

# (Quantitative) Imaging for Adaptive Radiotherapy

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# Disclosure: No Conflict, No Interest

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- Modus Medical Devices Inc.
- Shelley Medical Imaging Technologies

# Understanding Imaging: Overview

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- Adaptive Imaging Specifications
- Functional Imaging Techniques
- Validation and Standardization

# Imaging and Personalized Cancer Medicine

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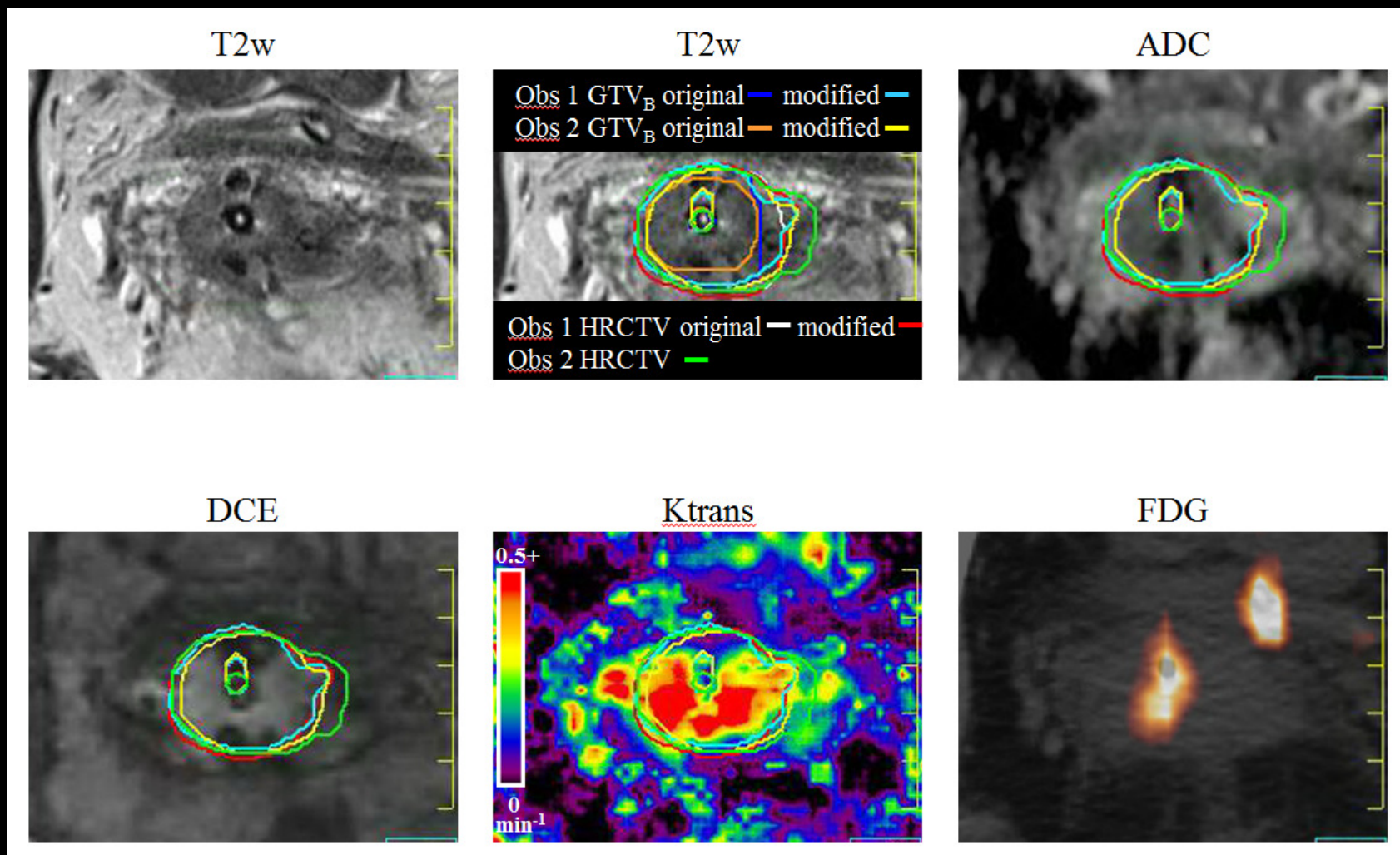
- Quantify individual tumor microenvironment
- Earlier physiological effect than volume change
- Response Assessment to adapt treatment where needed



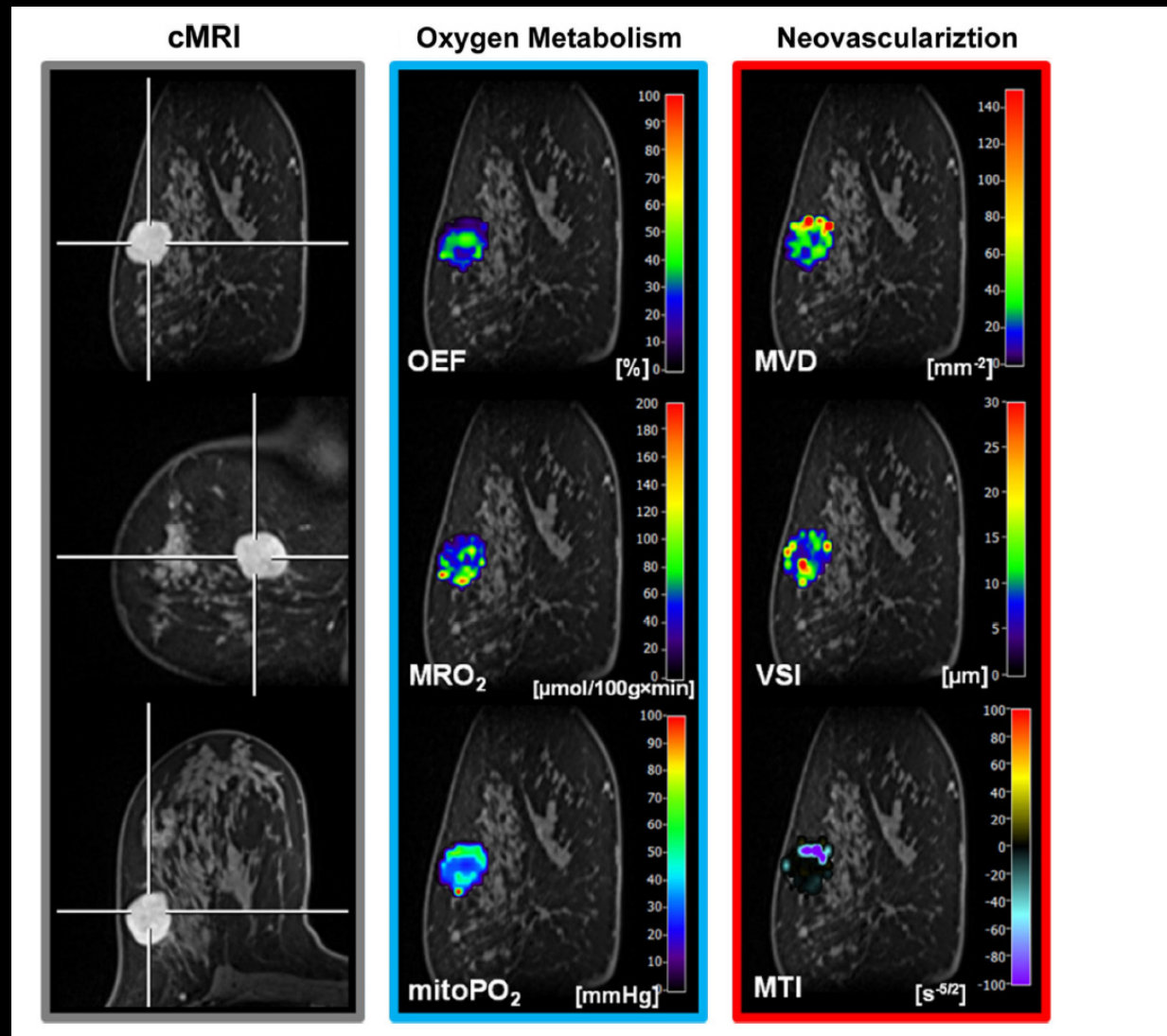
How interrogate the morphological and physiological status of the tumor before, during and after treatment?

