

Functional lung radiation therapy with 4DCT-Ventilation: from theory to clinical implementation

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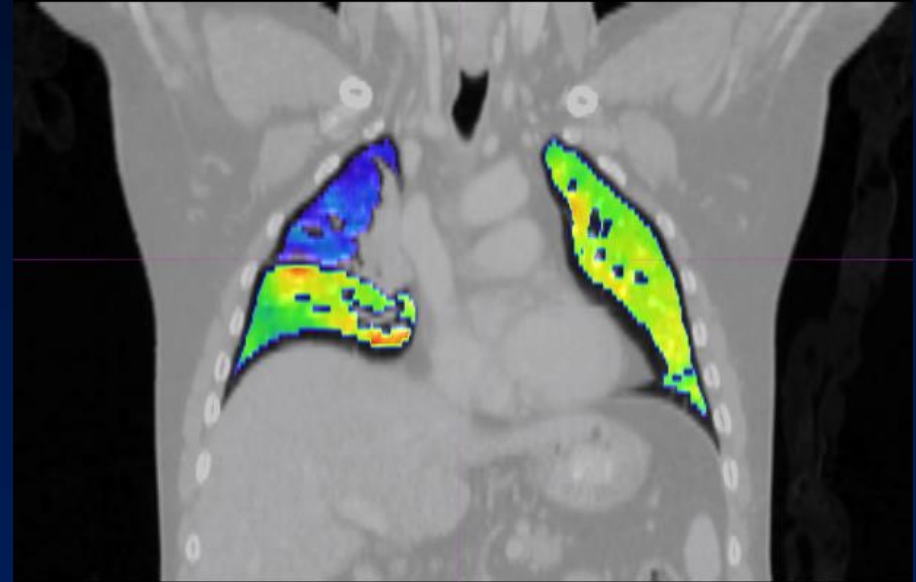
Background

4DCT-Ventilation Imaging

4DCT



4DCT-Ventilation



Background

4DCT-Ventilation Imaging

- Reduced cost
- Reduced dose
- Better spatial resolution
- Anatomical + Functional information
- Better quantification

Outline

4DCT-Ventilation Imaging

- Image formation
- Validation
- Clinical applications
- Clinical trial

Outline

4DCT-Ventilation Imaging

- **Image formation**
- Validation
- Clinical applications
- Clinical trial

Calculating Ventilation Images

Calculating ventilation maps

4DCT – 10 phases



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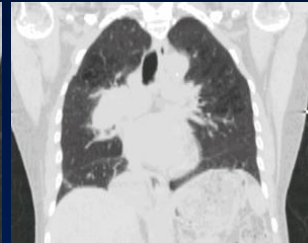
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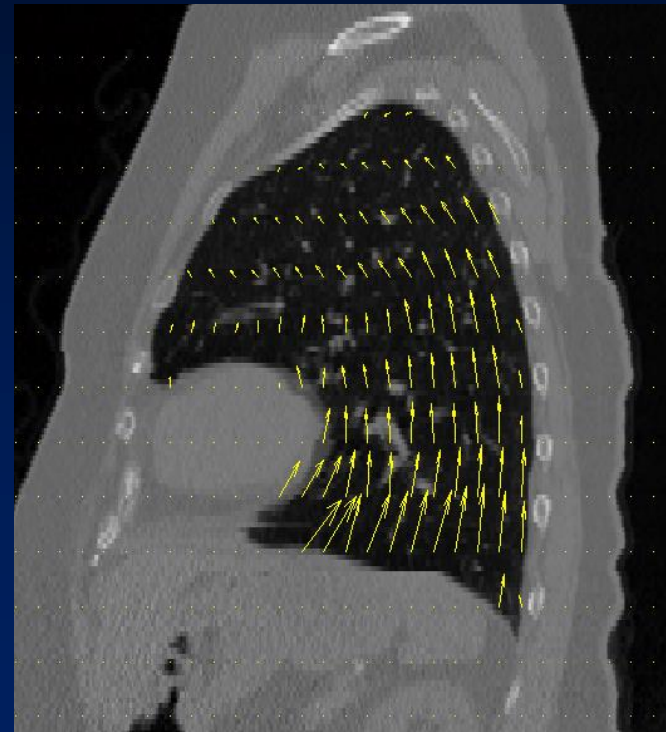


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Calculating Ventilation Images

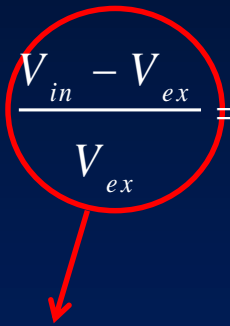
Link lung voxels from inhale phase to exhale phase
using deformable image registration

Deformable registration maps



4D deformable registration using trajectory modeling (Castillo et al., 2010)

Calculating Ventilation Images

$$\frac{V_{in} - V_{ex}}{V_{ex}} = 1000 \frac{\overline{HU}_{in}^{voi} - HU_{ex}}{HU_{ex} (1000 + \overline{HU}_{in}^{voi})}$$


