

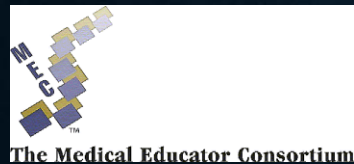
# **ARTA M. MONJAZEB, MD, PHD**

## **ADVANCES IN RADIATION ONCOLOGY**

**RELEVANT FINANCIAL RELATIONSHIPS IN THE PAST TWELVE MONTHS BY  
PRESENTER OR SPOUSE/PARTNER.**

**GRANT/RESEARCH SUPPORT: TRANSGENE, BMS, GENENTECH, INCYTE, DYNAVAX, MERCK  
CONSULTANT: INCYTE, DYNAVAX, ASTRA-ZENECA**

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**19th Annual Advances in Oncology – 2018  
September 28-29, 2018**

# Advances in Radiation Oncology

- **Advances in Treatment Delivery**
  - IGRT
  - IMRT
  - IMAT/VMAT
  - SBRT
  - **Particle beam therapy**
- **Advances in Treatment Planning**
  - Tumor Localization & Image fusion
  - Breathing adaptive radiotherapy
  - Functional treatment planning
- **Advances in Radiobiology**
  - Immunotherapy

# IGRT for prone pelvic therapy

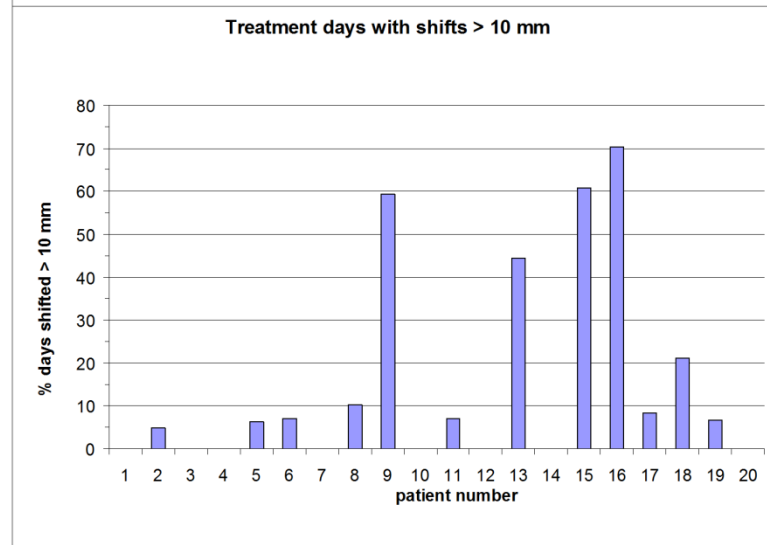
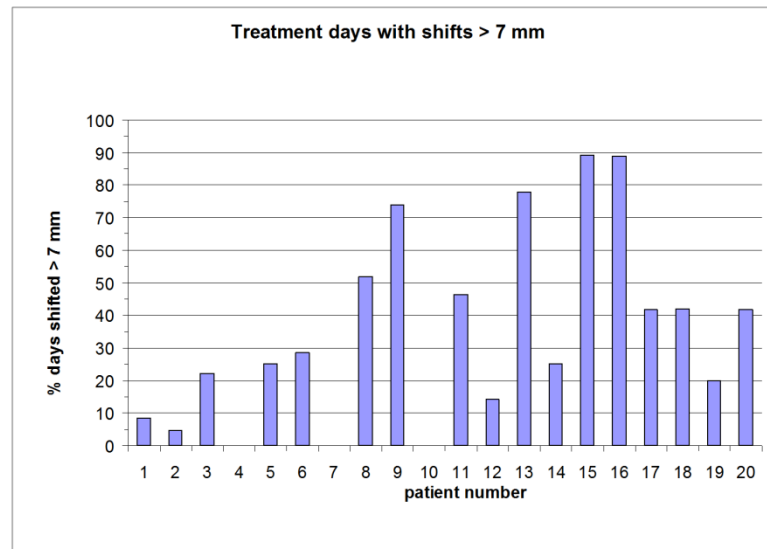
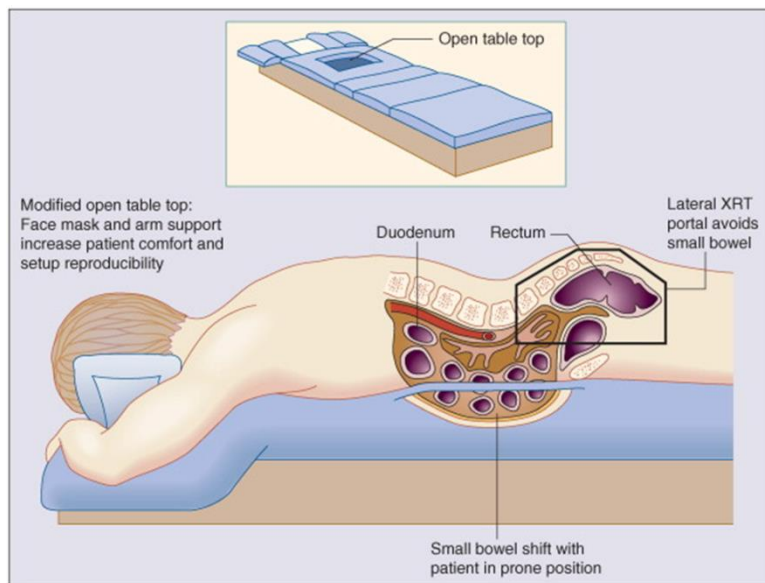
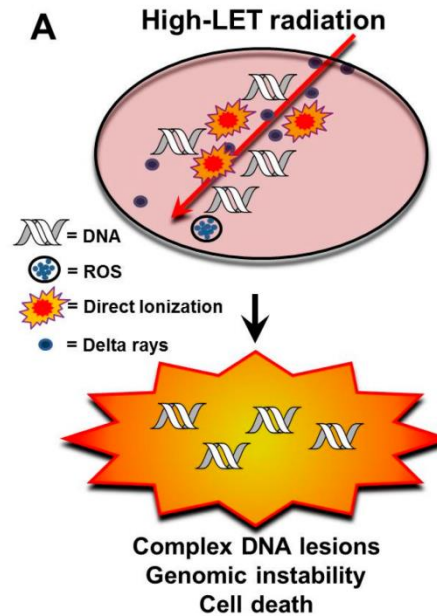
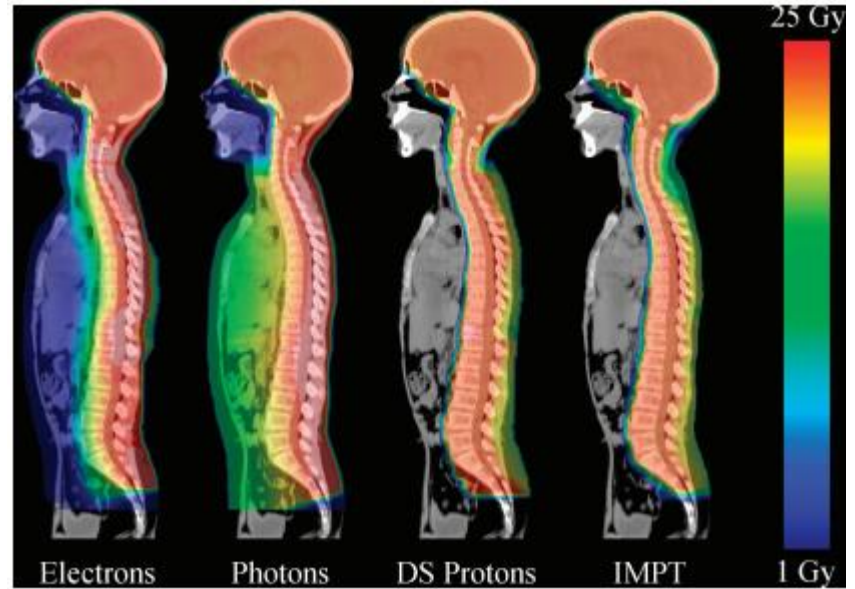
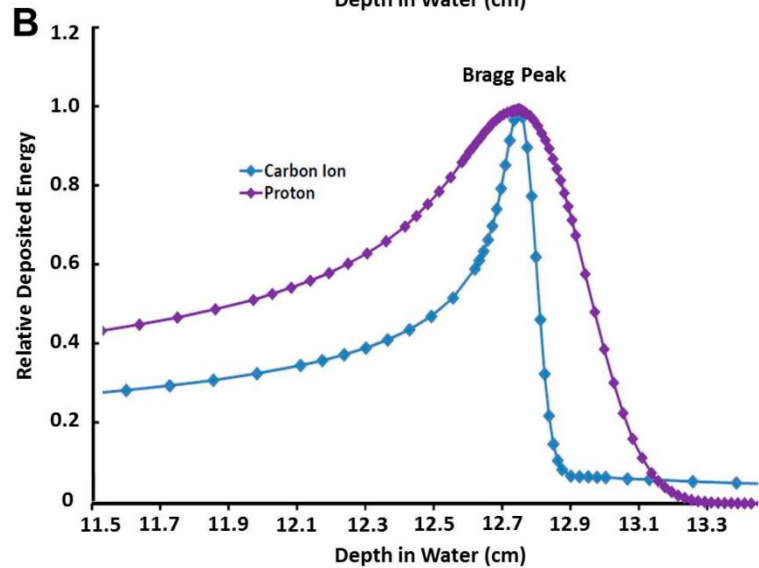
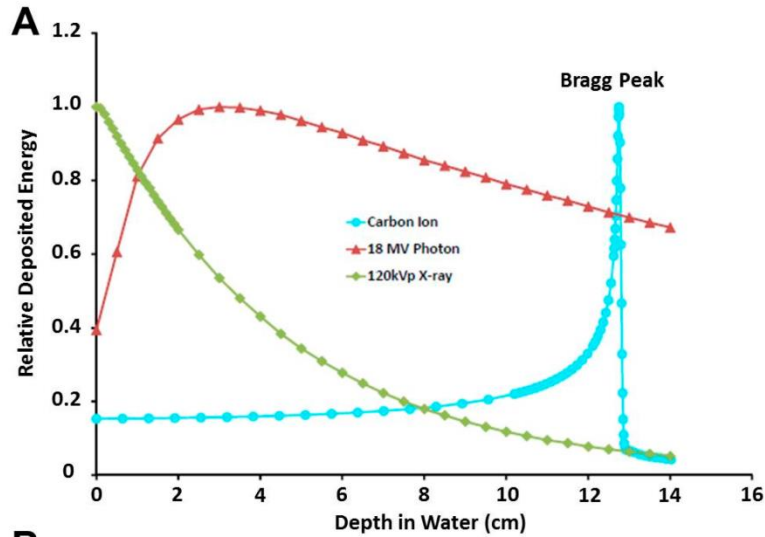


Figure 4: Proportion of days requiring IGRT-based shifts of > 7 mm, or > 10 mm, for the patient cohort.

