

Cancer Imaging-Focused Trials in the NCTN: ECOG-ACRIN Examples and Opportunities for Early Career Investigators

**Fred Hutch CTN early career faculty lecture series
August 26, 2026**

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**Matthew J Wilson, Jr Professor and Vice-Chair for Research, Department of Radiology
Associate Director for Education & Training and Breast Program Member, Abramson Cancer Center
University of Pennsylvania**



Penn Medicine
Department of Radiology



Penn Medicine
Abramson Cancer Center

Mankoff Disclosures

(Discussed in this Talk)

- **Consulting fee/advisory board:**
 - **Blue Earth Diagnostics**
- **Ownership Interest/scientific advisory board:**
 - **Trevarx**
- **Other (material transfer)**
 - **Calithera**
 - **Blue Earth Diagnostics**

Cancer Imaging-Focused Trials in the NCTN

- **My background and path NCTN participation and leadership**
- How Does Imaging Fit into the NCTN Portfolio?
- Types of Imaging trials in the NCTN
- Examples of ECOG-ACRIN Imaging Trials and Opportunities for Early Investigators
- Guidance & Future Directions

My Background

- MD/PhD at UPenn, PhD In Bioengineering
 - PhD thesis on applied physics of PET detectors
- Short stint in a start-up – UGM Medical Systems
 - Director of Engineering
- GME Training at UW (1990-1996)
 - Internal Medicine
 - Short-tracked to Nuclear Medicine
 - Research Fellowship in PET cancer imaging – quantitative imaging, PET proliferation imaging, and novel breast cancer response assessment
- Faculty in UW Radiology (1997-2012)
 - Moved to SCCA/FH in 2006 to focus on cancer imaging and thyroid cancer Rx
- Moved to UPenn in 2012
 - Nuclear Medicine Division Chief 2012-2016
 - Vice-Chair for Research 2016 – present
 - Abramson Cancer Center – Cancer Imaging Core Co-Lead; AD Education and Training

My Path to the NCTN

- Research focus on early translation of novel PET probes and methods; Br Ca focus; mechanistic studies (i.e., NIH-fundable)
 - PET Cell proliferation and Breast Cancer ER imaging (NCI P01 Project PI)
 - LABC response to neo-adjuvant systemic therapy – metabolism and blood flow (NCI R01)
 - Internal mammary lymph node mapping and metastasis detection (NCI R01)
 - Br CA bone metastases response (NCI R01)
 - CIP PET Translational Program – led FES PET/CT IND-enabling trial, helped with FMISO and FLT
- Exposure to translational/clinical trials leaders
 - Served as the imaging expert on Cancer Biomarkers Study Section
 - AACR/ASCO Vail Course – imaging expert
- NCTN
 - ACRIN (2007-2012). Inaugural Leader for Experimental Imaging Science Committee (EISC)
 - ECOG-ACRIN (2012-present)
 - Imaging Committee: EIWG Leader (2012-2016)
 - Scientific Planning Committee Co Chair (with Keith Flaherty) (2017-present)
 - Breast Committee senior member (2012-present)
 - National: CTEP Correlative Science Steering Committee (2018 – 2021); NCI CTAC 2014-2019

Cancer Imaging-Focused Trials in the NCTN

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- **How Does Imaging Fit into the NCTN Portfolio?**
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NCTN Structure for Imaging Studies

Group

Imaging Core Lab

(All: Guide Rx Trials; Review
Imaging Components)

Primary Imaging Trials

ECOG-ACRIN

Qualification
Image Analysis
Informatics/AI

Methodology
Detection/Diagnosis
Imaging Biomarkers
Theranostics

NRG

Qualification
Image Analysis
XRT Rx Planning
RP Dosimetry?

Theranostics

Alliance

Qualification
Image Analysis
RP Dosimetry?