# 1. Understanding Cancer: Target Definition

Reduced variation in delineation



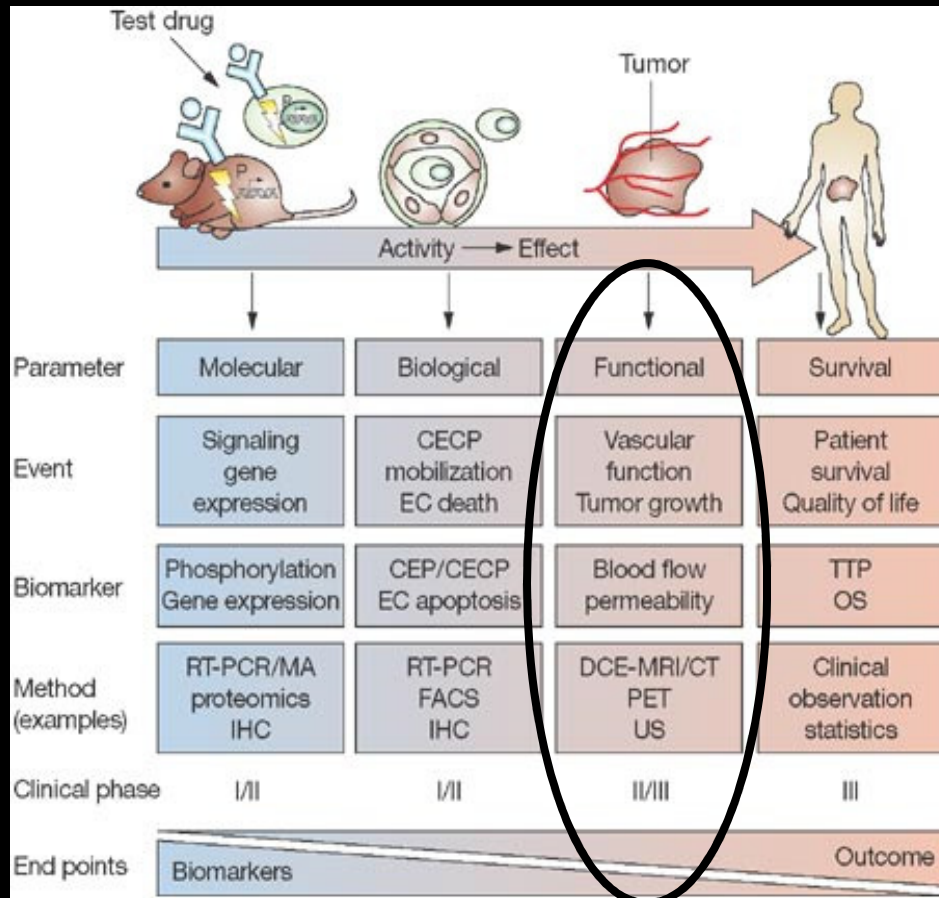
# 1. Understanding Cancer: Heterogeneity



Surrogate  
prognostic  
factor



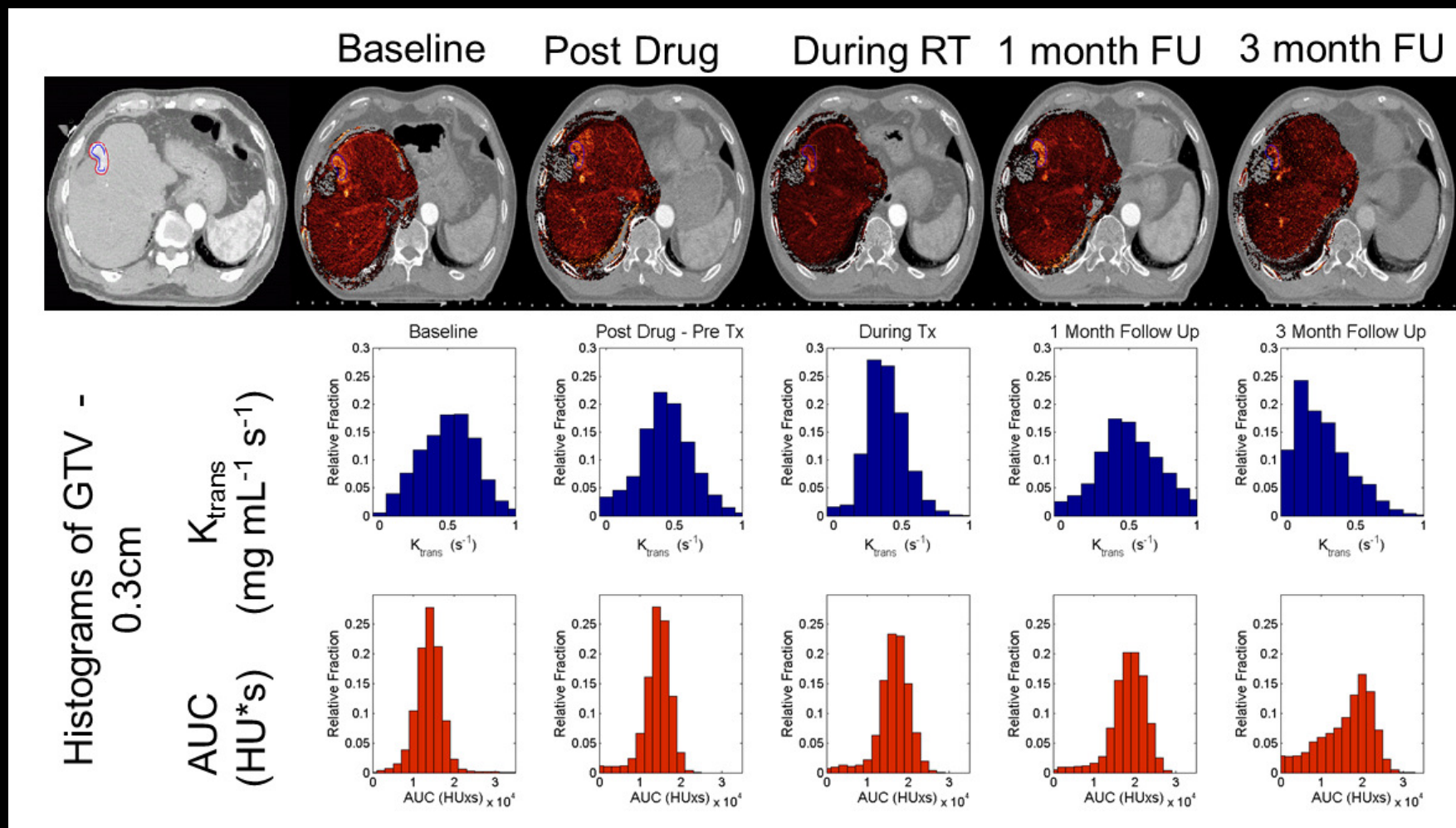
# 1. Understanding Cancer: Microenvironment



Source: Nat Clin Pract Oncol

- Angiogenesis
- Interstitial Fluid Pressure
- Metabolism
- Oxygenation/Hypoxia
- Cell density
- Vessel Permeability