# Particle beam therapy



# An evidence based review of proton beam therapy: The report of ASTRO's emerging technology committee

Aaron M. Allen<sup>a,\*</sup>, Todd Pawlicki<sup>b</sup>, Lei Dong<sup>c</sup>, Eugene Fourkal<sup>d</sup>, Mark Buyyounouski<sup>d</sup>, Keith Cengel<sup>e</sup>, John Plastaras<sup>e</sup>, Mary K. Bucci<sup>c</sup>, Torunn I. Yock<sup>f</sup>, Luisa Bonilla<sup>a</sup>, Robert Price<sup>d</sup>, Eleanor E. Harris<sup>g</sup>, Andre A. Koniski<sup>h</sup>

<sup>a</sup> Davidoff Center, Tel Aviv University, Israel; <sup>b</sup> University of California, San Diego, La Jolla, USA; <sup>c</sup> M.D. Anderson Cancer Center, University of Texas, Houston, USA; <sup>d</sup> Fox Chase Cancer Center, Philadelphia, USA; <sup>e</sup> University of Pennsylvania, Philadelphia, USA; <sup>f</sup> Massachusetts General Hospital, Boston, USA; <sup>g</sup> H. Lee Moffitt Cancer Center, Tampa, USA; <sup>h</sup> Wayne State University Medical Center, Detroit, USA

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## ABSTRACT

Proton beam therapy (PBT) is a novel method for treating malignant disease with radiotherapy. The purpose of this work was to evaluate the state of the science of PBT and arrive at a recommendation for the use of PBT. The emerging technology committee of the American Society of Radiation Oncology (ASTRO) routinely evaluates new modalities in radiotherapy and assesses the published evidence to determine recommendations for the society as a whole. In 2007, a Proton Task Force was assembled to evaluate the state of the art of PBT. This report reflects evidence collected up to November 2009. Data was reviewed for PBT in central nervous system tumors, gastrointestinal malignancies, lung, head and neck, prostate, and pediatric tumors. Current data do not provide sufficient evidence to recommend PBT in lung cancer, head and neck cancer, GI malignancies, and pediatric non-CNS malignancies. In hepatocellular carcinoma and prostate cancer and there is evidence for the efficacy of PBT but no suggestion that it is superior to photon based approaches. In pediatric CNS malignancies PBT appears superior to photon approaches but more data is needed. In large ocular melanomas and chordomas, we believe that there is evidence for a benefit of PBT over photon approaches. PBT is an important new technology in radiotherapy. Current evidence provides a limited indication for PBT. More robust prospective clinical trials are needed to determine the appropriate clinical setting for PBT.

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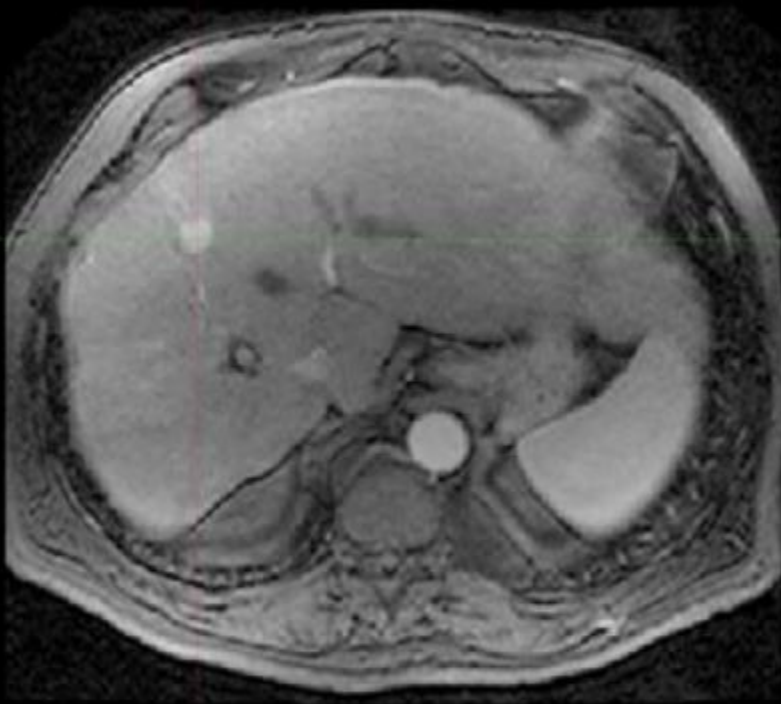
122 PBT trials ongoing with an expected accrual of > 42,000 patients. Only 5 of these trials randomize patients to protons vs. photons

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# Image Fusion for Tumor Localization

Liver: CT (No Contrast  $\Rightarrow$  No visible GTV)



Liver: MR (Visible GTV)

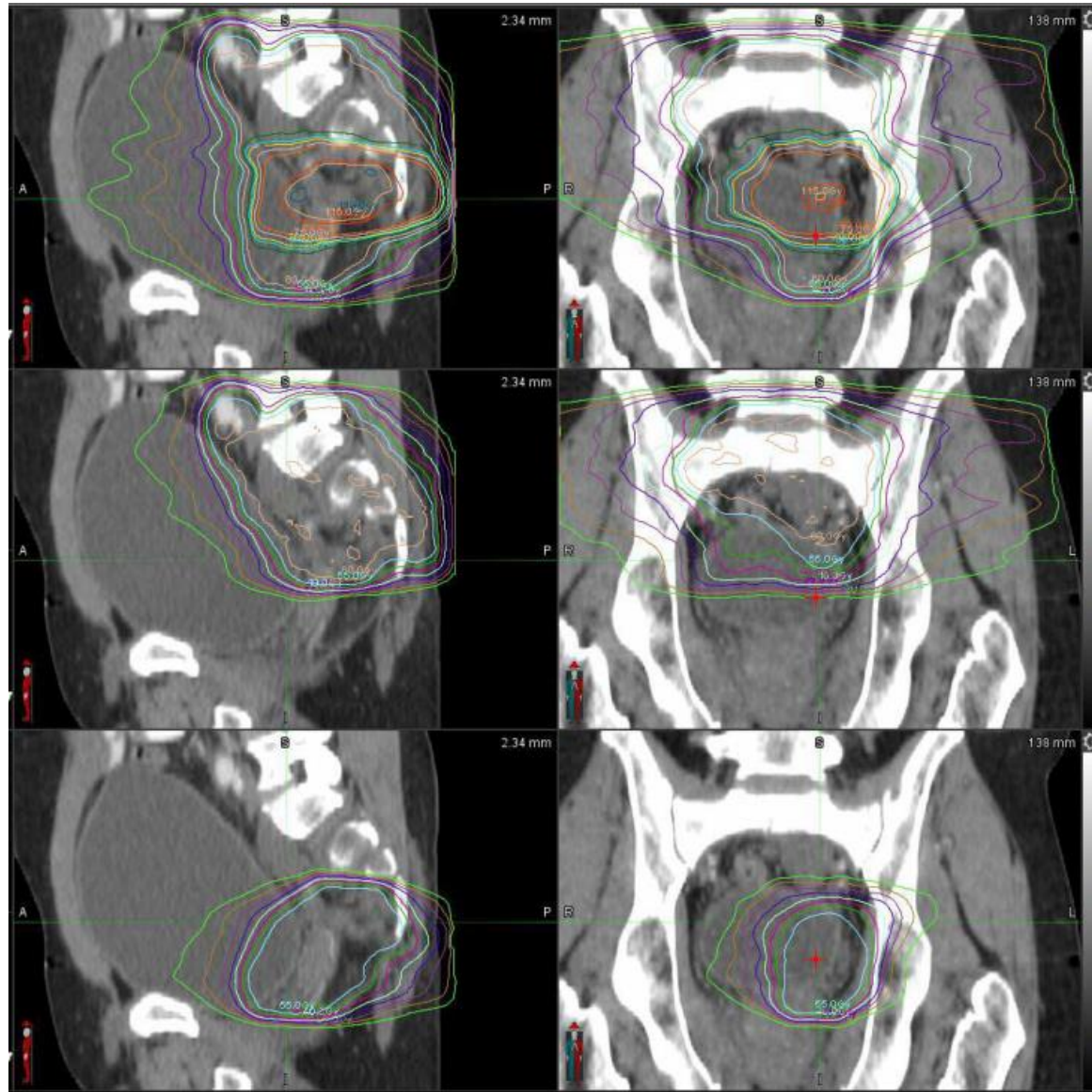
# Quality Improvement Challenges: Image Co-Registration (Fusion & DIR)