Theranostics

SWOG

Qualification
Image Analysis

COG

Qualification
Image Analysis

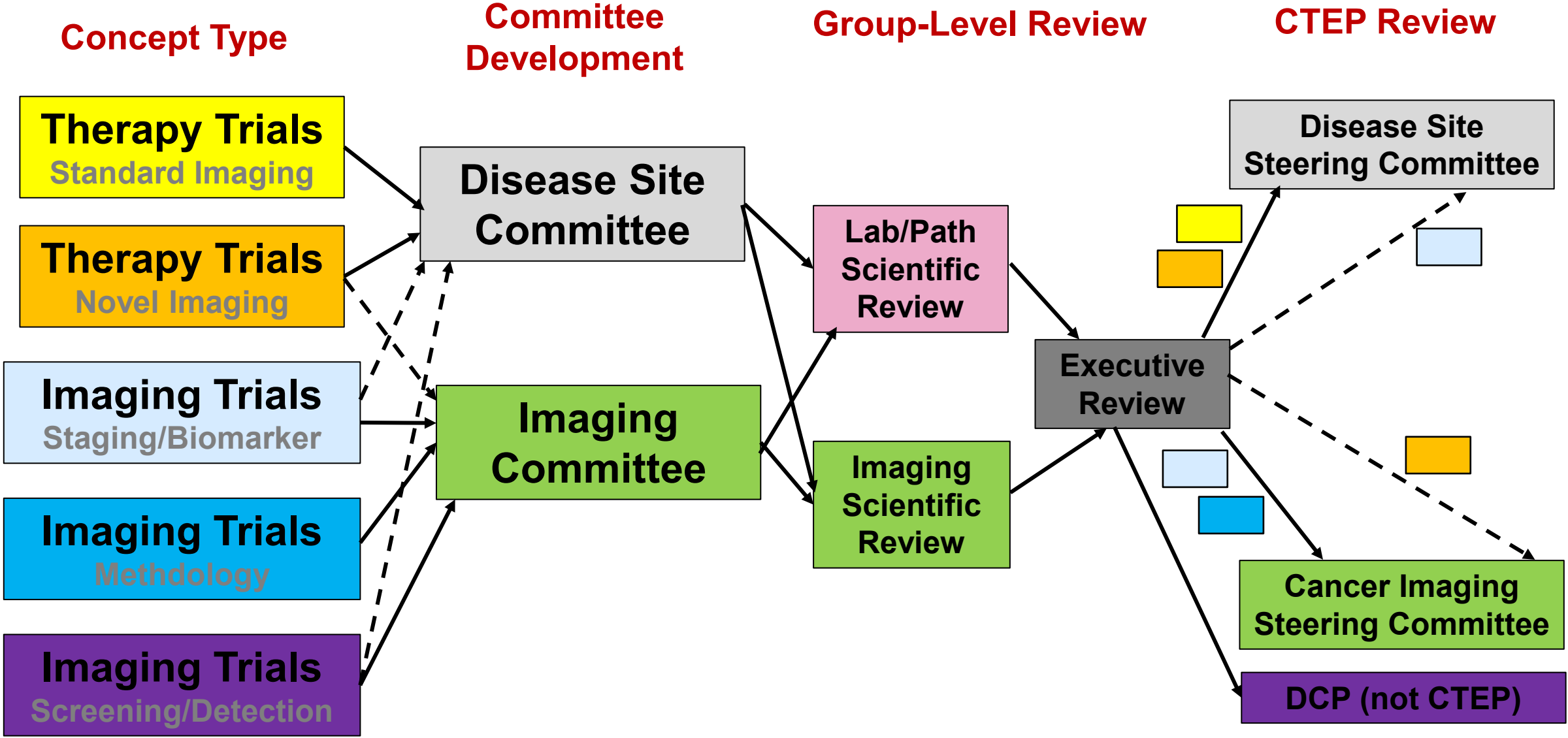
Detection
Biomarkers
Theranostics

IROC (core)

Archiving
QC



Development & Review Pathway For Imaging Studies



Cancer Imaging-Focused Trials in the NCTN

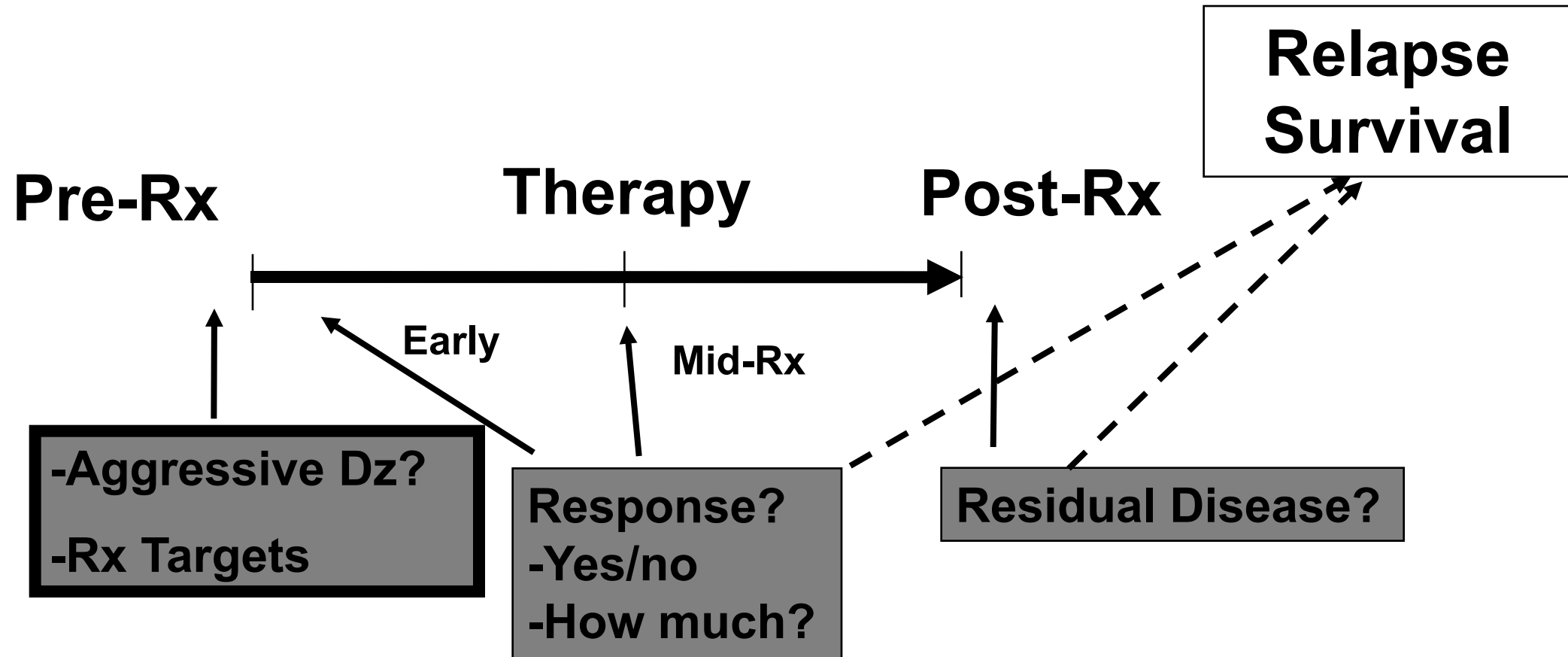
- My background and path NCTN participation and leadership
- How Does Imaging Fit into the NCTN Portfolio?
- Types of Imaging trials in the NCTN
- **Examples of ECOG-ACRIN Imaging Trials and Opportunities for Early Investigators**
- Guidance and Future Directions

