

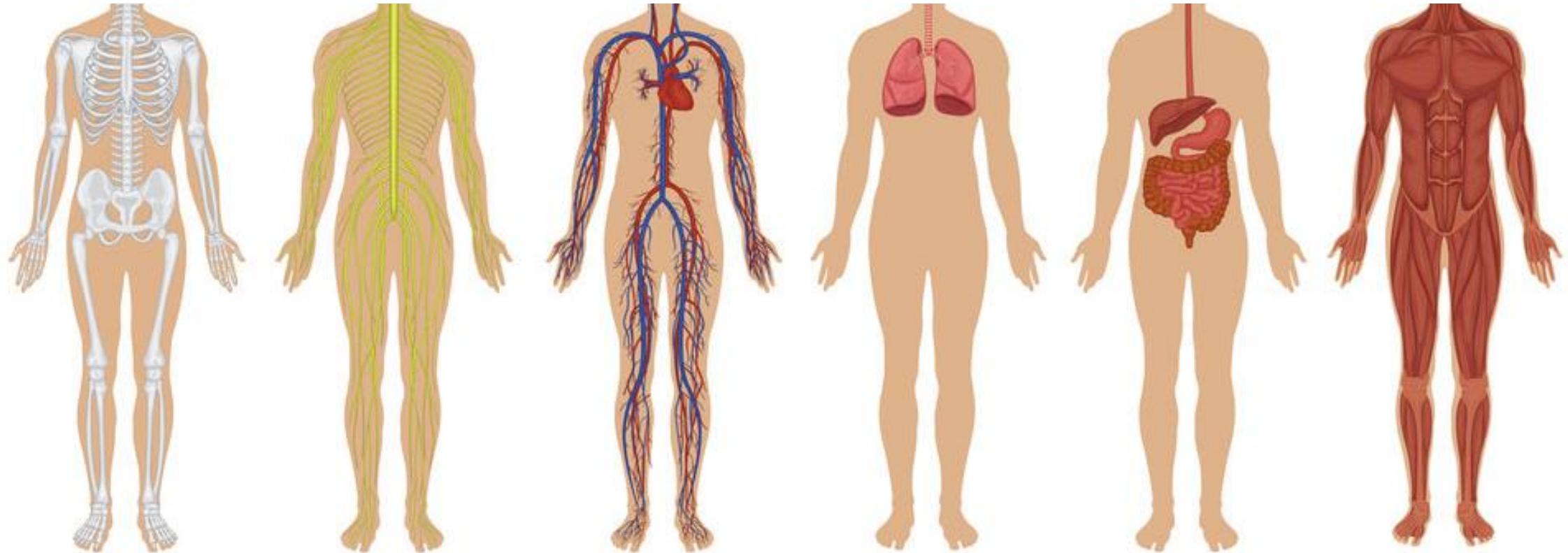
Unsupervised deep learning image registration: Beyond the cranial vault

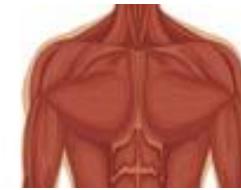
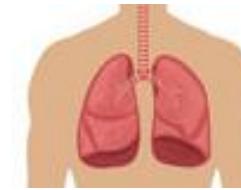
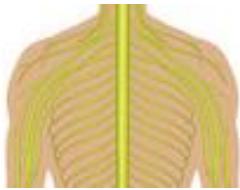
Bob de Vos

Postdoc Amsterdam UMC

Learn2Reg



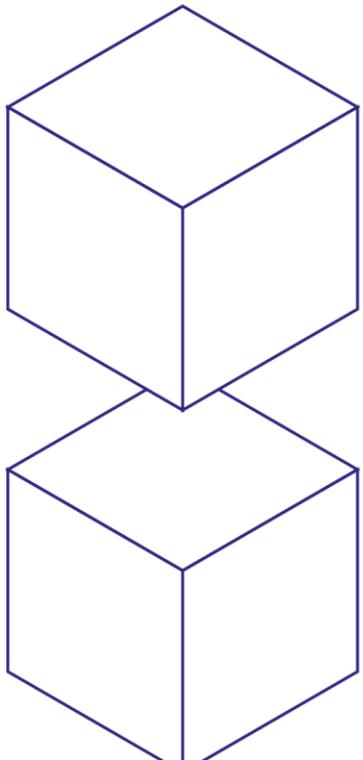




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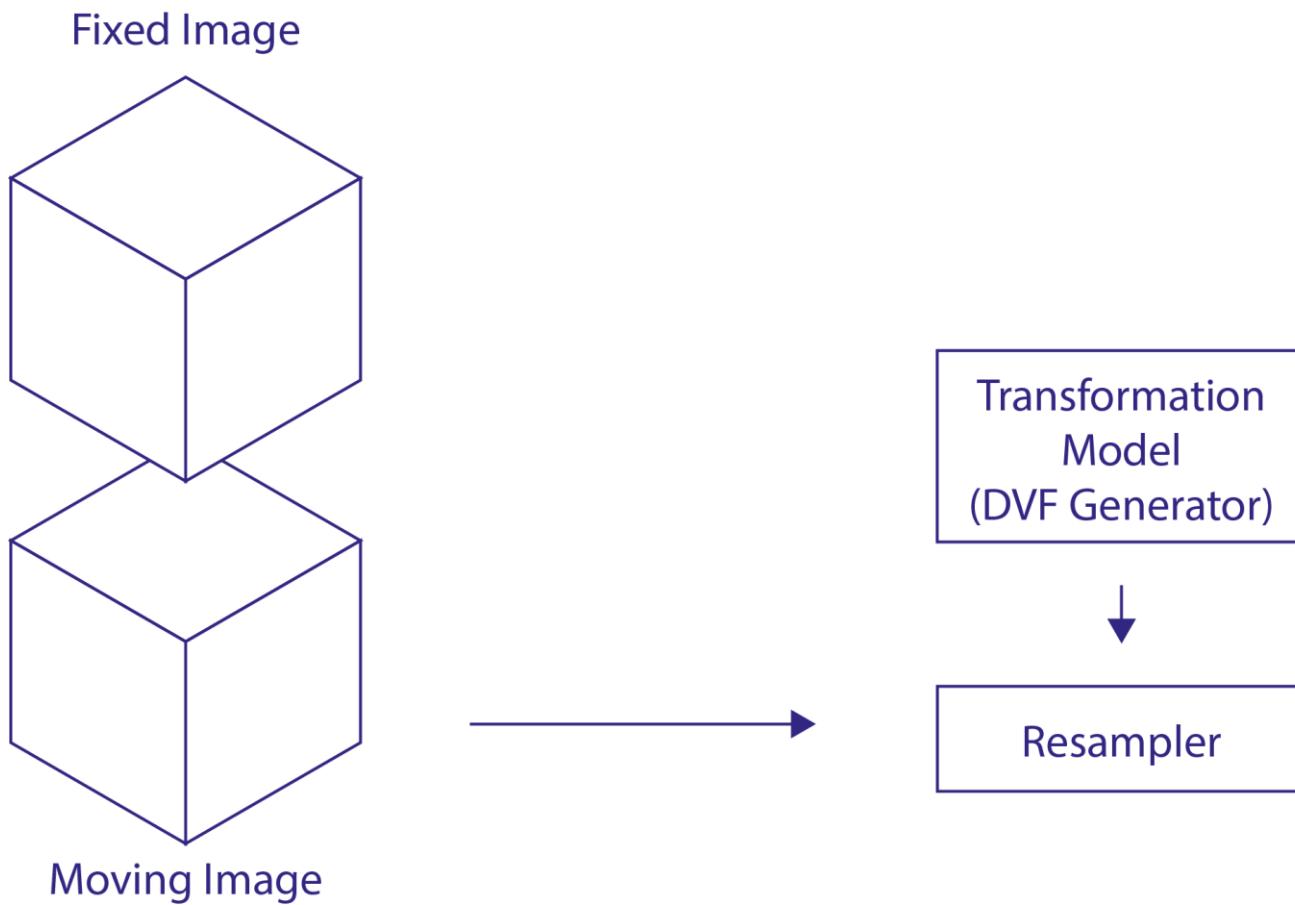