Specific ventilation

Local fractional change in air content

Specific ventilation of 0 = no volume change

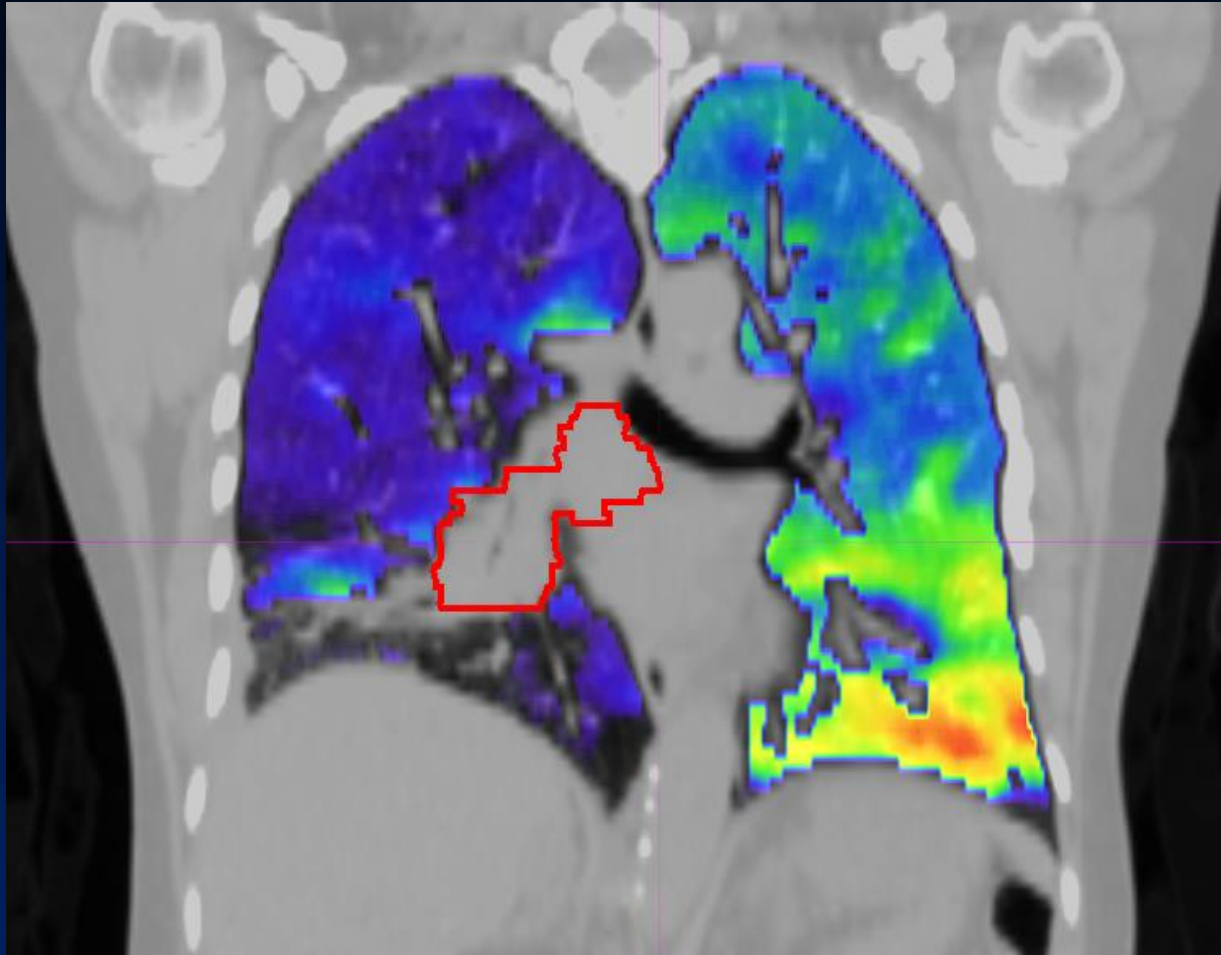
Specific ventilation of 1 = volume of air doubled

Calculating Ventilation Images

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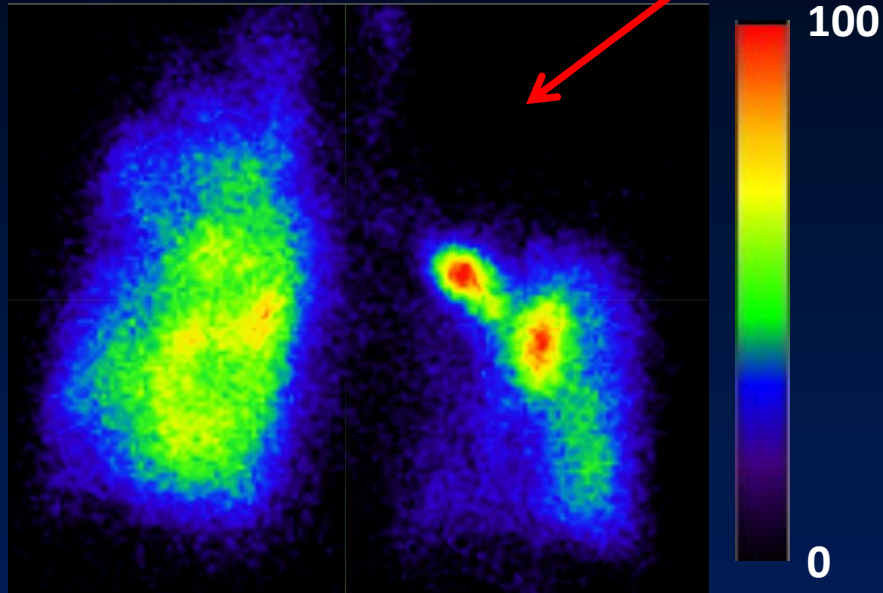
Outline

4DCT-Ventilation Imaging

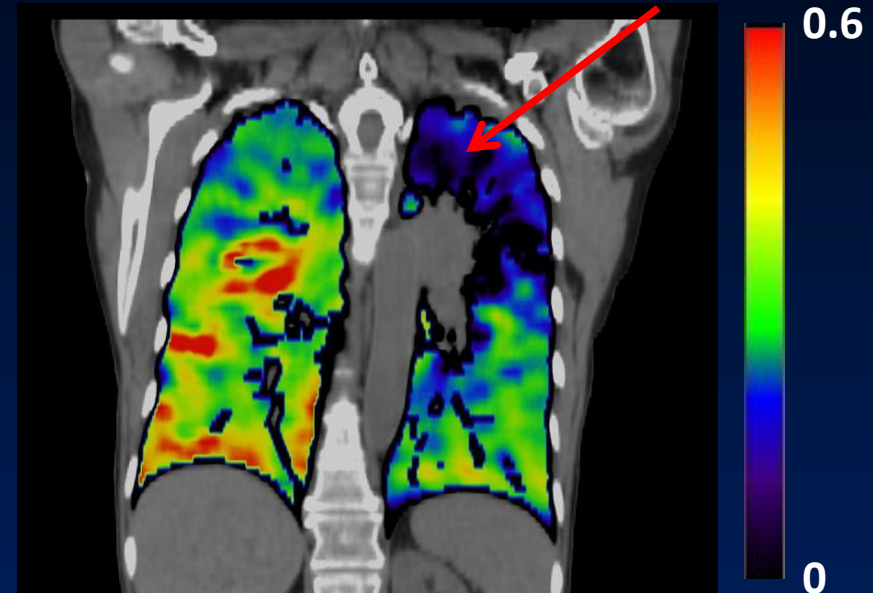
- Image formation
- **Validation**
- Clinical applications
- Clinical trial