Imaging Trials in ACRIN and ECOG-ACRIN

- **ACRIN 6684 - FMISO (hypoxia) PET and DCE/DSC MRI in glioma (Clin Cancer Res paper*; secondary analysis papers*)**
- **ACRIN 6687 - NaF PET to Assess Response to dasatinib (J Nucl Med paper*; secondary analysis papers*)**
- **ACRIN 6688 - FLT PET Early Response to Br Ca Neoadjuvant Therapy (J Nucl Med paper)**
- ACRIN 6691 - Diffuse Optical Spectroscopy Imaging Br Ca Response (Cancer Res paper, secondary paper*)
- LOCATE (ACR core lab) – Impact of Fluciclovone PET on treatment (J Urol paper; secondary papers*)
- EAI141 – FLT PET to measure early AML response (Leuk Lymphoma paper; secondary analysis planned*)
- **EAI142 - FES PET to predict 1st-line ER + MBC response (J Clin Onc in press; secondary analysis planned*)**
- **EA1183 - FDG PET/CT Br Ca bone met response (SABCS oral; primary paper pending; secondary analysis pending*)**
- EA1181 - FDG PET/CT for early HER2-target Rx response (trial design paper published*, accrual completed)
- EA1252 - Randomize Use of FES PET/CT to direct 2nd line ER+ Rx (under development)

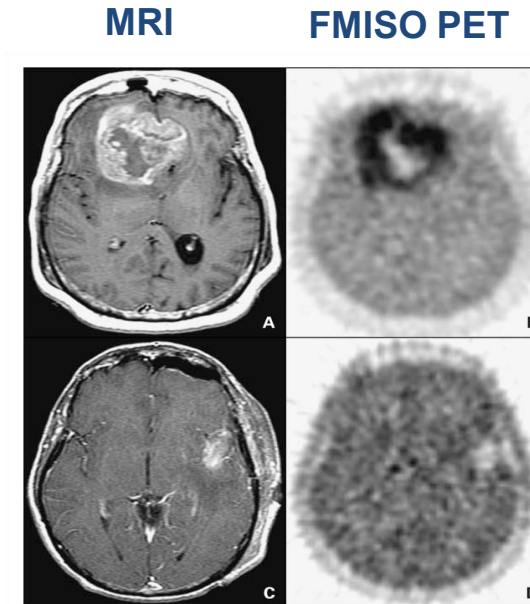
* Junior investigator authorship opportunity

Imaging to Guide Cancer Therapy: Clinical Needs



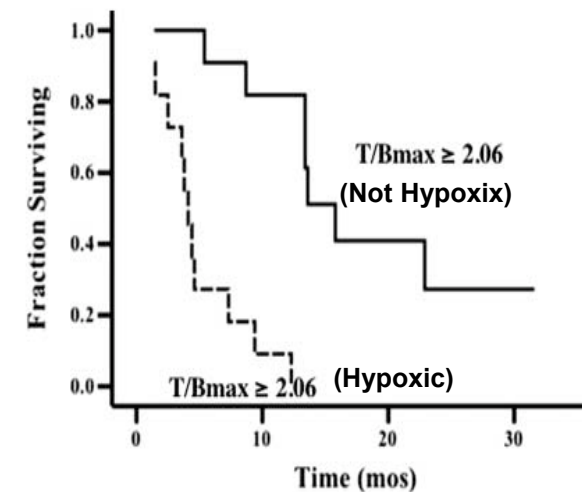
[¹⁸F]FMISO PET Predicts Outcome for GBM Patients

Spence, Clin Cancer Res 14:2623, 2008



Hypoxic

Not
Hypoxic



University of Washington

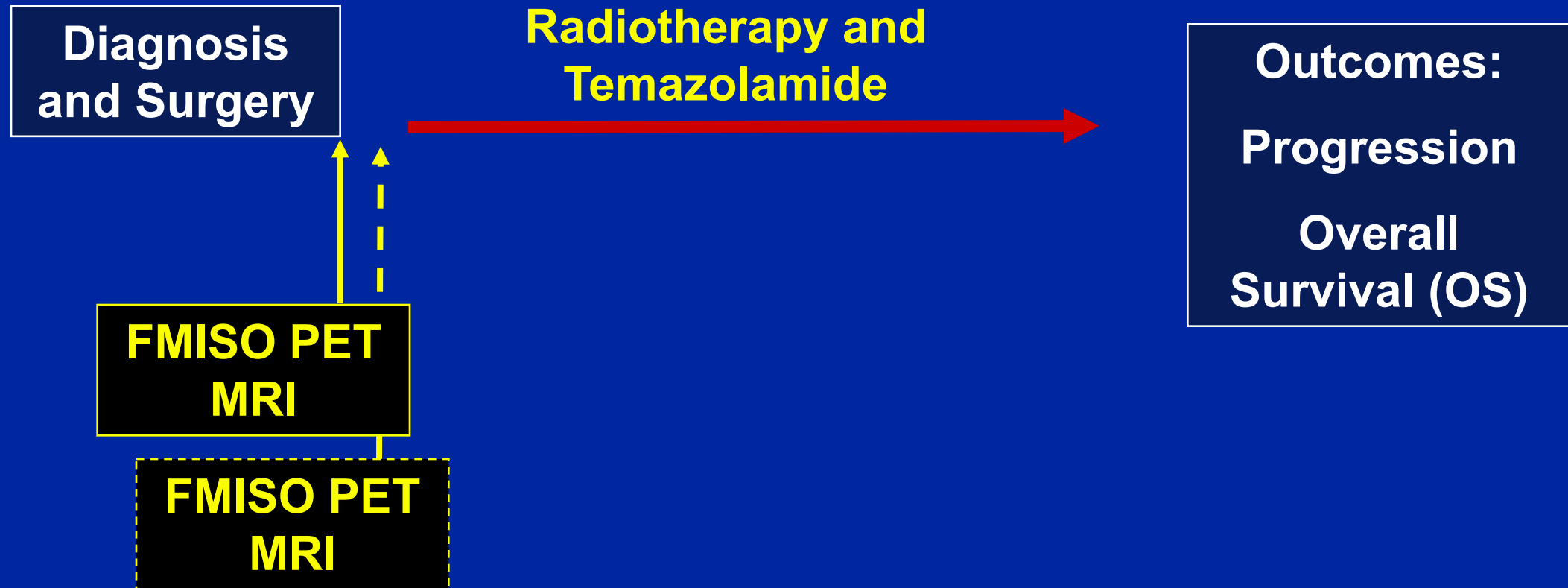


ACRIN 6684

MULTICENTER, PHASE II ASSESSMENT OF TUMOR HYPOXIA IN
GLIOBLASTOMA USING ^{18}F -FLUOROMISONIDAZOLE (FMISO)