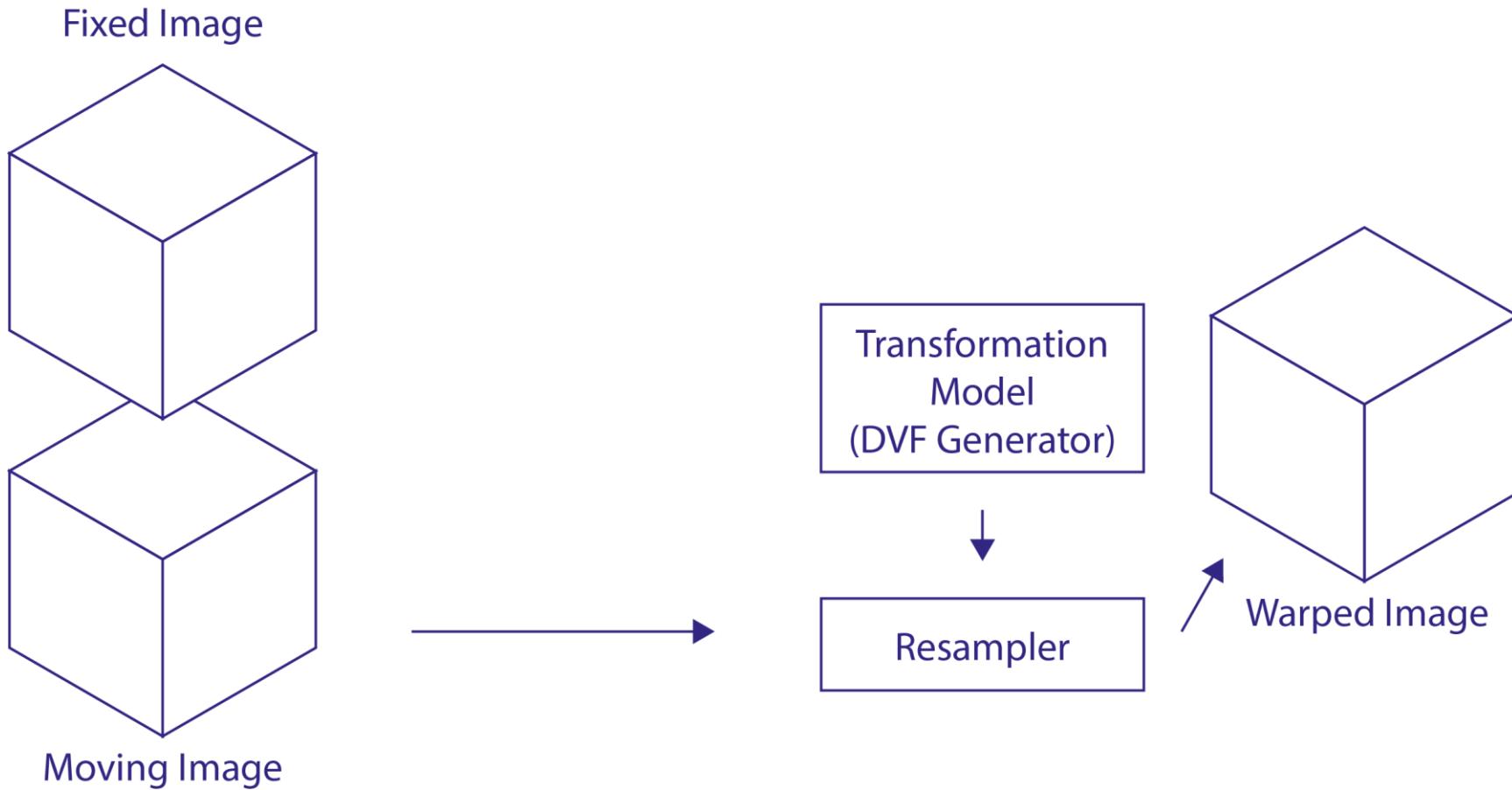
Fixed Image

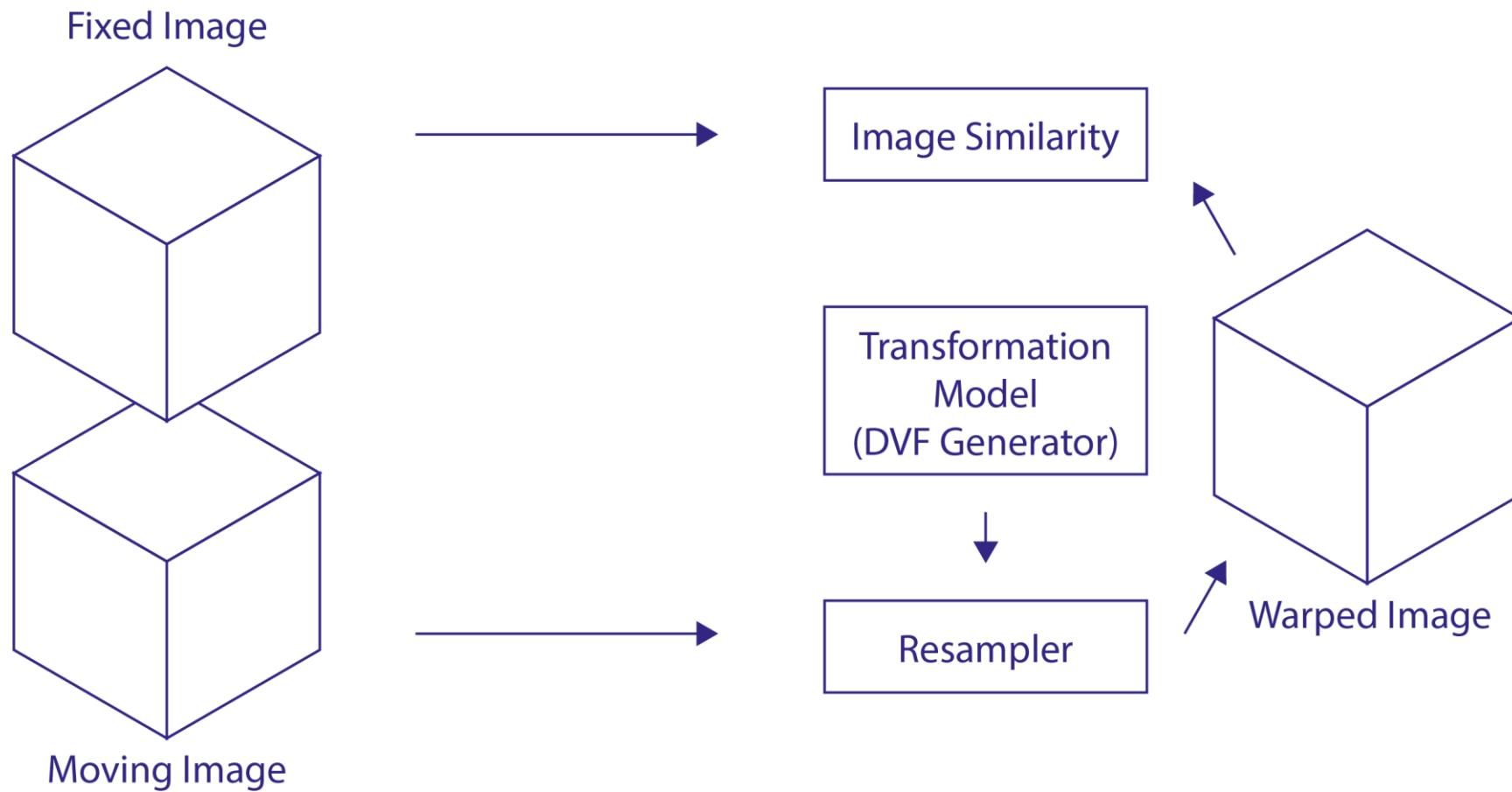


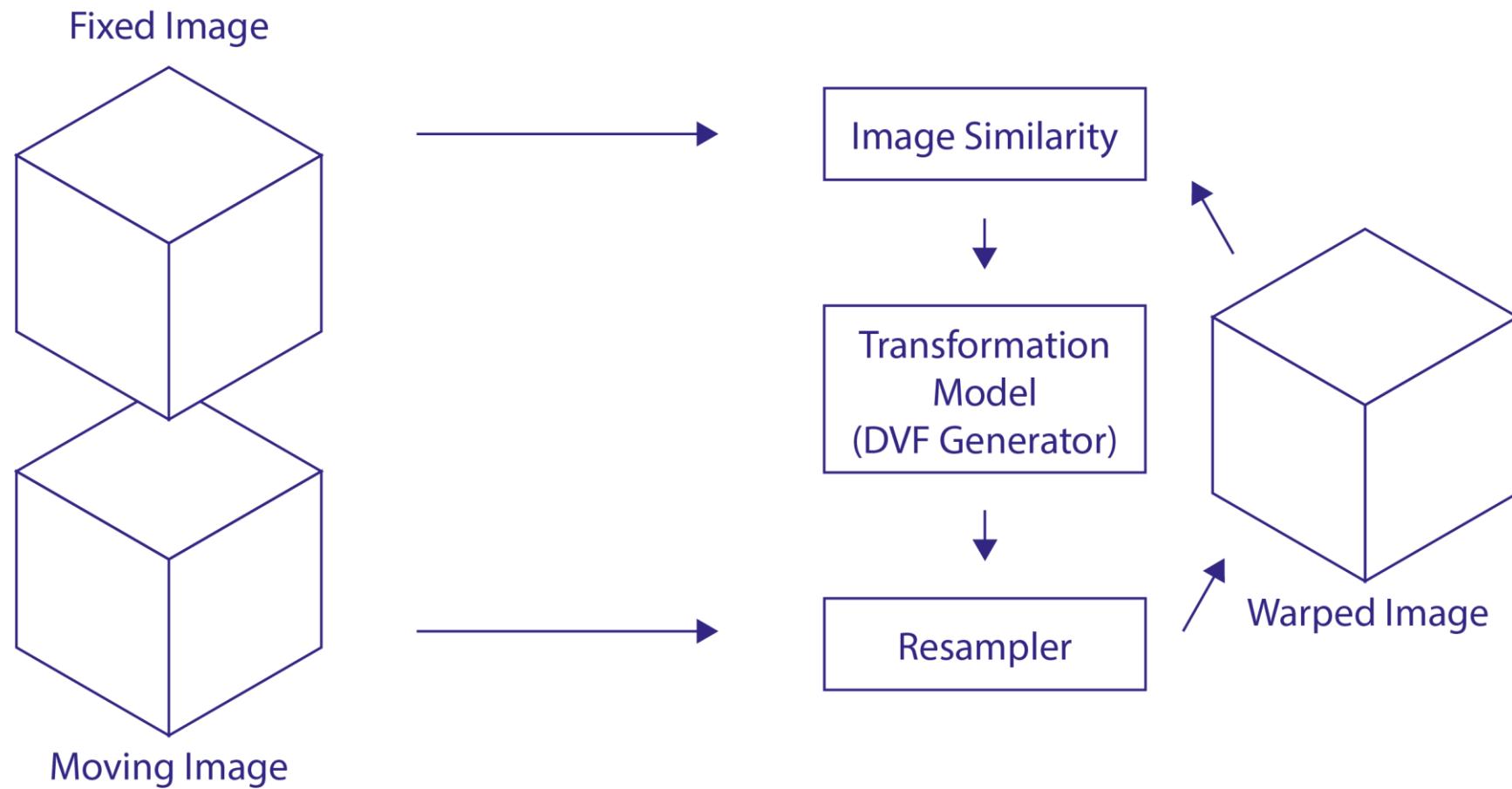
Moving Image

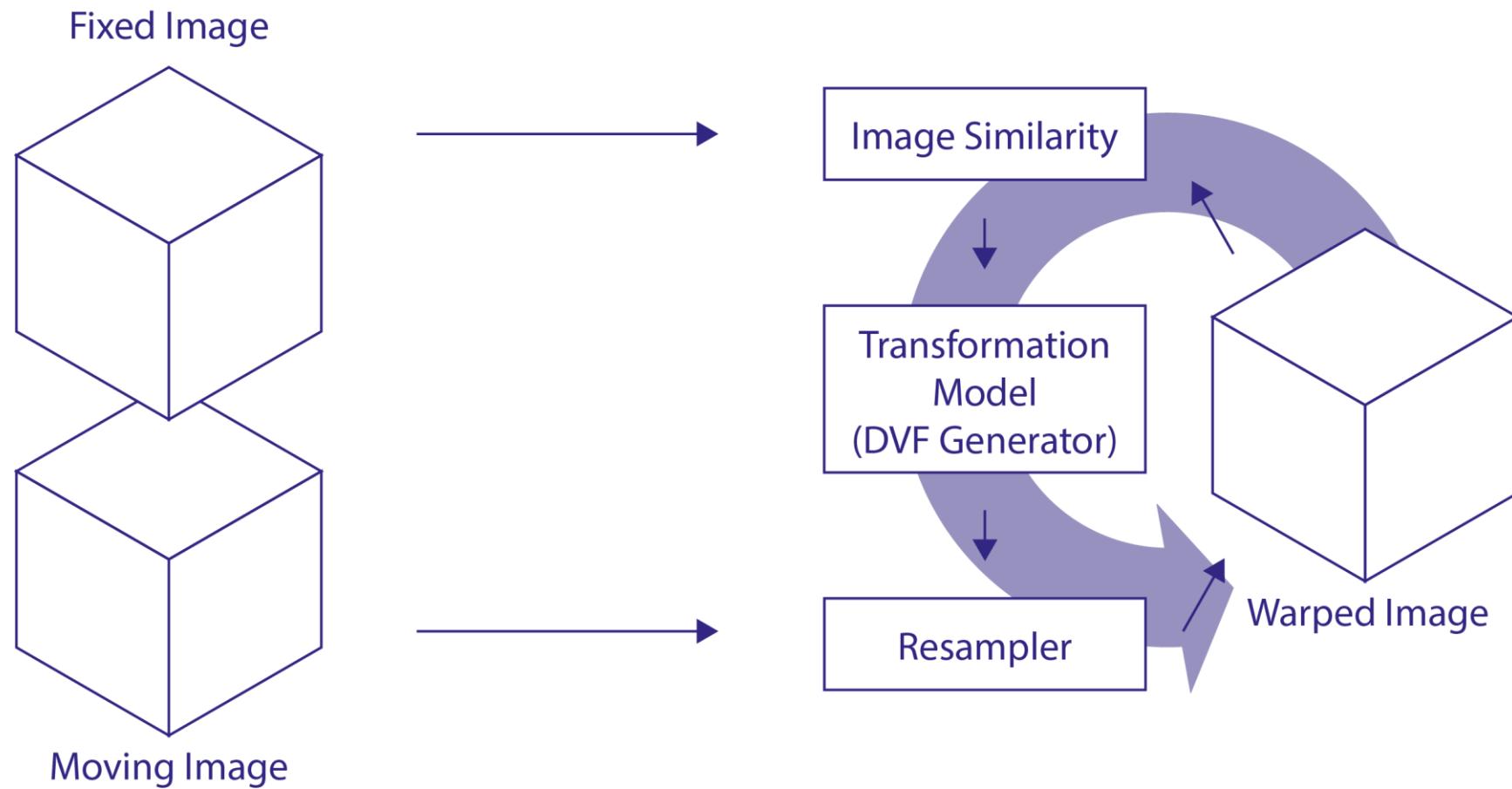
Transformation
Model
(DVF Generator)

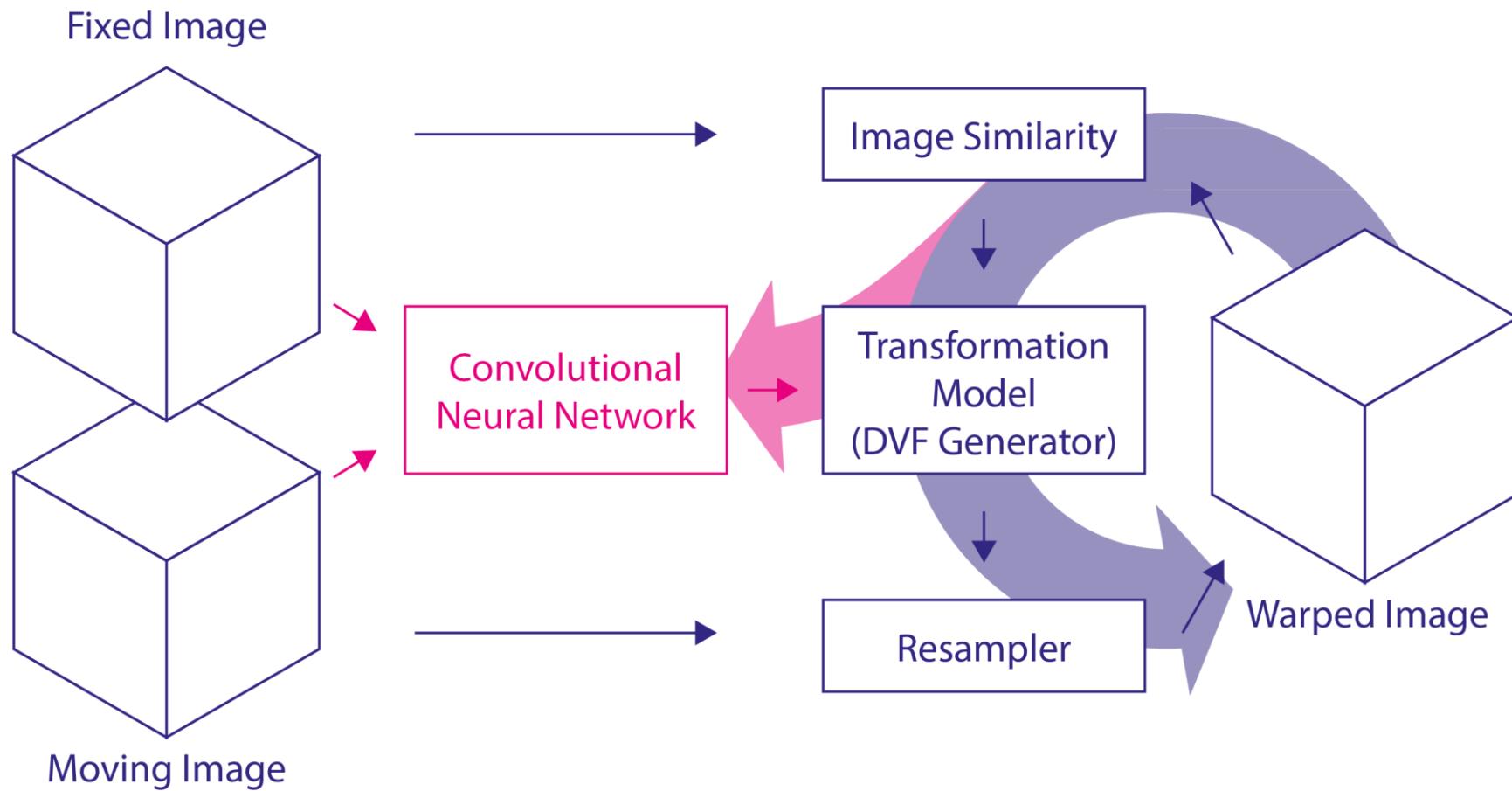














End-to-End Unsupervised Deformable Image Registration with a Convolutional Neural Network

Bob D. de Vos¹, Floris F. Berendsen², Max A. Viergever¹, Marius Staring², and Ivana Išgum¹

¹Image Sciences Institute, University Medical Center Utrecht, the Netherlands

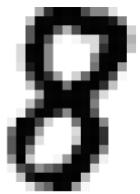
²Division of Image Processing, Leiden University Medical Center, the Netherlands

Image registration with unsupervised deep learning

Keywords: convolution neural network, deformable image registration, spatial transformer, cardiac MRI



Moving image

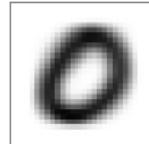


Fixed image





Average of 1,000 moving images from the test-set





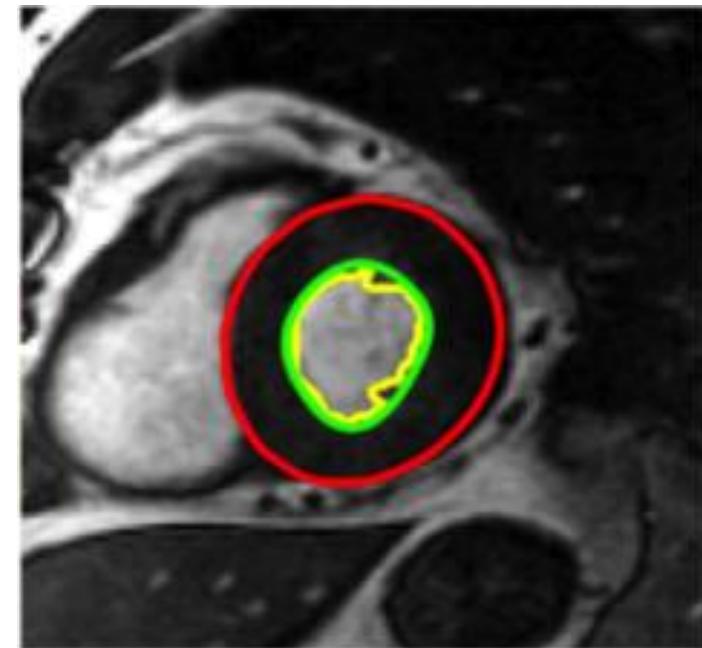
Sunnybrook Cardiac Data: cardiac cine MRI