Validation against nuclear medicine

VQ Ventilation Scan

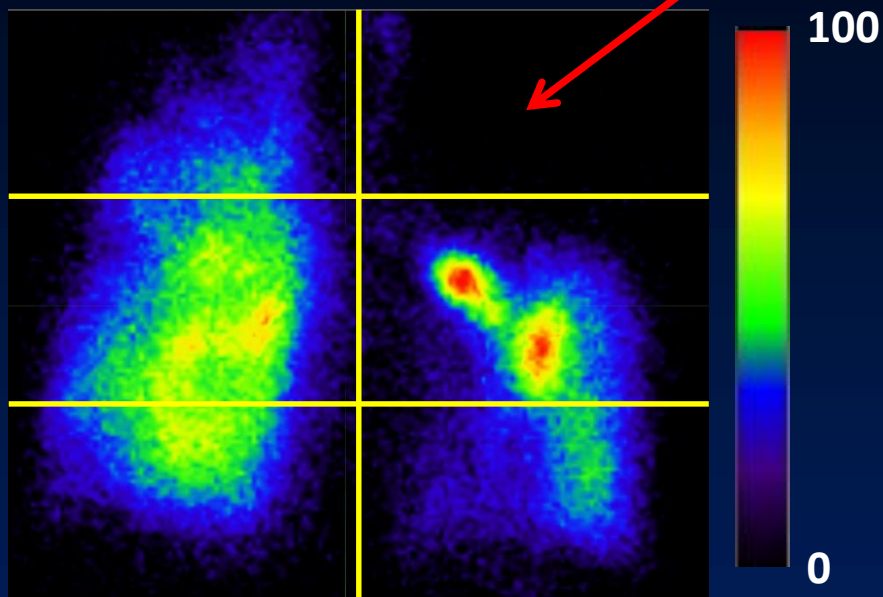


4DCT Ventilation Map

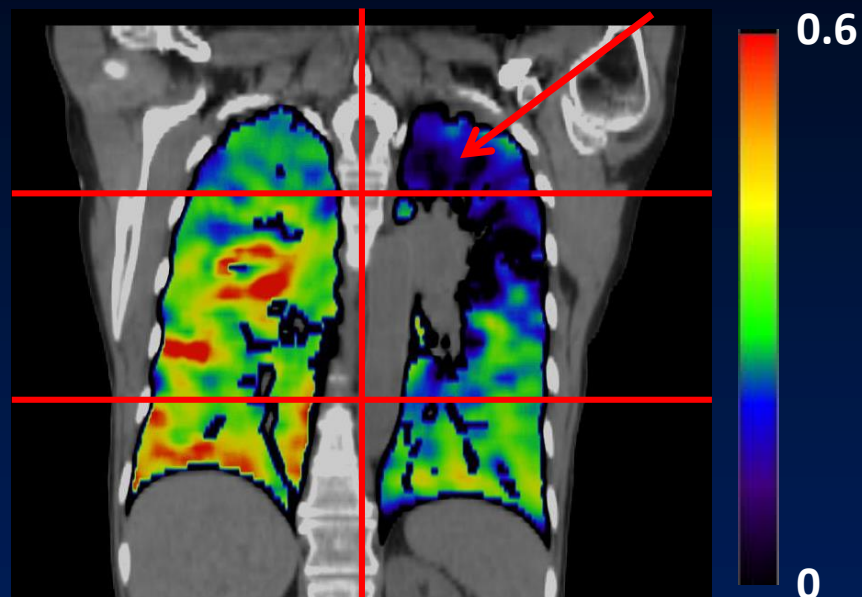


Validation against nuclear medicine

VQ Ventilation Scan



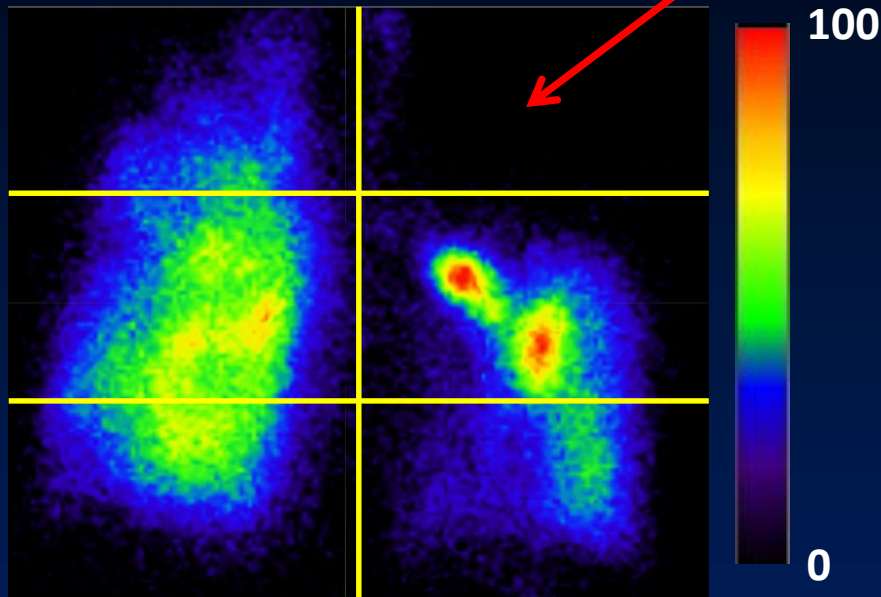
4DCT Ventilation Map



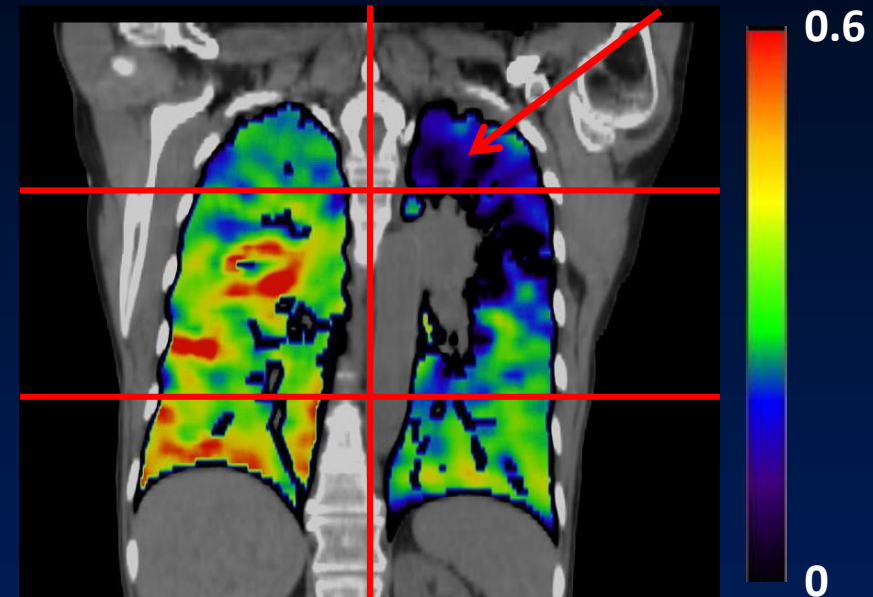
	SPECT Ventilation		4DCT Ventilation	
	Right (%)	Left (%)	Right (%)	Left (%)
Top	9.5	2.7	16.8	8.1
Middle	30.2	21.1	21.9	12.4
Lower	21.4	15.1	23.2	17.7
Total	61.1	38.9	61.8	38.2

Validation against nuclear medicine

VQ Ventilation Scan



4DCT Ventilation Map



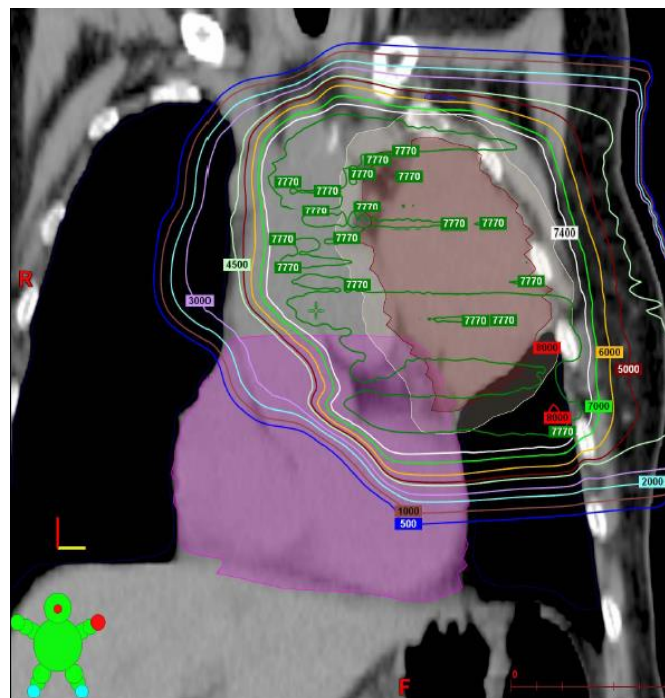
- Correlation coefficient = 0.65
- Radiologist observations: Sensitivity = 90%, Specificity = 64%, Accuracy = 81%

Outline

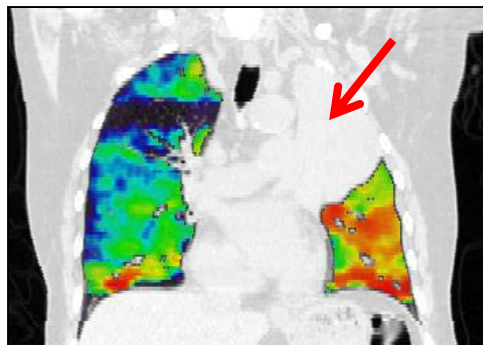
4DCT-Ventilation Imaging

- Image formation
- Validation
- **Clinical applications**
- Clinical trial

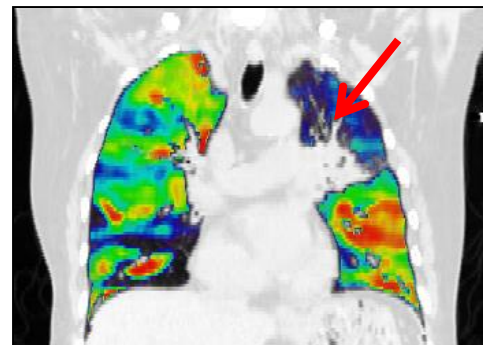
