



UNIVERSITY OF COPENHAGEN

Extraction of Airways from Volumetric Data

A graph refinement view

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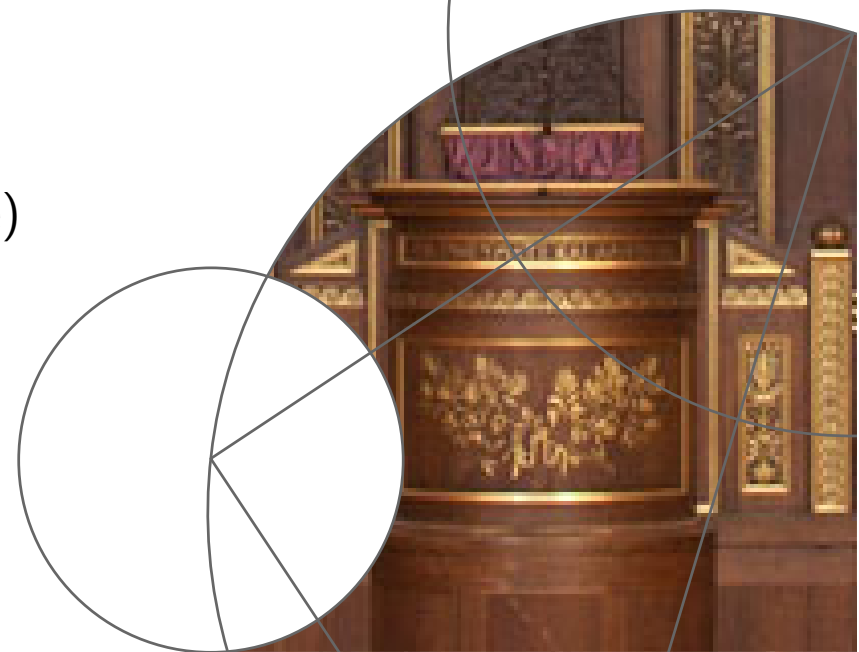
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[@raghavian](https://twitter.com/raghavian)

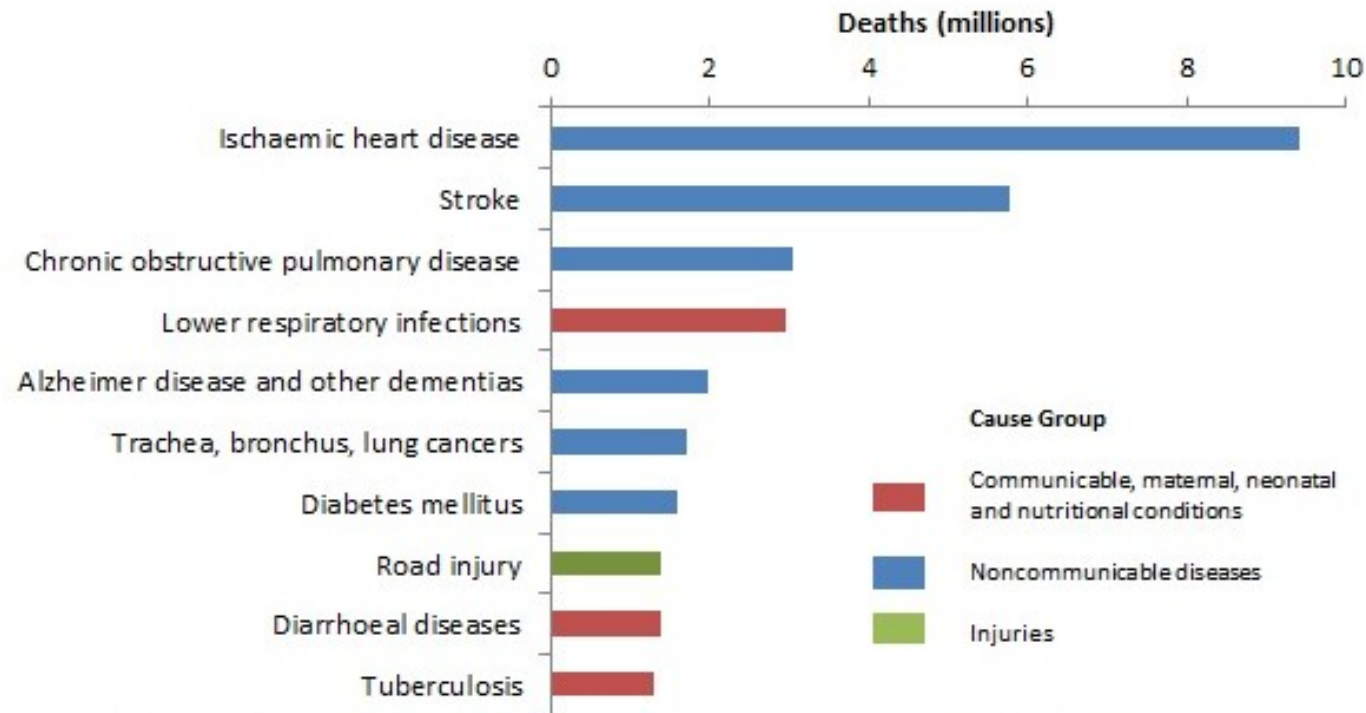
<https://raghavian.github.io>

December 16, 2020



Respiratory diseases: Major cause of morbidity & mortality

Top 10 global causes of deaths, 2016



Source: Global Health Estimates 2016, World Health Organization, 2018



Outline

- ① Objective of the study
- ② Data
- ③ Graph Refinement Models
- ④ Summary & Conclusions
- ⑤ Supplementary material



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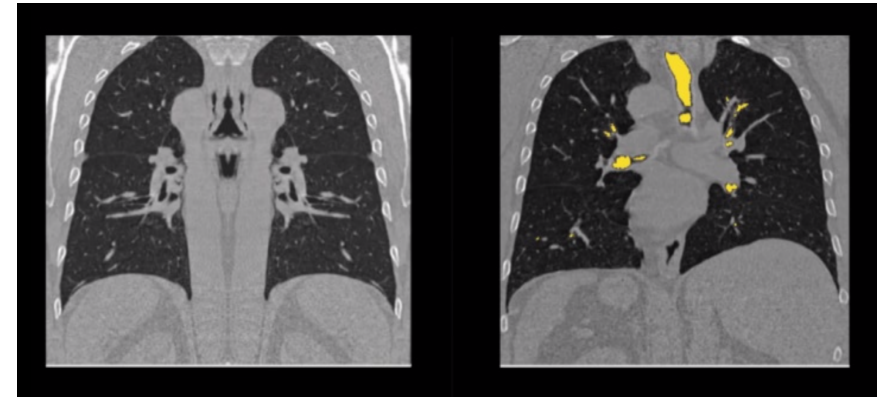
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Imaging based analysis of airways & challenges