45 MR images

Training/validation/test → 15/15/15

Image size 256x256x~10x20 voxels

Register 2D slices
from two time points
the same scan and slice location





Original



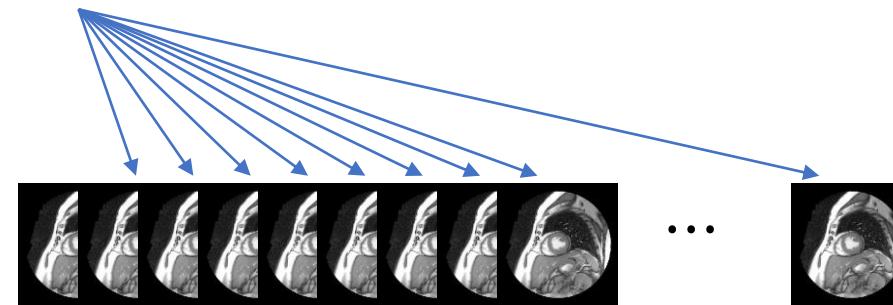


Original





Original





Original





Original



Warped

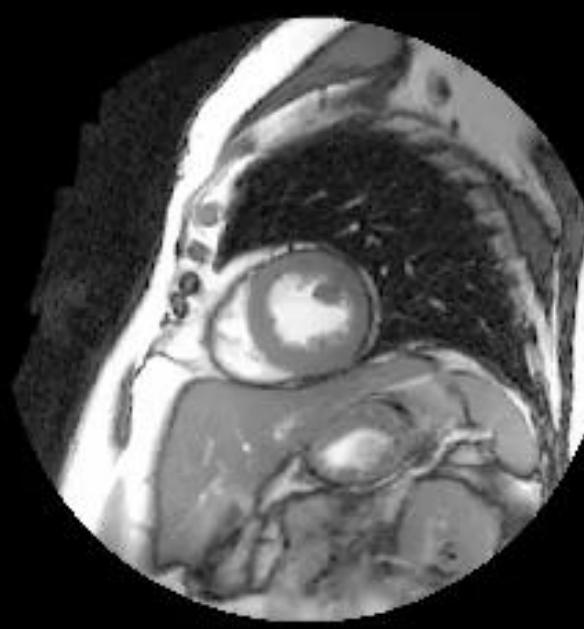




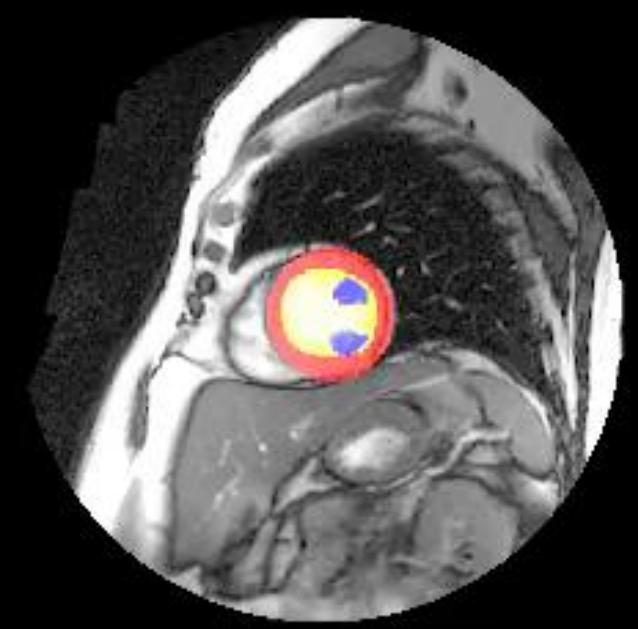
Original

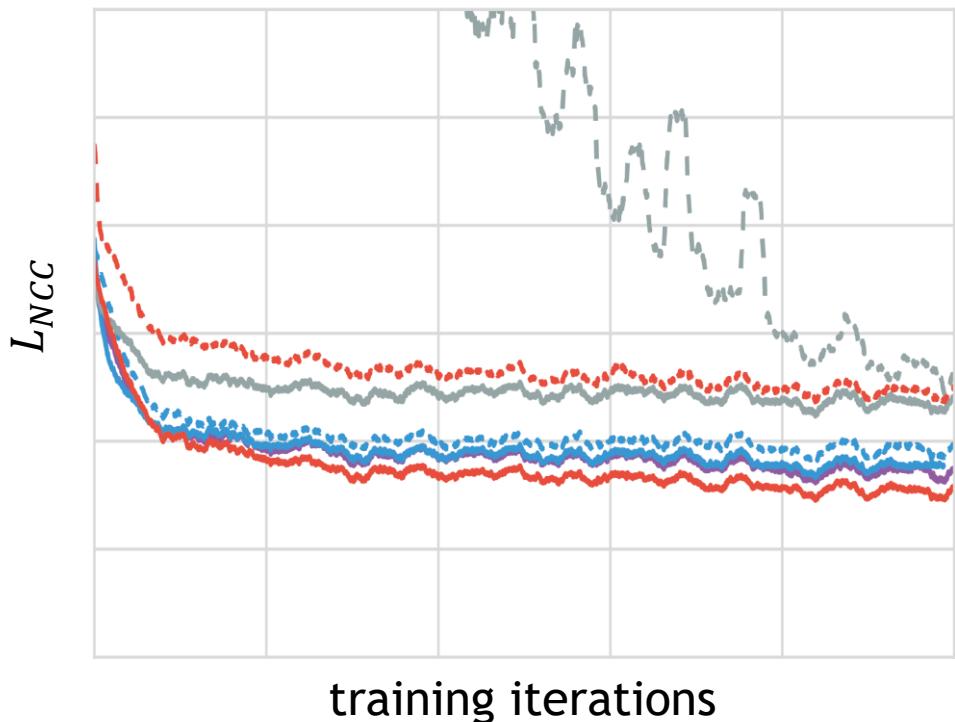


Warped



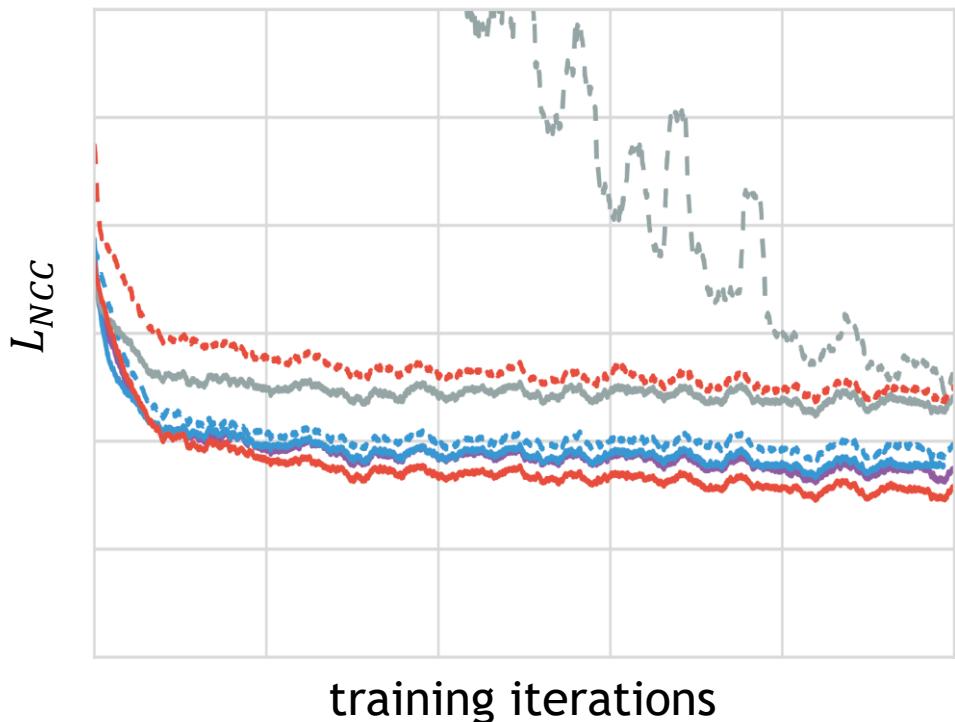
Propagated labels





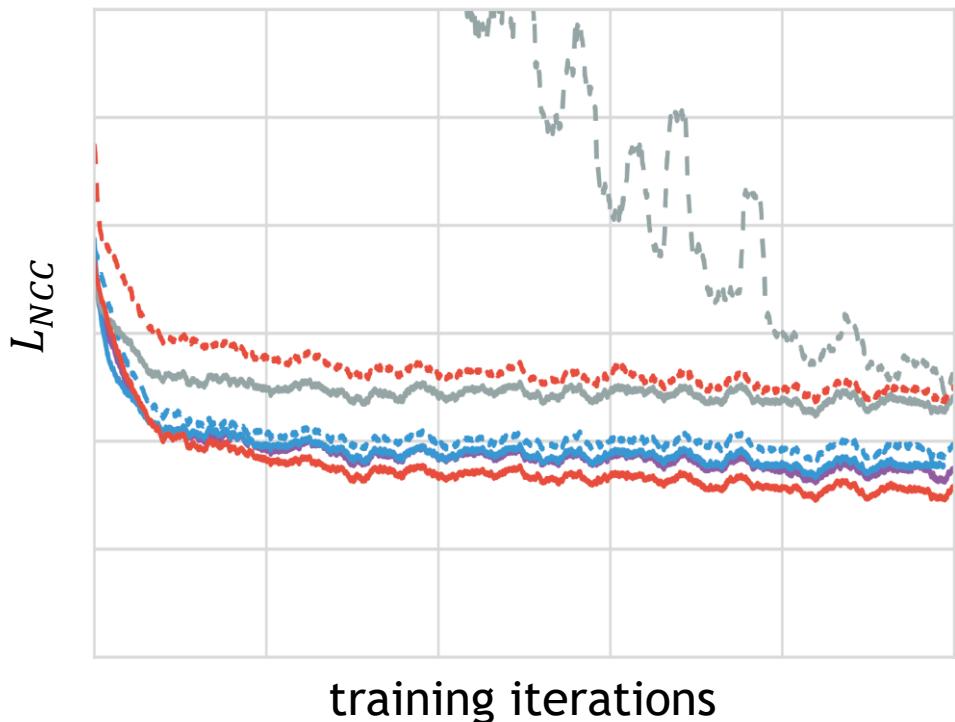
	Dice	95^{th}SD (mm)	MAD (mm)	Time (s)
No registration	0.62 ± 0.15	7.79 ± 2.92	2.89 ± 1.07	-
SimpleElastix 2 × 100	0.79 ± 0.08	5.09 ± 2.36	1.91 ± 0.94	0.51 ± 0.07
SimpleElastix 2 × 2000	0.81 ± 0.08	5.09 ± 7.25	1.75 ± 1.29	7.38 ± 0.94
Baseline	0.80 ± 0.08	5.03 ± 2.30	1.83 ± 0.89	0.049 ± 0.004
max pooling	0.78 ± 0.08	5.26 ± 2.16	1.95 ± 0.85	-
strided convolutions	0.78 ± 0.08	5.30 ± 2.28	1.97 ± 0.87	-
quadratic B-spline	0.72 ± 0.11	6.41 ± 2.61	2.40 ± 0.96	-
thin plate spline	0.78 ± 0.09	5.48 ± 2.36	2.01 ± 0.89	-
overlapping patches	0.79 ± 0.08	5.20 ± 2.30	1.92 ± 0.89	-
full image	0.76 ± 0.09	5.55 ± 2.24	2.10 ± 0.90	-

Evaluation: label propagation of left ventricle myocardium segmentation



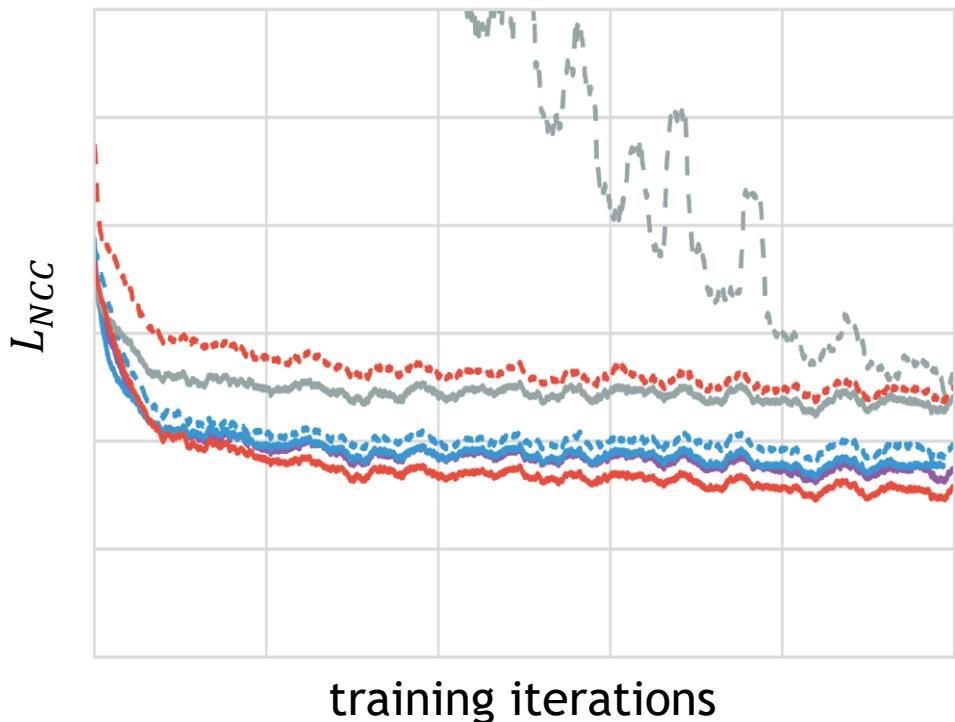
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SimpleElastix 2 × 2000	0.81 ± 0.08	5.09 ± 7.25	1.75 ± 1.29	7.38 ± 0.94
Baseline	0.80 ± 0.08	5.03 ± 2.30	1.83 ± 0.89	0.049 ± 0.004
max pooling	0.78 ± 0.08	5.26 ± 2.16	1.95 ± 0.85	-
strided convolutions	0.78 ± 0.08	5.30 ± 2.28	1.97 ± 0.87	-
quadratic B-spline	0.72 ± 0.11	6.41 ± 2.61	2.40 ± 0.96	-
thin plate spline	0.78 ± 0.09	5.48 ± 2.36	2.01 ± 0.89	-
overlapping patches	0.79 ± 0.08	5.20 ± 2.30	1.92 ± 0.89	-
full image	0.76 ± 0.09	5.55 ± 2.24	2.10 ± 0.90	-