# 1. Understanding Cancer: Response



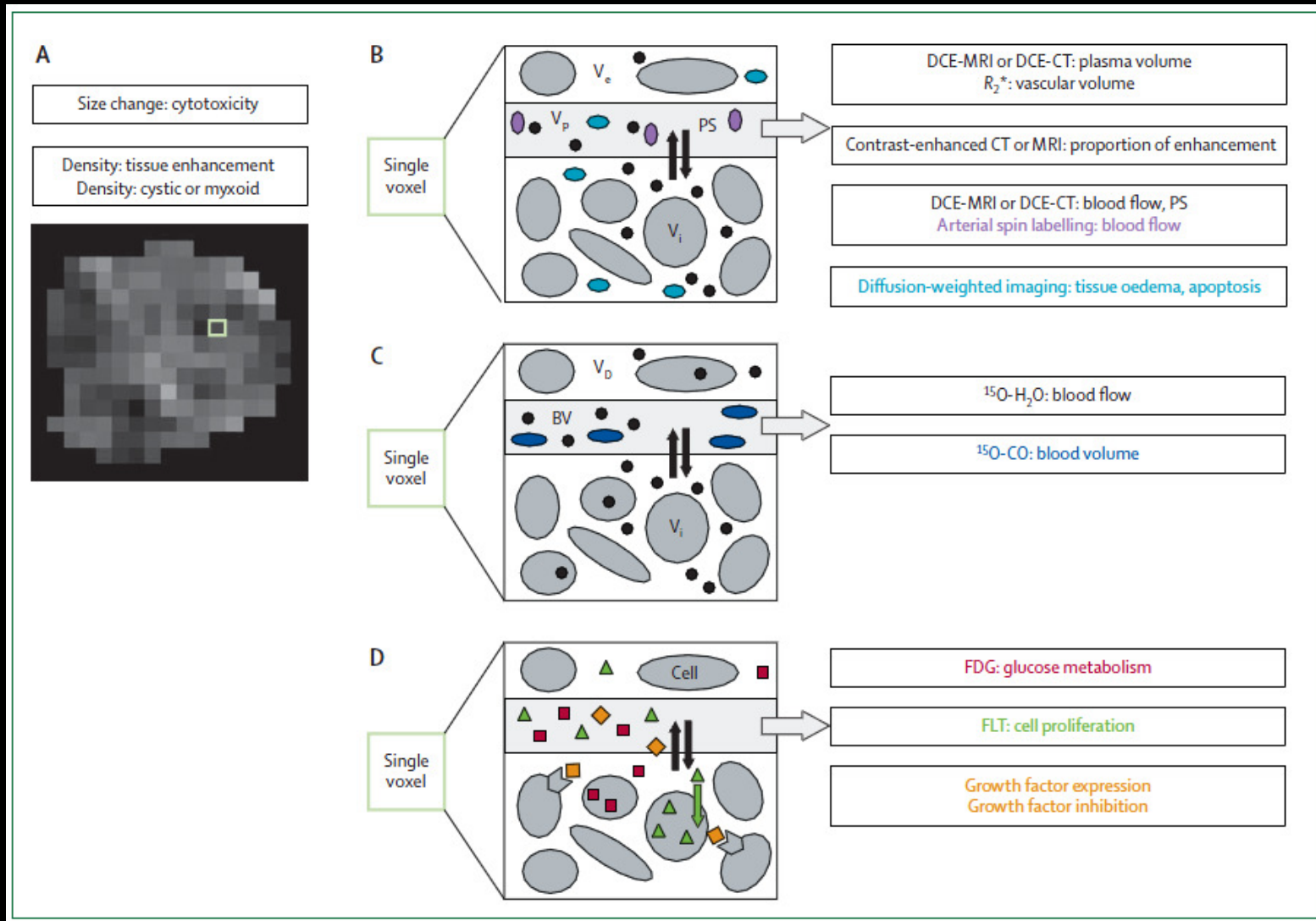
Only histogram moments analysis of parametric maps prognostic



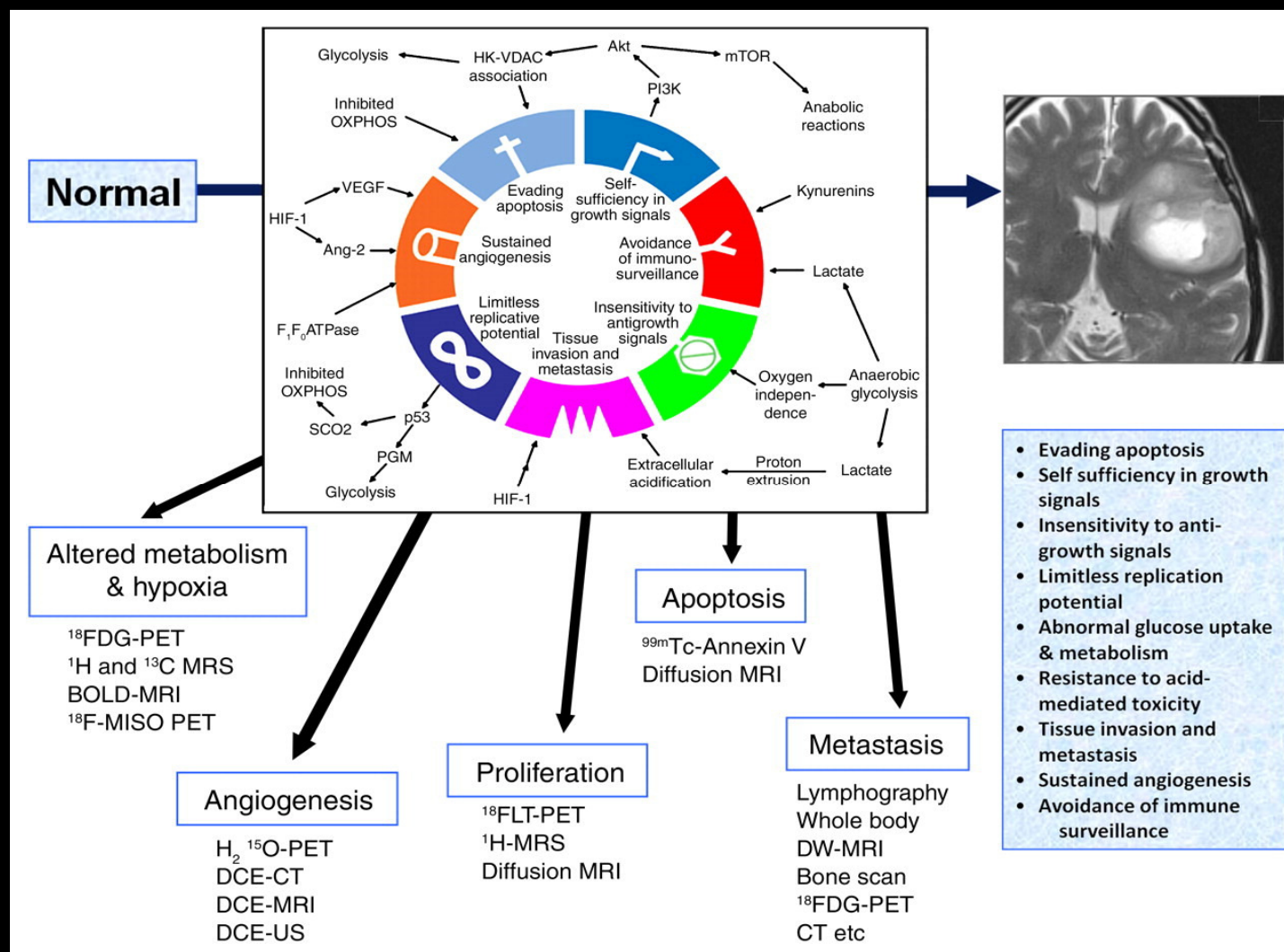
## 2. Functional Imaging: Requirements

- High spatial resolution
- High temporal resolution
- Clinical convenience
- Non-invasive (single or no bolus)
- (Direct) Quantification
- Biologically relevant surrogate
- Meaningful parametric model