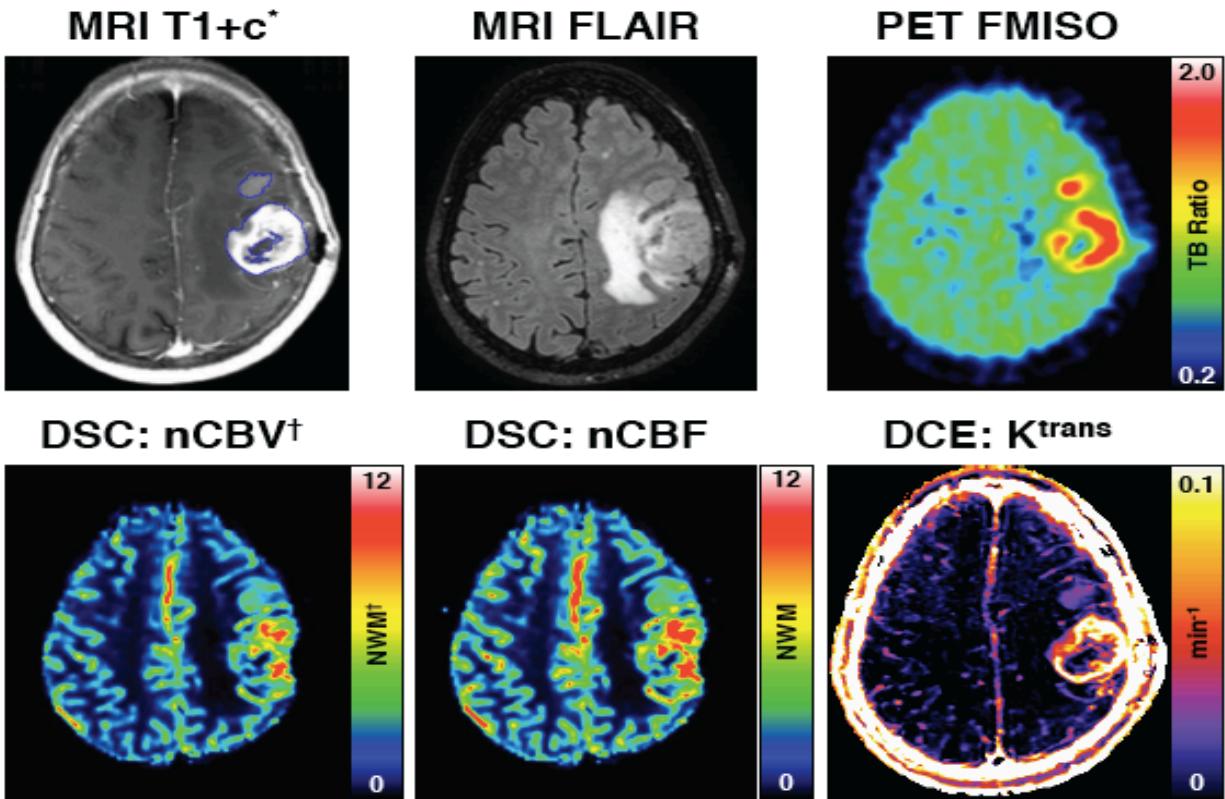
WITH PET AND MRI

Elizabeth Gerstner, MD, PI



ACIN 6684: Hypoxia PET and MRI Predict GBM PFS and OS

Gerstner, Clin Cancer Res, 22:5079, 2016



Personalized Medicine and Imaging

Clinical
Cancer
Research

ACRIN 6684: Assessment of Tumor Hypoxia in Newly Diagnosed Glioblastoma Using ¹⁸F-FMISO PET and MRI

Elizabeth R. Gerstner^{1,2}, Zheng Zhang³, James R. Fink⁴, Mark Muzi⁴, Lucy Hanna⁵, Erin Greco⁶, Melissa Prah⁶, Kathleen M. Schmainda⁶, Akiva Mintz⁶, Lale Kostakoglu⁷, Edward A. Eikman⁸, Benjamin M. Ellingson⁹, Eva-Maria Rataj^{2,10}, A. Gregory Sorensen¹¹, Daniel P. Barboriak¹², David A. Mankoff¹³, and for the ACRIN 6684 Trial Group

Table 3: Cox Regression Model

Parameter	Overall Survival Time			Progression Free Survival		
	Hazard Ratio	95% CI	p-value	Hazard Ratio	95% CI	p-value
<i>Univariate Model</i>						
SUVpeak	1.54	1.00, 2.36	0.048*	1.24	0.80, 1.91	0.33
TBmax	1.16	0.75, 1.81	0.50	0.93	0.61, 1.40	0.72
HV	1.00	0.97, 1.03	0.80	1.01	0.98, 1.04	0.38
Mean k ^{trans} _{tumor}	1.17	1.02, 1.34	0.024*	1.10	0.99, 1.23	0.074
Median k ^{trans} _{tumor}	1.32	1.01, 1.72	0.045*	1.30	1.04, 1.63	0.021*
nRCBV	1.11	0.90, 1.37	0.31	1.28	1.06, 1.54	0.0096*
nCBF	1.07	0.88, 1.29	0.51	1.18	1.01, 1.38	0.038*

