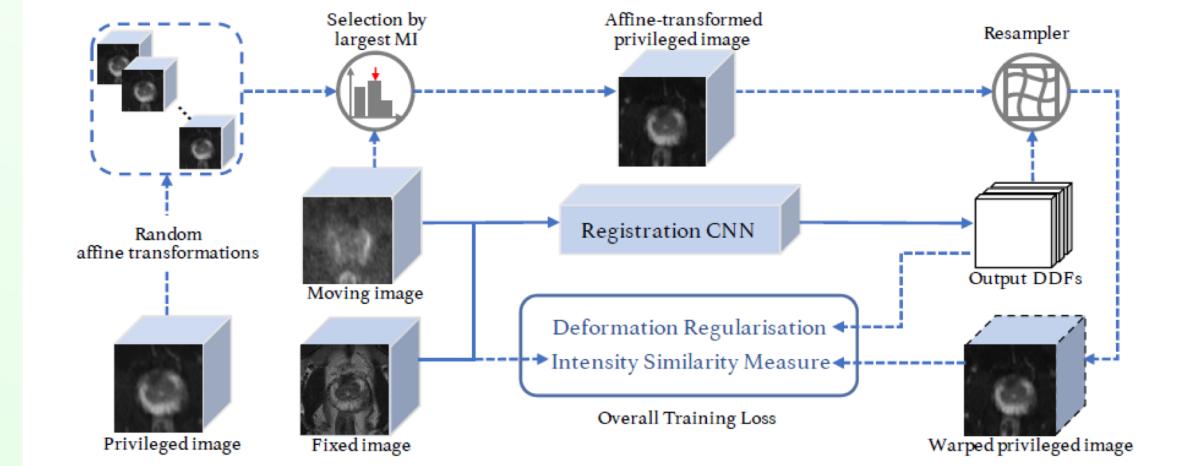
- Utilises privileged information during training:
  - Segmentations
  - Points
  - Images
  - ...

# Weakly-supervised training



## Cross-Modality Image Registration Using a Training-Time Privileged Third Modality

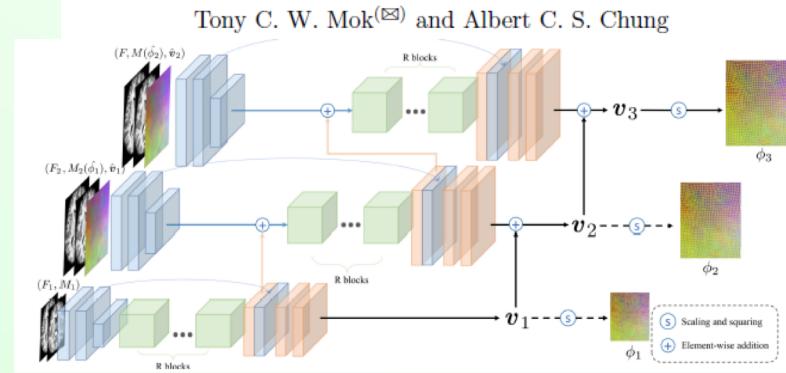
Qianye Yang <sup>b</sup>, David Atkinson <sup>b</sup>, Yunguan Fu <sup>b</sup>, Tom Syer <sup>b</sup>, Wen Yan <sup>b</sup>, Shonit Punwani <sup>b</sup>, Matthew J. Clarkson <sup>b</sup>, Dean C. Barratt <sup>b</sup>, Tom Vercauteren <sup>b</sup>, and Yipeng Hu <sup>b</sup>



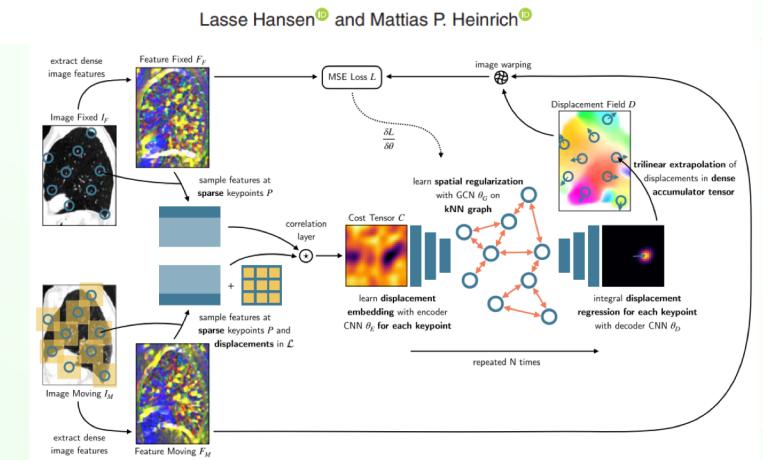
# Recent(ish) developments: Inspired by classical image registration