## 2. Functional Imaging: Overview



## 2. Frontiers: Multi-modal Imaging



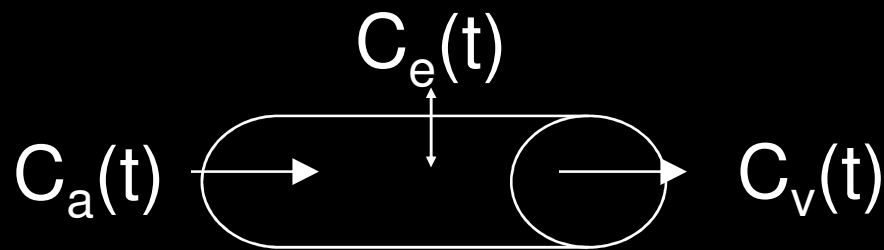
## 2. Functional Imaging: DCE Imaging

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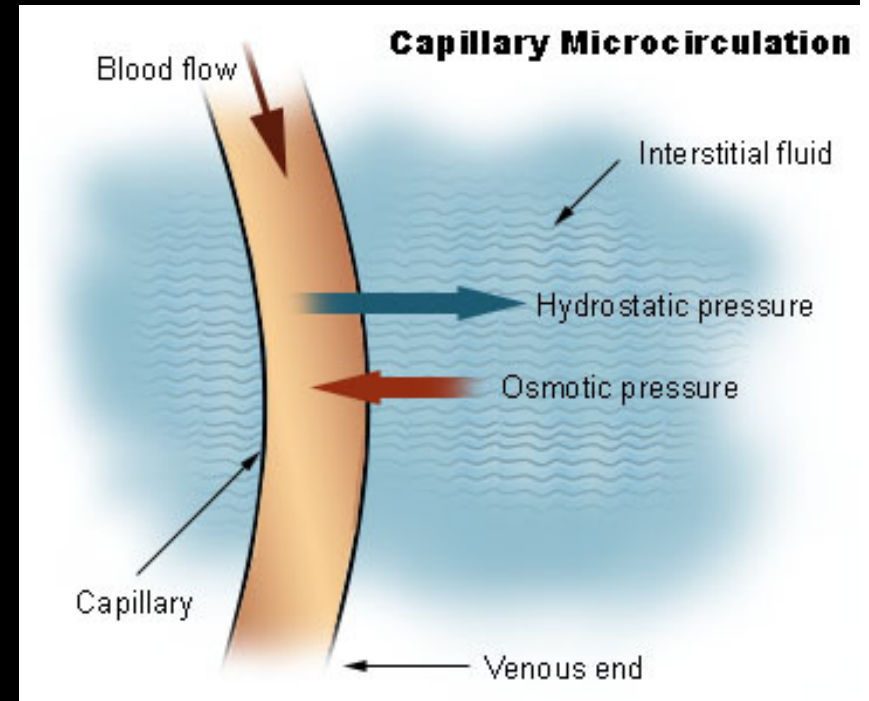
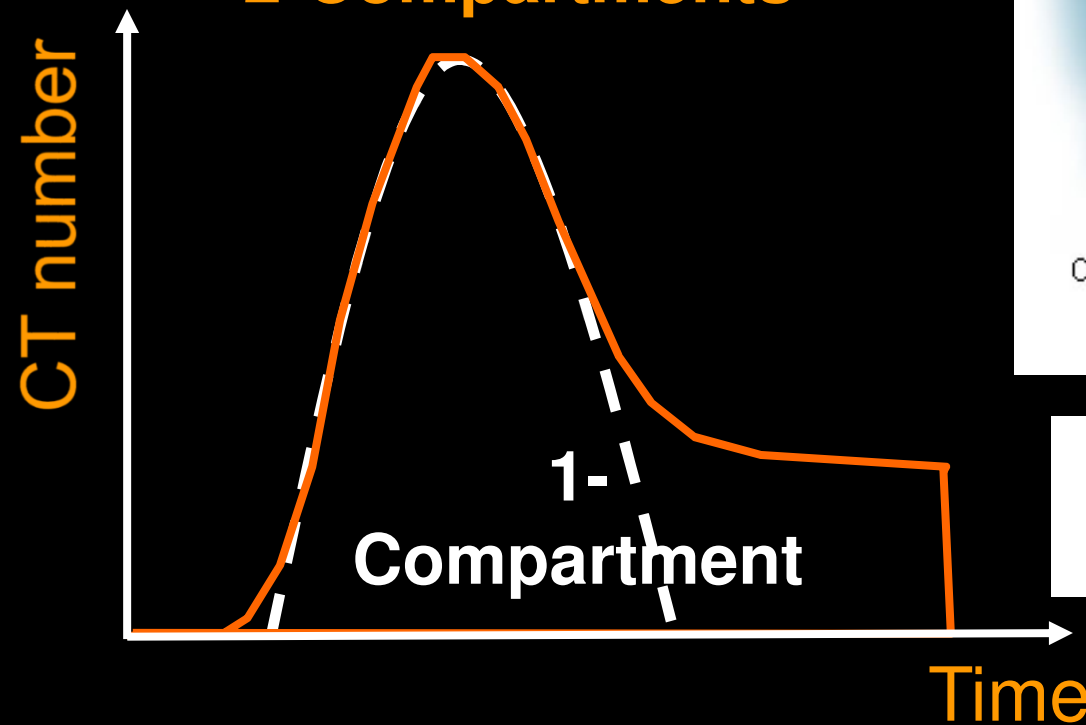


AAPM Webinar Feb 2019

## 2. Functional Imaging: Kinetic modeling



### 2-Compartment

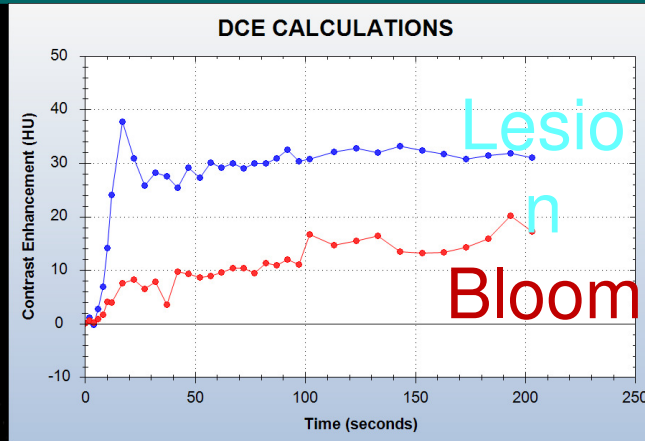
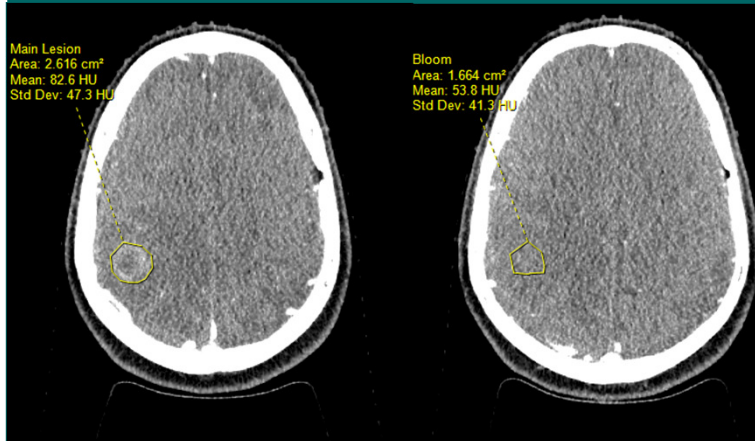


$$C_{\text{tiss}}(t) = \frac{\rho F}{1 - Hct} \int_0^t C_a(t - \tau) R(\tau) d\tau,$$