Changes in lung function during RT



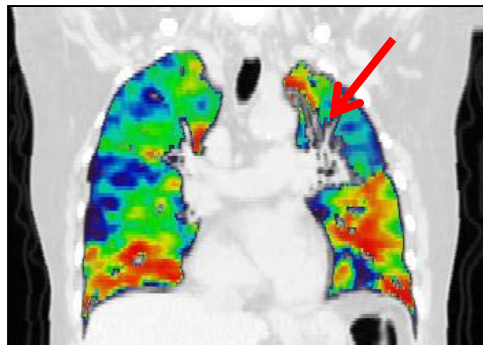
Week 0



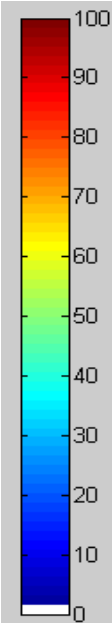
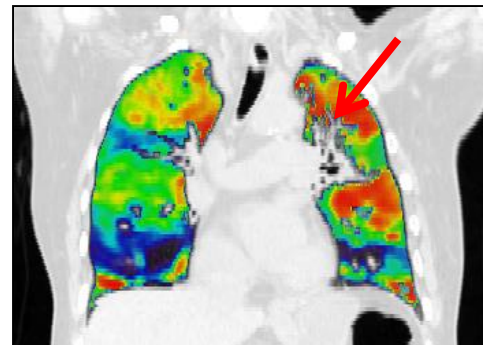
Week 3



Week 5



Week 7



Functional planning concept

- Avoid functional portions of the lung in favor of irradiating through less functioning lung tissue

Incorporation of functional imaging data in the evaluation of dose distributions using the generalized concept of equivalent uniform dose

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Volume 33, Issue 1, 30 August 1995, Pages 65–75



Clinical original contribution

The role of three dimensional functional lung imaging in radiation treatment planning: The functional dose-volume histogram

Lawrence B. Marks, M.D. , David P. Spencer, Ph.D.*, George W. Sherouse, Ph.D.*, Gunilla Bentel, R.N., R.T.T.*, Robert Clough*, Karen Vann, R.N.*, Ronald Jaszczak, Ph.D.†, R.Edward Coleman, M.D.†, Leonard R. Prosnitz, M.D.*