Three primary steps:

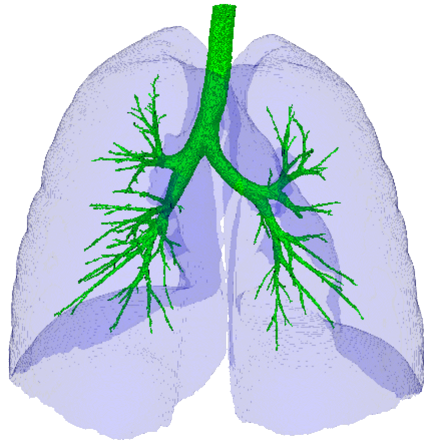
- ① **Detection** of airways
- ② Measurement of airway morphology
- ③ Deriving biomarkers



Coronal view of chest CT scan



Methods exist. Majority of them are sequential



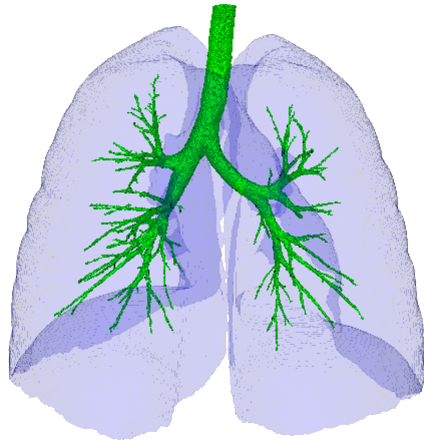
Sequential segmentation methods

Lo, P., et.al : Extraction of airways from CT (EXACT'09). IEEE Transactions on Medical Imaging, (2012)

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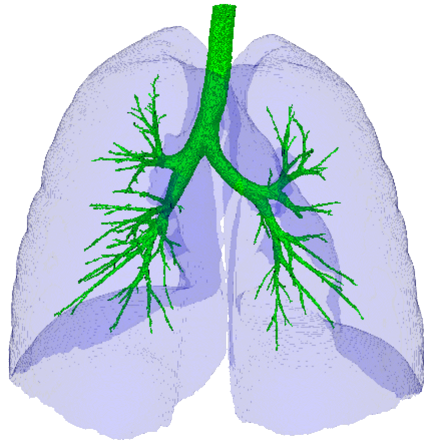
- Susceptible to occlusions in data
- Small branches are challenging
- EXACT'09 Study
 - Airway extraction challenge
 - Compares 15 methods
 - 10 use region growing!
- Quite recently some U-net based attempts on patches

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Objective of this work

Extraction of airways from volumetric data

With automatic methods that:

- Are exploratory
- Use more global information in local decisions



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Data from Danish Lung Cancer Screening Trial (DLCST)

- > 10,000 Low-dose CT from 2052 subjects
- Smoker or former smoker (> 20 pack years)
- Voxels $\sim 0.75 \times 0.75 \times 1 \text{ mm}^3$
- 32 scans with manual annotations for evaluation
- And additional 100 with automatic segmentations for hyperparameter tuning

Pedersen, J. H., et.al : The Danish randomized lung cancer CT screening trial – Overall design and results of the prevalence round. *Journal of Thoracic Oncology*, (2009)



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Graph Refinement Models

Work based on

- [1] Raghavendra Selvan, Thomas Kipf, Max Welling, Antonio GU Juarez, Jesper H. Pedersen, Jens Petersen, and Marleen de Bruijne. “Graph Refinement based airway extraction using Mean-Field Networks and Graph Neural Networks” Preprint/Medical Image Analysis (2018/2020).
- [2] Raghavendra Selvan, Max Welling, Jesper H. Pedersen, Jens Petersen, and Marleen de Bruijne. “Mean field network based graph refinement with application to airway tree extraction.” 21st Conference on Medical Image Computing & Computer Assisted Intervention (MICCAI 2018), pp. 750-758, Cham. Springer International Publishing.
- [3] Raghavendra Selvan, Thomas Kipf, Max Welling, Jesper H. Pedersen, Jens Petersen, and Marleen de Bruijne. “Extraction of Airways using Graph Neural Networks.” 1st Conference on Medical Imaging with Deep Learning (MIDL 2018), Amsterdam.



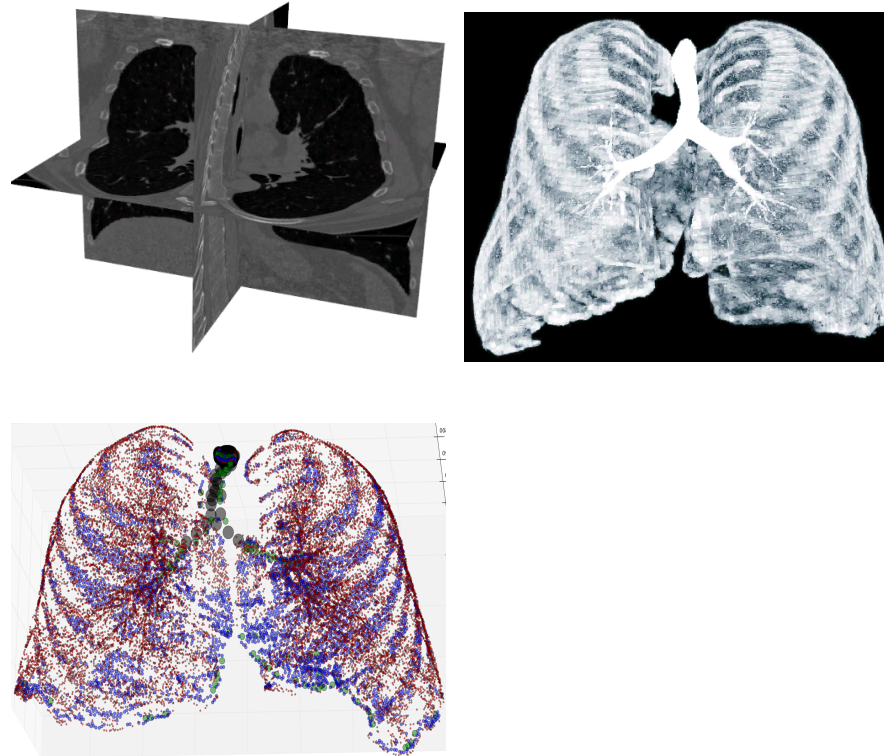
Graph Refinement Model for Airway Extraction