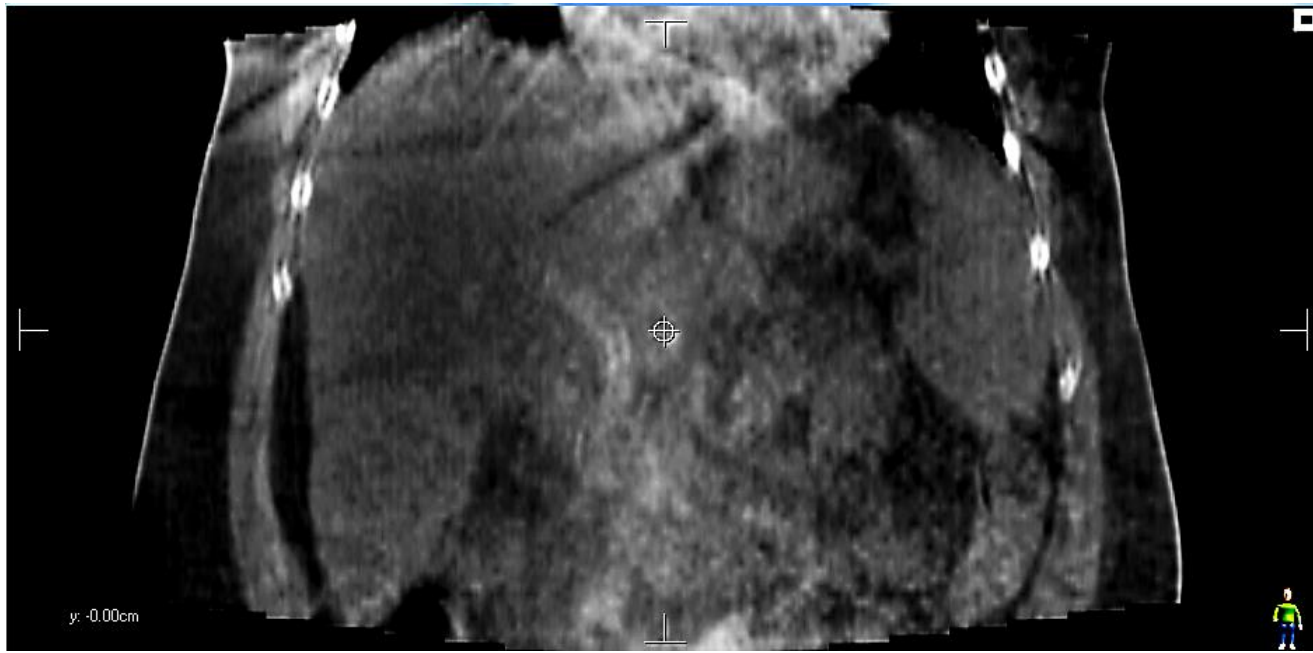
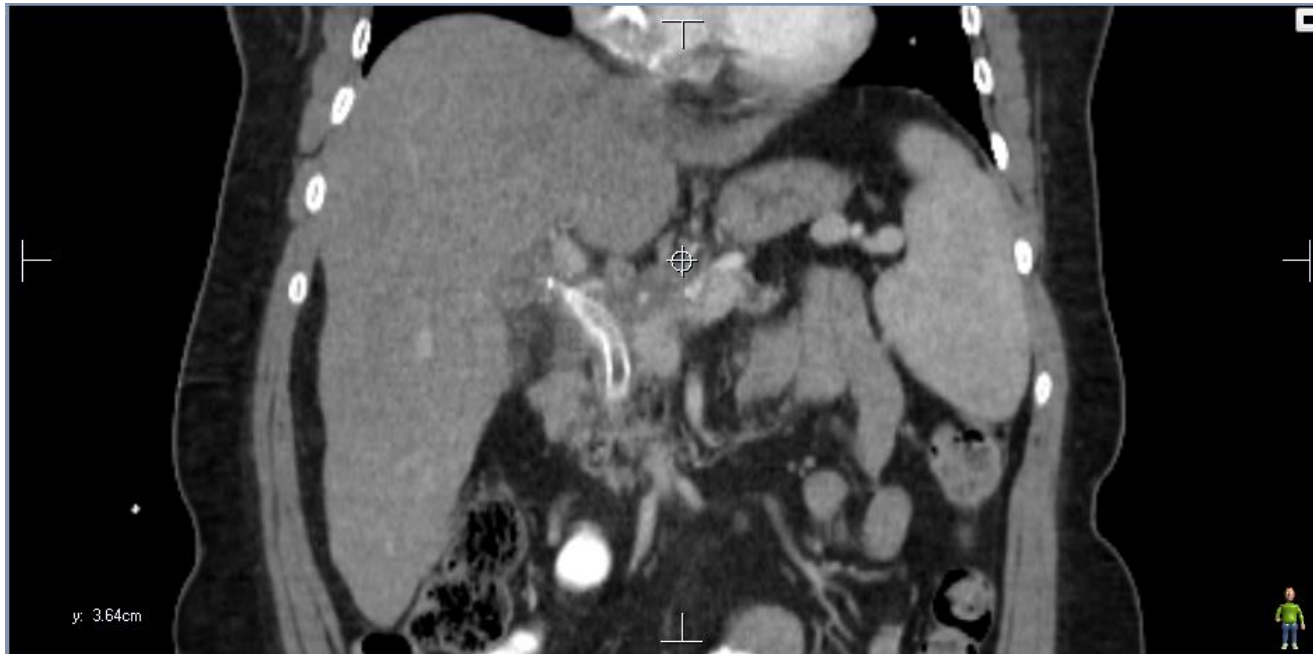
**Severus Snape & Harry Potter**



