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

Lasse Hansen, Mattias P. Heinrich

Ha Young Kim



### **Motivation**



Knowing airflow in the lung helps diagnose disease such as COPD (Chronic Obstructive Pulmonary Disease), Asthma, lung cancer

Current DL method doesn't reach an acceptable accuracy

Challenge :

Large motion estimation of the lung between inhale and exhale

Presents a lightweight U - Net like architecture to predict large deformation displacement field that allows inhale to exhale CT scan registration



### Method



#### Preprocessing



- Fixed Image (Inhale), Moving Image (Exhale)
  - 1. Preprocess the image
  - 2. **Sparse** Keypoint extraction using Foerstner interest operator + max pooling
  - 3. Feature extraction using MIND (Modality Independent Neighbourhood Descriptor)
  - 4. Cost tensor generation using the extracted features p : keypoints , l : displacement locations

$$C(\mathbf{p}, \mathbf{l}) = \frac{1}{12} \sum_{i=0}^{11} (F_F^i(\mathbf{p}) - F_M^i(\mathbf{p} + \mathbf{l}))^2$$

#### GraphRegNet



GraphRegNet takes Cost tensor as input and generate displacement vector for each keypoints

U - Net - like architecture consisted of CNN - GNN - CNN



#### **GraphRegNet - Encoder**



#### CNN

3 convolutional layers, instance normalization, leaky ReLU

Generate low dimensional displacement embedding

### **GraphRegNet - GNN**





Predicted low dimensional displacement embeddings are concatenated with coordinates of respective keypoints

The concatenated output is distributed across kNN graph of keypoints (k = 15)

Perform three graph convolutions defined as

$$\mathbf{f}'_i = ReLU(\underset{(i,j)\in E}{\operatorname{avg}} \mathbf{e}_{ij}) \qquad \mathbf{e}_{ij} = h_{\theta}(\mathbf{f}_i, \mathbf{f}_j - \mathbf{f}_i)$$

f : feature vector at point keypoint pi
h : inner product of parameters with the keypoint fi and neighbourhood features fj-fi
Achieves spatial regularization
Helps with smoothing the image



#### **GraphRegNet - Decoder**



#### CNN

Two Upconvolutions + single convolutional layer

Generates single channel feature map for each keypoint (Hp)

#### GraphRegNet

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#### **GraphRegNet - Sparse to Dense supervision**



Final displacement vector (**d**) obtained by integrating the generated feature map (Hp) over the displacement search region (l)

$$\mathbf{d} = \sum_{\mathbf{l}\in\mathcal{L}} \mathbf{l}\cdot\tilde{H}_{\mathbf{p}}(\mathbf{l}).$$

The sparse displacement vector (**d**) is accumulated in a dense tensor at respective keypoints using trilinear extrapolation which yields final displacement field **D** 





# Training

Dataset inhale / exhale lung CT scan datasets
 OIR - Lab 4D CT

 Normal resting breathing
 COPD gene dataset
 Breath - hold CT scans, have larger deformations

For training, additional dataset - Empire10, POPI - are added to have total of 45 training pair

5 - fold cross validation used. First fold for hyperparameter tuning



## **Evaluation**

Evaluation metric

- TRE (Target Registration Error) between expert annotated landmarks
- Jacobian determinants

Evaluation

- Compare the performance with recent DL approaches
   DLIR, Ep18, OSL, LRN, mlVN, BMRF, VM+, LapIRN, FE+, PDD+, MST
   + means the original architecture is modified to fit for both DIR and COPD dataset
- Ablation studies

RW (Random Walk)Noreg (GNN removed)Coords (k = 1 in kNN graph)SI (no refinement stage)Unif (uniformly sample keypoints, non - distinctive keypoints)