High level idea

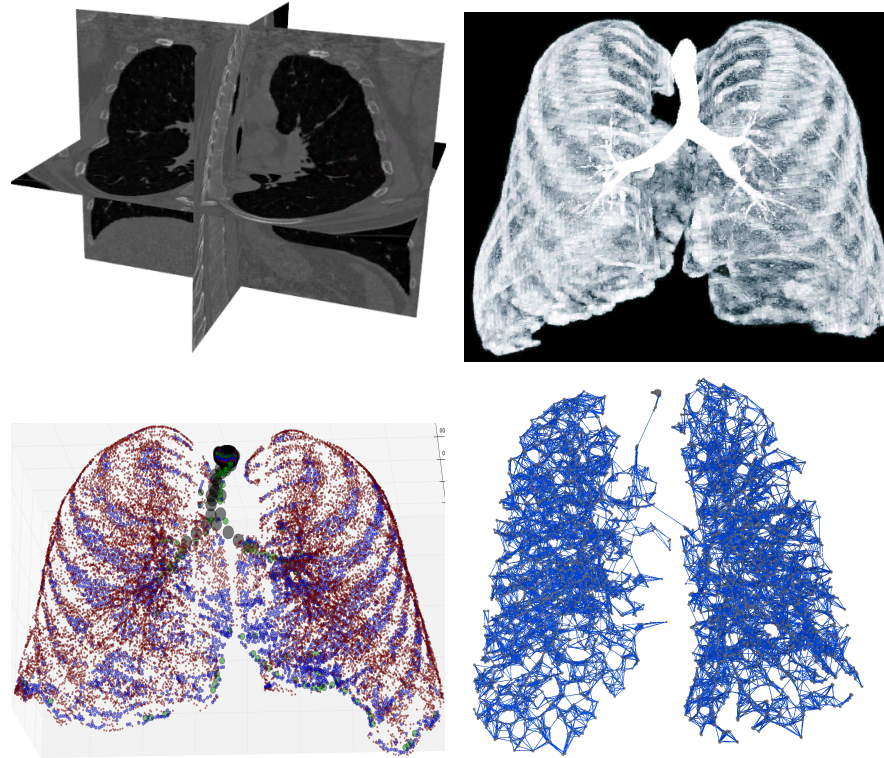
- Assume over-connected graphs with node attributes
- Optimise global connectivity, instead of qualifying individual branches



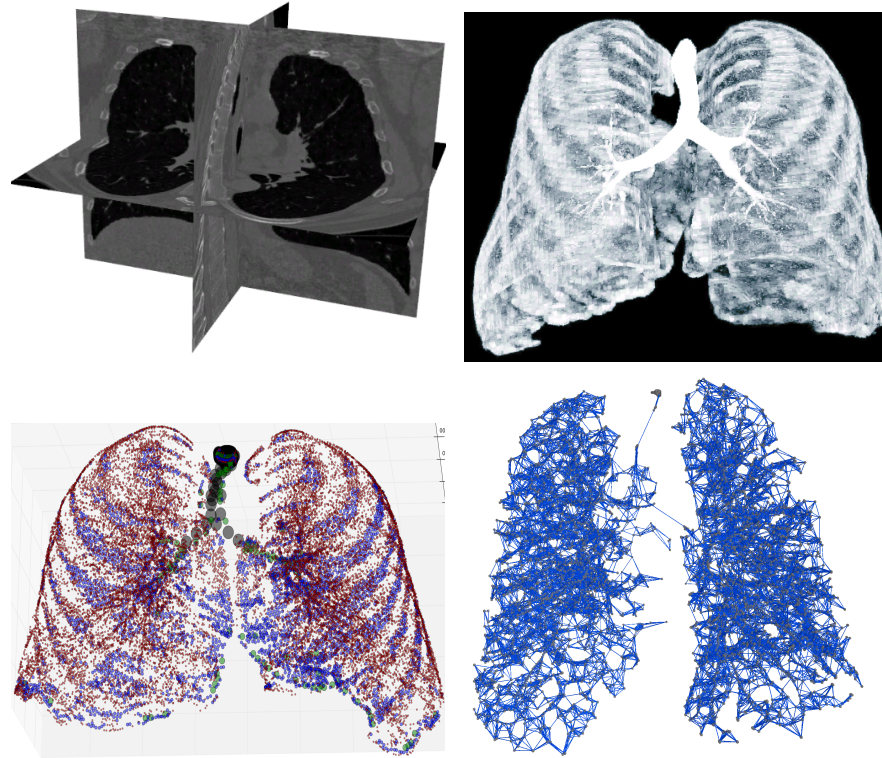
Volumetric data to Graph data



Volumetric data to Graph data



Volumetric data to Graph data



- Overconnected input graph: $\mathcal{G}_{in} : \{\mathcal{V}, \mathcal{E}_{in}\}$, with $|\mathcal{V}| = N$
- Node features: $\mathbf{X} \in \mathbb{R}^{F \times N}$
- Input adjacency: $\mathbf{A}_{in} \in \{0, 1\}^{N \times N}$