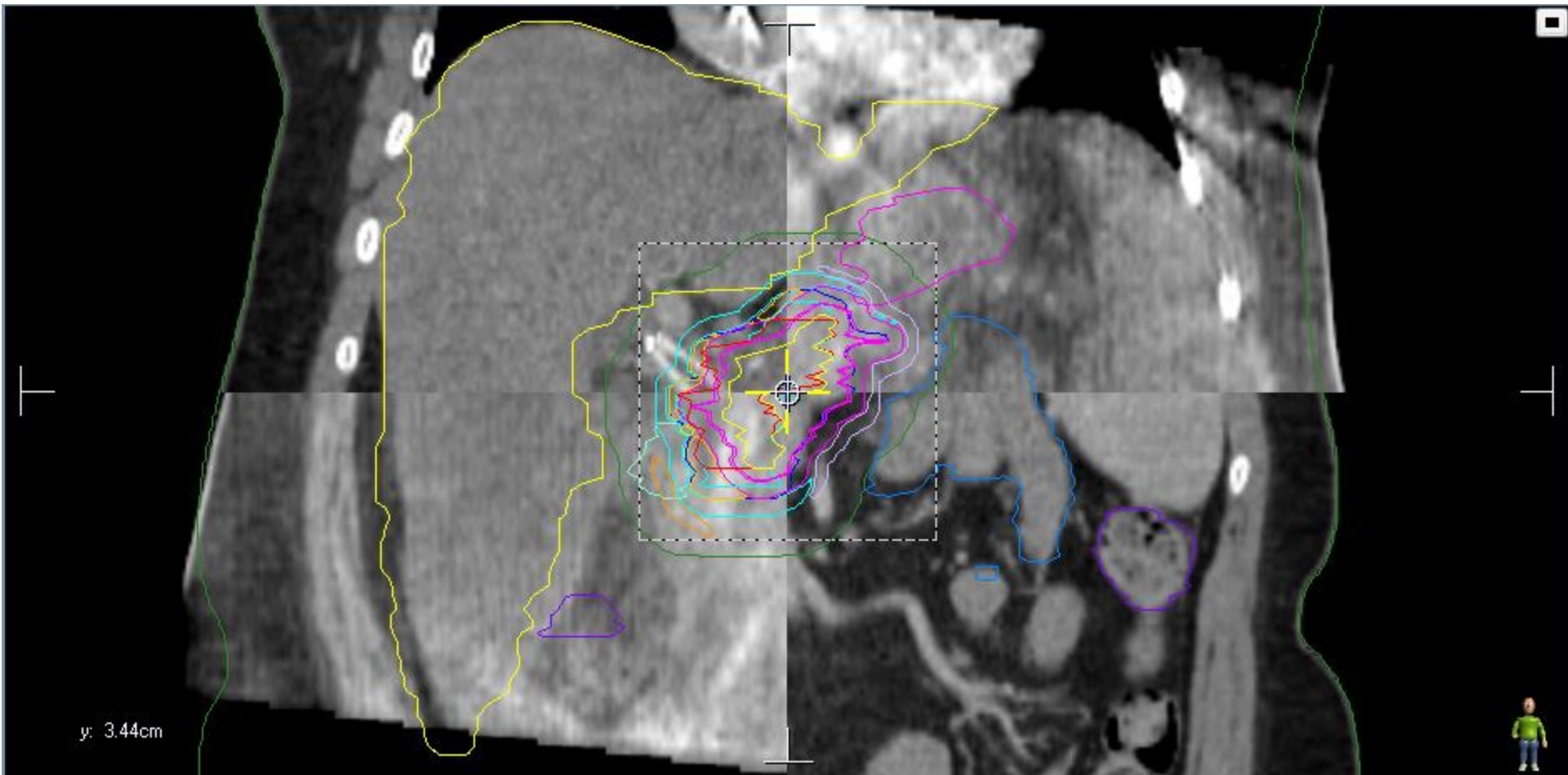
# Image and Dose Registration



# Tumor Localization for Pancreas SBRT



# Tumor Localization for Pancreas SBRT



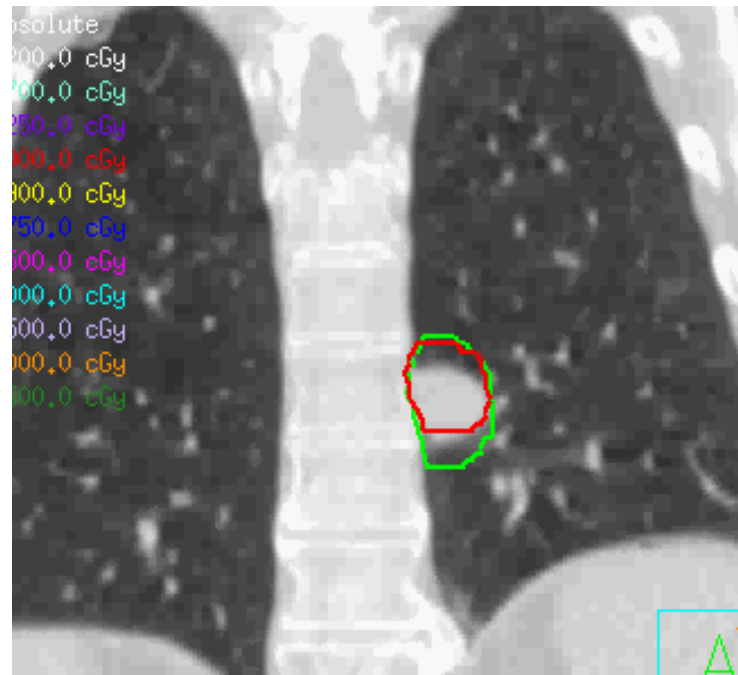
# Respiratory Motion and 4D-CT

Tokihiro Yamamoto/UC Davis

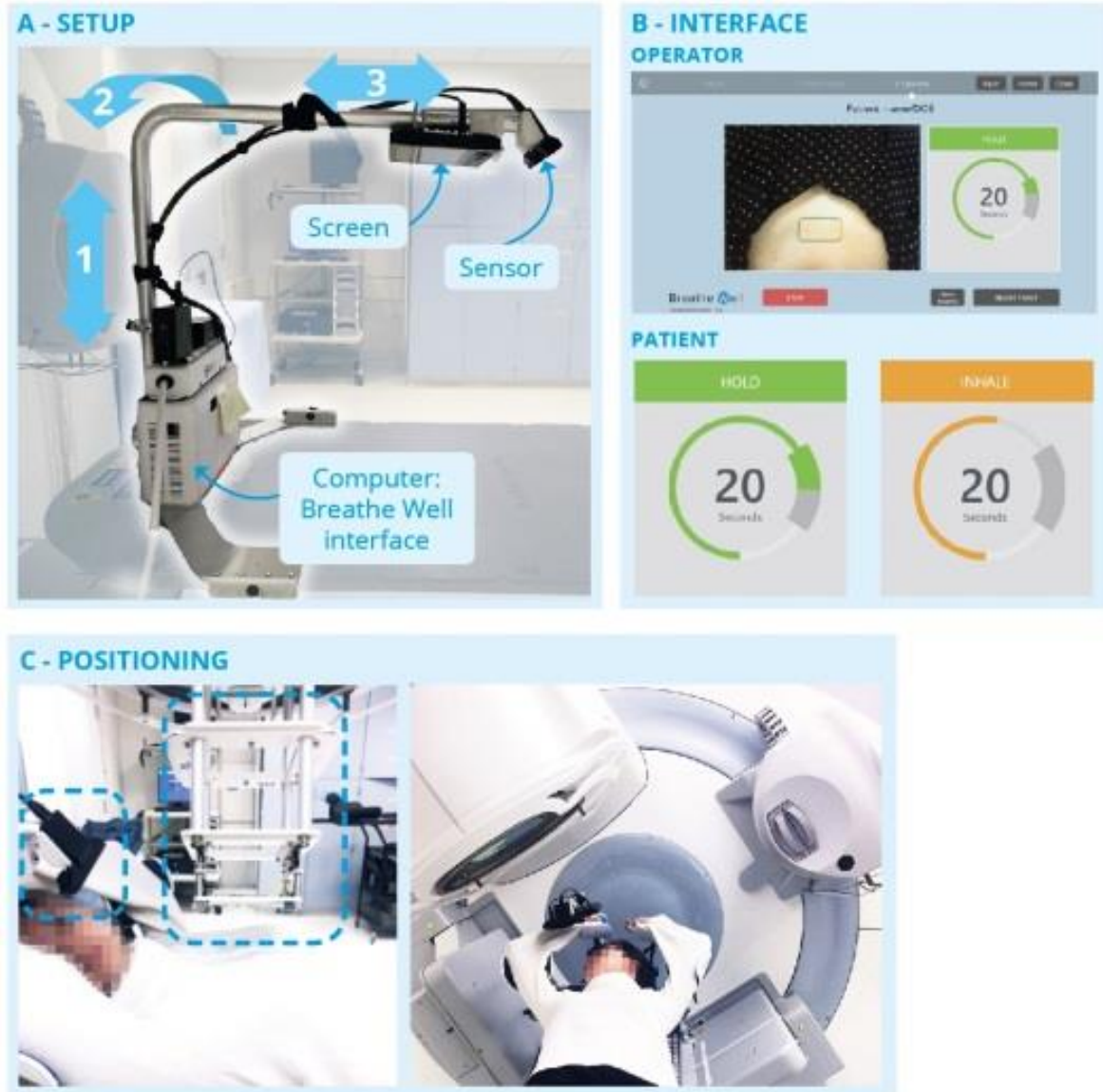


# ITV Delineation with Gating

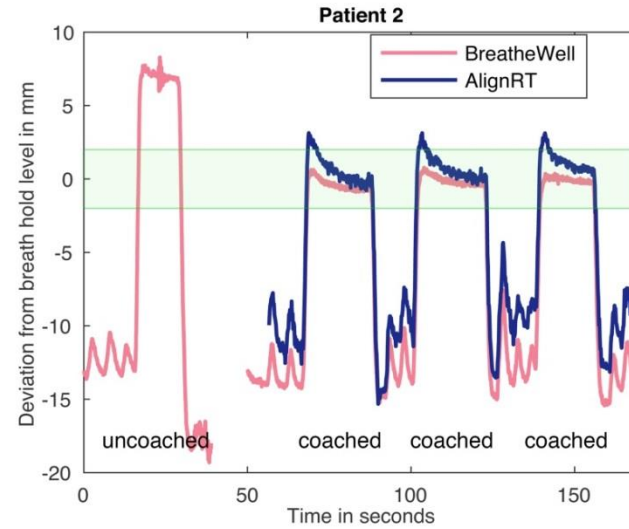
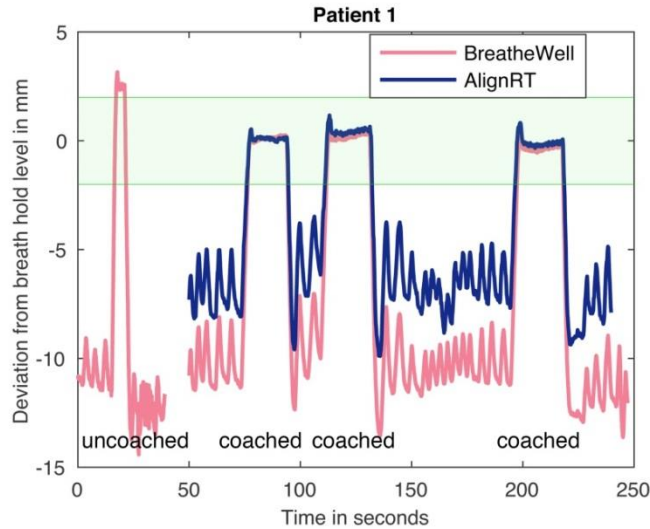
- Select a gating window based on tumor motion on 4DCT
- ITV will include tumor excursion and deformation within the selected gating window



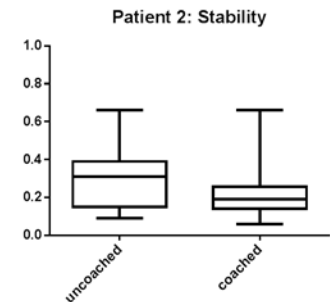
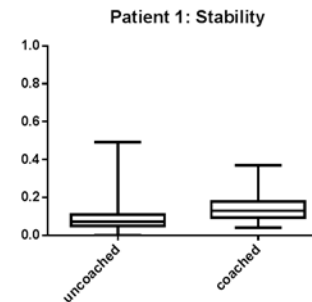
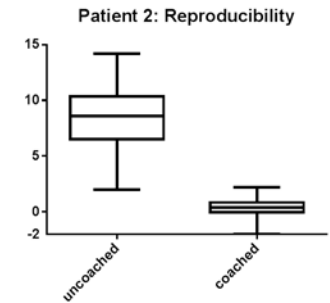
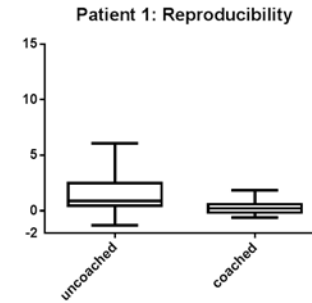
# Breathe Well System Prototype



# Patients' Breath Holds



- Patients instructed to hold an uncoached breath for 10 seconds before treatment
- Reproducibility and stability for first two patients on treatment
- Reproducibility improved significantly ( $p < 0.001$ )
- Stability did not change significantly ( $p = 0.96$ )