ACRIN 6684 Secondary Papers

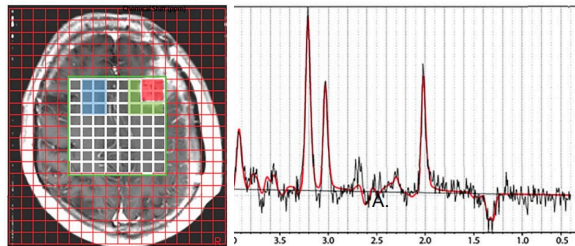
Planned Secondary Aim



RESEARCH ARTICLE

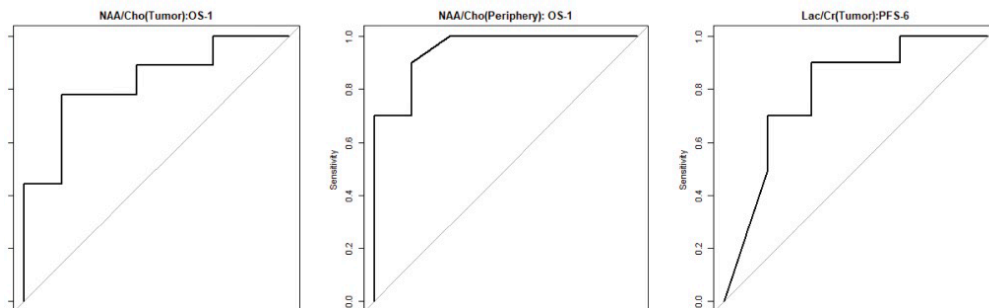
ACRIN 6684: Multicenter, phase II assessment of tumor hypoxia in newly diagnosed glioblastoma using magnetic resonance spectroscopy

Eva-Maria Ratai^{1,2*}, Zheng Zhang³, James Fink⁴, Mark Muzi⁴, Lucy Hanna³, Erin Greco^{3a}, Todd Richards⁴, Daniel Kim^{1,2}, Ovidiu C. Andronesi^{1,2}, Akiva Mintz^{5a,b}, Lale Kostakoglu⁶, Melissa Prah⁷, Benjamin Ellingson⁸, Kathleen Schmainda⁷, Gregory Sorensen^{1,2a,c}, Daniel Barboriak⁹, David Mankoff¹⁰, Elizabeth R. Gerstner^{2,11}, on behalf of the ACRIN 6684



MRS
Methodology

2⁰ Aim: MRS Accuracy in Predicting 1 year OS



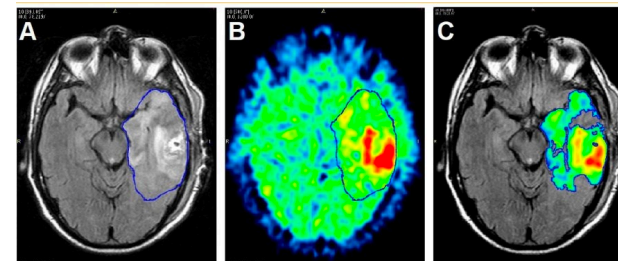
(Ratai, Plus One, June 2018)

Ad-Hoc Analysis: Data Request

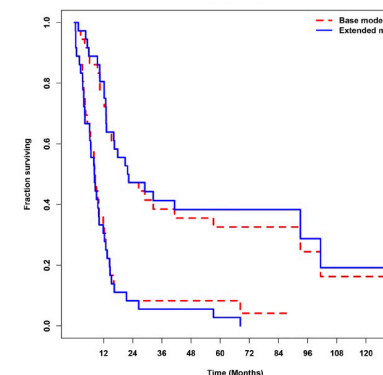
TOMOGRAPHY[®] RESEARCH ARTICLE

Assessment of the Prognostic Value of Radiomic Features in ¹⁸F-FMISO PET Imaging of Hypoxia in Postsurgery Brain Cancer Patients: Secondary Analysis of Imaging Data from a Single-Center Study and the Multicenter ACRIN 6684 Trial

Mark Muzi¹, Eric Wolsztynski^{2,4}, James R. Fink¹, Janet N. O'Sullivan², Finbarr O'Sullivan², Kenneth A. Krohn¹, and David A. Mankoff³



PET Radiomics
Methodology



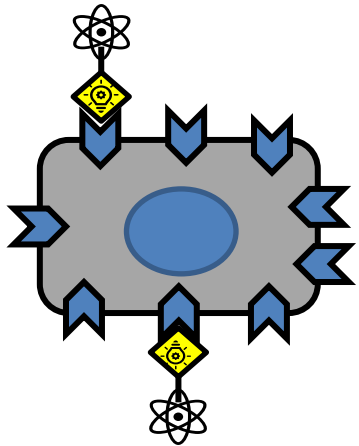
Predicting OS

(Muzii, Tomography, 6:14, 2020)

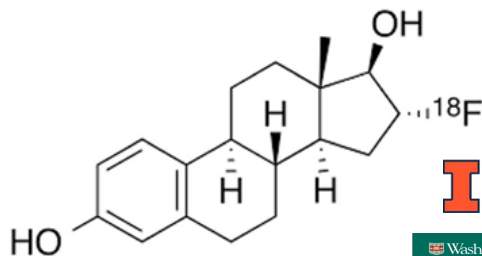
Radiopharmaceutical Diagnostics:

Quantify Target Expression by in vivo Radioligand Binding Assay

Measure Target
Expression



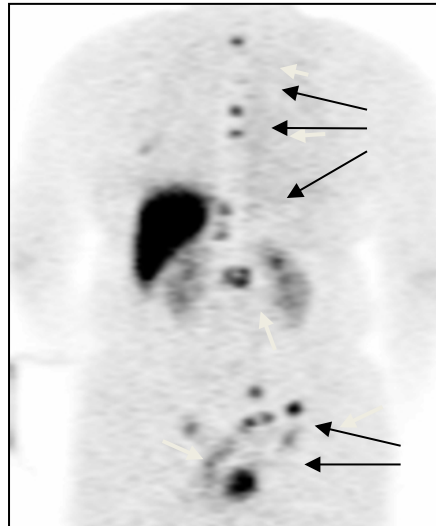
16α -[^{18}F]fluoroestradiolol (FES)