- Diffeomorphisms
  - Velocity fields
- Multiresolution
- Feature-driven
- Graph based
- 4D

## Large Deformation Diffeomorphic Image Registration with Laplacian Pyramid Networks

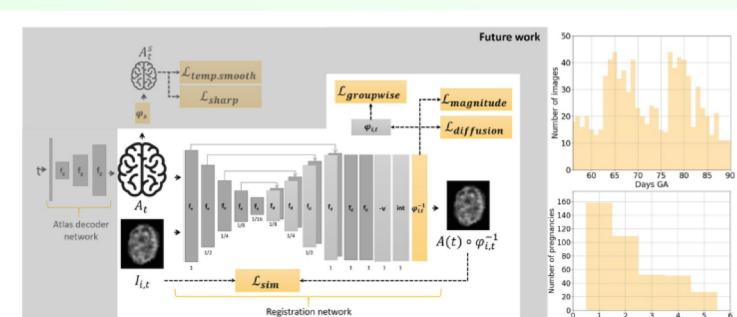


## GraphRegNet: Deep Graph Regularisation Networks on Sparse Keypoints for Dense Registration of 3D Lung CTs



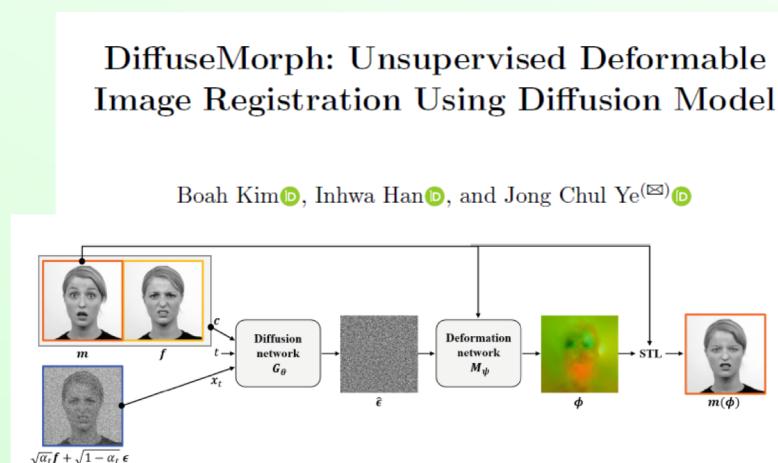
## Towards a 4D Spatio-Temporal Atlas of the Embryonic and Fetal Brain Using a Deep Learning Approach for Groupwise Image Registration

Wietske A. P. Bastiaansen<sup>1,2(✉)</sup>, Melek Rousian<sup>2</sup>,  
Régine P. M. Steegers-Theunissen<sup>2</sup>, Wiro J. Niessen<sup>1</sup>, Anton H. J. Koning<sup>3</sup>,  
and Stefan Klein<sup>1</sup>



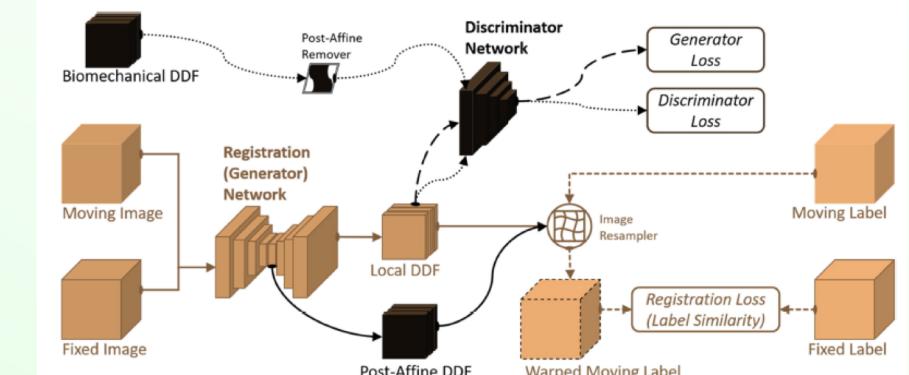
# Recent(ish) developments: Inspired by advances in DL