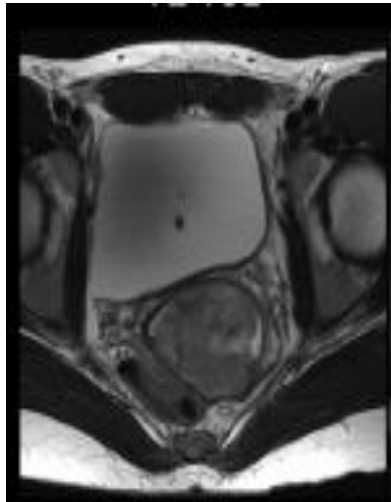
# Anatomic vs. Biological imaging

**ANATOMICAL**

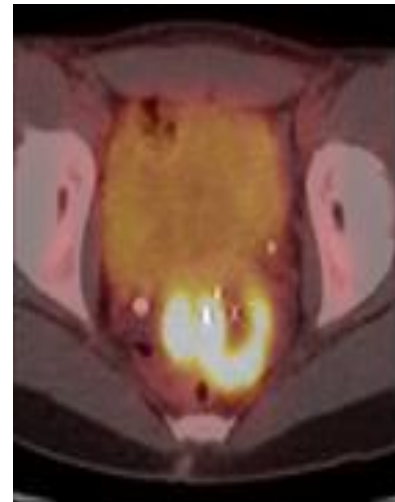
**BIOLOGICAL**



CT



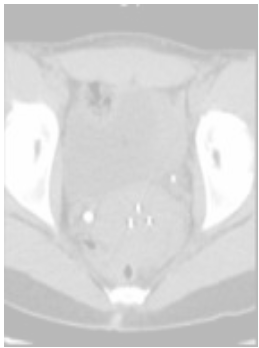
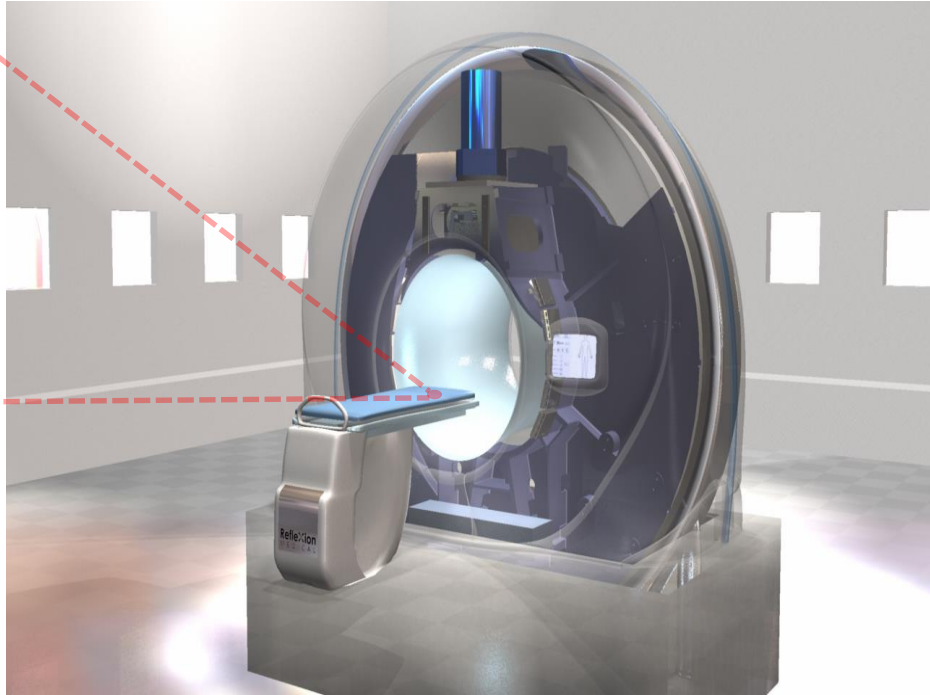
MRI



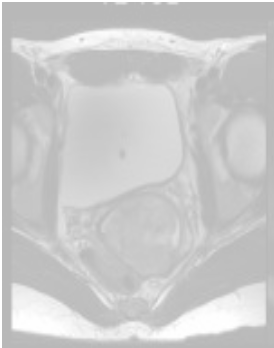
PET/CT



# Biology-guided

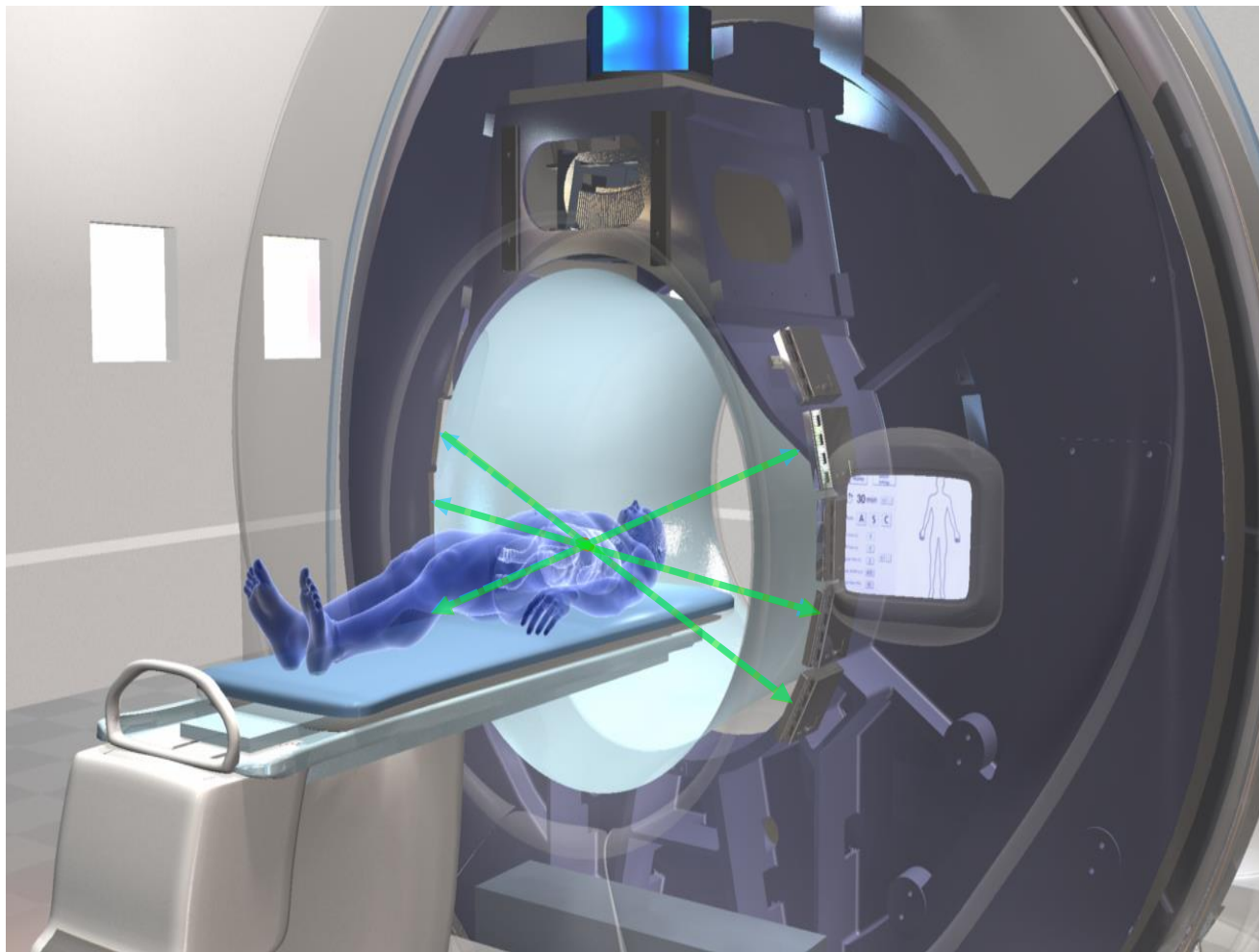


CT



MRI

Can guiding RT from “biological” signals improve or enable new applications?



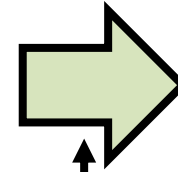
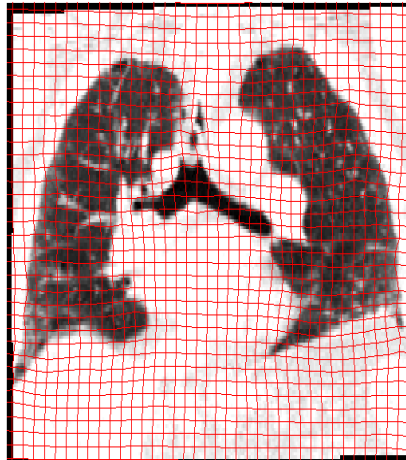
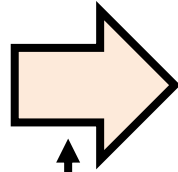
**Reflexion** - A novel radiation therapy system that responds to individual PET emissions in real-time to guide the treatment beam

# CT Ventilation Imaging

4D CT or  
exhale/inhale CT

Displacement  
vector field

Ventilation image



Deformable image  
registration (DIR)

Quantification of  
regional volume change

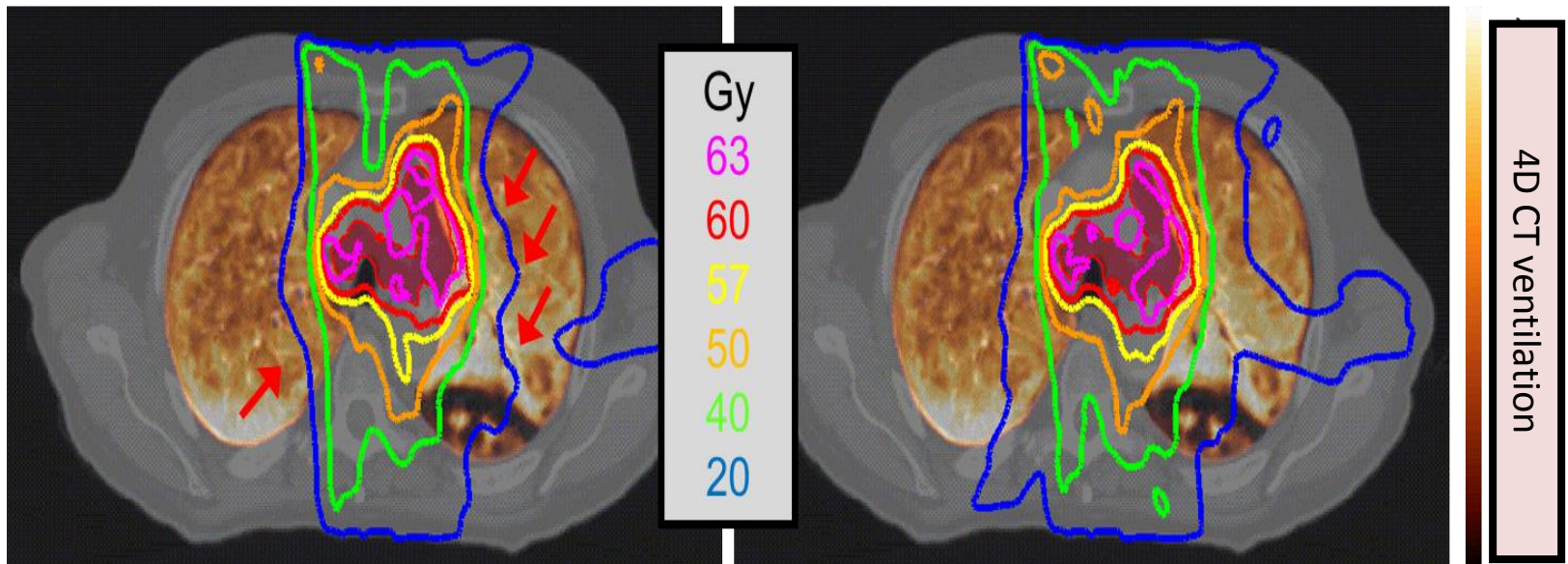
- Higher resolution, lower cost, or shorter scan time than other modalities