Evaluation: label propagation of left ventricle myocardium segmentation



	Dice	95 th SD (mm)	MAD (mm)	Time (s)
No registration	0.62 ± 0.15	7.79 ± 2.92	2.89 ± 1.07	-
SimpleElastix 2 × 100	0.79 ± 0.08	5.09 ± 2.36	1.91 ± 0.94	0.51 ± 0.07
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Baseline	0.80 ± 0.08	5.03 ± 2.30	1.83 ± 0.89	0.049 ± 0.004
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thin plate spline	0.78 ± 0.09	5.48 ± 2.36	2.01 ± 0.89	-
overlapping patches	0.79 ± 0.08	5.20 ± 2.30	1.92 ± 0.89	-
full image	0.76 ± 0.09	5.55 ± 2.24	2.10 ± 0.90	-

Evaluation: label propagation of left ventricle myocardium segmentation



	Dice	95 th SD (mm)	MAD (mm)	Time (s)
No registration	0.62 ± 0.15	7.79 ± 2.92	2.89 ± 1.07	-
SimpleElastix 2 × 100	0.79 ± 0.08	5.09 ± 2.36	1.91 ± 0.94	0.51 ± 0.07
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full image	0.76 ± 0.09	5.55 ± 2.24	2.10 ± 0.90	-

Evaluation: label propagation of left ventricle myocardium segmentation



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A deep learning framework for unsupervised affine and deformable image registration



Bob D. de Vos^{a,*}, Floris F. Berendsen^b, Max A. Viergever^a, Hessam Sokooti^b,
Marius Staring^b, Ivana Išgum^a

^aImage Sciences Institute, University Medical Center Utrecht and Utrecht University, Utrecht, The Netherlands

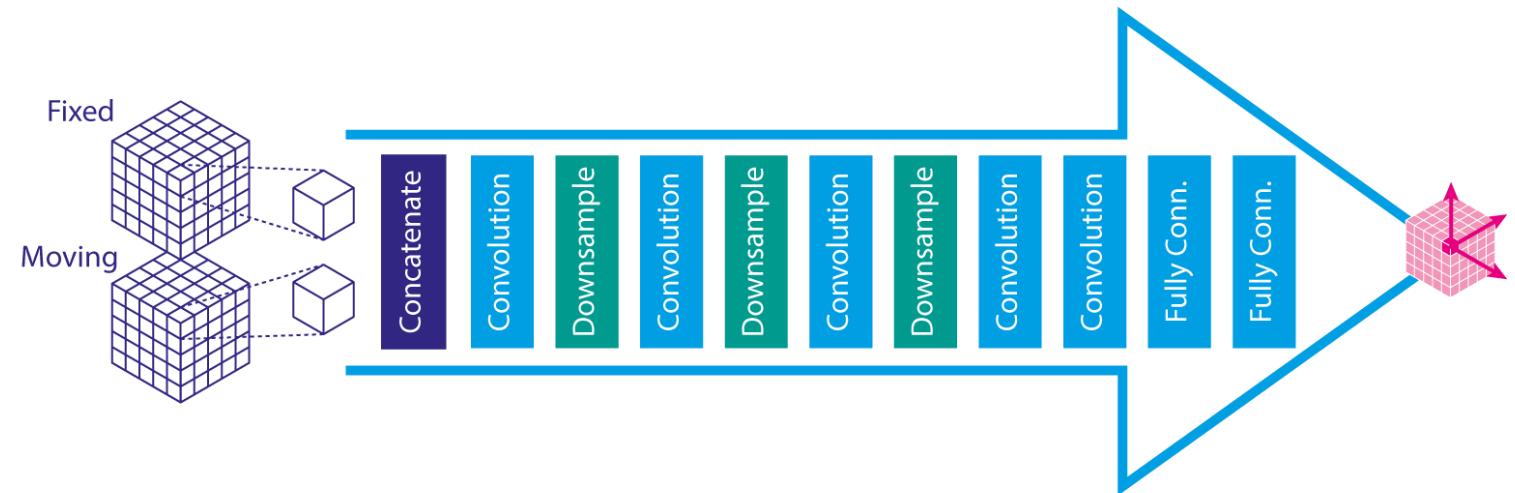
^bDivision of Image Processing of the Leiden University Medical Center, Leiden, The Netherlands

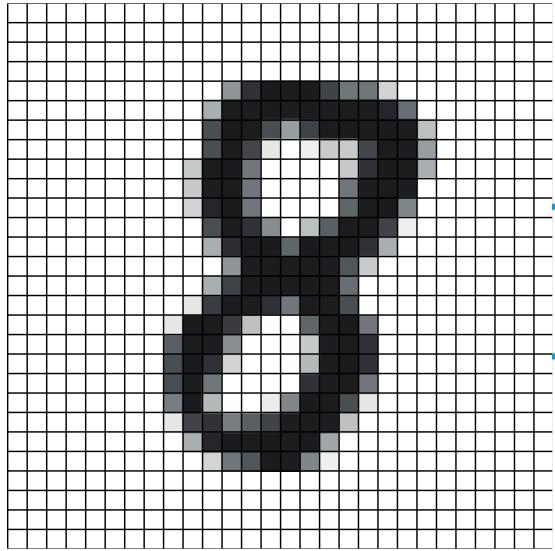
Deep learning image registration framework



Deformable image registration

- 3D
- B-splines
- Patch-based
- Fully convolutional
- Bending energy penalty
- 3x3x3 convolutions
- 2x2x2 avg. pooling
- 32 kernels/nodes per layer





Concatenate

Convolution

Downsample

Convolution

Downsample

Convolution

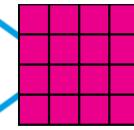
Downsample

Convolution

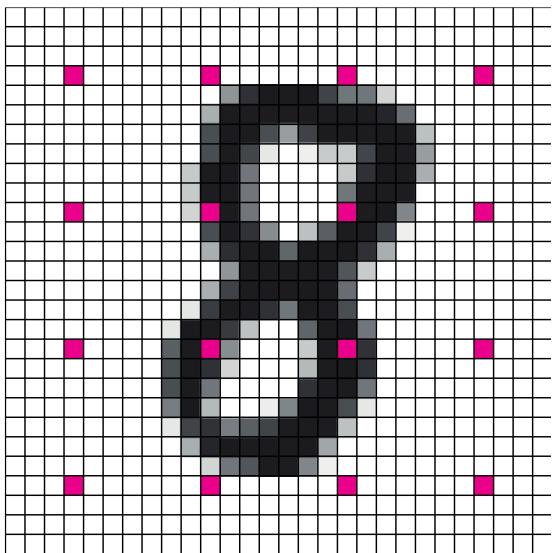
Convolution

Fully Conn.

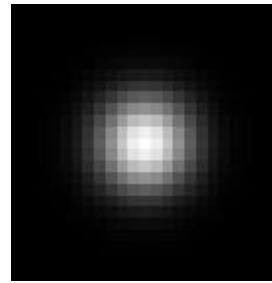
Fully Conn.



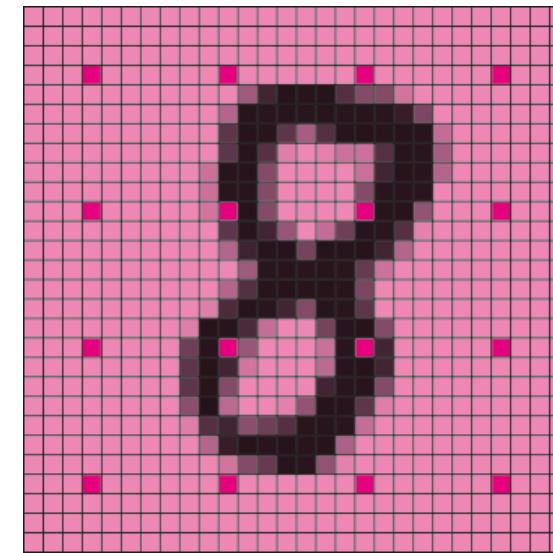
dense
DVF



*

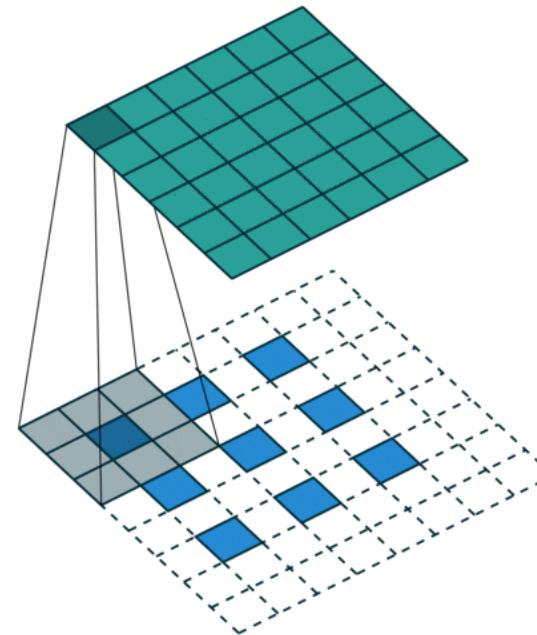


=





- Transposed convolutions for b-spline registration
- Available in every DL package (except the b-spline kernel)



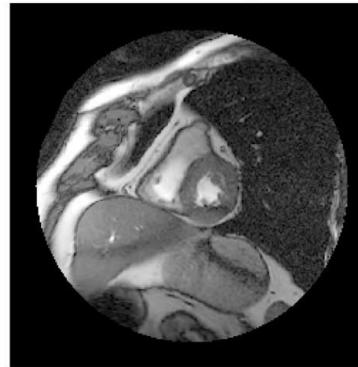


Regularization

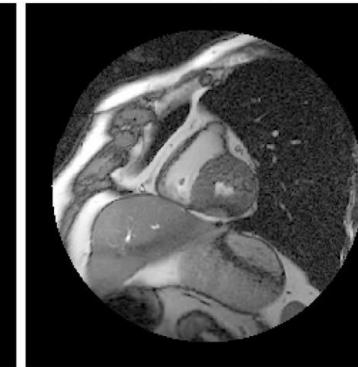
- Bending energy penalty

$$P = \frac{1}{V} \int_0^X \int_0^Y \int_0^Z \left[\left(\frac{\partial^2 \mathbf{T}}{\partial x^2} \right)^2 + \left(\frac{\partial^2 \mathbf{T}}{\partial y^2} \right)^2 + \left(\frac{\partial^2 \mathbf{T}}{\partial z^2} \right)^2 + 2 \left(\frac{\partial^2 \mathbf{T}}{\partial xy} \right)^2 + 2 \left(\frac{\partial^2 \mathbf{T}}{\partial xz} \right)^2 + 2 \left(\frac{\partial^2 \mathbf{T}}{\partial yz} \right)^2 \right] dx dy dz,$$

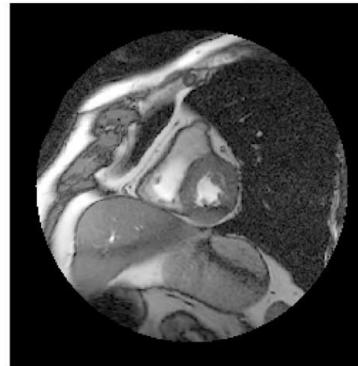
Fixed Image



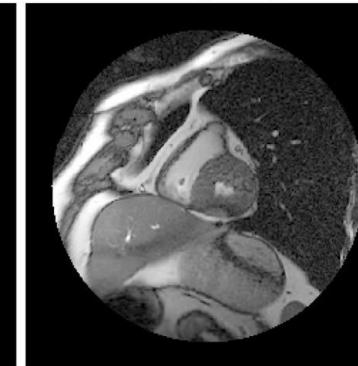
Moving Image



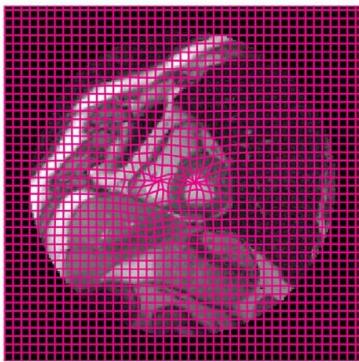
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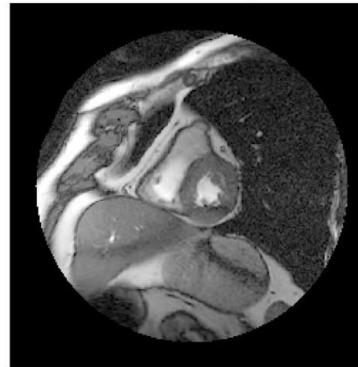
Moving Image



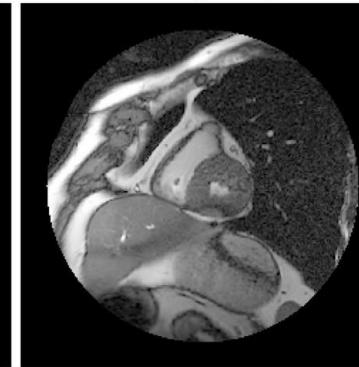
SE



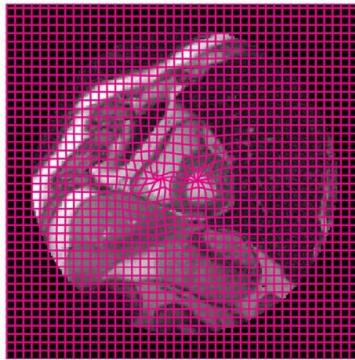
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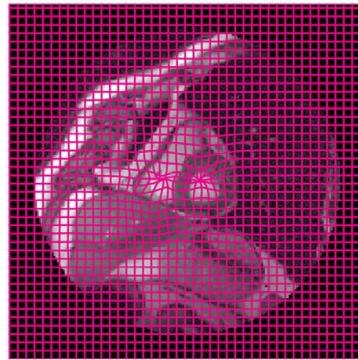
Moving Image



