



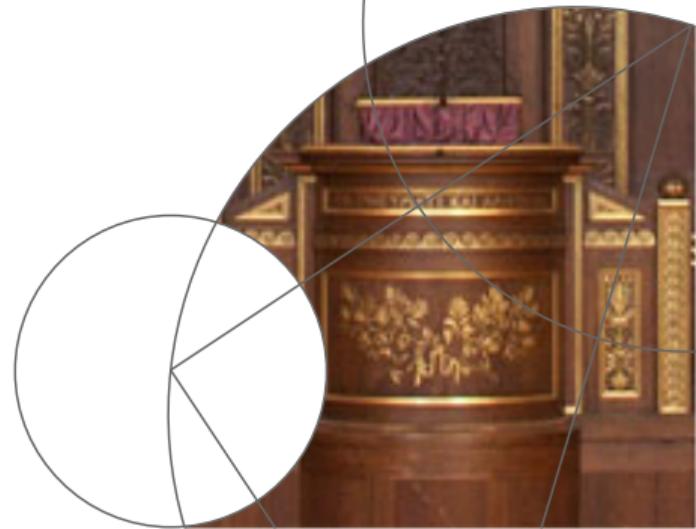
UNIVERSITY OF COPENHAGEN

Graph Refinement based Airway Extraction Using Mean-field and Graph Neural Networks

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Outline

① Airway Diseases

② Understanding the Data

③ Methods

Existing Methods

Graph Refinement Model

Mean-Field Networks

Graph Neural Networks

④ Experiments



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COPD: Leading Factor of Morbidity & Mortality

- Tobacco Smoking
- Indoor & Outdoor Air pollution



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- 174.5M affected; 3.2M deaths (2015)



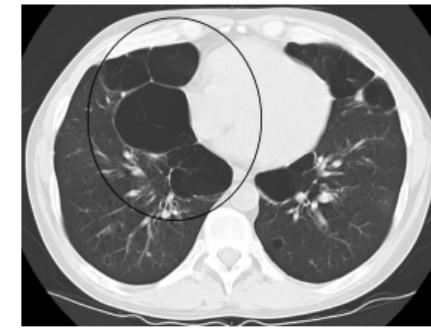
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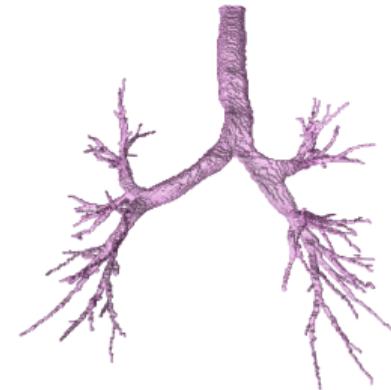
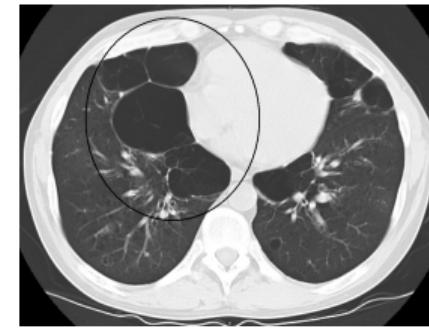
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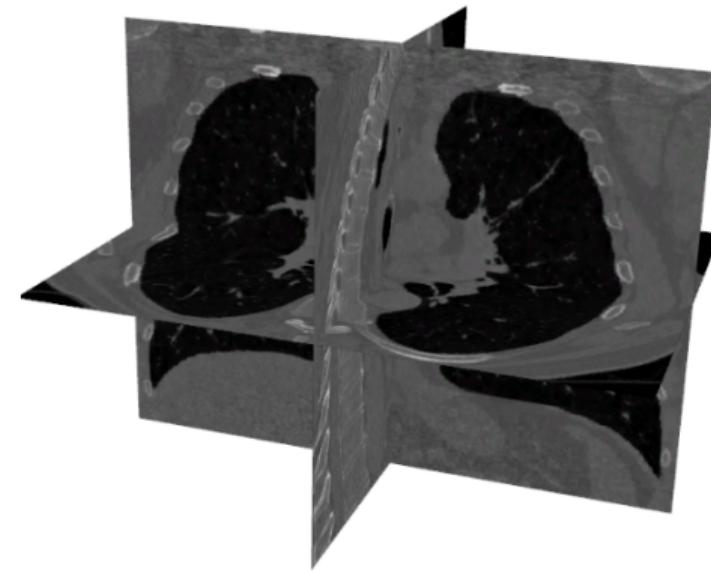
Existing Diagnostics are Rudimentary & Tedious