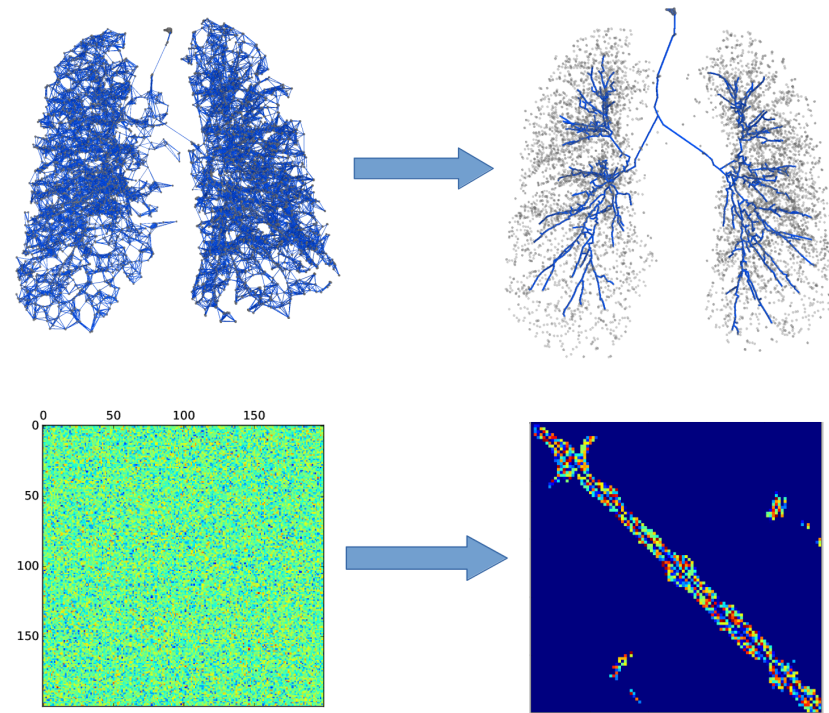


Airway extraction as Graph Refinement task

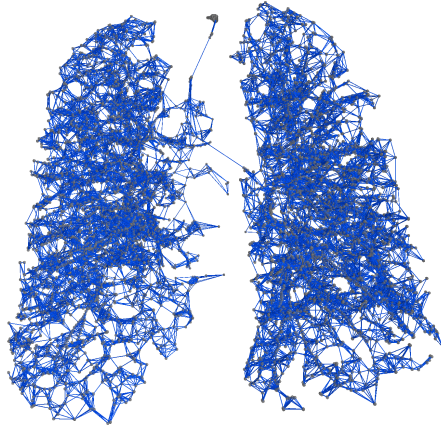
Graph Refinement Model

$$f(\cdot) : \mathcal{G}_{in} \mapsto \mathcal{G}$$

Output subgraph \mathcal{G} with $\mathcal{E} \subset \mathcal{E}_{in}$; $\mathbf{A} \in \{0, 1\}^{N \times N}$



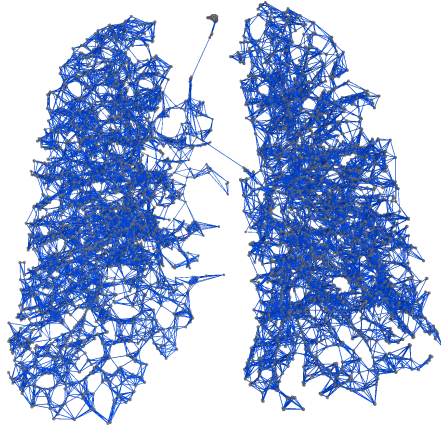
Probabilistic Graphical Model for MFN



- Binary random variable $s_{ij} \in \{0, 1\}$ with prob. $\alpha_{ij} \in [0, 1]$



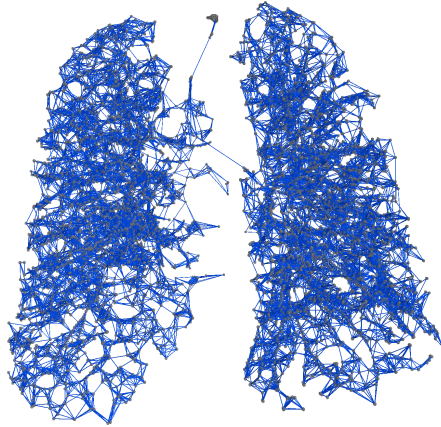
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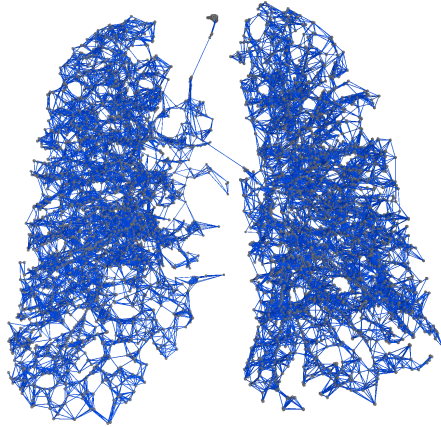
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- Instances of \mathbf{S} are $N \times N$ adjacency matrices



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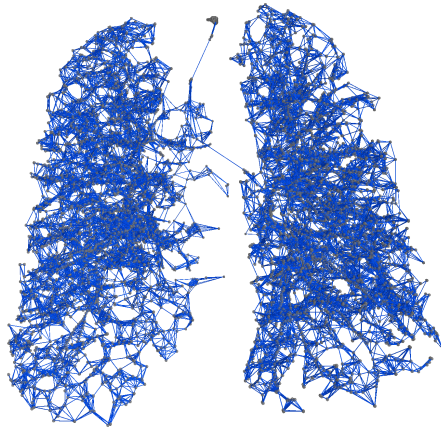


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$$\begin{aligned} \ln p(\mathbf{S} | \mathbf{X}, \mathbf{A}_{in}) &\propto \ln p(\mathbf{S}, \mathbf{X}, \mathbf{A}_{in}) \\ &= \sum_{i \in \mathcal{N}} \phi_i(\mathbf{s}_i) + \sum_{(i,j) \in \mathcal{E}} \phi_{ij}(\mathbf{s}_i, \mathbf{s}_j) - \ln Z, \end{aligned}$$



Node and Pairwise Potentials for MFA

Node Potential: For each node $i \in \mathcal{V}$

$$\phi_i(\mathbf{s}_i) = \sum_{v=0}^D \beta_v \mathbb{I} \left[\sum_j s_{ij} = v \right] + \mathbf{a}^T \mathbf{x}_i \sum_j s_{ij}, \quad (5)$$

Pairwise Potential: For each edge, $(i, j) \in \mathcal{E}_{in}$

$$\phi_{ij}(\mathbf{s}_i, \mathbf{s}_j) = \lambda(1 - 2|s_{ij} - s_{ji}|) + (2s_{ij}s_{ji} - 1) \left[\eta^T |\mathbf{x}_i - \mathbf{x}_j| + \nu^T (\mathbf{x}_i \mathbf{x}_j) \right]. \quad (6)$$

Parameters = $[\cdot]$



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Parameters = $[\boldsymbol{\beta}, \mathbf{a}, \lambda]$



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Parameters = $[\boldsymbol{\beta}, \mathbf{a}, \lambda, \boldsymbol{\eta}, \boldsymbol{\nu}]$



Approximate posterior density with a simpler one



Approximate posterior density with a simpler one

Mean-Field Factorisation: $q(\mathbf{S}) \in \mathcal{Q}$

$$q(\mathbf{S}) = \prod_{i=1}^N \prod_{j=1}^N q_{ij}(s_{ij}), \quad (1)$$

Implication: Node connectivities are independent.