UNIVERSITY OF
ILLINOIS
URBANA-CHAMPAIGN

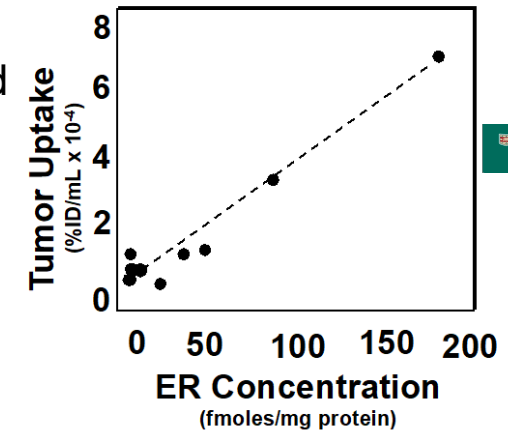
Washington University in St. Louis
SCHOOL OF MEDICINE

Estrogen Receptor PET
(Breast Cancer)



Uptake (SUV) vs Tissue Assay

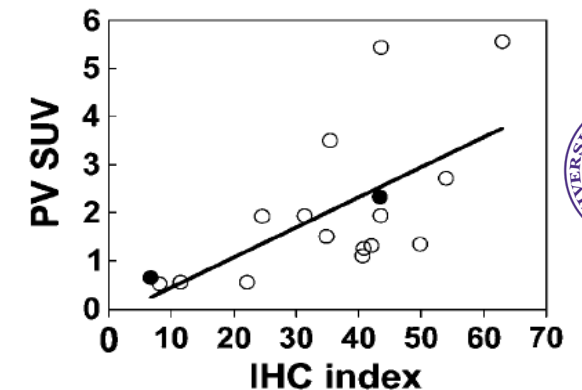
Radioligand
Binding



Washington University in St. Louis
SCHOOL OF MEDICINE

(Mintun, Radiology 169:45, 1988)

Immunohisto-
chemistry

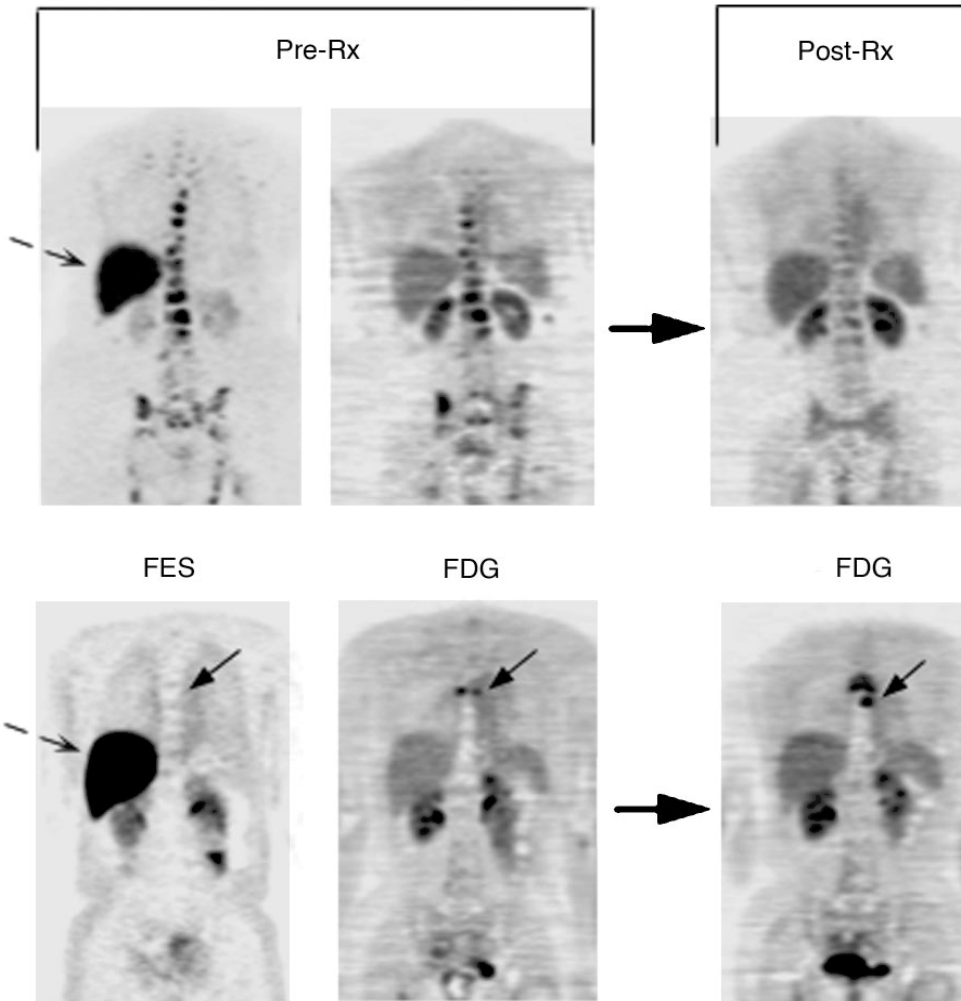


(Peterson, J Nucl Med 49: 367, 2008)