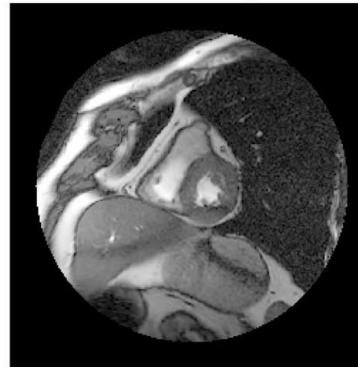
SE



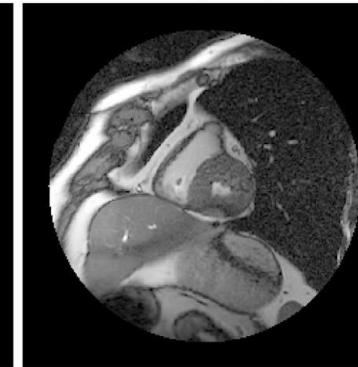
SE + BP



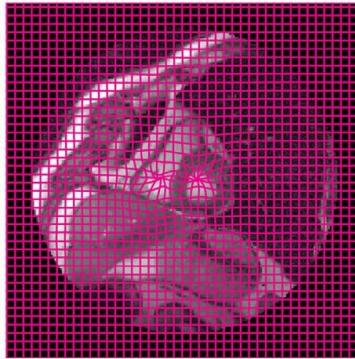
Fixed Image



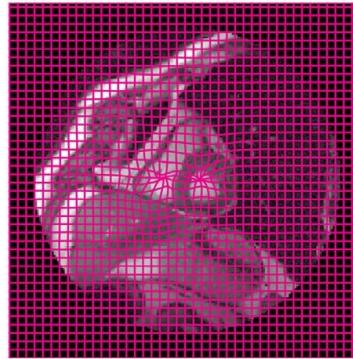
Moving Image



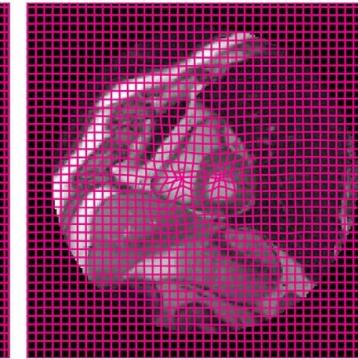
SE



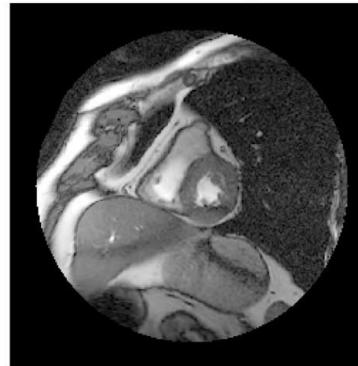
SE + BP



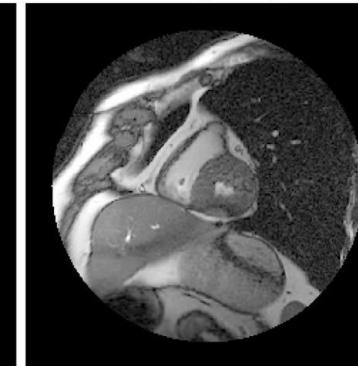
DLIR



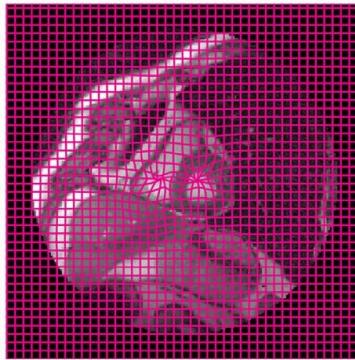
Fixed Image



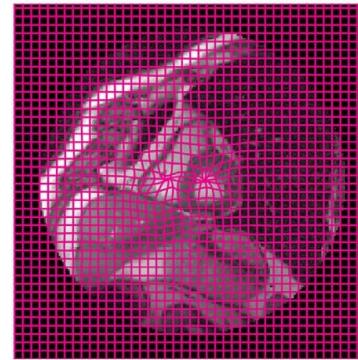
Moving Image



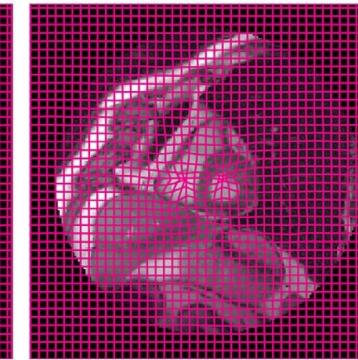
SE



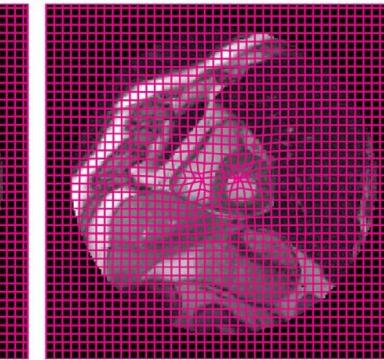
SE + BP



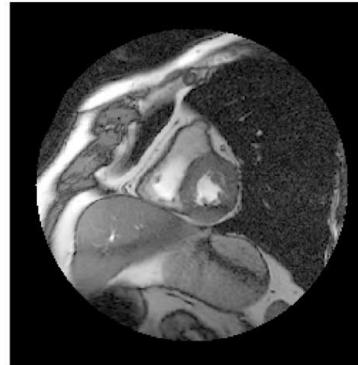
DLIR



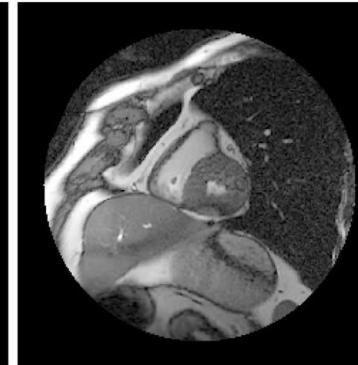
DLIR + BP



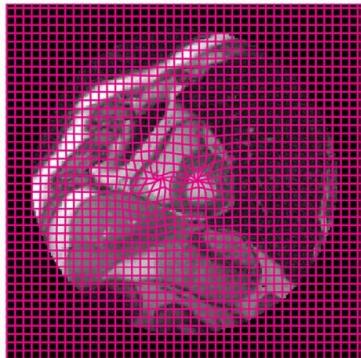
Fixed Image



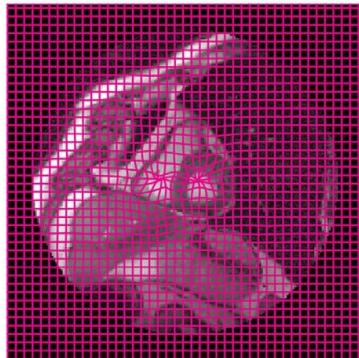
Moving Image



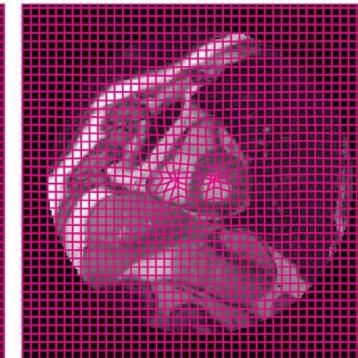
SE



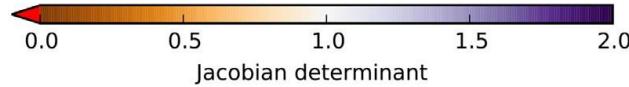
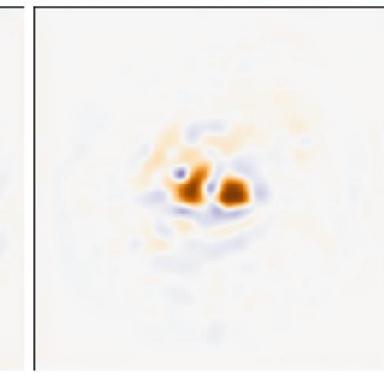
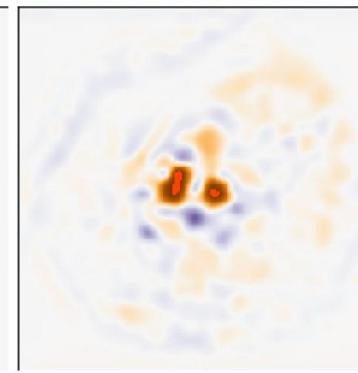
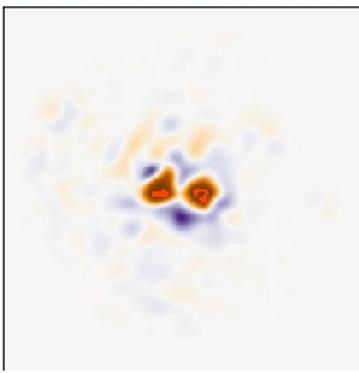
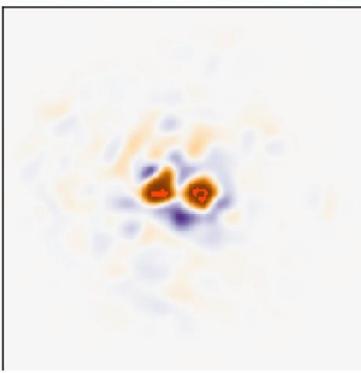
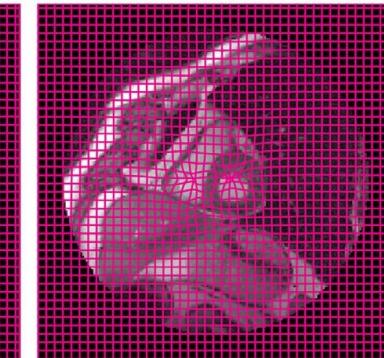
SE + BP



DLIR



DLIR + BP





	Dice	HD	ASD	Fraction folding	Std. dev. Jacobian	CPU time (s)	GPU time (s)
Before registration	0.70 ± 0.30	15.46 ± 4.50	4.66 ± 4.26	–	–	–	–
Single stage	SE	0.86 ± 0.18	9.76 ± 4.78	1.14 ± 1.40	0.08 ± 0.16	0.15 ± 0.08	$13.49(3.27)$
	SE + BP	0.86 ± 0.17	9.64 ± 4.15	1.13 ± 1.38	0.07 ± 0.15	0.15 ± 0.08	$14.89(3.07)$
	DLIR	0.87 ± 0.18	9.47 ± 5.26	0.98 ± 1.12	0.03 ± 0.06	0.14 ± 0.04	$1.71(0.45)$
	DLIR + BP	0.86 ± 0.18	9.10 ± 4.26	1.01 ± 1.42	0.00 ± 0.01	0.09 ± 0.03	0.03 ± 0.01

SE = conventional IR with SimpleElastix

DLIR = deep learning image registration

BP = bending energy penalty



		Dice	HD	ASD	Fraction folding	Std. dev. Jacobian	CPU time (s)	GPU time (s)
Before registration		0.70 ± 0.30	15.46 ± 4.50	4.66 ± 4.26	–	–	–	–
Single stage	SE	0.86 ± 0.18	9.76 ± 4.78	1.14 ± 1.40	0.08 ± 0.16	0.15 ± 0.08	13.49(3.27)	–
	SE + BP	0.86 ± 0.17	9.64 ± 4.15	1.13 ± 1.38	0.07 ± 0.15	0.15 ± 0.08	14.89(3.07)	–
DLIR	DLIR	0.87 ± 0.18	9.47 ± 5.26	0.98 ± 1.12	0.03 ± 0.06	0.14 ± 0.04	1.71(0.45)	0.03 ± 0.01
	DLIR + BP	0.86 ± 0.18	9.10 ± 4.26	1.01 ± 1.42	0.00 ± 0.01	0.09 ± 0.03		

SE = conventional IR with SimpleElastix

DLIR = deep learning image registration

BP = bending energy penalty



		Dice	HD	ASD	Fraction folding	Std. dev. Jacobian	CPU time (s)	GPU time (s)
Before registration		0.70 ± 0.30	15.46 ± 4.50	4.66 ± 4.26	–	–	–	–
Single stage	SE	0.86 ± 0.18	9.76 ± 4.78	1.14 ± 1.40	0.08 ± 0.16	0.15 ± 0.08	13.49(3.27)	–
	SE + BP	0.86 ± 0.17	9.64 ± 4.15	1.13 ± 1.38	0.07 ± 0.15	0.15 ± 0.08	14.89(3.07)	–
	DLIR	0.87 ± 0.18	9.47 ± 5.26	0.98 ± 1.12	0.03 ± 0.06	0.14 ± 0.04	1.71(0.45)	0.03 ± 0.01
	DLIR + BP	0.86 ± 0.18	9.10 ± 4.26	1.01 ± 1.42	0.00 ± 0.01	0.09 ± 0.03		

SE = conventional IR with SimpleElastix

DLIR = deep learning image registration

BP = bending energy penalty



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SE = conventional IR with SimpleElastix