Tofts et al 1999 JMRI

## 2. DCE Imaging: Iodine vs Gd-DTPA

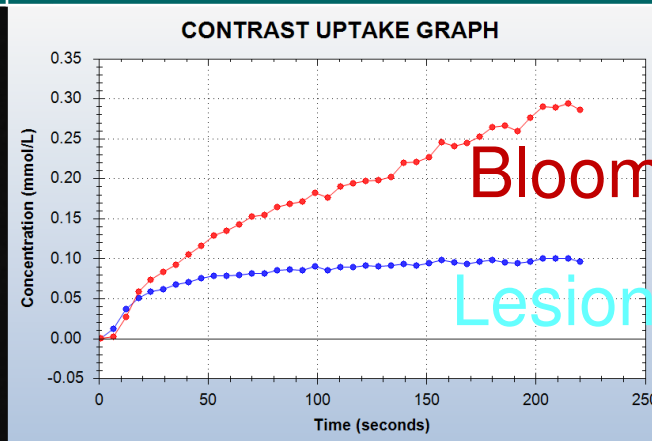
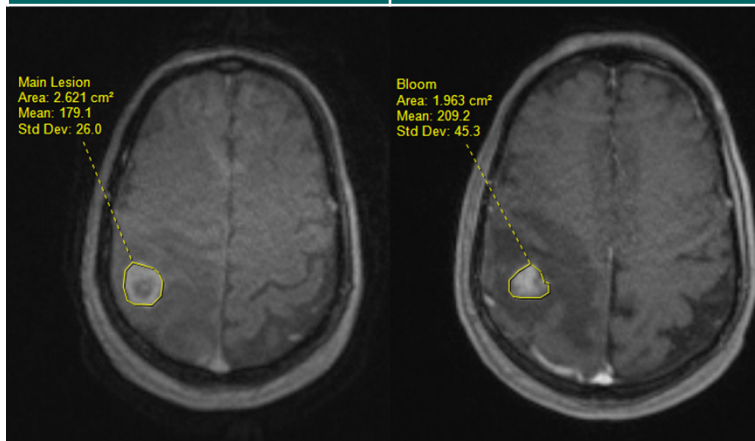
- Main Lesion has stayed roughly the same volume
- Contrast Bloom evident in MR but not CT



Lesion

Contrast Bloom

Time Enhancement Curve



Visipaque

- 320 mgI/mL
- MW **1550.2**

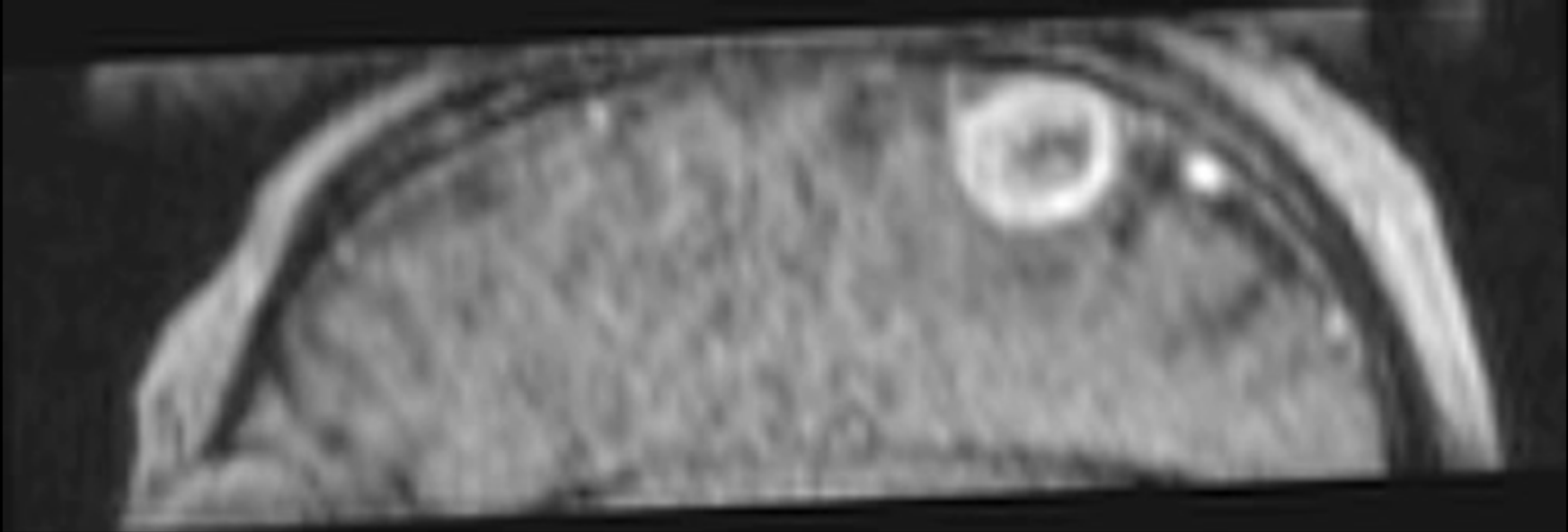
Gadovist

- 1.0 mmol Gd/mL
- MW **604.7**



## 2. Functional Imaging: Transport Flow

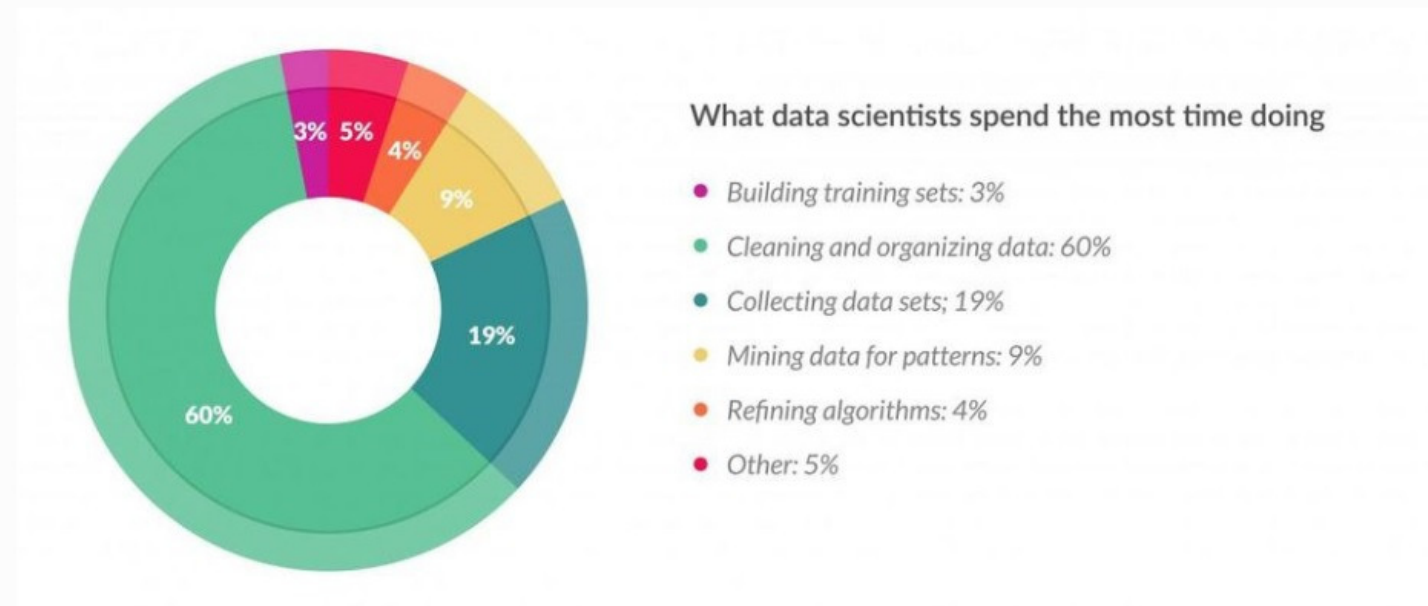
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Case Study (DCE-MRI) courtesy of T. Hompland

# III. Robustness and Standardization

*Data preparation accounts for about 80% of the work of data scientists*



Data scientists spend 60% of their time on cleaning and organizing data. Collecting data sets comes second at 19% of their time, meaning data scientists spend around 80% of their time on preparing and managing data for analysis.