- GANs and adversarial learning
- Diffusion networks
- Vision transformers
- Uncertainty



## Adversarial Deformation Regularization for Training Image Registration Neural Networks

Yipeng Hu<sup>1,2(✉)</sup>, Eli Gibson<sup>1</sup>, Nooshin Ghavami<sup>1</sup>, Ester Bonmati<sup>1</sup>,  
Caroline M. Moore<sup>3</sup>, Mark Emberton<sup>3</sup>, Tom Vercauteren<sup>1</sup>,

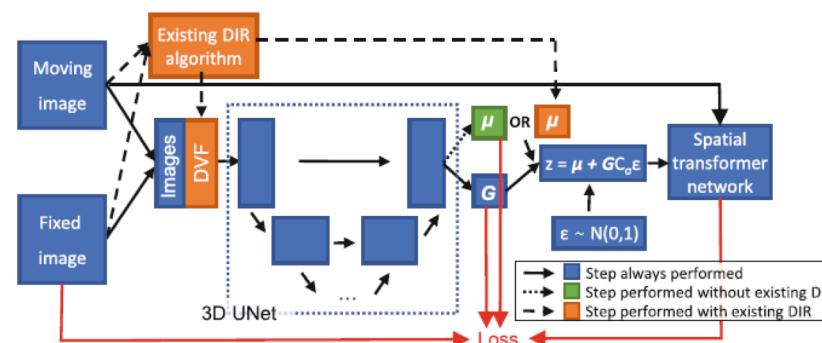


# Recent(ish) developments: Inspired by advances in DL

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- Diffusion networks
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- Uncertainty

## Deformable Image Registration Uncertainty Quantification Using Deep Learning for Dose Accumulation in Adaptive Proton Therapy

A. Smolders<sup>1,2</sup> (✉), T. Lomax<sup>1,2</sup>, D. C. Weber<sup>1</sup>, and F. Albertini<sup>1</sup>



TransMorph: Transformer for unsupervised medical image registration

Junyu Chen<sup>a,b,\*</sup>, Eric C. Frey<sup>a,b</sup>, Yufan He<sup>c</sup>, William P. Segars<sup>c</sup>, Ye Li<sup>d</sup>, Yong Du<sup>a</sup>

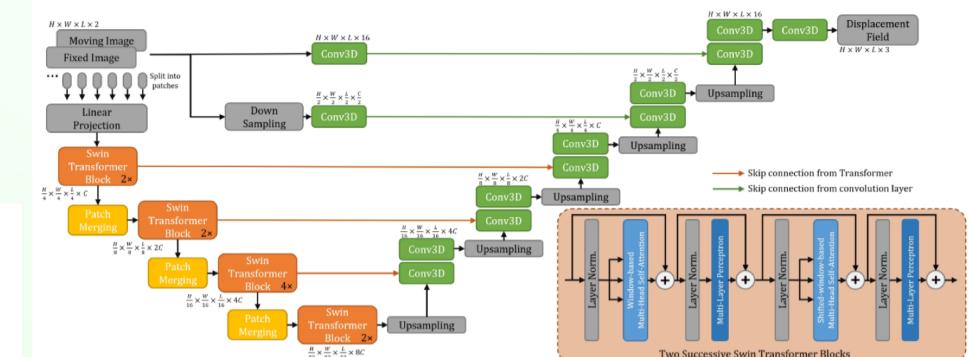


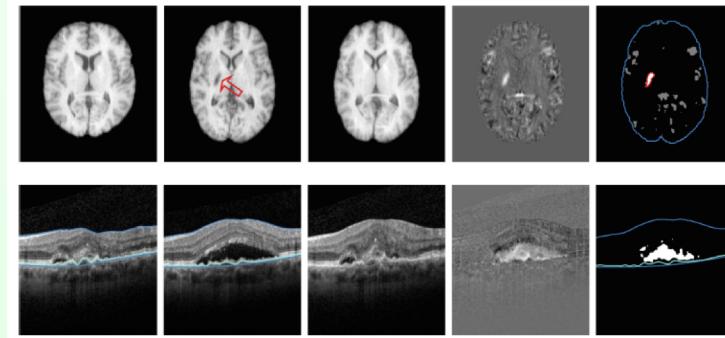
Fig. 1: The architecture of the proposed TransMorph registration network.