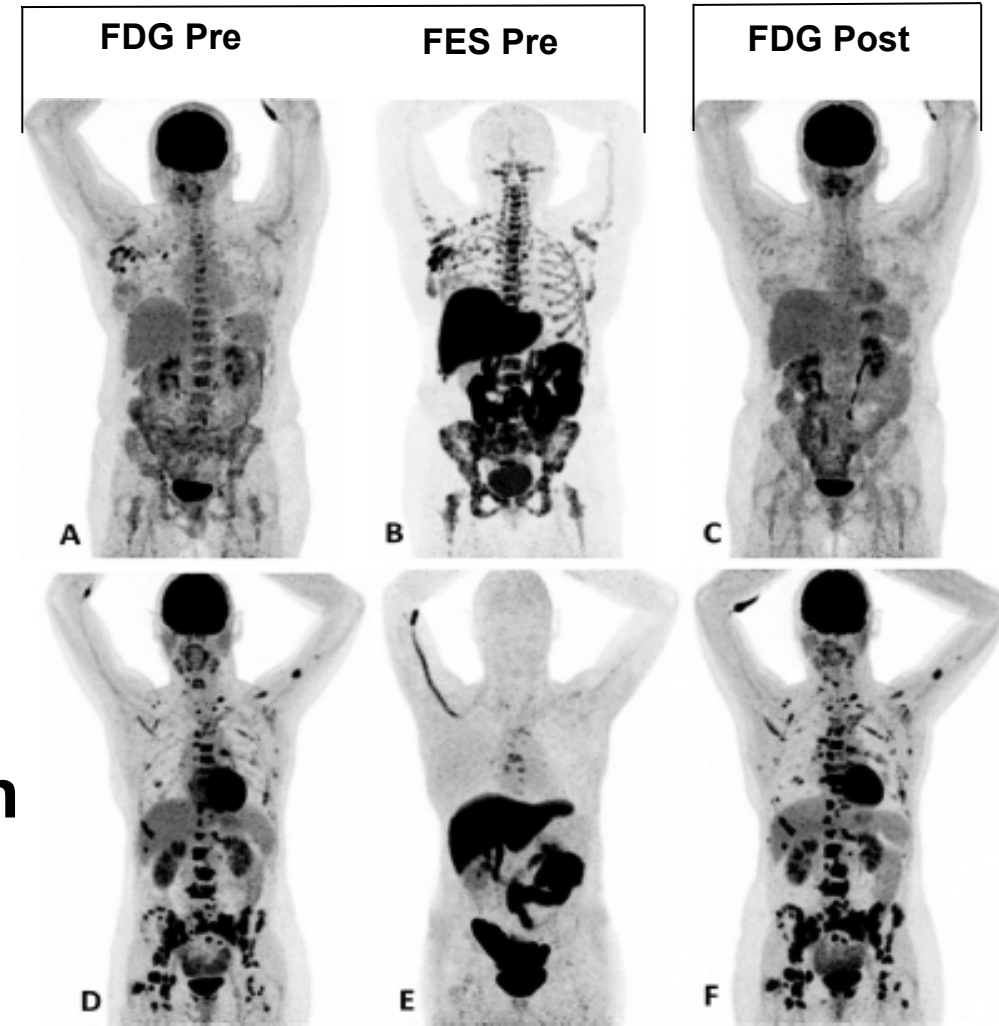
(Katzenellenbogen, Nucl Med Biol 92:24, 2021)

Is the Target Present?

FES Uptake Predicts Breast Cancer Response to Hormonal Therapy



Response



Progression

(Linden, J Clin Onc, 24:2793, 2006)

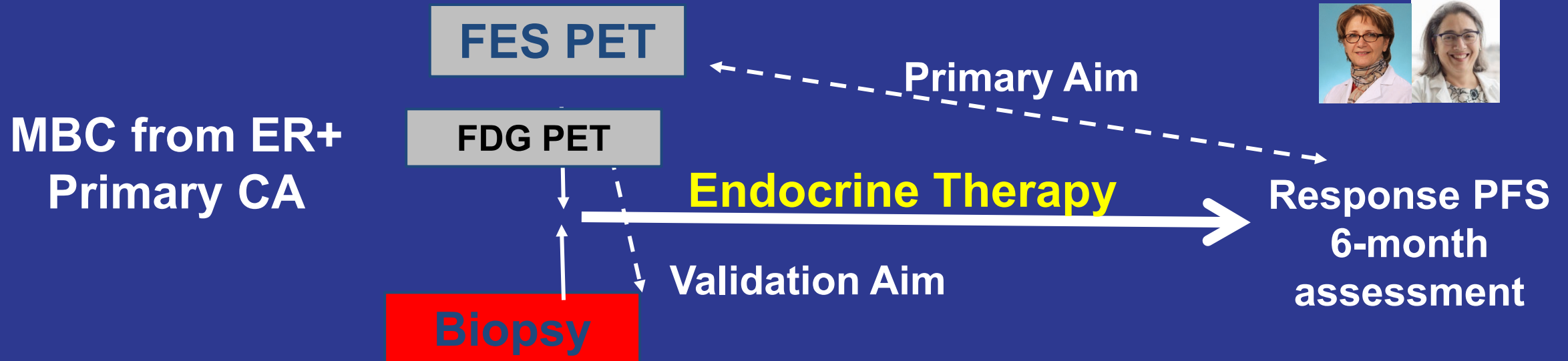


(Boers, Eur J Cancer 126:11e20, 2020)