- Lung Function Tests
 - + Simple and inexpensive
 - Patient dependent
 - Low reproducibility
 - Mild cases go unnoticed



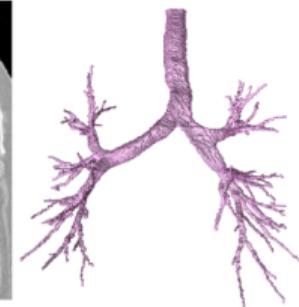
Existing Diagnostics are Rudimentary & Tedious

- Lung Function Tests
 - + Simple and inexpensive
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- 3D CT Scans
 - + Provide more information
 - Arduous to read the data; even for experts
 - Low inter-observer agreement



Objective

Automatic Airway Segmentation and obtain **useful** COPD biomarkers



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Primary Data from Danish Lung Cancer Study

- Danish Lung Cancer Screening Trial ¹
- Low-dose CT
- > 10,000 scans
- Age 50-70 years.
- Smoker or former smoker (> 20 pack years)
- 32/10,000 have segmentations verified by an expert user!

¹Pedersen, J. H., et.al : The Danish randomized lung cancer CT screening trialoverall design and results of the prevalence round. Journal of Thoracic Oncology, (2009)



CT Images are noisy, low contrast & low-res.



- Volume resolution $\sim 300 \times 250 \times 275$
- Voxels $\sim 0.75\text{mm} \times 0.75\text{mm} \times 1\text{mm}$
- Challenges
 - Acquisition noise
 - Inter-patient variability
 - Several “interfering” structures
 - Labels/Annotations



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Most Existing Methods handle occlusions poorly

- State-of-the-art: Region-growing (!) based methods
- EXACT Study² compares 15 methods; No clear winner
- Small airways are challenging
- Challenging to overcome occlusions

²Lo, P., et.al : Extraction of airways from CT (EXACT'09). IEEE Transactions on Medical Imaging, (2012)



Graph Refinement Model for Airway Extraction



Graph Refinement Model for Airway Extraction

Desired Properties

- Exploratory (to overcome occlusions)
- Detect small airways
- Uncertainty estimates



Preprocess Image to Graph Model

One possibility



Preprocess Image to Graph Model

One possibility



Figure 1: Visualisation of the pre-processing carried out to transform the input image (left) into a probability image (center) and then into graph format (right). Nodes in the graph are shown in scale to capture the variations in their local radius.



Graph Refinement Model

- Input graph: $\mathcal{G}_i : \{\mathcal{N}, \mathcal{E}_i\}$
- Node features: $\mathbf{X} \in \mathbb{R}^{F \times N}$
- Input adjacency: $\mathbf{A}_i \in \{0, 1\}^{N \times N}$



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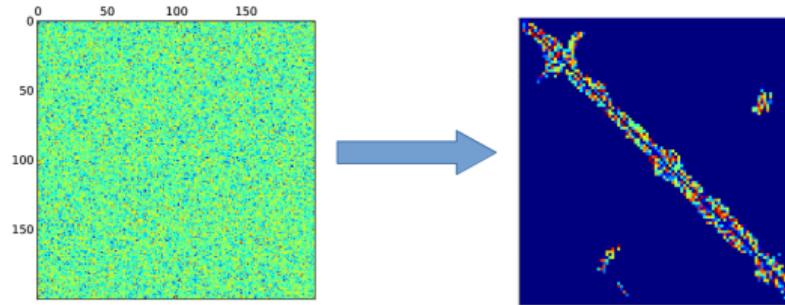
Graph Refinement Task