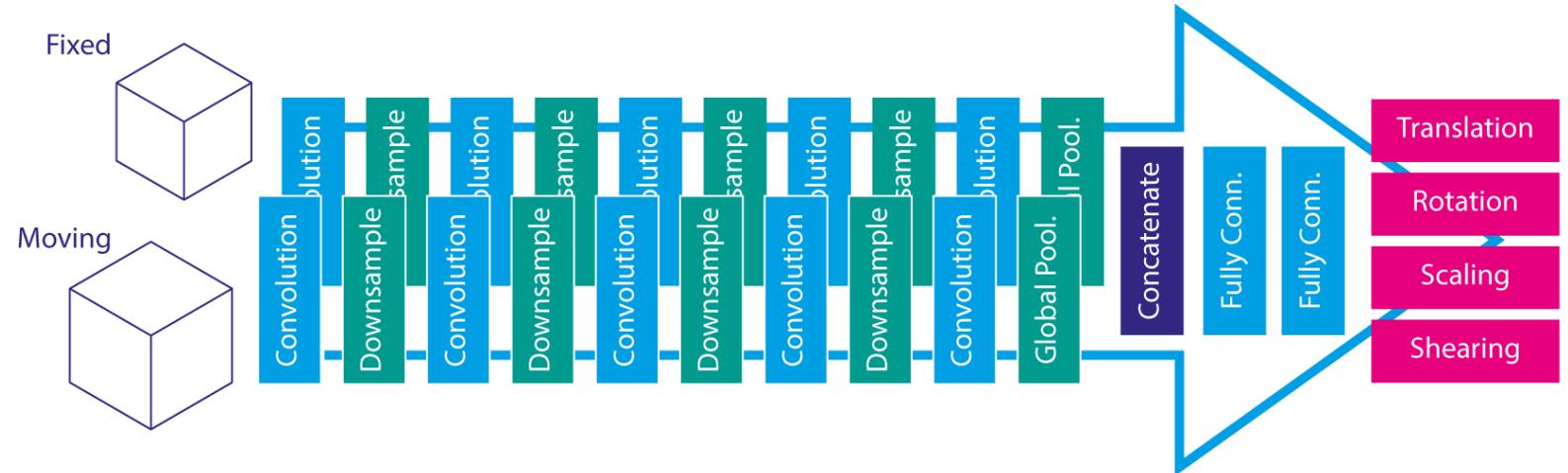
DLIR = deep learning image registration

BP = bending energy penalty



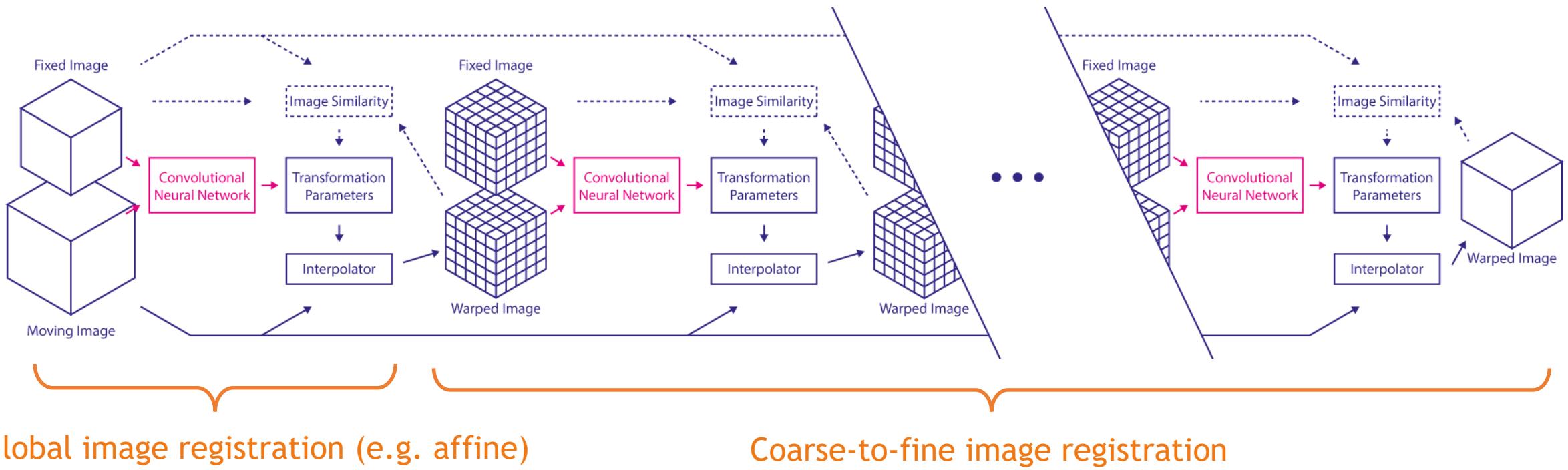
Global image registration

- Affine registration
- Different size input
- 3x3x3 convolutions
- 2x2x2 avg. Pooling
- 32 kernels per layer





Multi-stage image registration





Moving

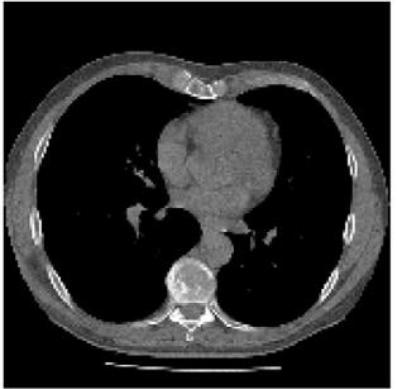


Fixed

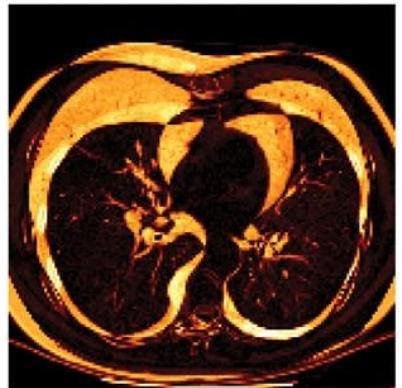




Moving



AIR

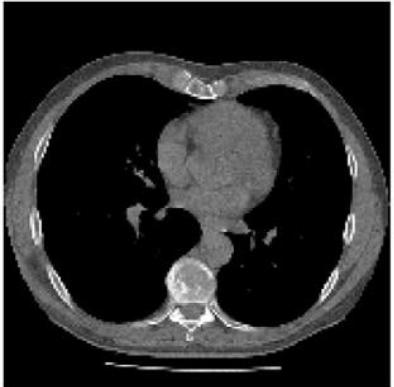


Fixed





Moving



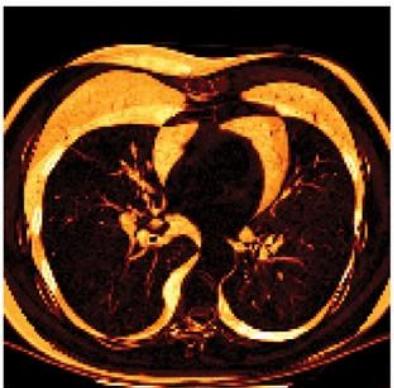
AIR



DIR-1

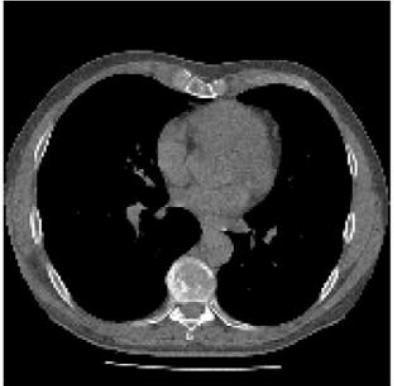


Fixed





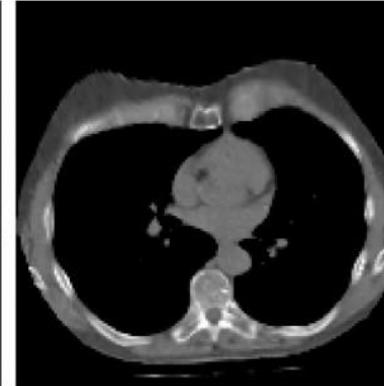
Moving



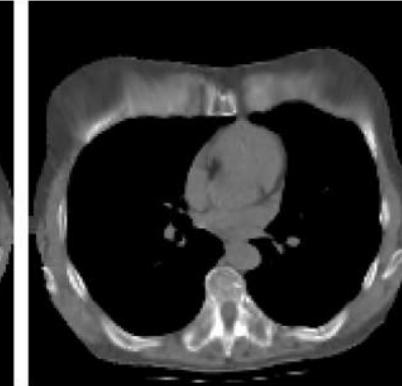
AIR



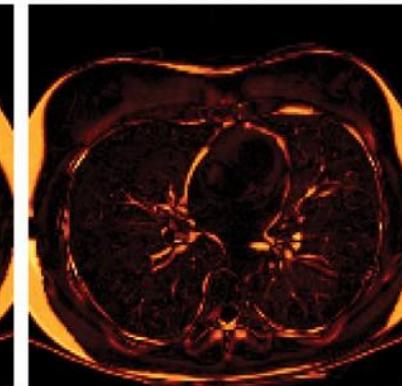
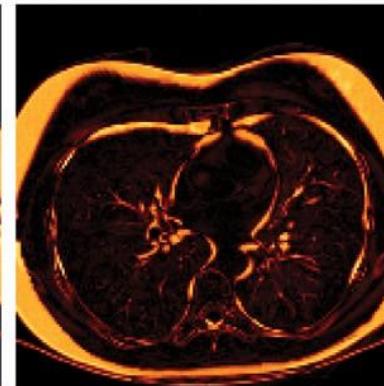
DIR-1



DIR-2

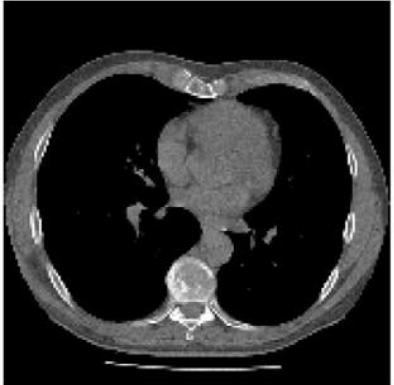


Fixed





Moving



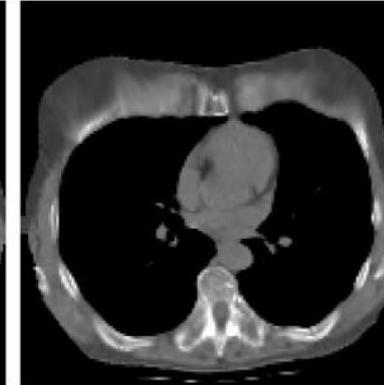
AIR



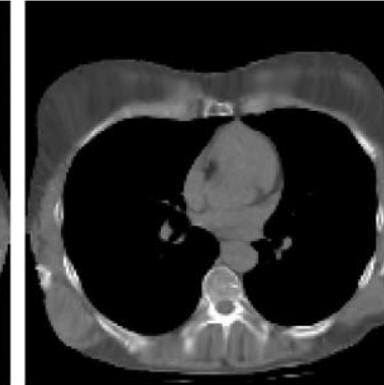
DIR-1



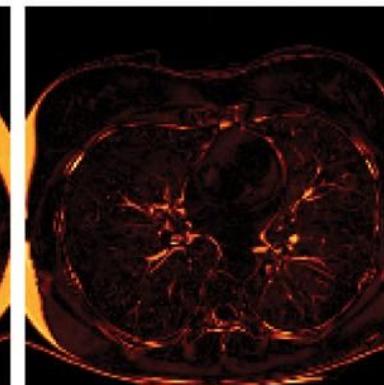
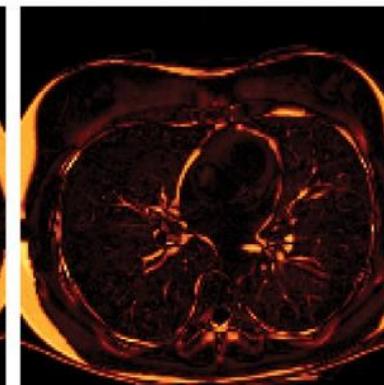
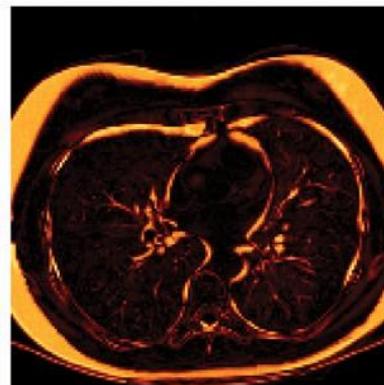
DIR-2