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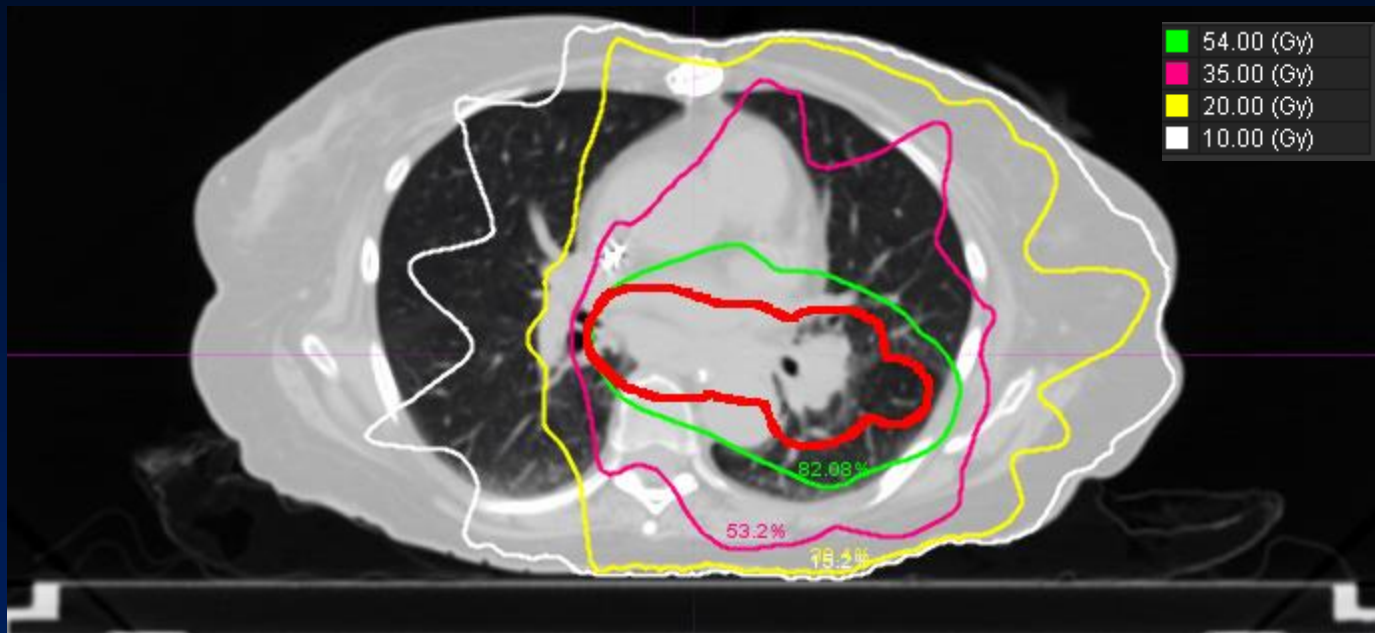
SPECT in treatment planning

The incorporation of SPECT functional lung imaging into inverse radiotherapy planning for non-small cell lung cancer

Judith A. Christian^a, , Mike Partridge^a, Elena Nioutsikou^a, Gary Cook^a, Helen A. McNair^a, Bernadette Cronin^a, Frederic Courbon^b, James L. Bedford^a, Michael Brada^a