$$f(\mathcal{G}_i) \rightarrow \mathcal{G}$$

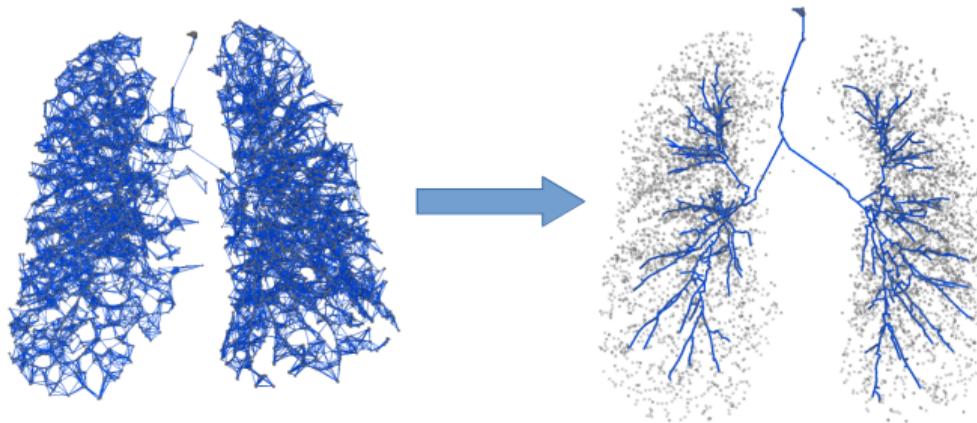
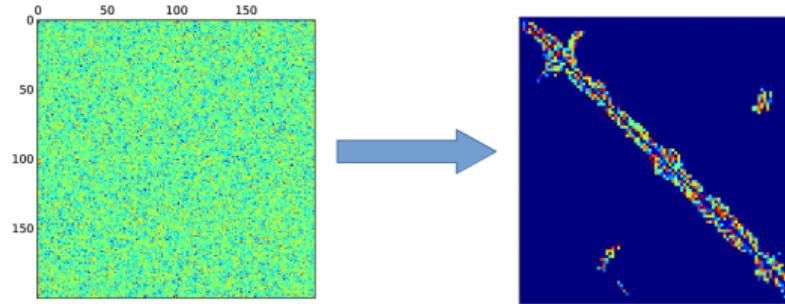
Output subgraph \mathcal{G} with $\mathcal{E} \subset \mathcal{E}_i$; $\mathbf{A} \in \{0, 1\}^{N \times N}$



Visualise Graph Refinement Task



Visualise Graph Refinement Task



Updating Probabilistic Graphical Model

- Binary random variable to capture existence of edge between nodes
- $s_{ij} \in \{0, 1\}$ with prob. $\alpha_{ij} \in [0, 1]$



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



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Variational Approximate Inference

Minimize ELBO to obtain $q(\mathbf{S}) \in \mathcal{Q}$

$$\mathcal{F}(q_{\mathbf{S}}) = \ln Z + \mathbb{E}_{q_{\mathbf{S}}} \left[\ln p(\mathbf{S}|\mathbf{X}) - \ln q(\mathbf{S}) \right]$$



Simpler approximation using MFA



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Factorisable $q(\mathbf{S})$

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

$$\text{where, } q_{ij}(s_{ij}) = \begin{cases} \alpha_{ij} & \text{if } s_{ij} = 1 \\ (1 - \alpha_{ij}) & \text{if } s_{ij} = 0 \end{cases},$$



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Assumes edges to be independent.



Node and Pairwise Potentials for MFA



Node and Pairwise Potentials for MFA

Node Potential

For each node $i \in \mathcal{N}$,

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



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Pairwise Potential

For each edge, $(i, j) \in \mathcal{E}$,

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

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



Minimize ELBO to get MFA Iterations

MFA Iterations

$$\alpha_{kl}^{(t+1)} = \sigma(\gamma_{kl}) = \frac{1}{1 + \exp^{-\gamma_{kl}}} \quad \forall k = \{1 \dots N\}, l \in \mathcal{N}_k,$$

where $\sigma(\cdot)$ is sigmoid activation, \mathcal{N}_k are neighbours of node k , and



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



MFA to Mean-Field Networks



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- MFA Iterations resemble feed-forward operations in neural network