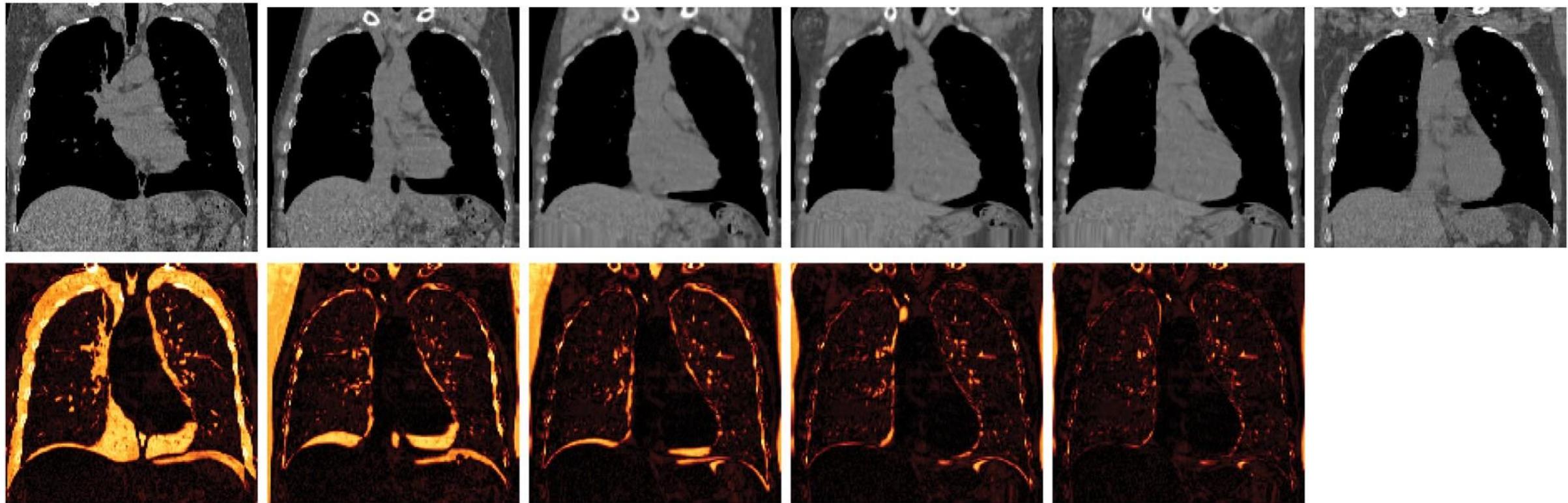


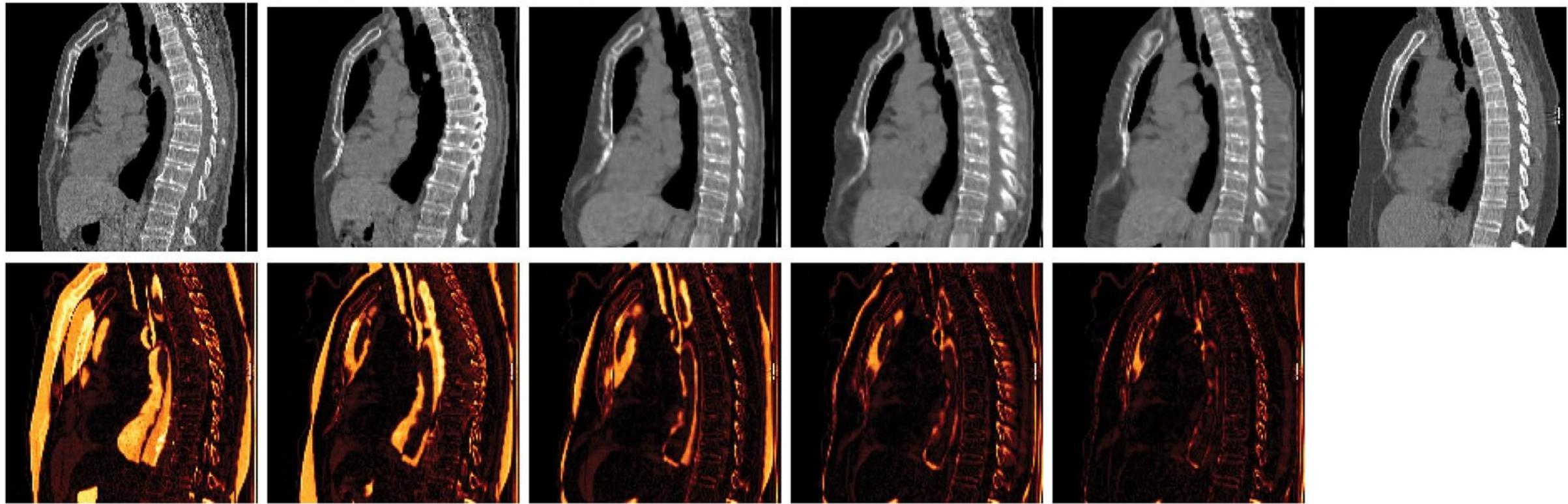
DIR-3



Fixed

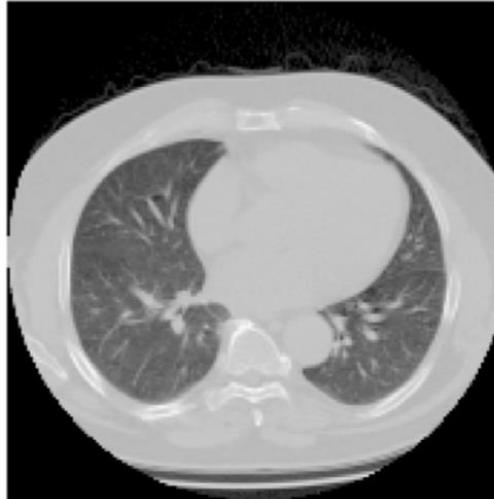




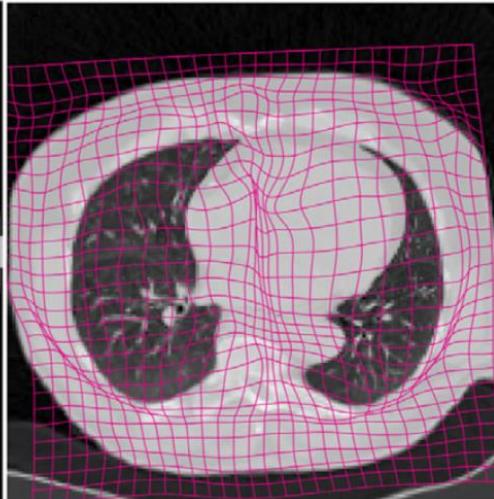




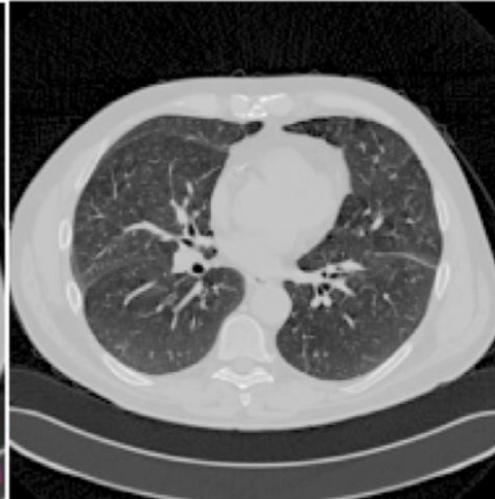
Fixed



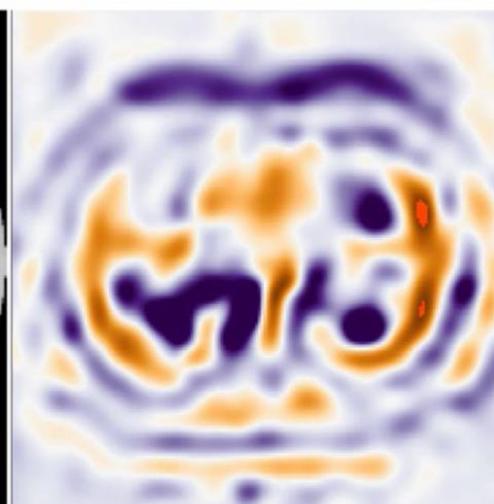
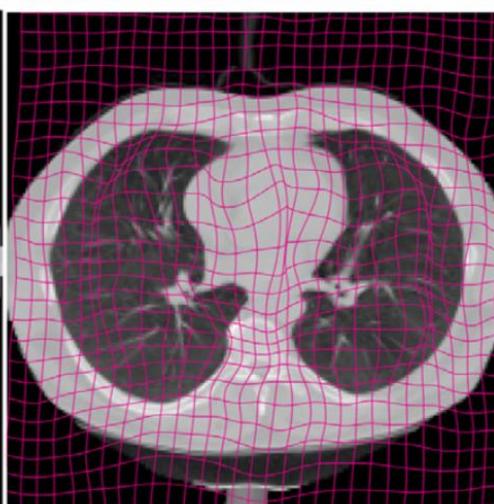
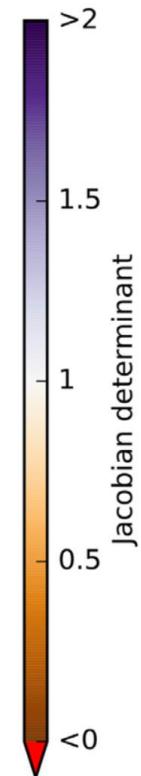
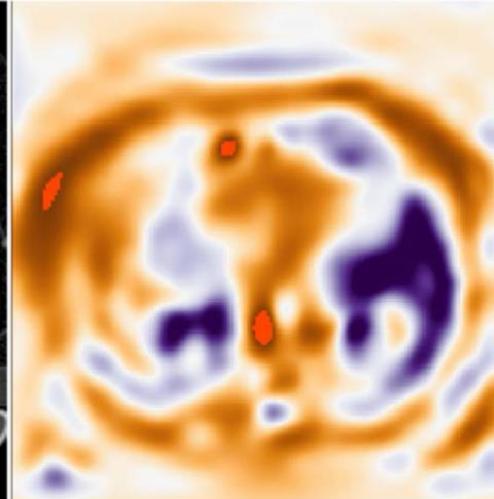
Warped



Moving



Jacobian





	Dice	HD	ASD	Fraction folding (%)	Std. dev. Jacobian	CPU time (s)	GPU time (s)
Before registration	0.31 ± 0.21	32.62 ± 12.21	9.21 ± 4.53	-	-	-	-
SE	AIR	0.60 ± 0.19	25.81 ± 15.34	4.89 ± 2.36	-	$3.73(0.26)$	-
	DIR-1	0.69 ± 0.11	20.30 ± 13.26	3.39 ± 1.11	0.00 ± 0.00	0.19 ± 0.11	$11.67(1.07)$
	DIR-2	0.75 ± 0.08	21.26 ± 11.31	2.67 ± 0.87	0.00 ± 0.08	0.27 ± 0.13	$14.83(3.37)$
	DIR-3	0.77 ± 0.08	20.83 ± 11.81	2.45 ± 0.89	0.04 ± 0.19	0.30 ± 0.15	$20.36(8.41)$
DLIR	AIR	0.58 ± 0.16	26.79 ± 13.05	5.24 ± 2.19	-	$1.02(0.29)$	$0.17(0.05)$
	DIR-1	0.64 ± 0.11	21.68 ± 13.09	3.86 ± 1.74	0.00 ± 0.00	0.16 ± 0.09	$3.85(0.99)$
	DIR-2	0.70 ± 0.10	19.95 ± 13.30	3.21 ± 1.15	0.00 ± 0.00	0.19 ± 0.10	$8.18(2.03)$
	DIR-3	0.75 ± 0.08	19.34 ± 13.41	2.46 ± 0.80	0.75 ± 1.08	0.45 ± 0.21	$15.41(4.38)$
							$0.43(0.10)$



	Dice	HD	ASD	Fraction folding (%)	Std. dev. Jacobian	CPU time (s)	GPU time (s)
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	DIR-1	0.69 ± 0.11	20.30 ± 13.26	3.39 ± 1.11	0.00 ± 0.00	0.19 ± 0.11	$11.67(1.07)$
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	DIR-1	0.64 ± 0.11	21.68 ± 13.09	3.86 ± 1.74	0.00 ± 0.00	0.16 ± 0.09	$3.85(0.99)$
	DIR-2	0.70 ± 0.10	19.95 ± 13.30	3.21 ± 1.15	0.00 ± 0.00	0.19 ± 0.10	$8.18(2.03)$
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SE	AIR	0.60 ± 0.19	25.81 ± 15.34	4.89 ± 2.36	–	$3.73(0.26)$	–
	DIR-1	0.69 ± 0.11	20.30 ± 13.26	3.39 ± 1.11	0.00 ± 0.00	0.19 ± 0.11	$11.67(1.07)$
	DIR-2	0.75 ± 0.08	21.26 ± 11.31	2.67 ± 0.87	0.00 ± 0.08	0.27 ± 0.13	$14.83(3.37)$
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DLIR	AIR	0.58 ± 0.16	26.79 ± 13.05	5.24 ± 2.19	–	$1.02(0.29)$	$0.17(0.05)$
	DIR-1	0.64 ± 0.11	21.68 ± 13.09	3.86 ± 1.74	0.00 ± 0.00	0.16 ± 0.09	$3.85(0.99)$
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	DIR-3	0.75 ± 0.08	19.34 ± 13.41	2.46 ± 0.80	0.75 ± 1.08	0.45 ± 0.21	$15.41(4.38)$



Evaluation on DIRLab data



- Inspiration-expiration chest CT
- Evaluation: TRE of manually annotated landmarks
- Coarse-to-fine image registration:
 - Stage 1: 5.12 (4.64) mm
 - Stage 2: 3.40 (4.17) mm
 - Stage 3: 2.64 (4.32) mm



Direct Automatic Coronary Calcium Scoring in Cardiac and Chest CT

Bob D. de Vos^{ID}, Jelmer M. Wolterink^{ID}, Tim Leiner, Pim A. de Jong,
Nikolas Lessmann^{ID}, and Ivana Išgum^{ID}

Abstract—Cardiovascular disease (CVD) is the global leading cause of death. A strong risk factor for CVD events is the amount of coronary artery calcium (CAC). To meet the demands of the increasing interest in quantification of CAC, i.e., coronary calcium scoring, especially as an unrequested finding for screening and research, automatic methods have been proposed. The current automatic cal-

need for early detection and treatment of individuals with CVD or those who are at high cardiovascular risk due to the presence of one or more risk factors [2]. A strong and independent risk factor for CVD events, e.g. myocardial infarction, is the quantity of coronary artery calcium (CAC) [3]–[5]. Quantification of CAC, i.e. calcium scoring, is typically per-

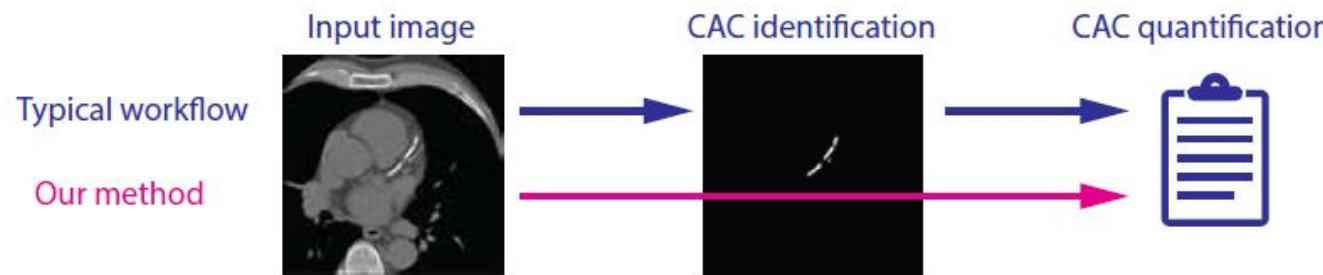
Group-wise image registration

weighted Cohen's kappa of 0.95 in cardiac CT and 0.93 in chest CT. Performance is similar to that of the state-of-the-art methods, but the proposed method is hundreds of times faster. By providing visual feedback, insight is given in the decision process, making it readily implementable in clinical

manually identify CAC in CT image slices. This is a tedious process of finding and selecting high density voxels in the coronary arteries—commonly defined as two or more connected voxels above 130 Hounsfield Units (HU). In scans not dedicated to calcium scoring, this can be particularly