Approximate posterior density with a simpler one

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Variational Inference to approximate $p(\mathbf{S}|\mathbf{X}, \mathbf{A}_{in})$

$$p(\mathbf{S}|\mathbf{X}, \mathbf{A}_{in}) \approx q(\mathbf{S}) \quad (2)$$



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$$p(\mathbf{S}|\mathbf{X}, \mathbf{A}_{in}) \approx q(\mathbf{S}) \quad (2)$$

Minimize KL Divergence \equiv Maximize Evidence Lower Bound (ELBO)

$$\text{ELBO}(q) = -\text{KLD}(q(\mathbf{S})||p(\mathbf{S}|\mathbf{X}, \mathbf{A}_{in})) + \ln Z \quad (3)$$



Maximising ELBO wrt $q_{ij}(s_{ij})$ yields MFA Iterations

MFA Iterations

$$\begin{aligned}\alpha_{kl}^{(t+1)} &= q_{kl}^{(t+1)}(s_{kl} == 1) \\ &= \frac{1}{1 + \exp^{-\gamma_{kl}}}\end{aligned}$$

$$\forall k = \{1 \dots N\}, l \in \mathcal{N}_k$$

α : Global connectivity prediction



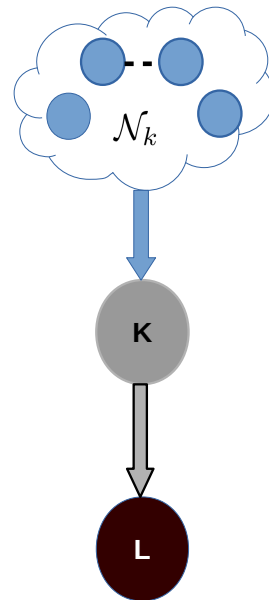
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$$\begin{aligned}\gamma_{kl} &= \prod_{j \in \mathcal{N}_k \setminus l} (1 - \alpha_{kj}^{(t)}) \left\{ \sum_{m \in \mathcal{N}_k \setminus l} \frac{\alpha_{km}^{(t)}}{(1 - \alpha_{km}^{(t)})} [(\beta_2 - \beta_1) \right. \\ &\quad \left. - \beta_2 \sum_{n \in \mathcal{N}_k \setminus l, m} \frac{\alpha_{kn}^{(t)}}{(1 - \alpha_{kn}^{(t)})}] + (\beta_1 - \beta_0) \right\} + \mathbf{a}^T \mathbf{x}_i \\ &\quad + (4\alpha_{lk}^{(t)} - 2)\lambda + 2\alpha_{lk}^{(t)} (\boldsymbol{\eta}^T |\mathbf{x}_i - \mathbf{x}_j| + \boldsymbol{\nu}^T (\mathbf{x}_i \mathbf{x}_j))\end{aligned}$$

$$\alpha_{kl}^{(t+1)} = \sigma(\gamma_{kl}) = \frac{1}{1 + \exp^{-\gamma_{kl}}}$$