Functional planning concept

Clinical plan



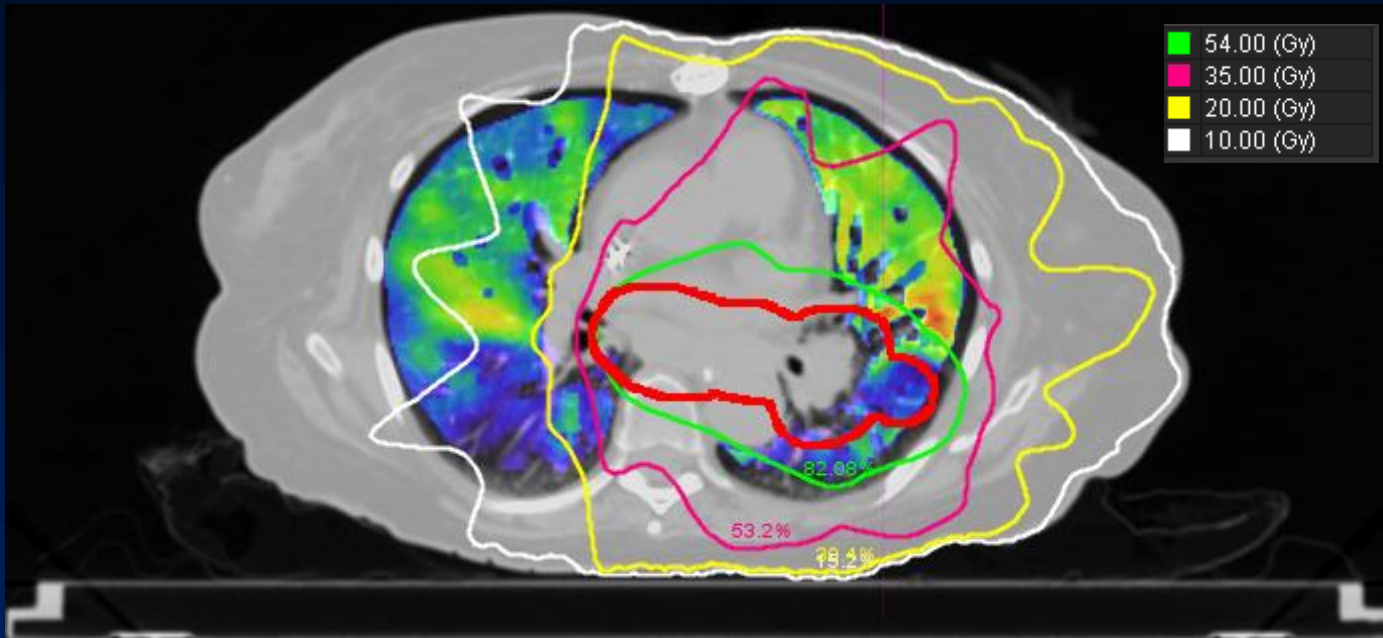
Functional planning concept

Clinical plan with functional imaging

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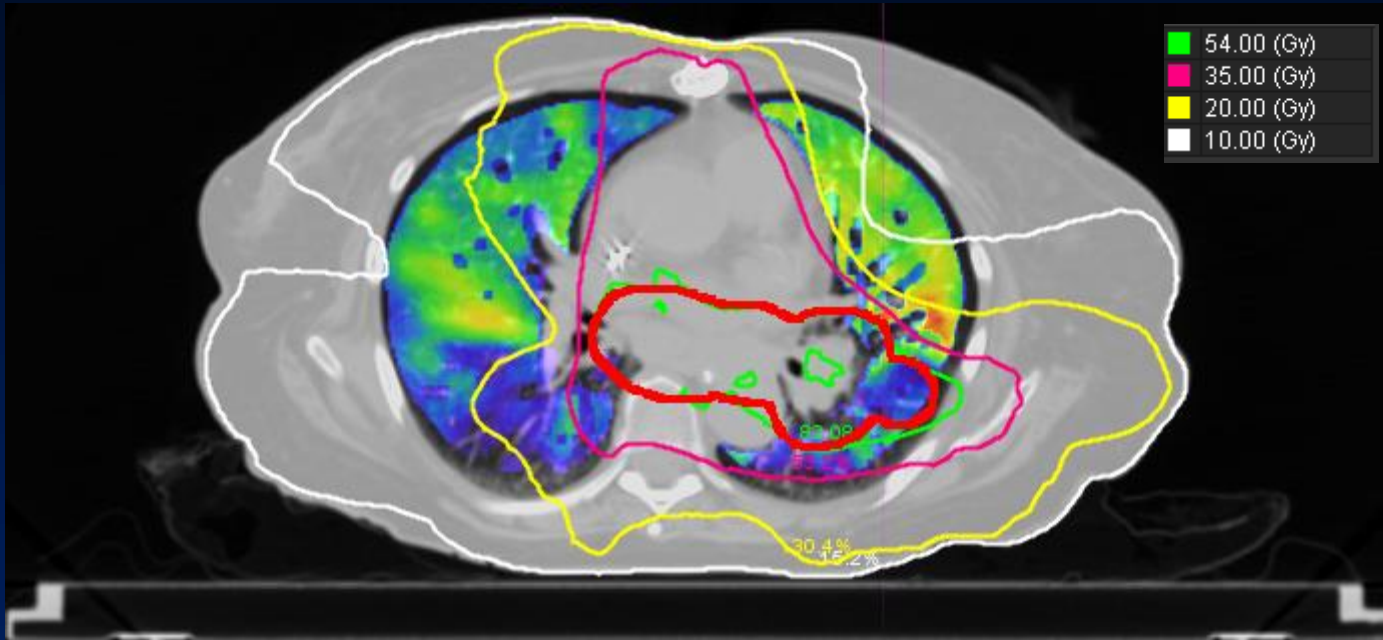
Functional planning concept

Functional plan with functional imaging

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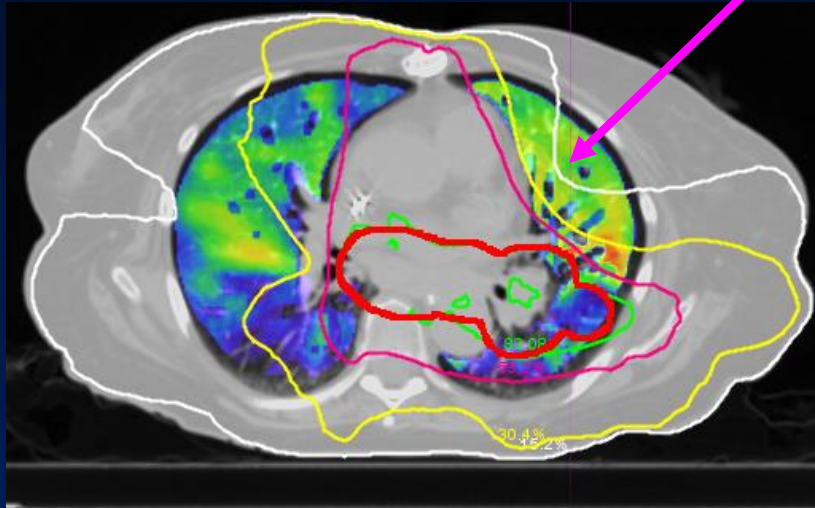
Functional planning concept

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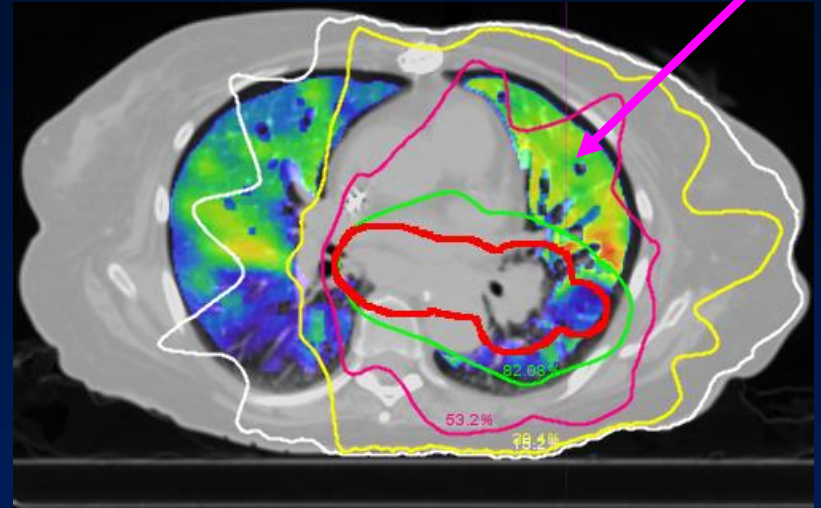


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Functional plan



Clinically used plan



Functional planning – Will it work?