# Functional Imaging Validation Framework

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- Standardization of image acquisition and analysis
- Correlation with outcome
- Correlation with pathology or tissue biomarkers
- Unified Transport modeling

# MULT-SITE DCE CHALLENGE: A CLUSTER FAILURE

## SCIENTIFIC REPORTS

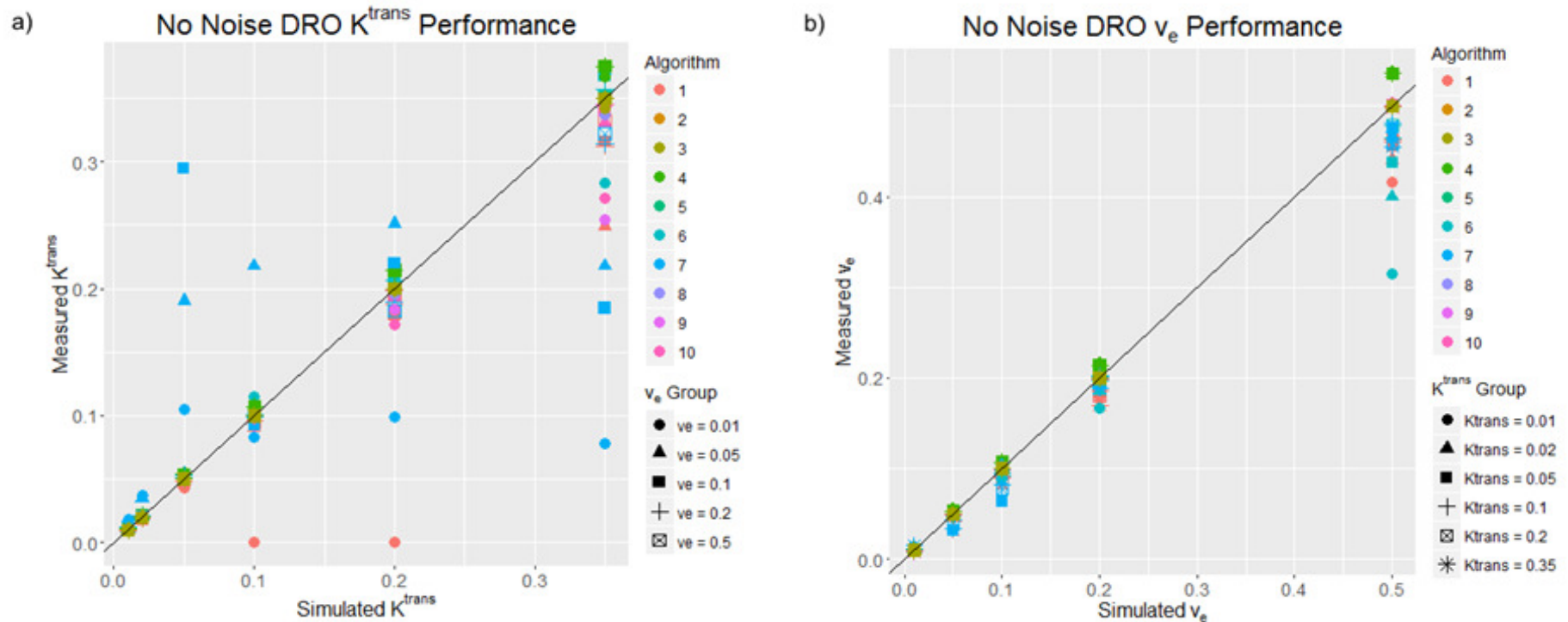
OPEN

### A Multi-Institutional Comparison of Dynamic Contrast-Enhanced Magnetic Resonance Imaging Parameter Calculations

1: 5 May 2017  
1: 18 August 2017

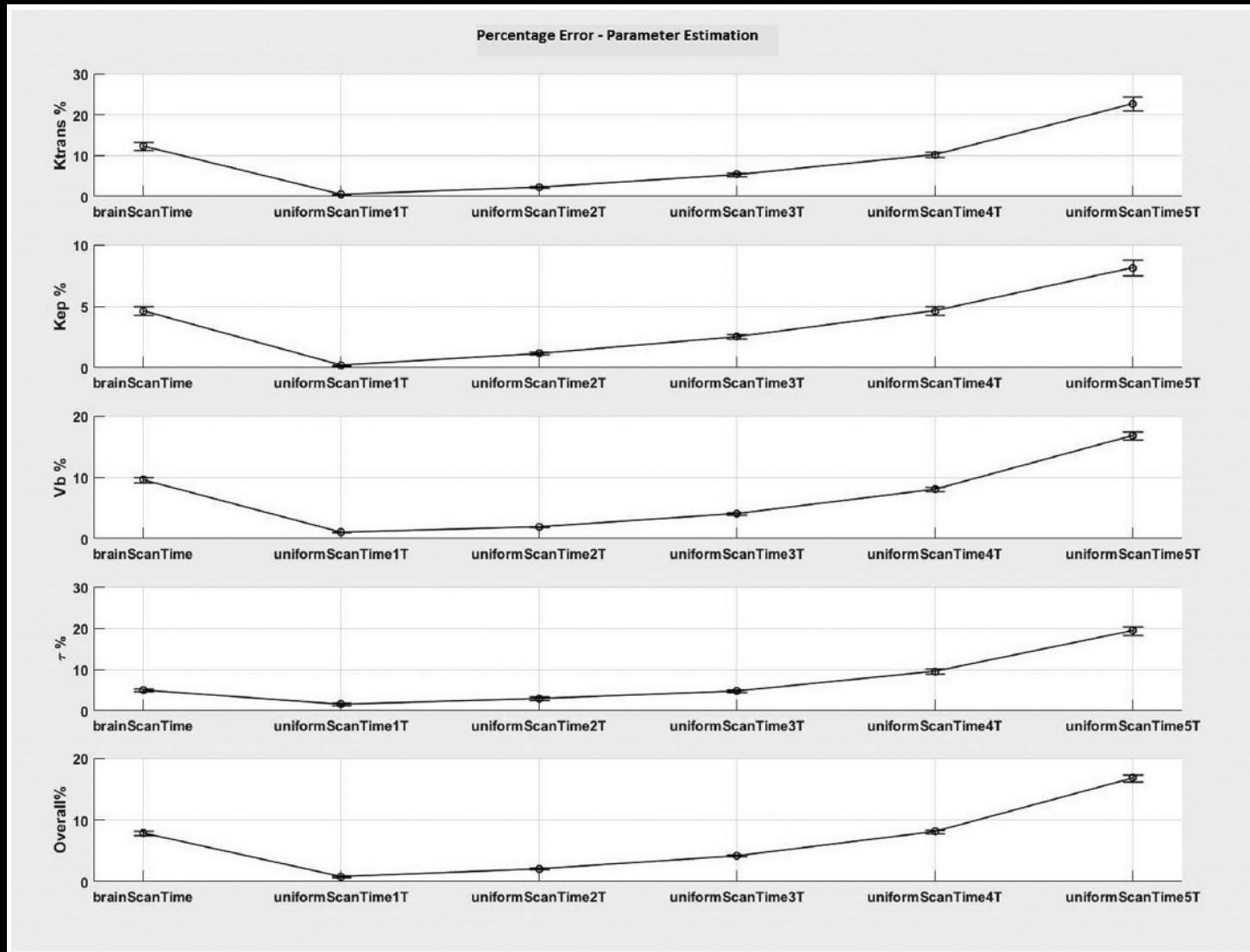
Rachel B. Ger<sup>1,2</sup>, Abdallah S. R. Mohamed<sup>3,4</sup>, Musaddiq J. Awan<sup>5,6</sup>, Yao Ding<sup>7</sup>, Kimberly Li<sup>8,9</sup>, Xenia J. Fave<sup>1,2</sup>, Andrew L. Beers<sup>10</sup>, Brandon Driscoll<sup>11</sup>, Hesham Elhalawani<sup>3</sup>, David A. Hormuth II<sup>12</sup>, Petra J. van Houdt<sup>13</sup>, Renjie He<sup>14</sup>, Shouhao Zhou<sup>15</sup>, Kelsey B. Mathieu<sup>7</sup>, Heng Li<sup>1,2</sup>, Catherine Coolens<sup>11,16,17</sup>, Caroline Chung<sup>3,11</sup>, James A. Bankson<sup>2,7</sup>, Wei Huang<sup>8</sup>, Jihong Wang<sup>1,2</sup>, Vlad C. Sandulache<sup>18</sup>, Stephen Y. Lai<sup>19,20</sup>, Rebecca M. Howell<sup>1,2</sup>, R. Jason Stafford<sup>2,7</sup>, Thomas E. Yankeelov<sup>12</sup>, Uulke A. van der Heide<sup>13</sup>, Steven J. Frank<sup>3</sup>, Daniel P. Barboriak<sup>21</sup>, John D. Hazle<sup>2,7</sup>, Laurence E. Court<sup>1,2,7</sup>, Jayashree Kalpathy-Cramer<sup>10</sup> & Clifton D. Fuller<sup>2,3</sup>