# Recent(ish) developments: Others

- Changing topology
- Instance optimization
- Implicit neural representations

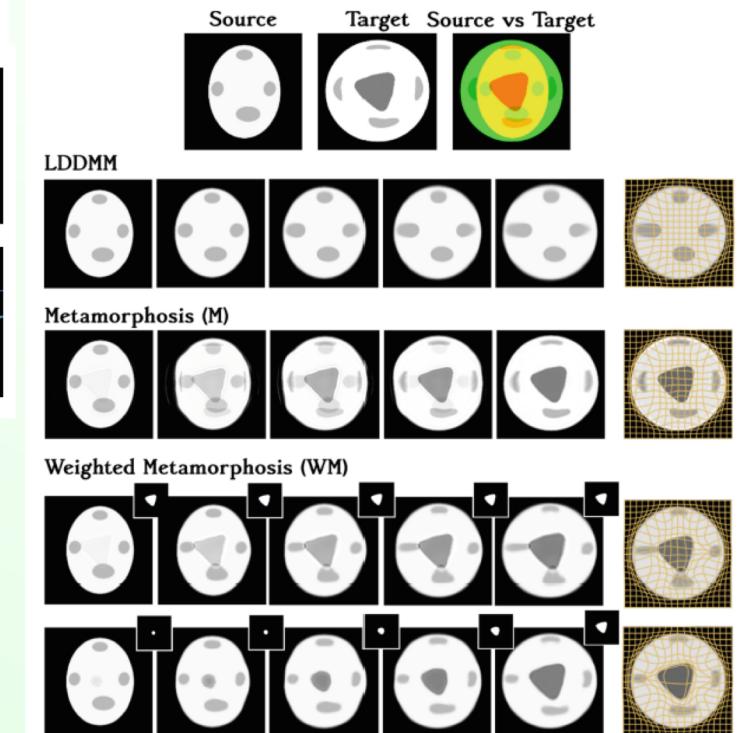
## Unsupervised Non-correspondence Detection in Medical Images Using an Image Registration Convolutional Neural Network

Julia Andresen<sup>1(✉)</sup>, Timo Kepp<sup>1</sup>, Jan Ehrhardt<sup>1,2</sup>,  
Claus von der Burchard<sup>3</sup>, Johann Roider<sup>3</sup>, and Heinz Handels<sup>1,2</sup>



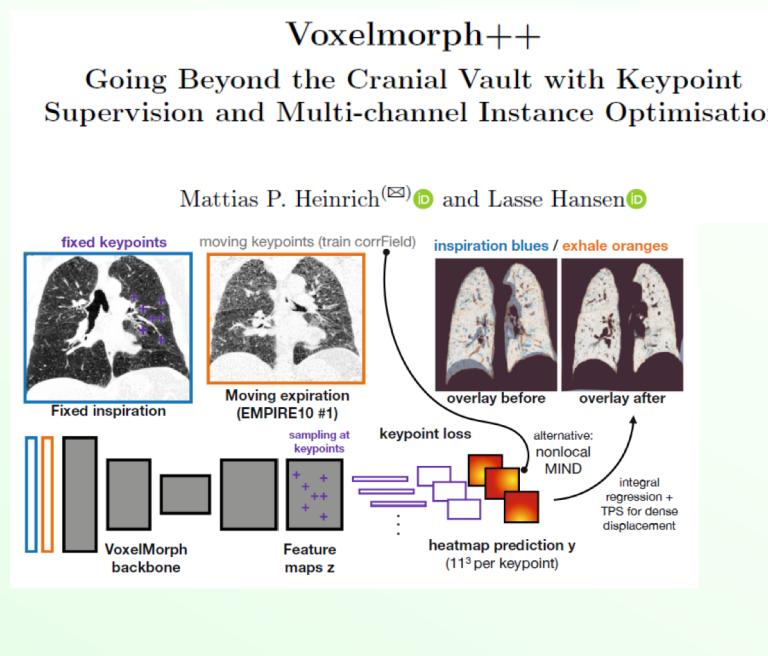
## Weighted Metamorphosis for Registration of Images with Different Topologies