# CT Ventilation Image-guided RT for Lung Functional Avoidance

CT ventilation

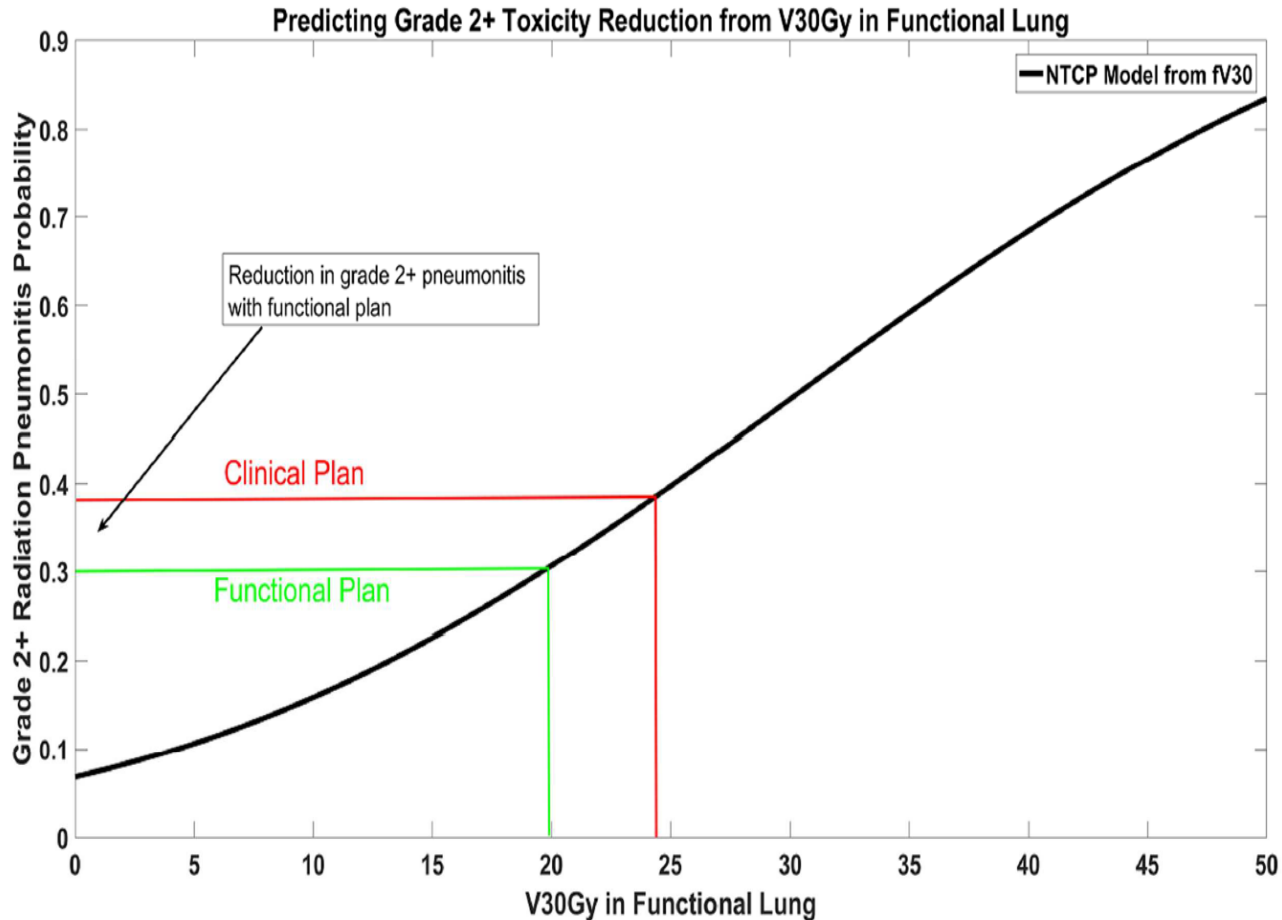
functional image-guided plan

Anatomic image-guided plan



Yamamoto *et al.* (*Radiother Oncol* 2016)

# Lung functional avoidance RT may reduce toxicity



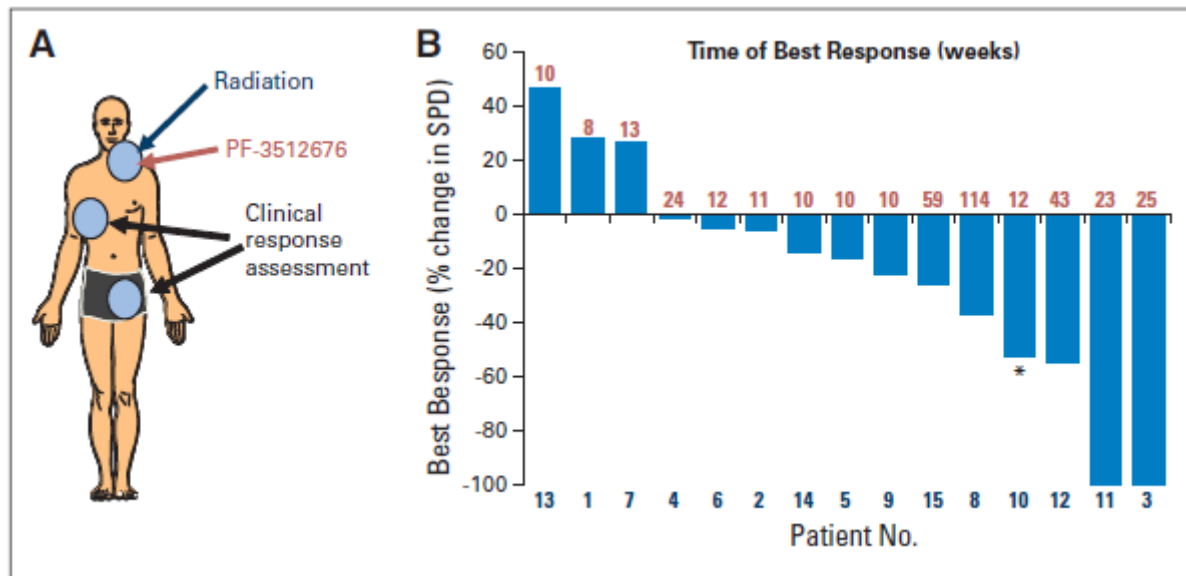
Faught/Yamamoto *et al.* (IJROBP In Press)

# Advances in Radiation Oncology

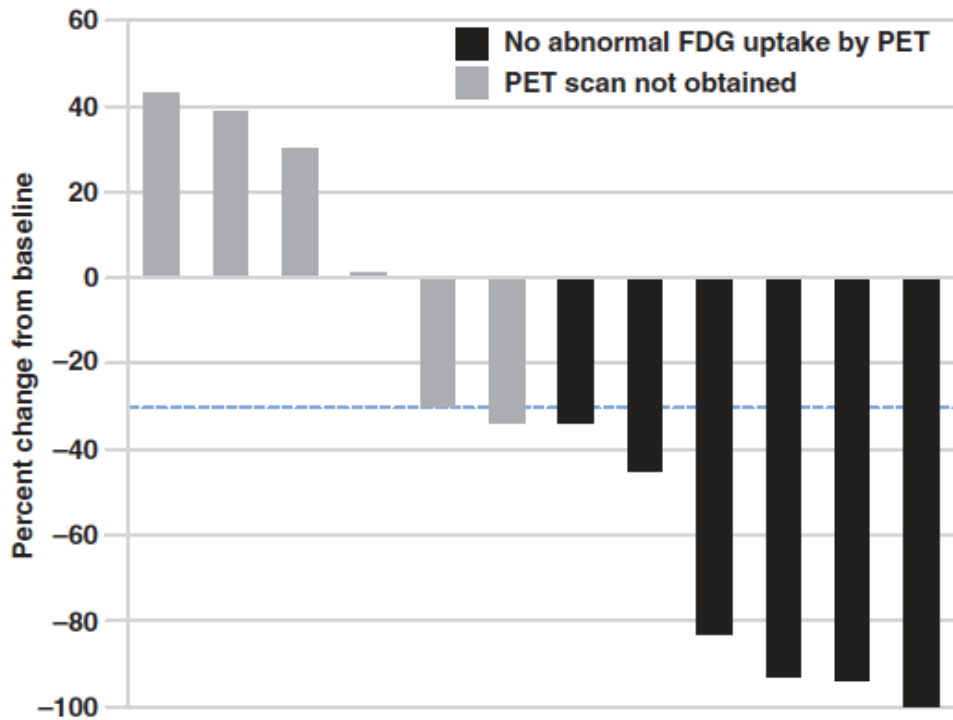
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# In Situ Vaccination With a TLR9 Agonist Induces Systemic Lymphoma Regression: A Phase I/II Study