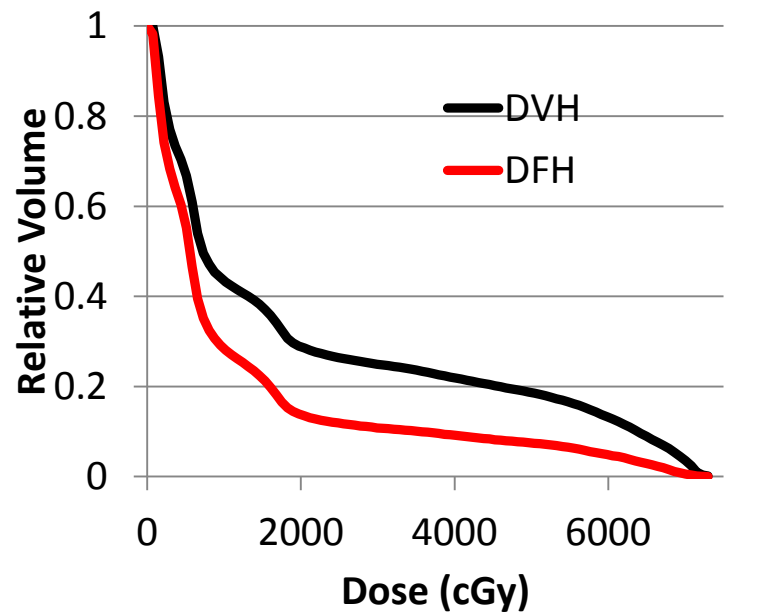
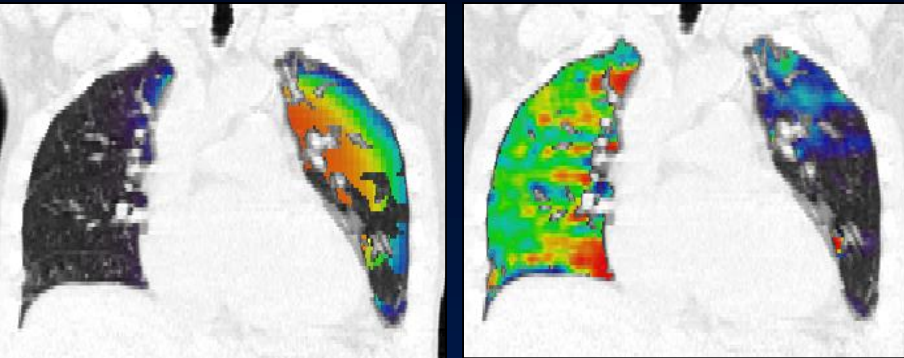
- 96 NSCLC patients
- Radiation pneumonitis toxicity information using CTCAE grading
- Calculated dose metrics
- Calculated dose + function metrics
- Is dose + function a better predictor of toxicity than dose alone

Functional planning

MLD = 22.9 Gy
No pneumonitis

Dose

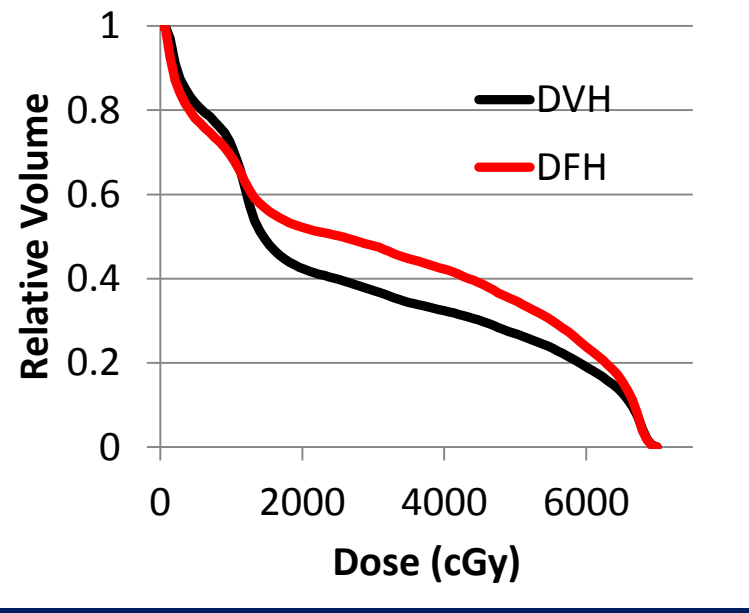
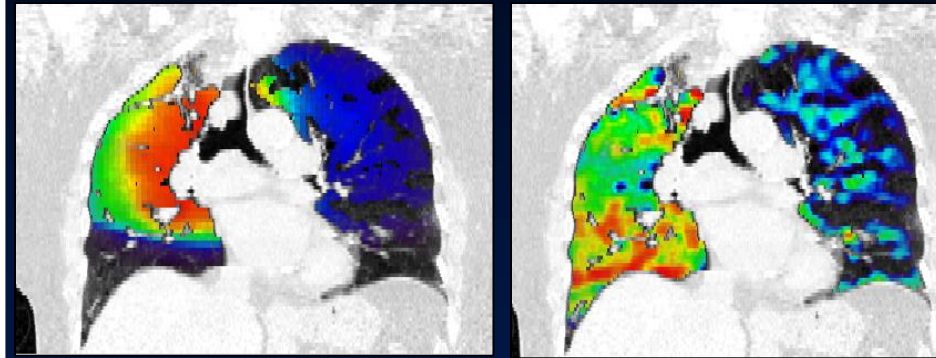
Ventilation



MLD = 23.2 Gy
Grade 3 pneumonitis

Dose

Ventilation



Functional planning

- Area under the curve (AUC) and logistic regression p value

MLD	fMLD	V20	fV20
0.55 (p=0.29)	0.62 (p=0.07)	0.57 (p=0.23)	0.66 (p=0.04)

Outline

4DCT-Ventilation Imaging

- Image formation
- Validation
- Clinical applications
- **Clinical trial**

4DCT-Ventilation Clinical Trial

- 70 lung cancer patients between 2 institutions
- Use 4DCT to calculate ventilation imaging
- Use 4DCT-ventilation to design functional radiation plans
- Hypothesis: 4DCT-ventilation functional planning results in less pulmonary toxicity than toxicity with current standard of care techniques
- Assess lung function in a variety of ways
 - CTCAE Toxicity (Pneumonitis, esophagitis)
 - QOL Questionnaires
 - PFTs
 - CT/4DCT-Ventilation imaging
 - Nuclear Medicine VQ Imaging
 - PET Imaging