Anton François<sup>1,2(✉)</sup>, Matthias Maillard<sup>2</sup>, Catherine Oppenheim<sup>3</sup>,  
Johan Pallud<sup>3</sup>, Isabelle Bloch<sup>2,4</sup>, Pietro Gori<sup>2(✉)</sup>, and Joan Glaunès<sup>1(✉)</sup>



# Recent(ish) developments: Others

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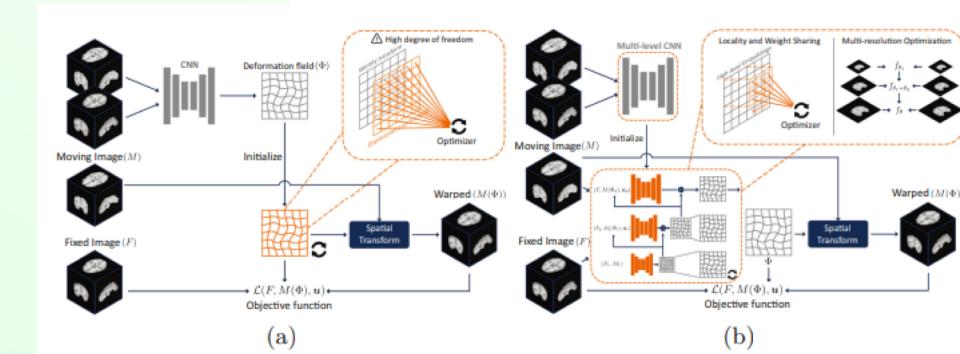


## Deformable Medical Image Registration Under Distribution Shifts with Neural Instance Optimization

Tony C. W. Mok<sup>1,2</sup><sup>(✉)</sup>, Zi Li<sup>1,2</sup>, Yingda Xia<sup>1</sup>, Jiawen Yao<sup>1,2</sup>, Ling Zhang<sup>1</sup>, Jingren Zhou<sup>1,2</sup>, and Le Lu<sup>1</sup>

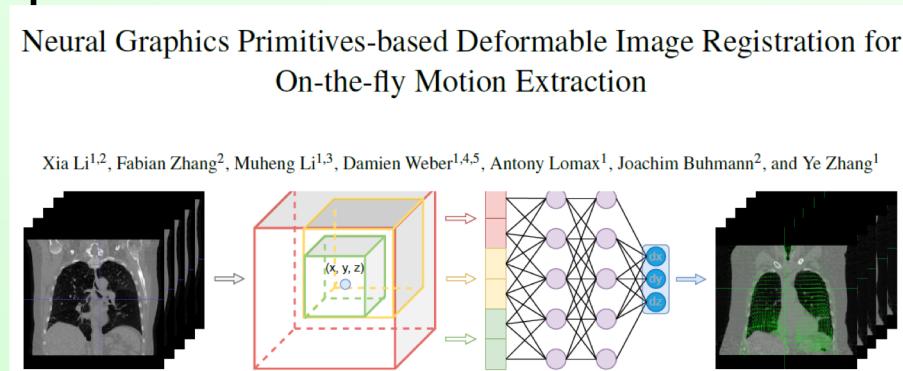
<sup>1</sup> DAMO Academy, Alibaba Group, Hangzhou, China  
mokchi-wing.mcw@alibaba-inc.com