Note: MFA iterations resemble feed-forward operations in neural nets



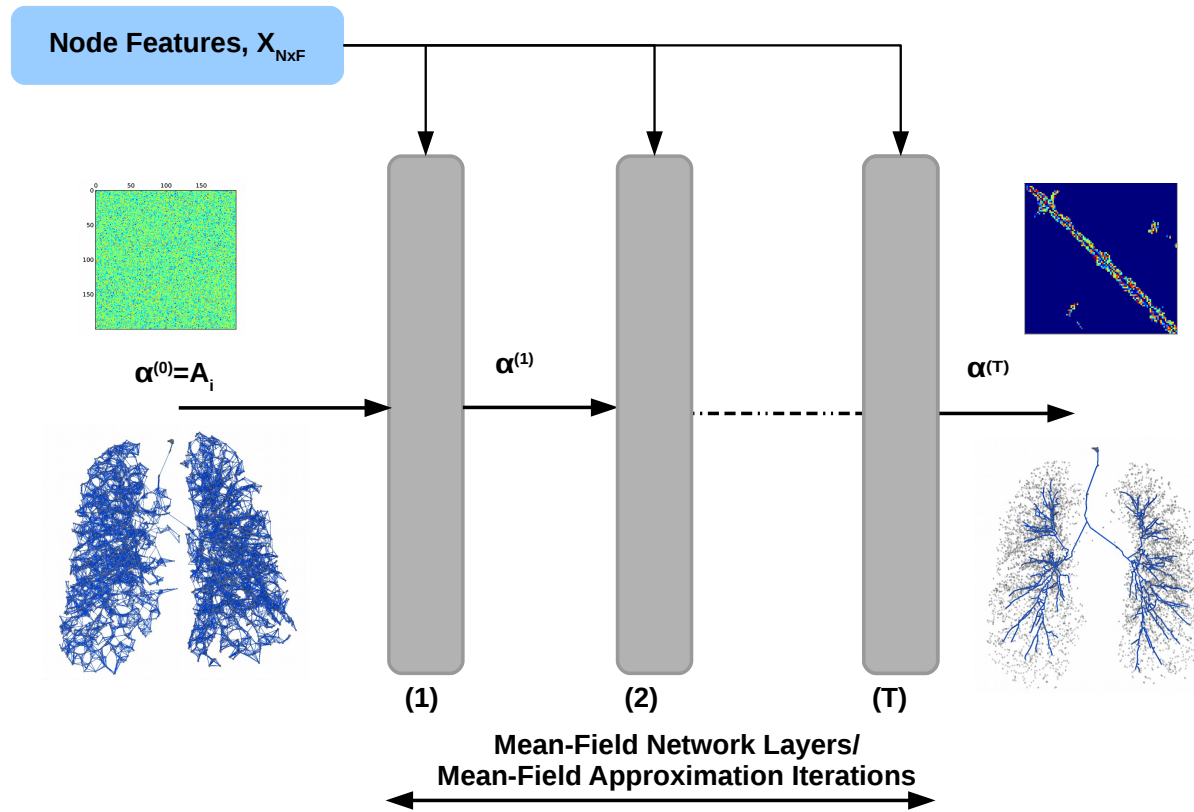
MFA as Mean-Field Networks

- T -iterations as a T -layered network

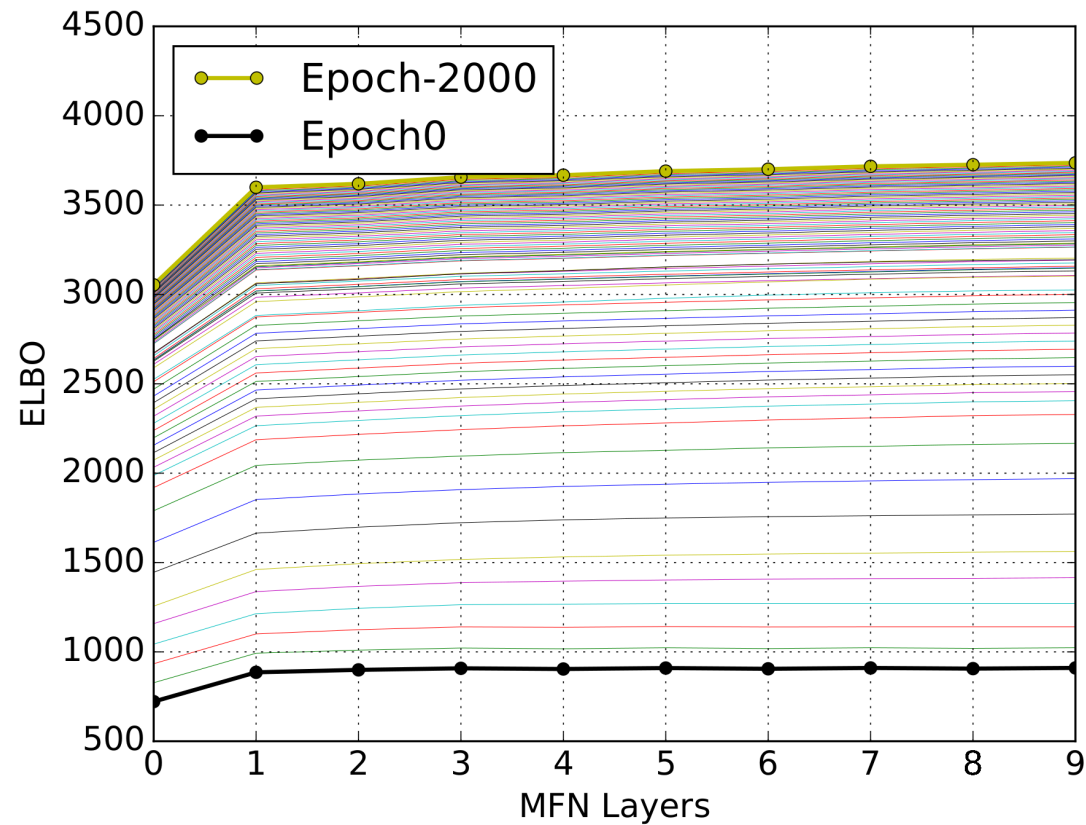


MFA as Mean-Field Networks

- T –iterations as a T –layered network
- Gradient descent to learn model parameters: $\mathcal{L}(\boldsymbol{\alpha}, \mathbf{A}_r)$



Increasing ELBO \implies Better approximation



Experiments

- **Baseline:** a) Region growing on probability images b) Bayesian smoothing merged with region growing for evaluation
- Pretraining dataset used to tune hyperparameters
- Eight-fold cross validation



Experiments

- **Baseline:** a) Region growing on probability images b) Bayesian smoothing merged with region growing for evaluation
- Pretraining dataset used to tune hyperparameters
- Eight-fold cross validation
- **Error measures:**
 - Average centerline distance: $d_{err} = (d_{FP} + d_{FN})/2$
 - $d_{FP} \equiv$ Specificity
 - $d_{FN} \equiv$ Sensitivity
 - Percentage of tree length (TL)
 - False positive rate (FPR)



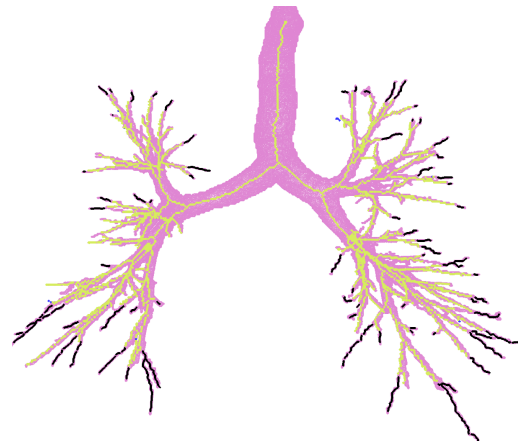
Performance comparison

	d_{FP} (mm)	d_{FN} (mm)	d_{err} (mm)	TL(%)	FPR(%)
Vox+RG	3.624 ± 0.776	5.155 ± 0.580	4.389 ± 0.441	79.6 ± 7.2	5.0 ± 3.9
BS+RG	3.921 ± 0.612	4.218 ± 0.334	4.069 ± 0.476	82.3 ± 6.1	8.7 ± 3.4
MFN	3.599 ± 0.583	3.491 ± 0.295	3.595 ± 0.321	83.1 ± 6.7	8.6 ± 5.3

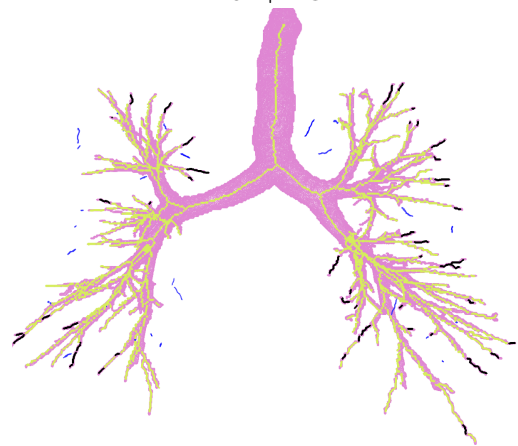
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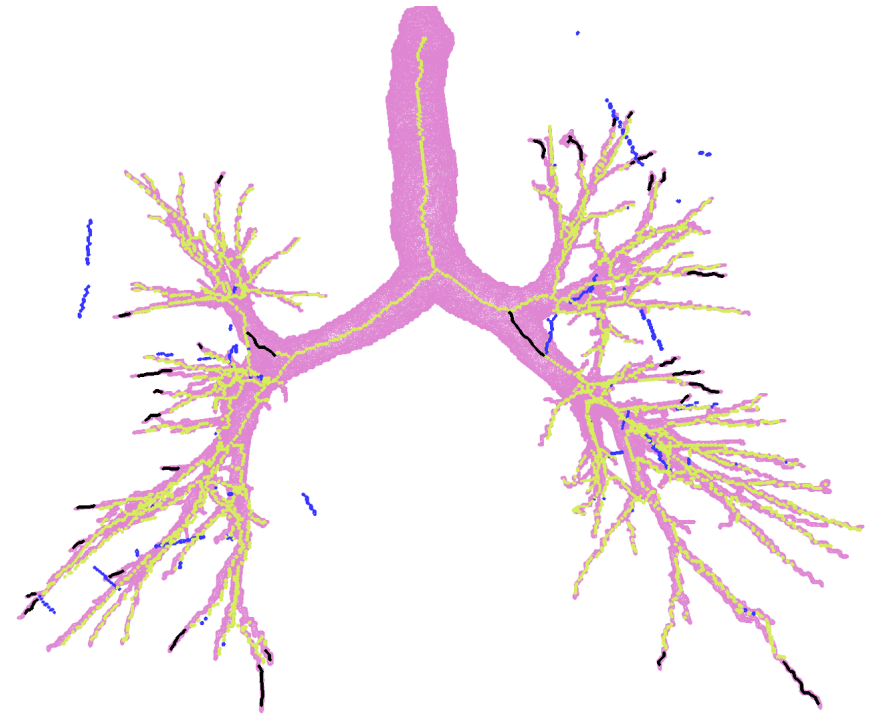
Visualisation of extracted airways