### **Results - TRE** Comparison

	init.	DLIR [51]	Ep18 [52]	OSL [53]	LRN [21]	mlVN [20]	BMRF [54]	VM+ [50]	LapIRN [6]	FE+ [55]	PDD+ [16]	MST	ours
4DCT 01 4DCT 02 4DCT 03 4DCT 04 4DCT 05 4DCT 06 4DCT 07 4DCT 08 4DCT 09 4DCT 10	03.89 04.34 06.94 09.83 07.48 10.89 11.03 14.99 07.92 07.30	$\begin{array}{c} 1.27 \\ 1.20 \\ 1.48 \\ 2.09 \\ 1.95 \\ 5.16 \\ 3.05 \\ 6.48 \\ 2.10 \\ 2.09 \end{array}$	$1.45 \\ 1.46 \\ 1.57 \\ 1.95 \\ 2.07 \\ 3.04 \\ 3.41 \\ 2.80 \\ 2.18 \\ 1.83$	$\begin{array}{c} 1.21 \\ 1.13 \\ 1.32 \\ 1.84 \\ 1.80 \\ 2.30 \\ 1.91 \\ 3.47 \\ 1.47 \\ 1.79 \end{array}$	0.98 0.98 1.14 <b>1.39</b> <b>1.43</b> 2.26 <b>1.42</b> 3.13 <b>1.27</b> 1.93	1.33 1.33 1.48 1.85 1.84 3.57 2.61 2.62 2.70 2.63		1.46 1.51 2.31 2.72 2.69 3.07 3.01 6.22 2.94 3.00	$\begin{array}{c} 1.00\\ 1.28\\ 2.18\\ 3.05\\ 2.36\\ 1.78\\ 2.24\\ 2.24\\ 2.26\\ 1.90 \end{array}$	2.20 3.89 2.71 2.95 3.03 3.36 3.10 2.94 2.86 2.99	$\begin{array}{c} 0.90\\ 0.91\\ 1.06\\ 1.66\\ 1.68\\ 1.86\\ 1.94\\ 1.79\\ 1.94\\ 2.03\\ \end{array}$	<b>0.82</b> <b>0.87</b> 1.09 1.63 1.58 1.71 1.73 1.55 1.85 1.90	0.86 0.90 1.06 1.45 1.60 <b>1.59</b> 1.74 1.46 1.58 <b>1.71</b>
avg std sig. level	08.46 06.58 ***	2.64 4.32	2.17 1.89	1.83 2.35	1.59 1.58	2.19 1.62		2.89 2.21 ***	2.03 1.89 ***	3.00 1.70 ***	1.57 1.36 ***	1.47 1.26 ***	<b>1.39</b> 1.29
COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 06 COPD 07 COPD 08 COPD 09 COPD 10	26.33 21.79 12.64 29.58 30.08 28.46 21.60 26.46 14.86 21.81						$ \begin{array}{r} 1.51\\ 2.27\\ 1.39\\ 1.86\\ 1.46\\ 1.40\\ 1.46\\ 1.53\\ 1.34\\ 1.71\\ 1.50\\ \end{array} $	9.95 9.96 4.41 7.08 9.19 8.12 7.10 7.92 6.93 9.16	6.85 6.90 1.51 6.38 6.81 4.19 2.73 4.32 3.60 6.59	4.89 7.30 2.89 5.46 5.19 5.53 4.40 3.94 3.57 4.44	2.57 4.01 1.46 2.19 2.22 1.89 1.62 1.72 1.51 2.43	1.42 3.42 1.32 <b>1.48</b> 1.44 1.47 1.37 1.33 1.22 1.55	1.38 2.09 1.22 1.58 1.37 1.10 1.19 1.19 0.99 1.38
avg std sig. level	23.36 11.86 ***						1.59 0.27	7.98 3.75 ***	4.99 3.94 ***	4.76 4.06 ***	2.16 2.63 ***	1.60 2.04 ***	<b>1.34</b> 1.44

DIR - Lab 4D CT

Improves LRN by 13% (1.59→1.39)