Direct calcium scoring



For real-time automatic cardiovascular disease (CVD) risk categorization



Cardiac CT



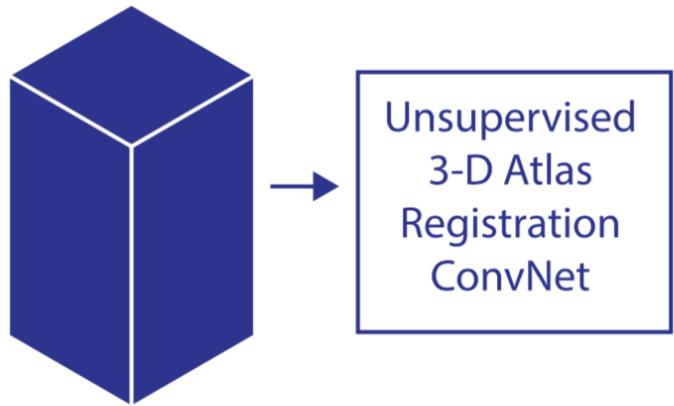
Cardiac CT



Chest CT

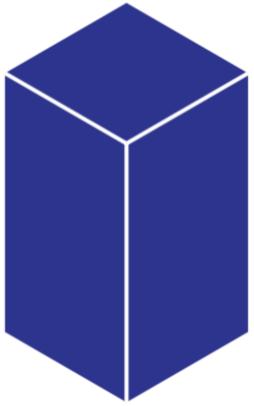


Input Image

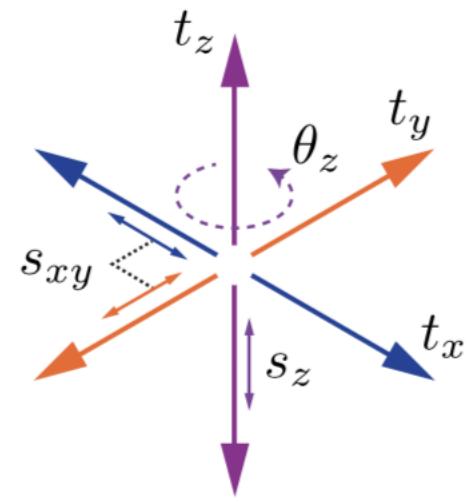




Input Image

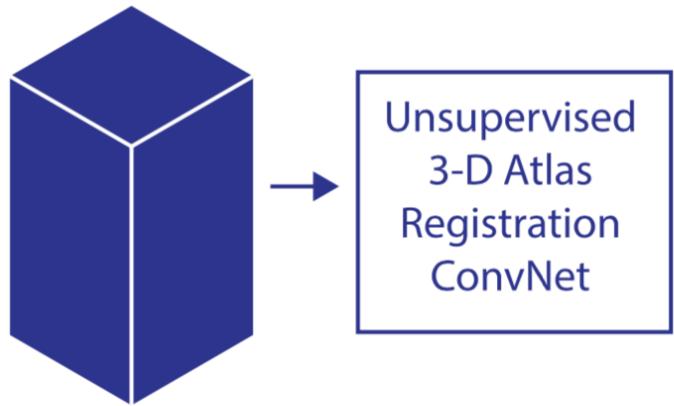


Unsupervised
3-D Atlas
Registration
ConvNet



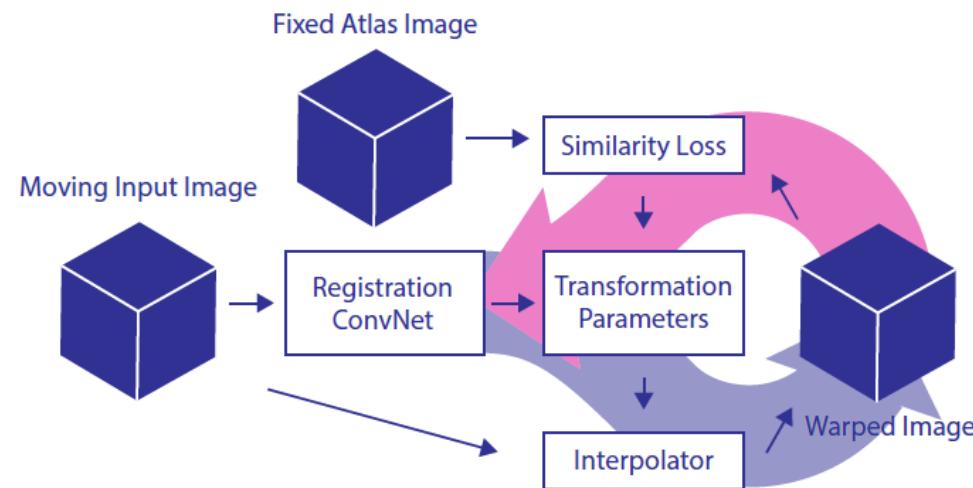


Input Image



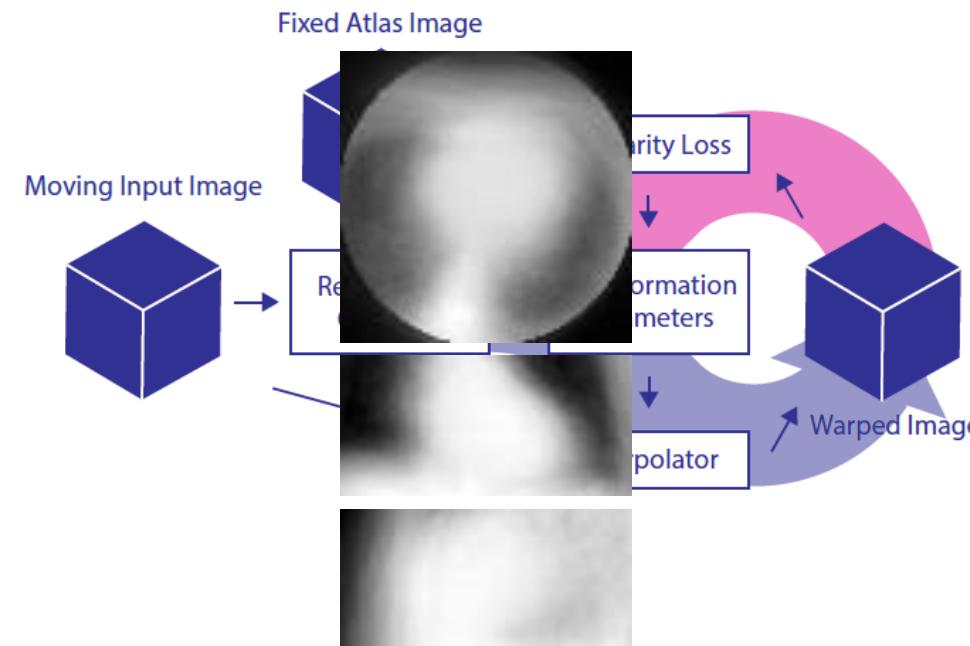
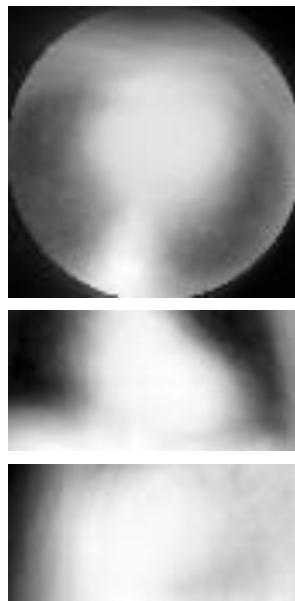


Group-wise image registration framework





Creating an atlas-image from cardiac CT



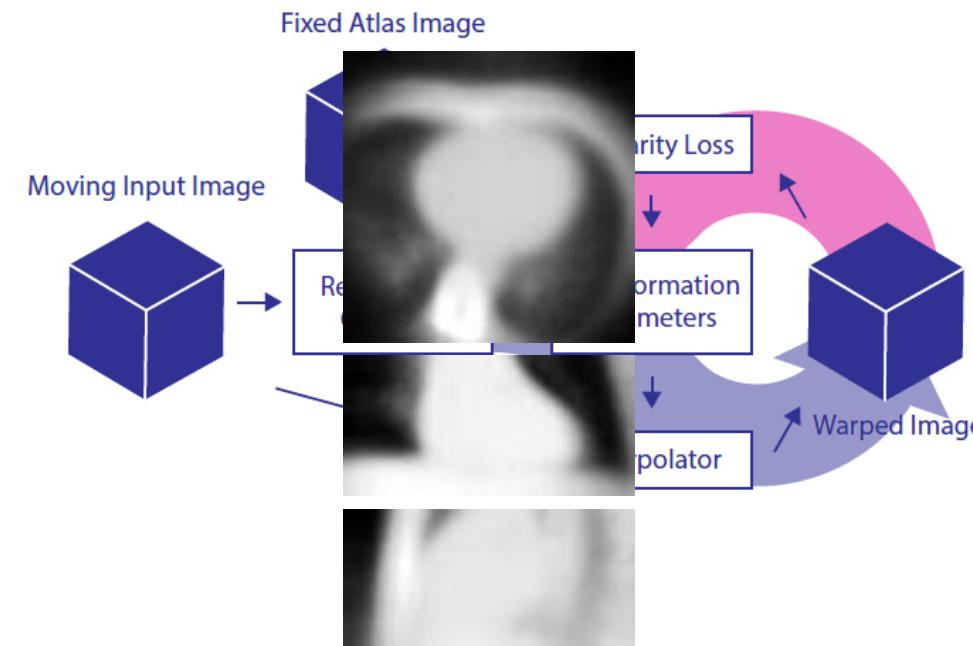
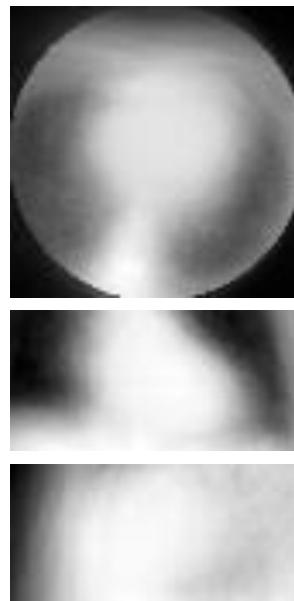
Average of training set
before registration

Fixed image

Average of training set
after registration



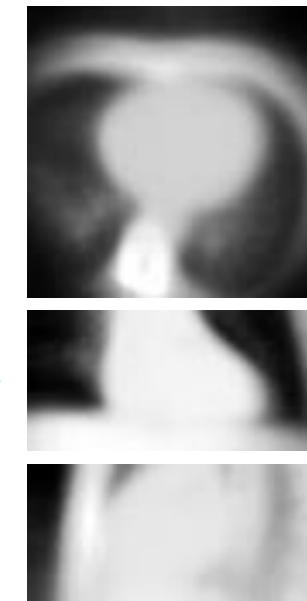
Creating an atlas-image from cardiac CT



Average of training set
before registration

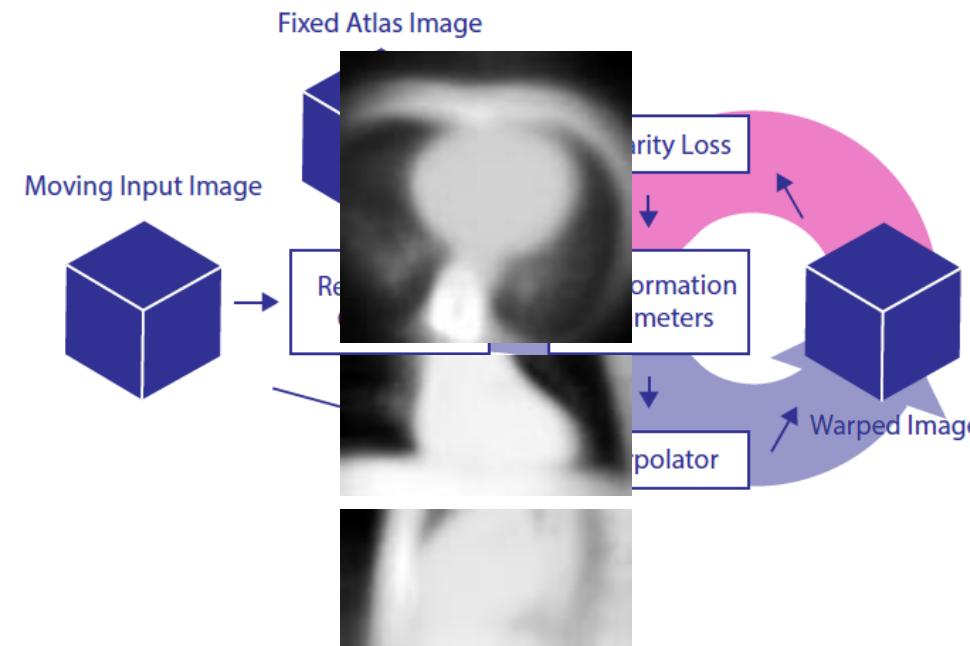
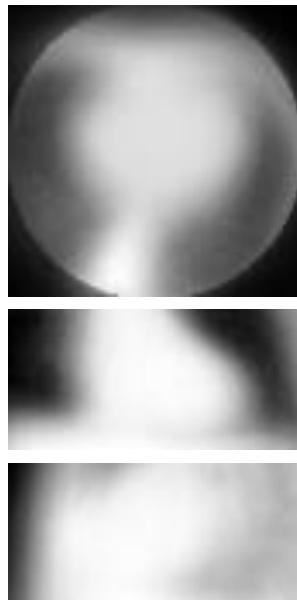
Refined atlas

Average of training set
after registration





Group-wise cardiac CT registration



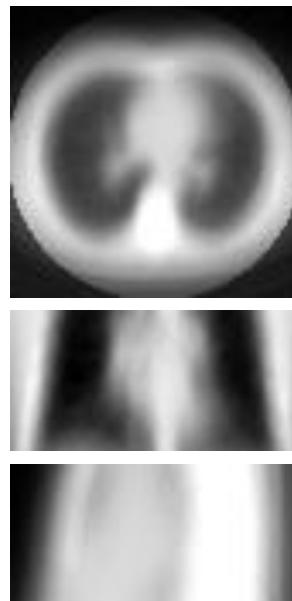
Average of test-set
before registration

Refined atlas

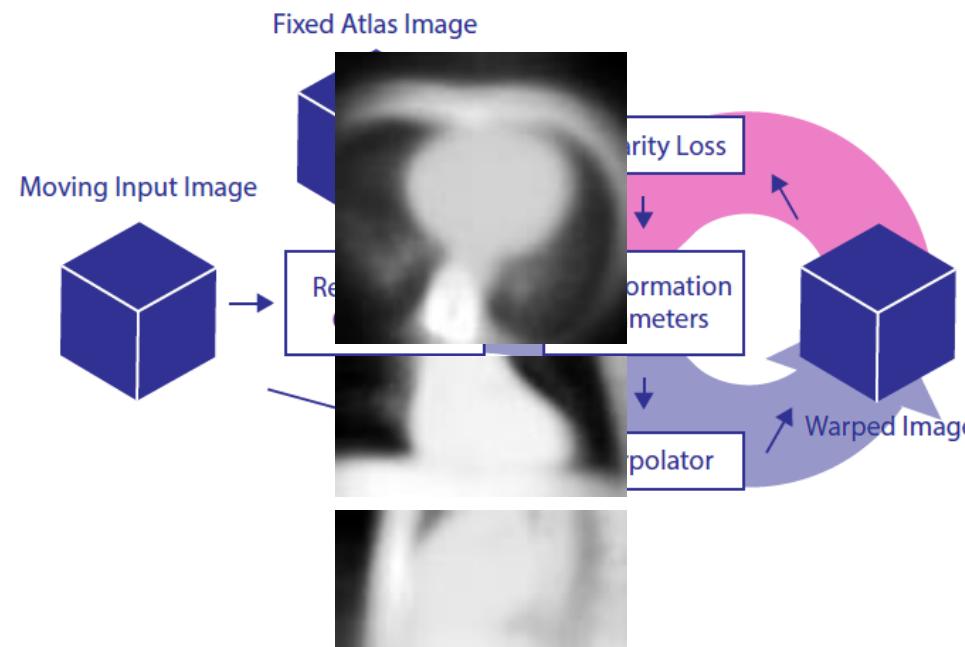
Average of test-set
after registration



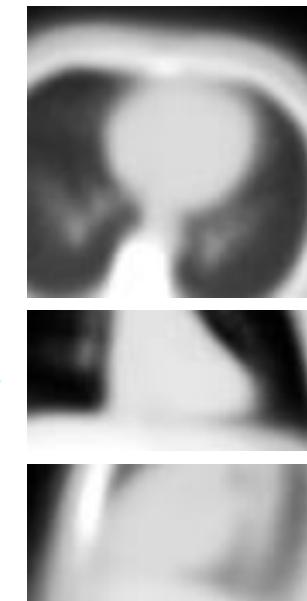
Group-wise chest CT registration



Average of moving images
before registration



Refined atlas



Average of moving images
after registration



Registration speed

- Current deep learning-based image registration is fast
 - Cardiac MR (256x256) 2D → ~10 ms
 - Cardiac MR (256x256x10) 3D single stage → ~30 ms
 - Chest CT (resampled 110³ - 220³) → ~0.43 s



Take-home messages

- There is not yet a one-size-fits-all solution
- Training takes some time, but inference is quick
- Inference time is mainly limited by GPU I/O-time
- High GPU-memory requirement for native resolution
- Large set of training data required, but training is unsupervised
- Easily implementable using existing building blocks from DL libraries
- Techniques from conventional image registration can be readily applied to unsupervised deep learning image registration