# Variation in modeling implementation



Ger et al. 2017. A Multi-Institutional Comparison of Dynamic Contrast-Enhanced Magnetic Resonance Imaging Parameter Calculations. *Scientific Reports*, 7(1), [11185]

# Parametric Sensitivity to Acquisition



Svistoun *et al.* QIN Special Issue. Tomography 5(1), 209-219, 2019.



# DCE and DWI Imaging Validation

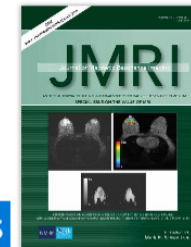


Review Article

## Quantitative imaging biomarkers alliance (QIBA) recommendations improved precision of DWI and DCE-MRI derived biomarkers in multicenter oncology trials

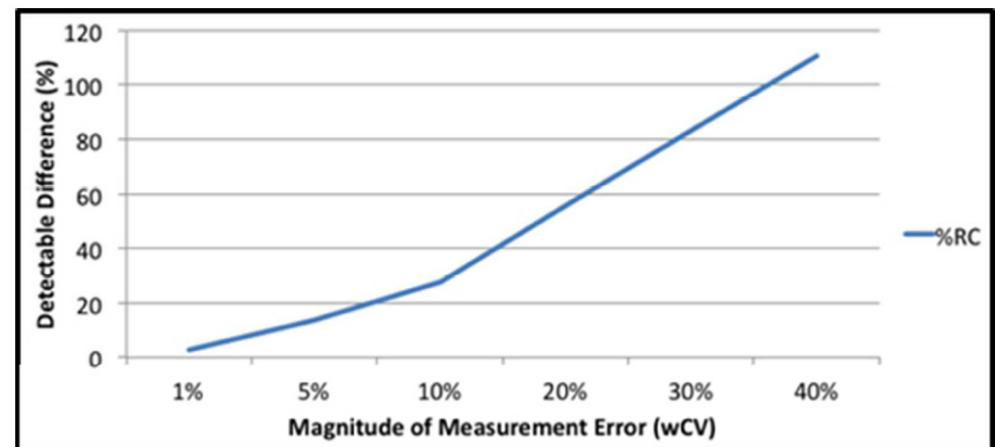
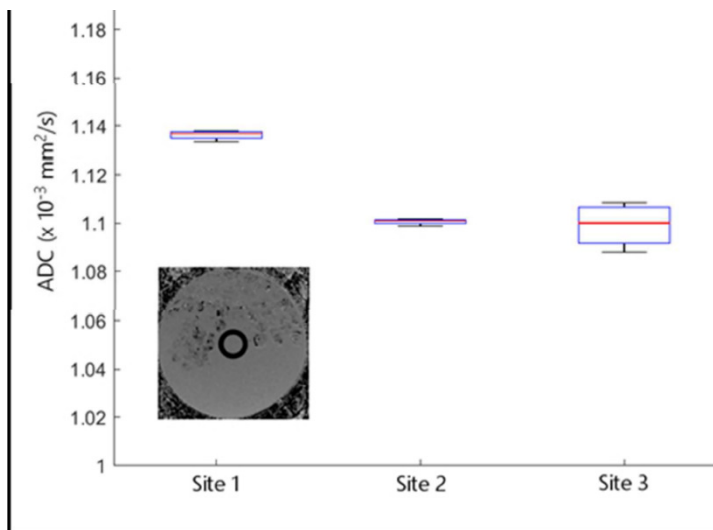
Amita Shukla-Dave PhD, Nancy A. Obuchowski PhD, Thomas L. Chenevert PhD, Sachin Jambawalikar PhD, Lawrence H. Schwartz MD, Dariya Malyarenko PhD, Wei Huang PhD ... [See all authors](#) ▾

First published 19 November 2018 <https://doi.org/10.1002/jmri.26518> Cited by: 5