<sup>2</sup> Hupan Lab, Hangzhou 310023, China



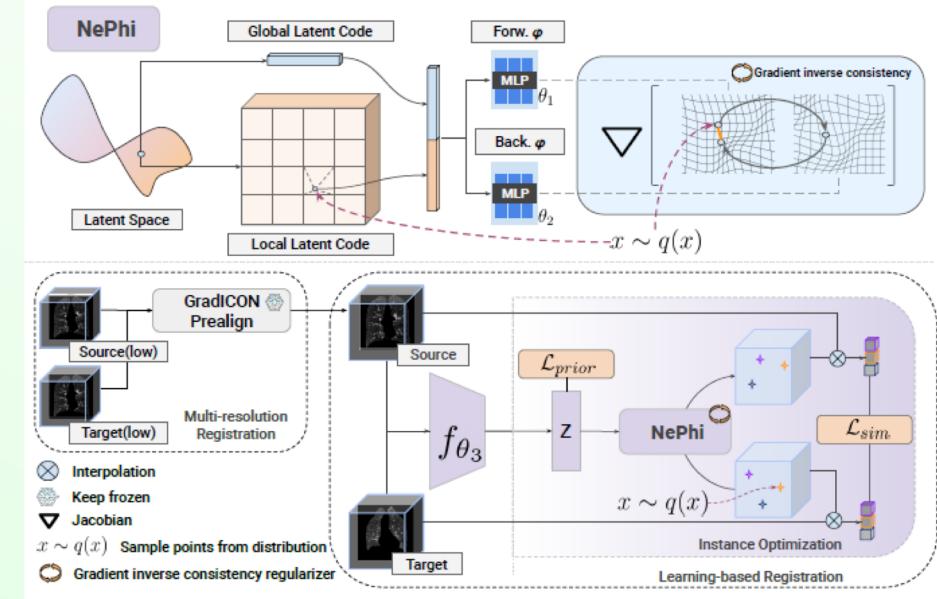
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- Changing topology
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**NePhi: Neural Deformation Fields for Approximately Diffeomorphic Medical Image Registration**

Lin Tian<sup>1</sup>, Hastings Greer<sup>1</sup>, Raúl San José Estépar<sup>2</sup>, Soumyadip Sengupta<sup>1</sup>, and Marc Niethammer<sup>1</sup>

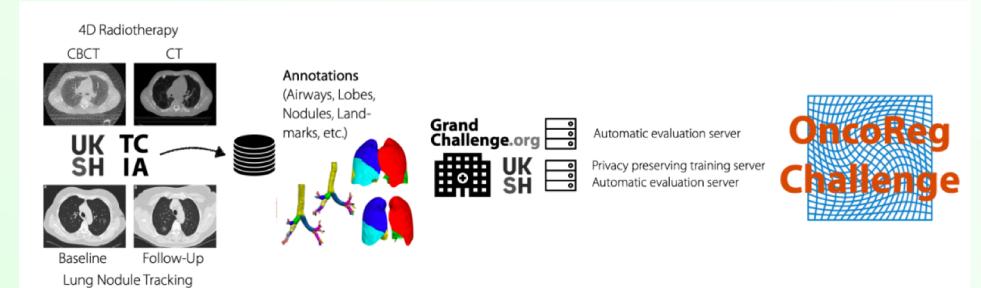


# The future of medical image registration...

- Less focus on novel methods, more focus on clinical translation
  - What are the important clinical applications?
  - What do we want the registration to do?
  - How do we know if it's doing it?
- Classical or learning based?

# Open datasets and challenges

- Super useful resources
- Drive innovation
- Fair comparisons between methods
- May not reflect current clinical challenges
- Sharing tools and data for evaluation



## What do we really want the registration to do?

- Application specific
  - Clear for some, not for others
  - How will registration be used?
- Requires thought
- Need to be precise

## And how do we know if it's doing it?

- Appropriate metrics for specific application
- Commissioning vs QA
- Applications specific cost/loss functions

# Classical vs learning-based registrations

- Speed
- Computational resources
- Data requirements
- Flexibility
- Generalizability
- Use of structures for guiding registrations
- Hybrid approaches

# Acknowledgments

- Present and former members of



- Colleagues from

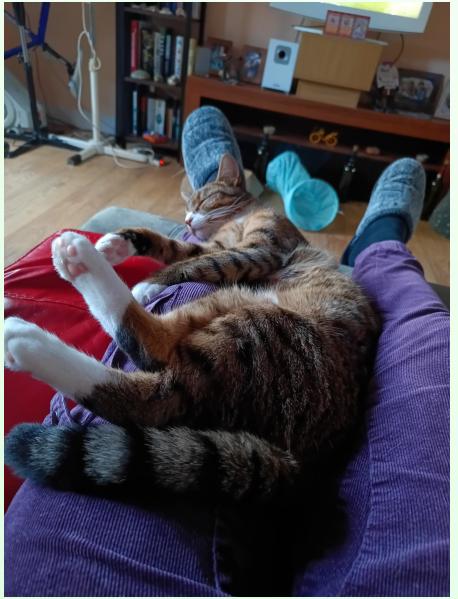


and



- Collaborators and Funders:





Thanks for listening...  
...any questions?