Joshua D. Brody, Weiyun Z. Ai, Debra K. Czerwinski, James A. Torchia, Mia Levy, Ranjana H. Advani, Youn H. Kim, Richard T. Hoppe, Susan J. Knox, Lewis K. Shin, Irene Wapnir, Robert J. Tibshirani, and Ronald Levy

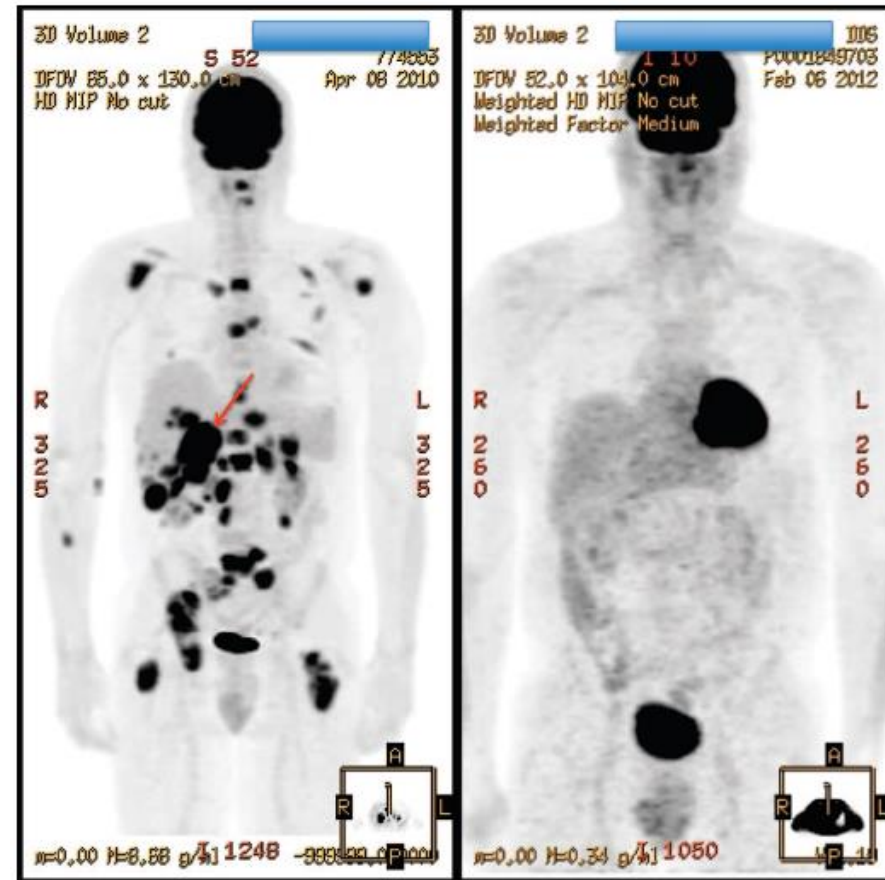


# RT + HD systemic IL-2



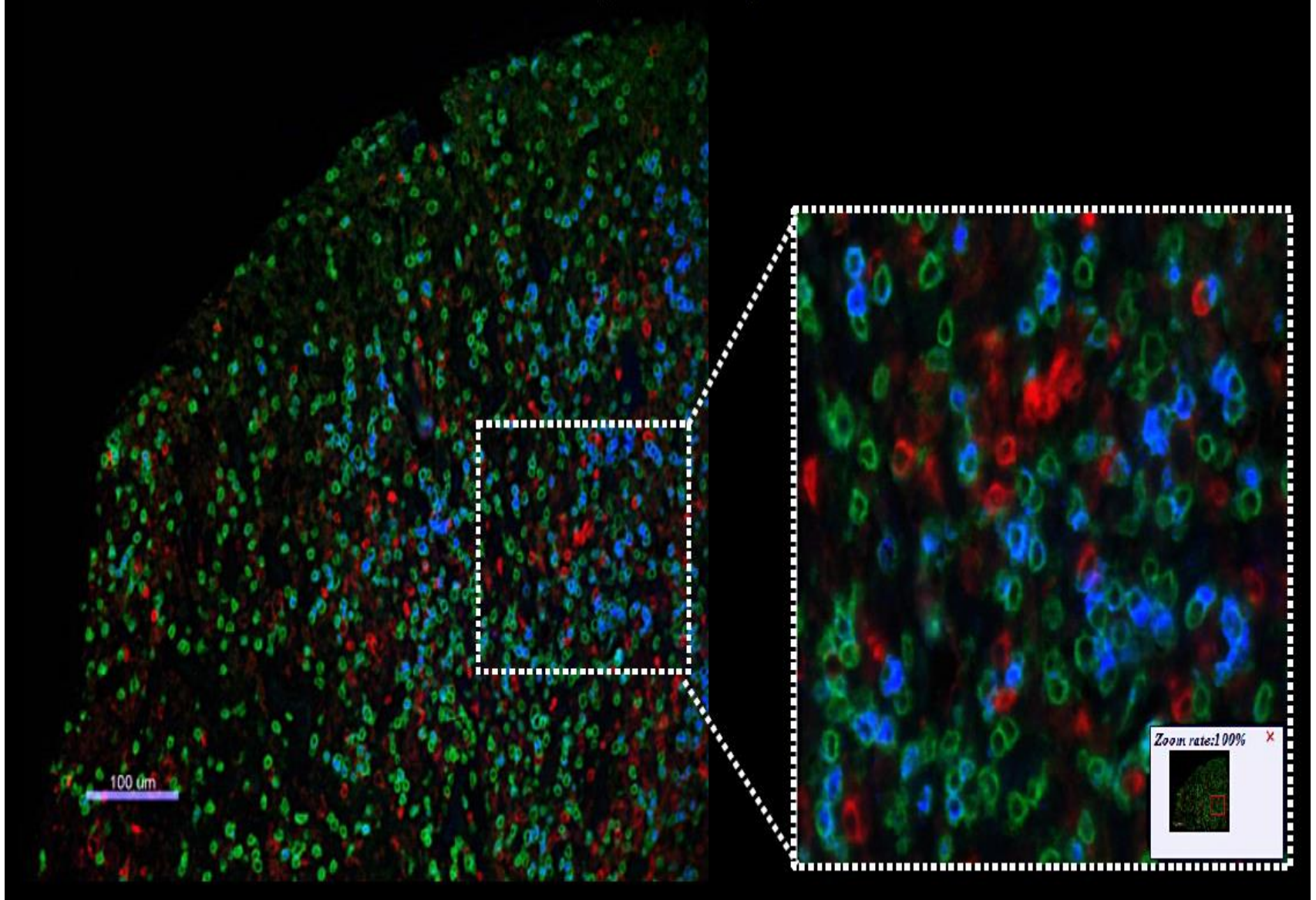
ORR: 66%

Historical response rate: 10-15%



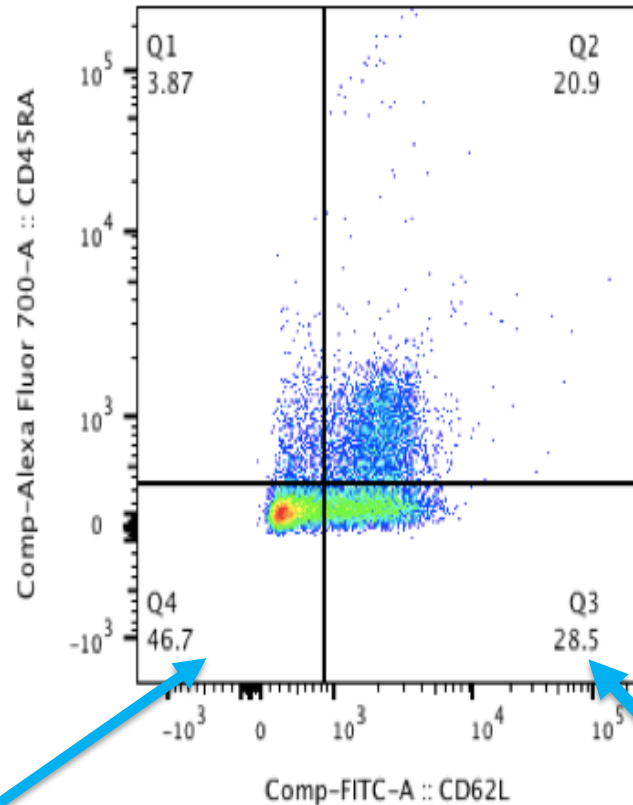


# CD4/CD8/CD20

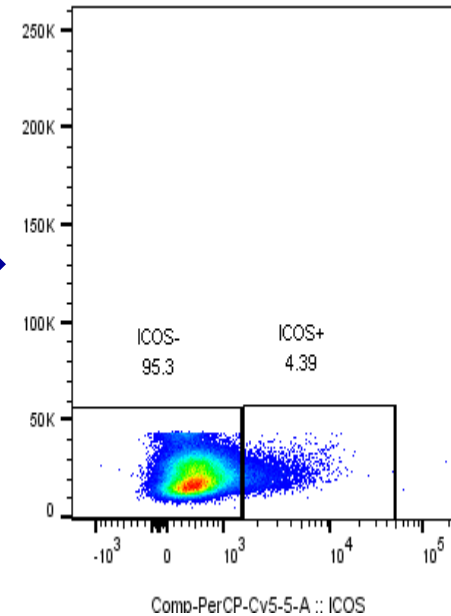


# FACS Analysis for Memory T cell populations

CD4+ Memory Cell Subsets