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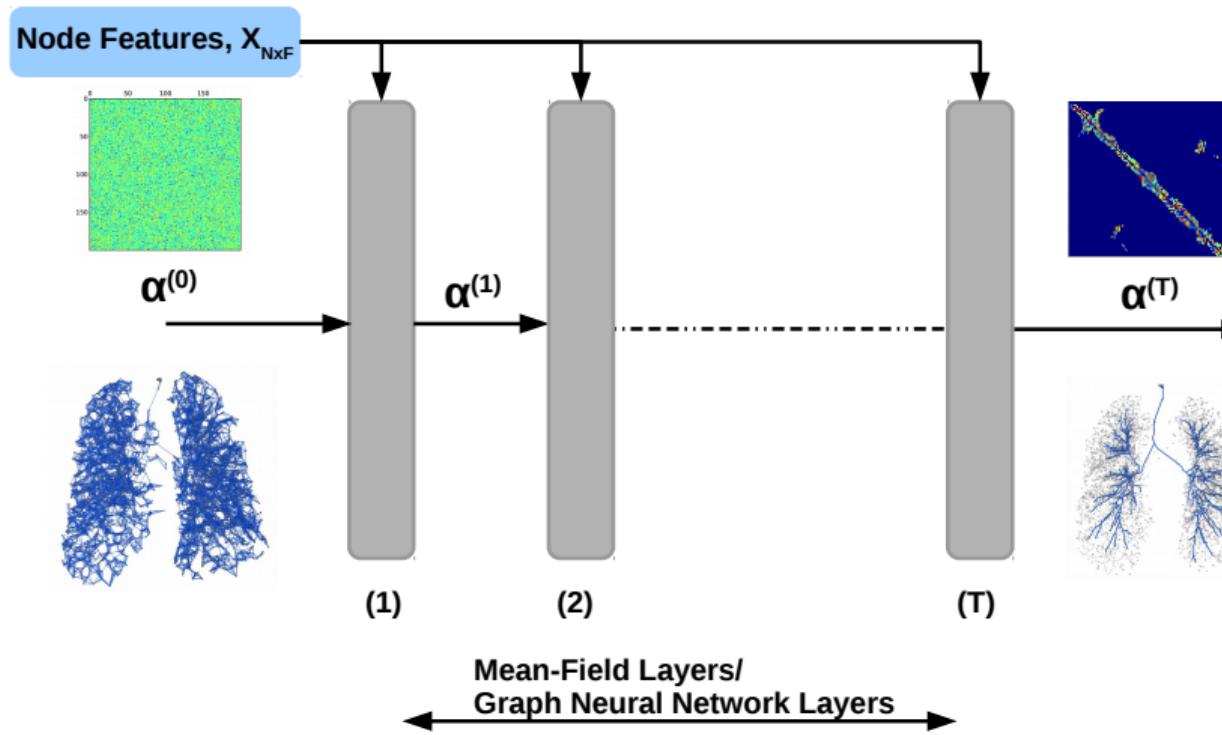
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- T -iterations as a T -layered network
- Automatic differentiation to learn parameters: $\mathcal{L}(\alpha, \mathbf{A}_r)$, where \mathbf{A}_r is reference adjacency.



Summarising MFN



Summarising MFN

- Yields approximation to underlying posterior
- Simple factors
- Handful of parameters
- Easy to optimise
 - Hand-crafting potentials is cumbersome
 - Might not generalise across applications



Graph Neural Networks



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- Neural networks operating directly on graph structured data



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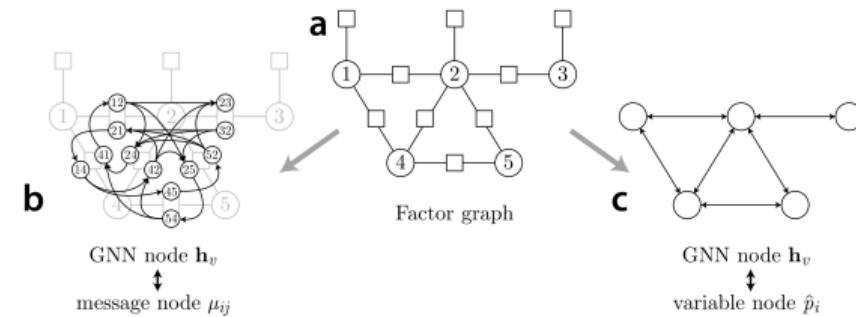


Figure: Two mappings of Factor Graphs into GNNs

Figure from Yoon et al. "Inference in probabilistic graphical models by Graph Neural Networks" (2018)



Graph Auto Encoder (GAE) for Graph Refinement



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- Encoder comprises of GNNs; Message passing between nodes



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GAE Model for Graph Refinement: Encoder



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Encoder:

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

$$\text{Node-to-Edge mapping: } \mathbf{h}_{(i,j)}^1 = g_{n2e}([\mathbf{h}_i^1, \mathbf{h}_j^1])$$

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$g_{...}(\cdot)$ are 2-layered MLPs, with ReLU, dropout and layer normalisation



GAE Model for Graph Refinement: Decoder

Decoder:

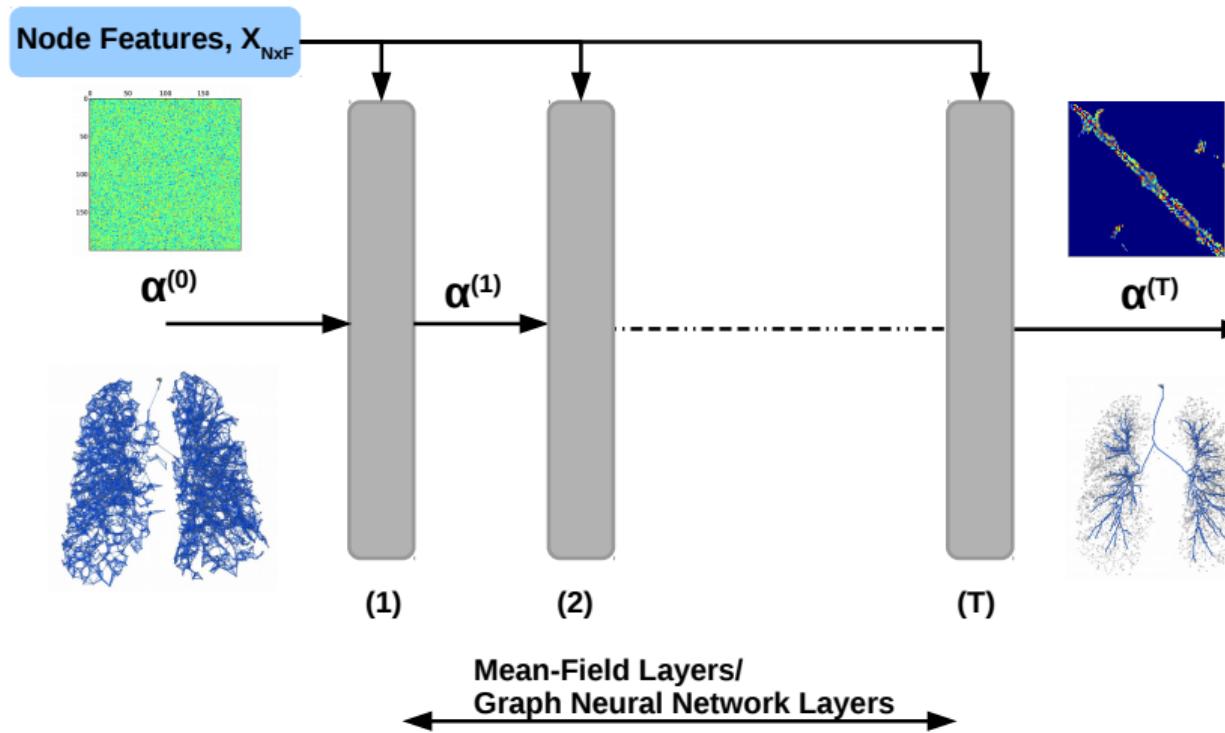
$$\alpha_{ij} = \sigma(g_{dec}(\mathbf{h}_{(i,j)}^2)) \quad (2)$$

g_{dec} is a 1D convolutional layer with one output channel

Model can be trained by computing the loss $\mathcal{L}\alpha, \mathbf{A}_r$



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Training both models: Dice Loss

To tackle Class Imbalance:



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To tackle Class Imbalance:

$$\mathcal{L}(\boldsymbol{\alpha}, \mathbf{A}_r) = 1 - \frac{2 \sum_{i,j=1}^N \alpha_{ij} A_{ij}}{\sum_{i,j=1}^N \alpha_{ij}^2 + \sum_{i,j=1}^N A_{ij}^2},$$

A_{ij} are individual binary entries \mathbf{A}_r



Data

- Danish Lung Cancer Screening Trial
- Low-dose Chest CT scans
- 32 scans with “Reference” annotations
- 100 scans with automatic segmentation



Results

- Error Measure based on centerline distances
- Average of two distances, $d_{err} = (d_{FP} + d_{FN})/2$
- Compared with Top Performer on Airway Extraction Challenge

Table I: Performance comparison

Method	d_{FP} (mm)	d_{FN} (mm)	d_{err} (mm)
Voxel Classifier	3.871	5.108	4.489 ± 0.754
MFN	3.716	3.992	3.845 ± 0.415
GNN	3.513	2.890	3.202 ± 0.386