Vox+RG



BS+RG



MFN

Legend: Reference (pink), True Positive (Yellow), False Negative (Black), False Positive (Blue)



Summary

- Airway extraction as graph refinement
- Novel use of Mean-Field Approximation
- Proposed expressive node and pairwise potentials
- Mean-Field Network interpretation
- Few parameters (46 scalar weights)
- Easy to optimise using gradient descent
 - Might not generalise across applications
 - Hand-crafting potentials is cumbersome



Graph Neural Networks

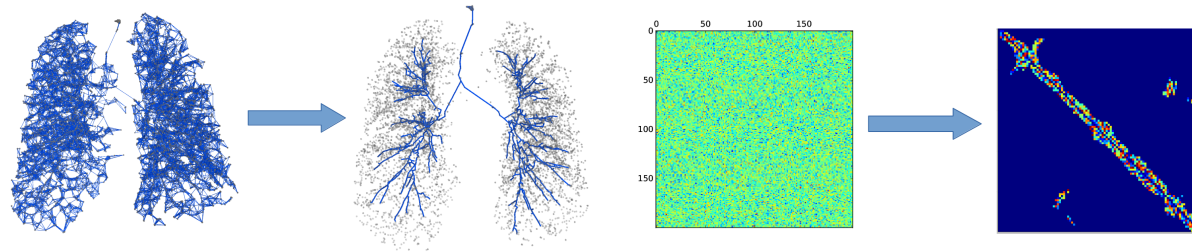


Graph Neural Networks

- Neural nets with graph input
- Step towards non-Euclidean (geometric) Deep Learning
- Generalisation of message passing algorithms
- Complex task-specific messages can be learnt
- End-to-end trainable inference systems



GNN based Graph Refinement



- Graph refinement task: $f(\cdot) : \mathcal{G}_{in} \mapsto \mathcal{G}$
- GNN based encoder-decoder pair
- Encoder comprises stacks of GNNs; Message passing between nodes
- Joint training of encoder-decoder pair to learn useful embeddings
- Simple decoder predicts graph connectivity



GNN Model for Graph Refinement



GNN Model for Graph Refinement

Consider node j with neighbours \mathcal{N}_j ,

$$\text{Node Embedding: } \mathbf{h}_j^1 = g_n(\mathbf{x}_j) \quad (8)$$

$$\text{N2E mapping: } \mathbf{h}_{(i,j)}^1 = g_{n2e}([\mathbf{h}_i^1, \mathbf{h}_j^1]) \quad (9)$$

$$\text{E2E mapping: } \mathbf{h}_j^2 = g_{e2n}(\sum_{i \in \mathcal{N}_j} \mathbf{h}_{(i,j)}^1) \quad \forall i \in \mathcal{N}_j \quad (10)$$

$$\text{N2E mapping: } \mathbf{h}_{(i,j)}^2 = g_{n2e}([\mathbf{h}_i^2, \mathbf{h}_j^2]) \quad (11)$$

$$\text{Decoder: } \alpha_{ij} = \sigma(g_{dec}(\mathbf{h}_{(i,j)}^2)) \quad (12)$$

$g_{\dots}(\cdot)$ are MLPs, g_{dec} is MLP with 1 output channel



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$$\text{N2E mapping: } \mathbf{h}_{(i,j)}^1 = g_{n2e}([\mathbf{h}_i^1, \mathbf{h}_j^1]) \quad (9)$$

$$\text{E2N mapping: } \mathbf{h}_j^2 = g_{e2n}(\sum_{i \in \mathcal{N}_j} \mathbf{h}_{(i,j)}^1) \quad \forall i \in \mathcal{N}_j \quad (10)$$

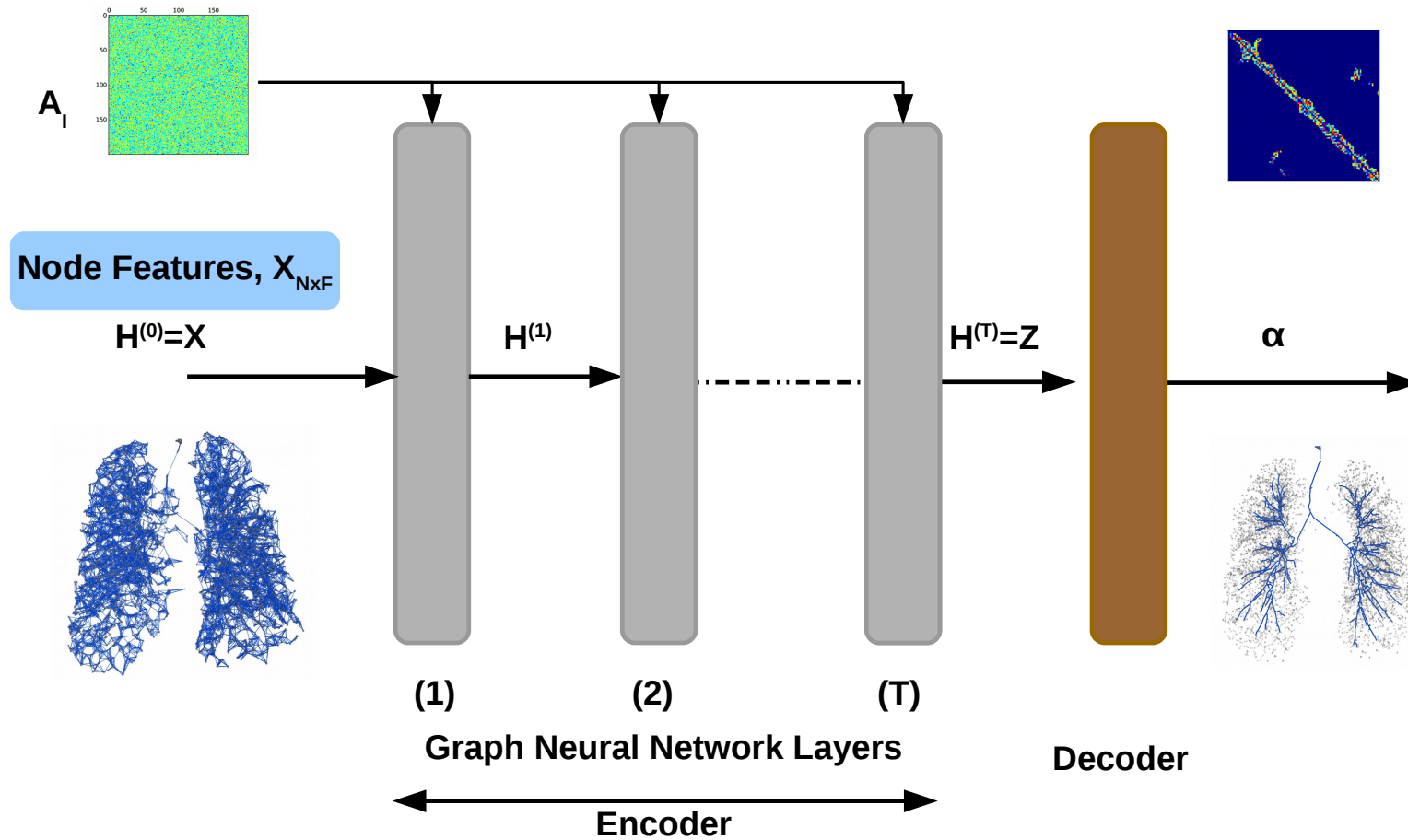
$$\text{N2E mapping: } \mathbf{h}_{(i,j)}^2 = g_{n2e}([\mathbf{h}_i^2, \mathbf{h}_j^2]) \quad (11)$$