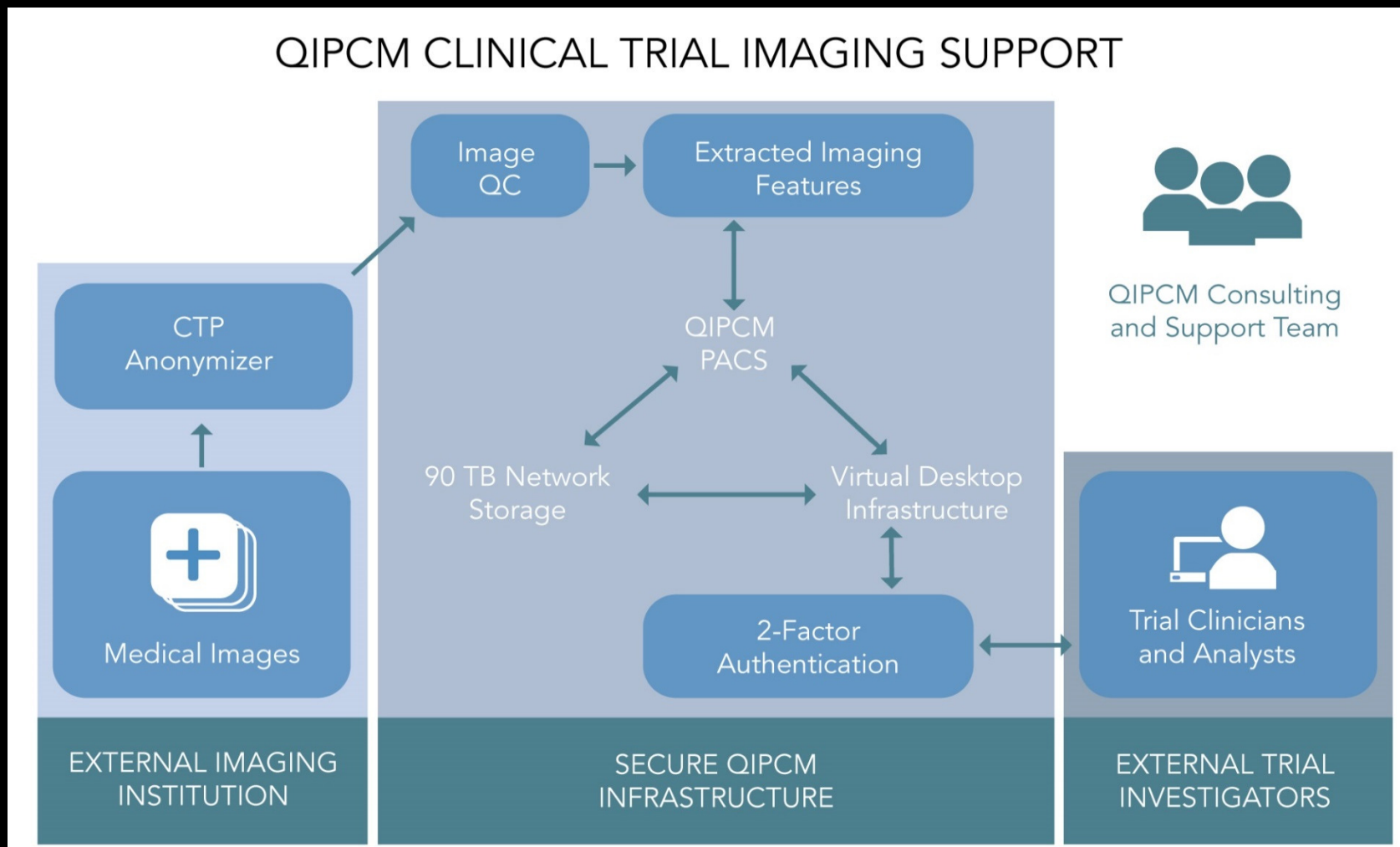


Volume 49, Issue 7  
[Special Issue: Special Issue on the Value of MRI](#)  
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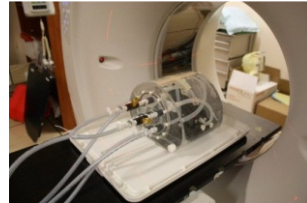
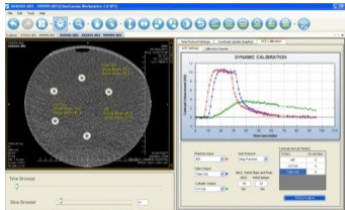


# Imaging Core Lab: Imaging Trial Support



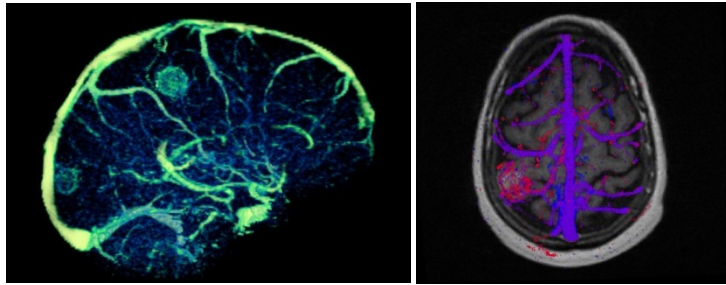
<https://technainstitute.com/qipcm/>

# QIPCM: Standardized Tools



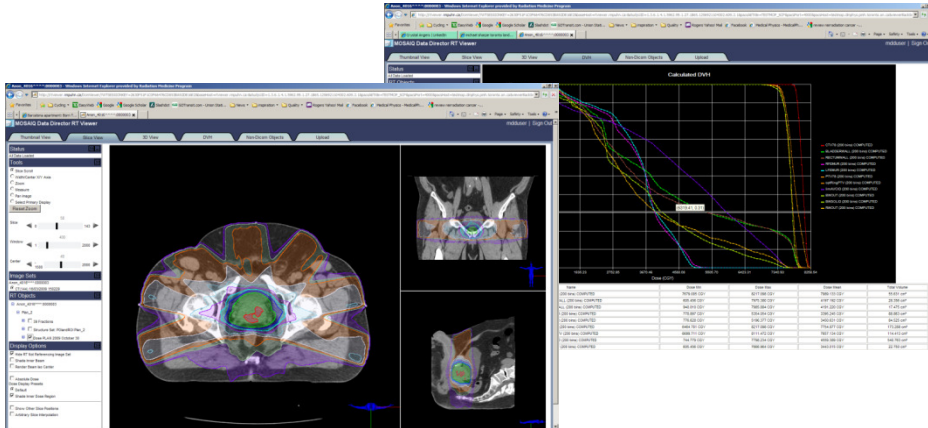
## The DCE QA Tool

- Using a validated dynamic flow phantom perform DCE QA for any CT scanner.



## 4D TDA CT/MRI

- A robust method for automatic 3D vasculature segmentation and unified parametric voxel-based analysis of DCE CT & DCE-MR, DWI scans without the need for manual tissue ROI delineation.



## Web Based RT Viewer

- Browser-based RT-specific viewer for visualization, review and workflow (automated navigation and quality assurance tools)

# Standards & Guidelines: RSNA

## ❖ Quantitative Imaging Biomarker Alliance (QIBA)

### QIBA organization

QIBA has advanced through the generous commitment of volunteer committee members from academia, medical device, pharmaceutical and other business sectors and government.

There are four modality-based Coordinating Committees, including Q-CT, Q-MR, Q-NM and Q-US. Nineteen Biomarker Committees include:

- CT
  - CT Angiography
  - CT Volumetry
  - Lung Density
  - Small Lung Nodule
- MR
  - Arterial Spin Labeling (ASL)
  - Dynamic Contrast-Enhanced (DCE) MRI
  - Dynamic Susceptibility Contrast (DSC) MRI
  - Diffusion Weighted Imaging (DWI) MRI
  - Functional Magnetic Resonance Imaging (fMRI)
  - Magnetic Resonance Elastography (MRE)
  - Proton Density Fat Fraction (PDFF)
  - Musculoskeletal (MSK)
- NM
  - FDG-PET/CT
  - Quantifying Dopamine Transporters with 123Iodine Labeled Ioflupane in Neurodegenerative Diseases (I-123)
  - PET-Amyloid
  - Technetium-99m for body, oncology and immunology (TC99m)
- US
  - Contrast Enhanced Ultrasound (CEUS)
  - Ultrasound Shear Wave Speed (SWS)
  - Ultrasound Volume Blood Flow (VBF)


Each committee has specific responsibilities for its respective modalities or disease-based approach, and is open to interested persons.

View the [QIBA organizational chart](#).




# Standards & Guidelines: NCI

## ❖ Quantitative Imaging for Evaluation of Responses to Cancer Therapies (QIN)

**NATIONAL CANCER INSTITUTE**  
DCTD Division of Cancer Treatment & Diagnosis

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**The next QIN Annual Meeting will be held May 4-5, 2020.**

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Programs & Resources | Quantitative Imaging Network (QIN)

Last Updated: 08/13/19

## Quantitative Imaging Network (QIN)

### About the Quantitative Imaging Network (QIN)

The Quantitative Imaging Network (QIN) promotes research, development and clinical validation of quantitative imaging tools and methods for the measurement or prediction of tumor response to therapies in clinical trial settings, with the overall goal of facilitating clinical decision making. Projects include the development and adaptation/implementation of quantitative imaging methods, imaging protocols, and software solutions/tools (using existing commercial imaging platforms and instrumentation) and application of these methods in current and planned clinical therapy trials. The research projects focus on image-derived quantitative measurements of responses to drugs and/or radiation therapy during clinical trials or standard of care. To achieve the **goals of the QIN**, multidisciplinary teams which include oncologists as well as clinical and basic imaging scientists are required. The involvement of industrial partners in the development and adaptation/implementation of quantitative imaging methods to aid cancer therapies are encouraged.





# Standards & Guidelines: AAPM

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- ❖ MRI - Molecular Imaging in
  - TG 294: Magnetic Resonance Biomarkers in Radiation Oncology
  - TG 284: MRI Simulation in Radiotherapy: Considerations for Clinical Implementation, Optimization, and Quality Assurance



# Summary

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1. Importance of Functional Imaging in Simulation and Treatment Response Assessment
2. Multi-modality Imaging Approach to support complementary Response Assessment describing broader micro-environment
3. Further standardization and validation of functional image acquisition and analysis for clinical trials is needed

# Acknowledgements

## Techna/QIPCM Team

### Medical Physics:

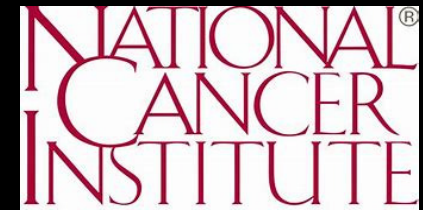
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Stephen Breen  
Jeff Winter  
David Jaffray

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Caroline Chung  
Laura Dawson  
Andrew Hope  
Mike Milosevic  
Kathy Han

### Medical Imaging

Narinder Paul  
Patrick Rogalla  
David Mikulis



# Thank You

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Questions?

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