COPD

Better performance in all cases



# **Results - TRE** Ablation studies

	RW	noreg	coords	sl	unif.	ours
4DCT 01	1.21	1.40	0.86	0.86	0.89	0.86
4DCT 02	1.17	1.64	0.98	0.90	0.93	0.90
4DCT 03	1.37	1.50	1.11	1.13	1.05	1.06
4DCT 04	1.52	2.05	1.65	1.61	1.51	1.45
4DCT 05	2.11	2.91	1.73	1.67	1.68	1.60
4DCT 06	1.83	2.19	1.60	1.64	1.59	1.59
4DCT 07	1.88	2.33	1.67	1.69	1.63	1.74
4DCT 08	1.77	2.88	2.28	1.58	1.43	1.46
4DCT 09	2.23	2.23	1.72	1.87	1.72	1.58
4DCT 10	1.97	2.43	1.75	1.97	2.26	1.71
avg	1.70	2.15	1.53	1.49	1.47	1.39
std	2.38	1.70	1.57	1.30	1.65	1.29
sig. level	***	***	***	***	*	
sig. level	***	***	***	***	*	1.38
sig. level COPD 01 COPD 02	*** 3.51 5.26	*** 4.32 7.27	*** 5.50 9.12	*** 1.71 2.75	* 1.80 <b>2.09</b>	1.38 2.09
sig. level COPD 01 COPD 02 COPD 03	*** 3.51 5.26 1.57	*** 4.32 7.27 1.42	*** 5.50 9.12 1.40	*** 1.71 2.75 1.42	* 1.80 <b>2.09</b> 1.18	<b>1.38</b> <b>2.09</b> 1.22
sig. level COPD 01 COPD 02 COPD 03 COPD 04	*** 3.51 5.26 1.57 2.51	*** 4.32 7.27 1.42 7.30	*** 5.50 9.12 1.40 4.46	*** 1.71 2.75 1.42 2.06	* 1.80 2.09 1.18 1.60	<b>1.38</b> <b>2.09</b> 1.22 1.58
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05	*** 3.51 5.26 1.57 2.51 3.33	*** 4.32 7.27 1.42 7.30 4.77	*** 5.50 9.12 1.40 4.46 3.44	*** 1.71 2.75 1.42 2.06 1.81	* 1.80 2.09 1.18 1.60 1.49	<b>1.38</b> <b>2.09</b> 1.22 1.58 <b>1.37</b>
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 06	*** 3.51 5.26 1.57 2.51 3.33 2.57	*** 4.32 7.27 1.42 7.30 4.77 3.58	*** 5.50 9.12 1.40 4.46 3.44 2.96	*** 1.71 2.75 1.42 2.06 1.81 1.43	* 1.80 2.09 1.18 1.60 1.49 1.31	<b>1.38</b> <b>2.09</b> 1.22 1.58 <b>1.37</b> <b>1.10</b>
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 06 COPD 07	*** 3.51 5.26 1.57 2.51 3.33 2.57 2.14	*** 4.32 7.27 1.42 7.30 4.77 3.58 2.68	*** 5.50 9.12 1.40 4.46 3.44 2.96 2.99	*** 1.71 2.75 1.42 2.06 1.81 1.43 1.64	* 1.80 2.09 1.18 1.60 1.49 1.31 1.23	<ol> <li>1.38</li> <li>2.09</li> <li>1.22</li> <li>1.58</li> <li>1.37</li> <li>1.10</li> <li>1.19</li> </ol>
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 06 COPD 07 COPD 08	*** 3.51 5.26 1.57 2.51 3.33 2.57 2.14 1.64	*** 4.32 7.27 1.42 7.30 4.77 3.58 2.68 4.21	*** 5.50 9.12 1.40 4.46 3.44 2.96 2.99 2.22	*** 1.71 2.75 1.42 2.06 1.81 1.43 1.64 1.54	* 1.80 2.09 1.18 1.60 1.49 1.31 1.23 1.44	<ul> <li>1.38</li> <li>2.09</li> <li>1.22</li> <li>1.58</li> <li>1.37</li> <li>1.10</li> <li>1.19</li> <li>1.19</li> </ul>
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 04 COPD 05 COPD 06 COPD 07 COPD 08 COPD 09	*** 3.51 5.26 1.57 2.51 3.33 2.57 2.14 1.64 2.79	*** 4.32 7.27 1.42 7.30 4.77 3.58 2.68 4.21 3.02	*** 5.50 9.12 1.40 4.46 3.44 2.96 2.99 2.22 1.68	*** 1.71 2.75 1.42 2.06 1.81 1.43 1.64 1.54 1.45	* 1.80 2.09 1.18 1.60 1.49 1.31 1.23 1.44 1.13	<ul> <li>1.38</li> <li>2.09</li> <li>1.22</li> <li>1.58</li> <li>1.37</li> <li>1.10</li> <li>1.19</li> <li>1.19</li> <li>0.99</li> </ul>
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 06 COPD 07 COPD 08 COPD 09 COPD 10	*** 3.51 5.26 1.57 2.51 3.33 2.57 2.14 1.64 2.79 2.62	*** 4.32 7.27 1.42 7.30 4.77 3.58 2.68 4.21 3.02 7.93	*** 5.50 9.12 1.40 4.46 3.44 2.96 2.99 2.22 1.68 6.95	*** 1.71 2.75 1.42 2.06 1.81 1.43 1.64 1.54 1.45 1.79	* 1.80 2.09 1.18 1.60 1.49 1.31 1.23 1.44 1.13 1.82	1.38 2.09 1.22 1.58 1.37 1.10 1.19 1.19 0.99 1.38
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 05 COPD 06 COPD 07 COPD 08 COPD 09 COPD 10	*** 3.51 5.26 1.57 2.51 3.33 2.57 2.14 1.64 2.79 2.62 2.79	*** 4.32 7.27 1.42 7.30 4.77 3.58 2.68 4.21 3.02 7.93 4.65	*** 5.50 9.12 1.40 4.46 3.44 2.96 2.99 2.22 1.68 6.95 4.07	*** 1.71 2.75 1.42 2.06 1.81 1.43 1.64 1.54 1.45 1.79 1.76	* 1.80 2.09 1.18 1.60 1.49 1.31 1.23 1.44 1.13 1.82 1.50	1.38 2.09 1.22 1.58 1.37 1.10 1.19 1.19 0.99 1.38 1.34
sig. level COPD 01 COPD 02 COPD 03 COPD 04 COPD 05 COPD 06 COPD 07 COPD 08 COPD 09 COPD 10 avg std	*** 3.51 5.26 1.57 2.51 3.33 2.57 2.14 1.64 2.79 2.62 2.79 4.51	*** 4.32 7.27 1.42 7.30 4.77 3.58 2.68 4.21 3.02 7.93 4.65 5.89	*** 5.50 9.12 1.40 4.46 3.44 2.96 2.99 2.22 1.68 6.95 4.07 5.57	**** 1.71 2.75 1.42 2.06 1.81 1.43 1.64 1.54 1.54 1.79 1.76 1.57	* 1.80 2.09 1.18 1.60 1.49 1.31 1.23 1.44 1.13 1.82 1.50 1.75	1.38 2.09 1.22 1.58 1.37 1.10 1.19 0.99 1.38 1.34 1.44