$$\text{Decoder: } \alpha_{ij} = \sigma(g_{dec}(\mathbf{h}_{(i,j)}^2)) \quad (12)$$

$g_{\dots}(\cdot)$ are MLPs, g_{dec} is MLP with 1 output channel



Summarising GNN Model



Experiments

- Same set-up as with MFNs
- Pretraining dataset used to tune hyperparameters
- Eight fold cross validation



Performance comparison

Table 1

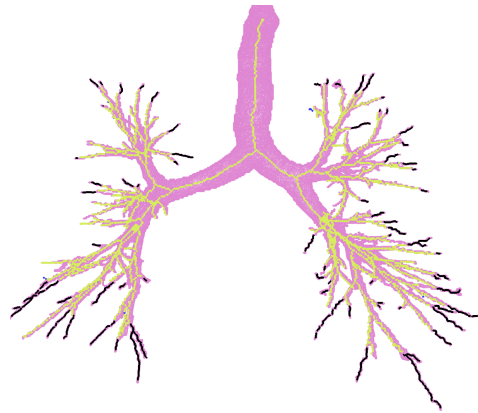
Performance comparison of five methods: Region growing on probability images (Vox+RG), Bayesian smoothing merged with Vox+RG (BS+RG), UNet, MFN and GNN models. Dice similarity, centerline distances (d_{FP} , d_{FN} , d_{err}), fraction of tree length detected (TL) and false positive rate (FPR) are reported based on 8-fold cross validation. Significant improvements when compared to other methods are shown in boldface. Additionally, we also report the running time to train each of the models in a single fold. Note that the MFN and GNN models require additional preprocessing that is performed only once when preparing the graphs..

	Dice(%)	d_{FP} (mm)	d_{FN} (mm)	d_{err} (mm)	TL(%)	FPR(%)	Time (m)
Vox+RG	–	2.937 ± 1.005	6.762 ± 2.1042	4.847 ± 2.527	73.2 ± 9.9	4.9 ± 3.9	90
BS+RG	–	2.827 ± 1.266	4.601 ± 2.002	3.714 ± 1.896	73.6 ± 6.1	7.9 ± 6.1	105
UNet	–	3.540 ± 1.316	3.525 ± 1.201	3.532 ± 1.259	75.6 ± 8.7	6.5 ± 3.3	5700
MFN	86.5 ± 2.5	3.608 ± 1.360	3.116 ± 0.632	3.362 ± 1.297	74.5 ± 6.7	8.6 ± 5.4	60 + 35
GNN	84.8 ± 3.3	2.216 ± 0.464	2.878 ± 0.505	2.547 ± 0.587	81.9 ± 7.3	7.8 ± 4.6	60 + 12

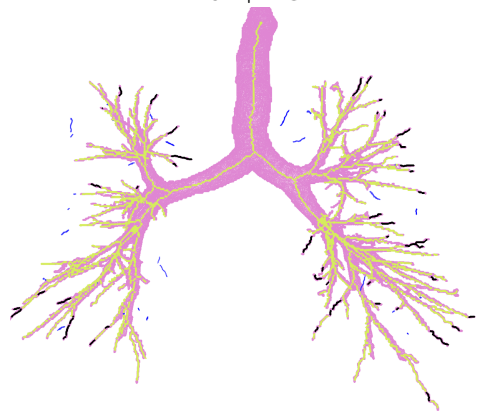
- $d_{FP} \equiv$ Specificity
- $d_{FN} \equiv$ Sensitivity
- Average centerline distance: d_{err}
- Percentage of tree length (TL)
- False positive rate (FPR)



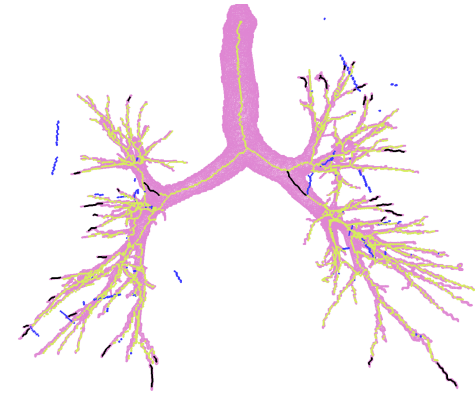
Visualisation of extracted airways



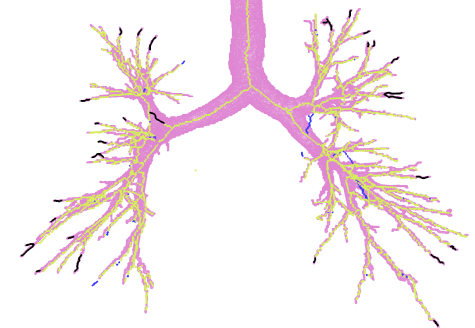
Vox+RG



BS+RG



MFN



GNN

Legend: Reference (pink), True Positive (Yellow), False Negative (Black), False Positive (Blue)



Summary

- GNN based supervised graph refinement
- Unique, inductive graph application of GNNs
- Edge embeddings used for prediction
- Competitive results with limited data
- Generalisations of MFNs
 - Disconnected trees
 - Relies on quality labelled training data



Outline

- ① Objective of the study
- ② Data
- ③ Graph Refinement Models
- ④ Summary & Conclusions
- ⑤ Supplementary material



Conclusions from the study

- Exploratory methods can extract more branches
- Graph based representations are less computationally intensive
- Using global information in local decisions is helpful
- Incorporating prior knowledge is valuable
- MFNs as structured neural networks
- GNNs as generalisations of message passing algorithms
- Bias-variance trade-off between MFNs and GNNs



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Questions?

Carbontracker: Tracking and Predicting the Carbon Footprint of Training Deep Learning Models

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