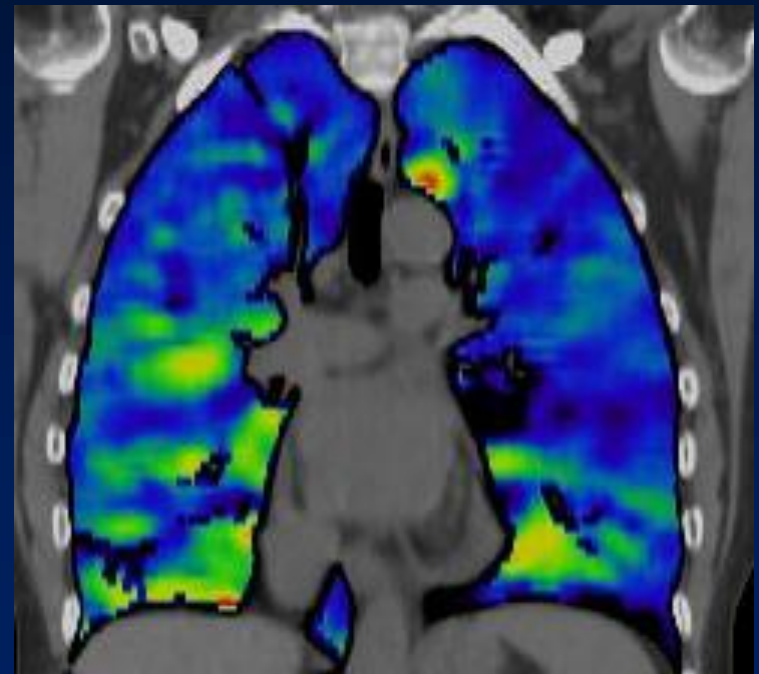
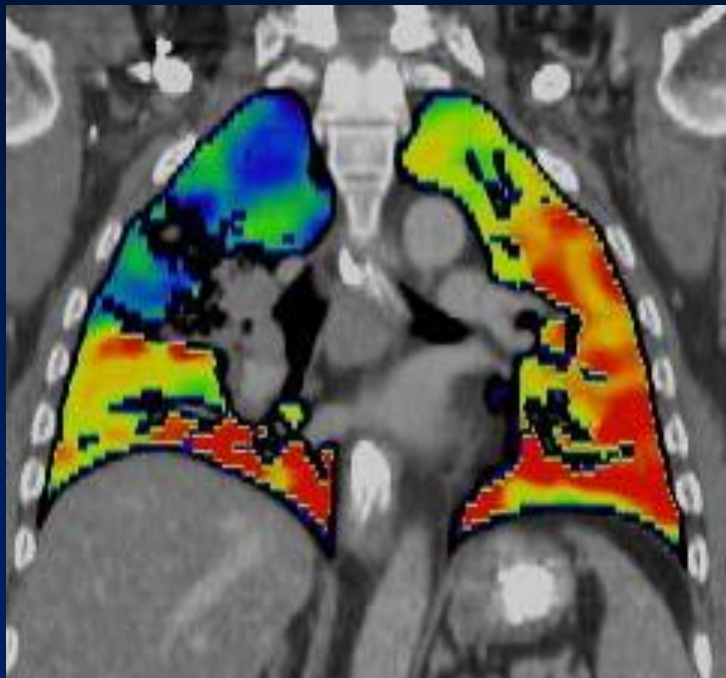
Should all patients be eligible?

Patient spatial lung function

Heterogeneous ventilation
Suitable for functional sparing

Homogenous ventilation
Not-suitable for functional sparing

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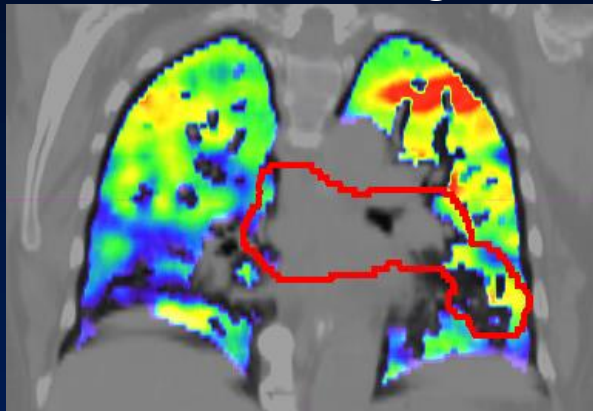


Protocol Basics

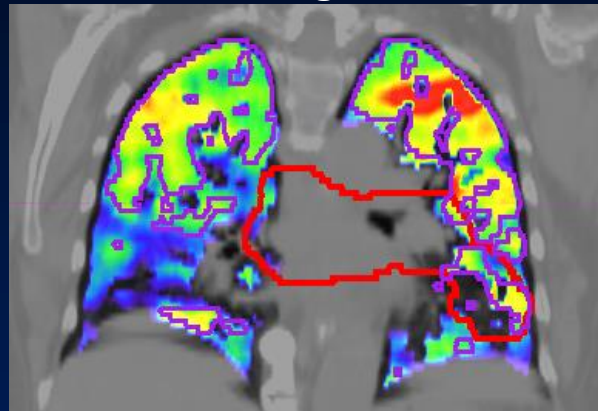
- **Functional planning**
 - **Structure based functional approach**

Planning techniques

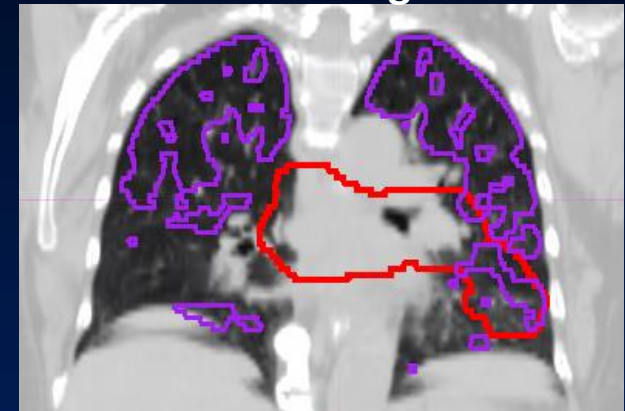
Functional Image



Functional Image + Structure



Functional Planning Structure



Protocol Basics

- **Functional planning**
 - **Structure based functional approach**
 - **Start with standard (non-functional plan)**
 - **Planning priorities 1) Target coverage 2) OAR constraints 3) Reducing dose to functional lung**

Conclusions

- **4DCT-Ventilation calculates lung ventilation maps from 4DCT data**
- **4DCT-Ventilation has been validated against established methods of measuring lung function**
- **Retrospective work suggests toxicity can be reduced with functional planning**
- **Clinical trials are underway to evaluate 4DCT-Ventilation based functional planning**

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