RW (Random walk)

- ~52% improved when using deep learning approach
- Noreg (GNN removed)
- ~71% improvement when using GNN
- $\circ$  Coords (k = 1 in kNN graph)
- ~67% improvement when exploiting neighborhood information
- SI (no refinement stage)
- ~24% improvement with two level approach
- Unif (non distinctive keypoints , uniformly sampled)
- ~11% improvement when using distinctive keypoint



#### **Results** Jacobian Determinant



	GraphRegNet	VM+	LapIRN	FE+	PDD+	MST
4D CT	0.13	0.11	0.12	0.15	0.10	0.10
COPD	0.21	0.20	0.17	0.32	0.19	0.18

Fraction of negative values Shows image foldings 4DCT : 0.02 - 0.21 % COPD : 0.15 - 0.83%

Standard deviation Smoothness of the transformation Closer to 0 : smooth transformation



Standard deviation of Jacobian determinant

#### Results

 Registration accuracy improves LungRegNet(LRN) by 13% but fails to reach the accuracy of Rühaak et al.

• Shows good improvement with COPD gene dataset

 $\circ$  Good U - Net like architecture design and keypoint based registration shows improvement in TRE by 70 %

 $\circ$  Outstanding improvement in computation time 2s for single registration 5min (Rühaak et al)  $\rightarrow$  2s

9 min for single scan pair, previously takes around 3h



#### Take Home message & Personal Review

• By using sparse keypoint, can decrease computation time

 GraphRegNet shows great potential of fast large deformation estimation algorithm

 $\circ$  By adding a GNN, it helps with smooth transform and spatial regularization

Future research sounds promising, trying different keypoint/feature extraction might improve the accuracy with a fast registration time
Code are given on Github, well drawn diagram



#### **Discussion**

Why is COPD performing better?

Is it a reliable comparison method of COPD since the architecture is changed?