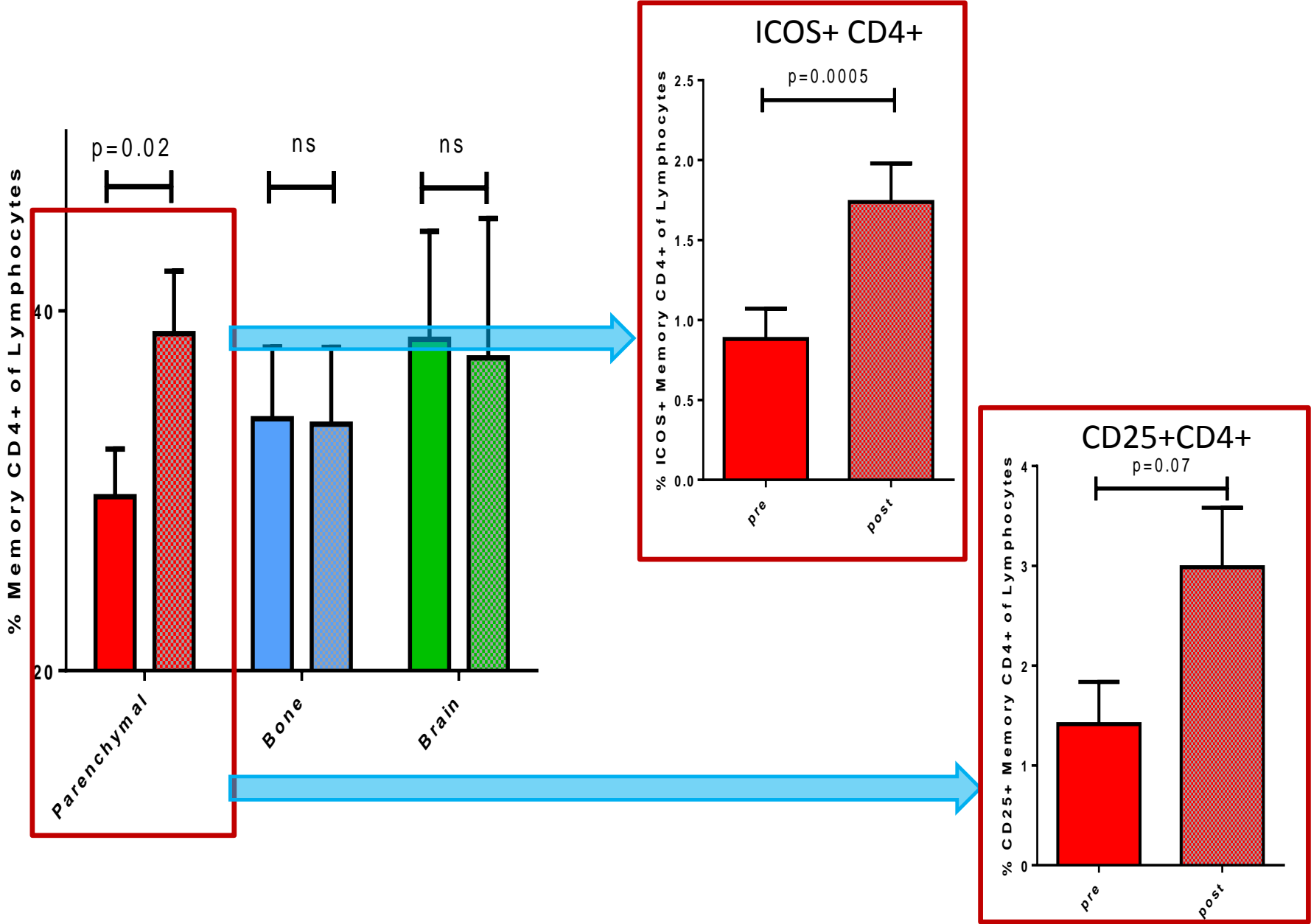
ICOS+ CD4+ Cells



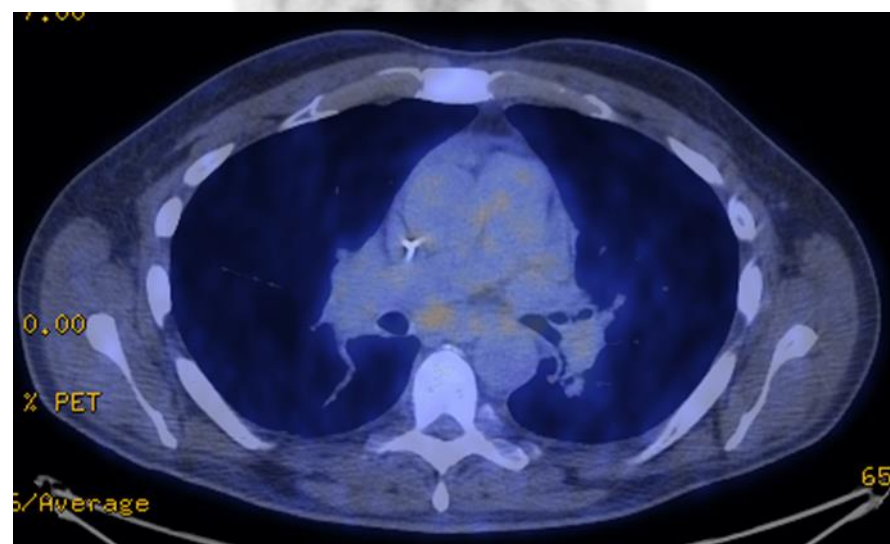
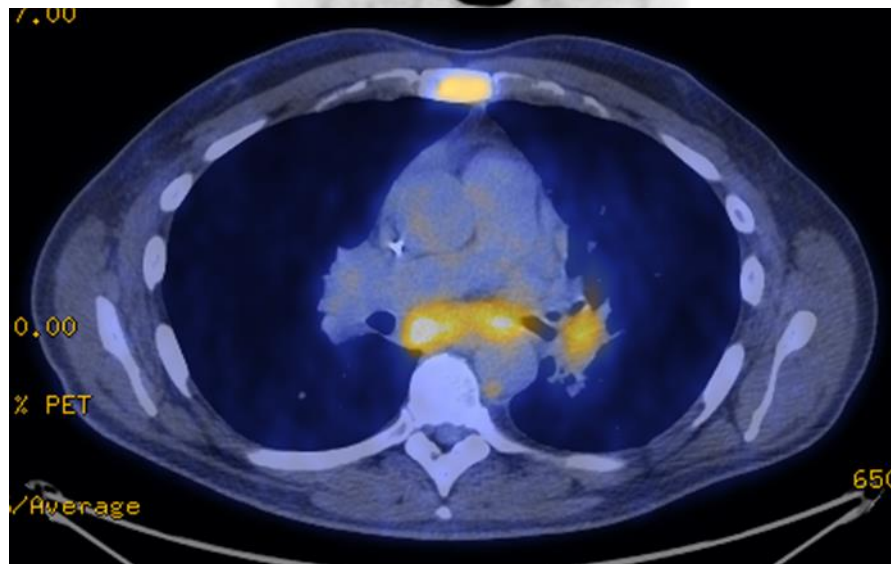
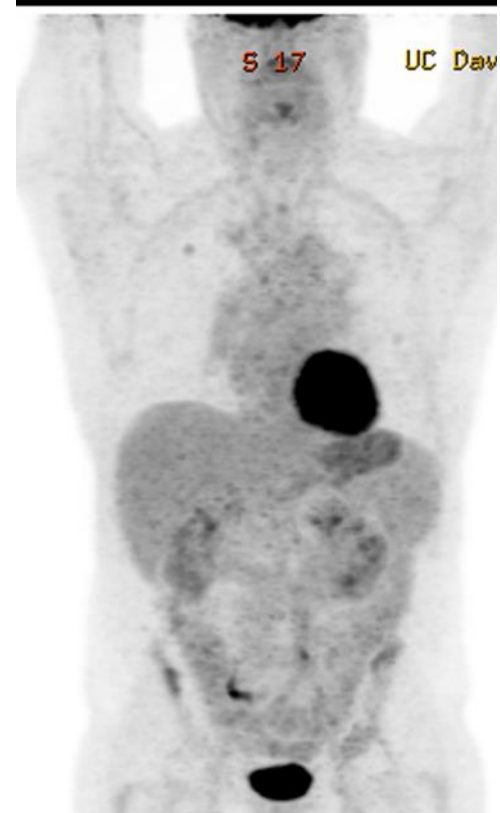
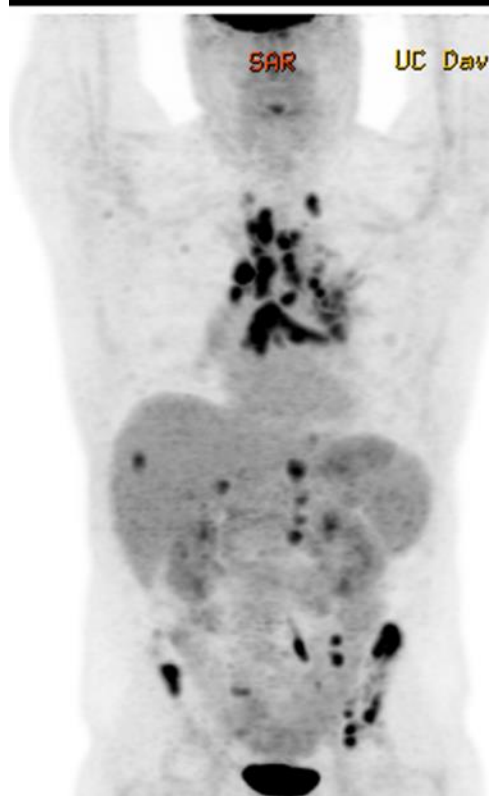
Effector Memory  
CD4+  
T cells (CD45RA-  
CD62L-)

Central Memory  
CD4+  
T cells (CD45RA-  
CD62L+)

# Systemic Increase in Activated Memory CD4+ T-cells







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- **UC Davis Laboratory of Cancer Immunology**
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