ECOG-ACRIN Biomarker Trial of FES PET: EAI142

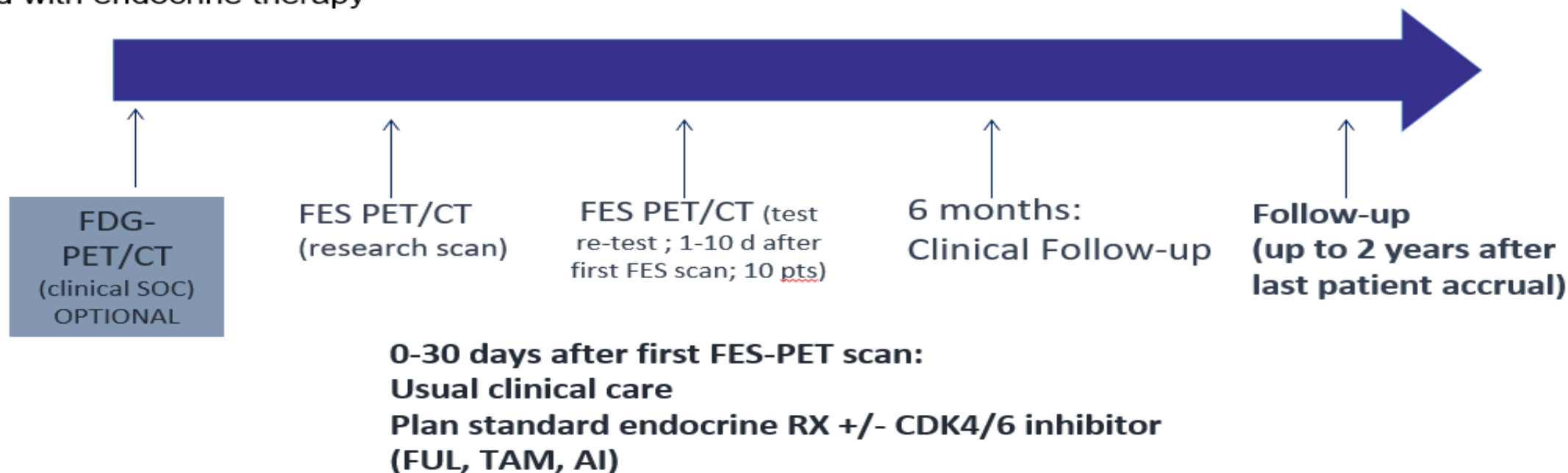
Dehdashti & Linden



- First line therapy
- Stand-alone imaging trial:
 - Clinical indication for endocrine therapy
 - Standard Rx allowed (AI, FUL, TAM)
 - Measurable and non-measurable disease

EAI142 Schema

ER+/HER2- Metastatic tumor
Untreated with endocrine therapy*



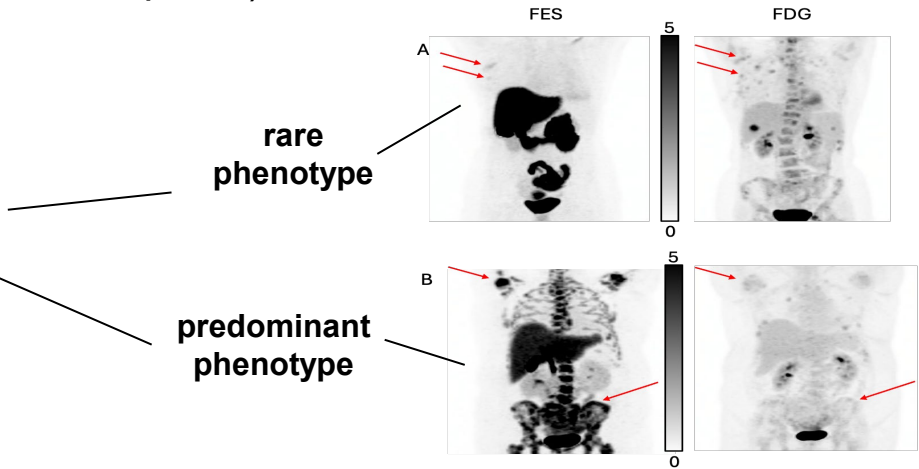
*Adjuvant endocrine therapy allowed, 1 line chemo in metastatic setting allowed,
off blocking therapy X 60 days

[¹⁸F]16α-fluoro-17β-estradiol (FES)-PET as a predictive measure for first-line endocrine therapy in patients with newly diagnosed metastatic breast cancer (EAI142)

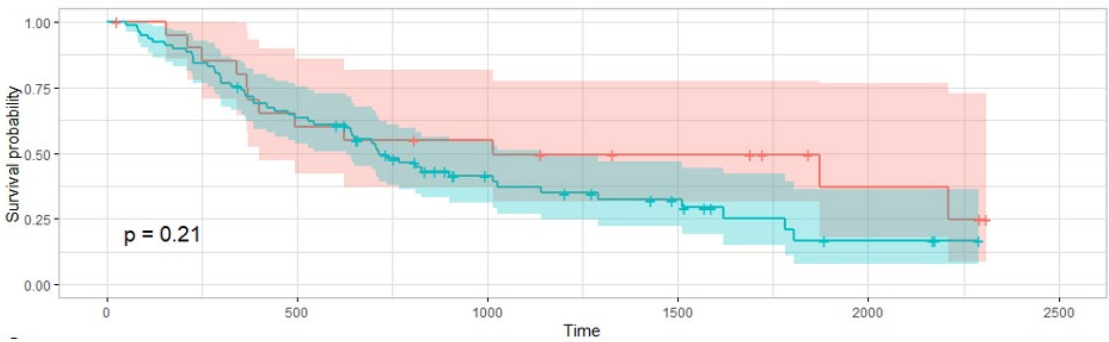
Linden, Dehdashti et al (J Clin Oncol, in press)



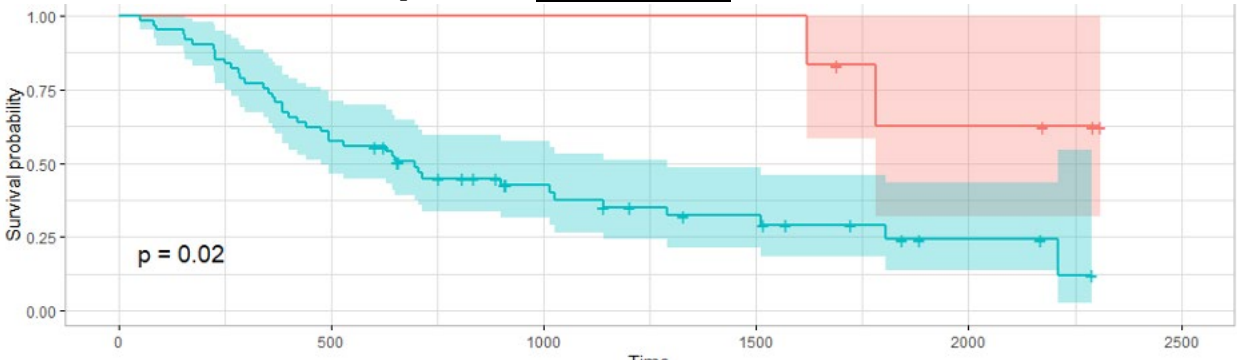
Visual Assessment	Number of Lesions	Number of Patients ¹	Number of Patients ²
No uptake	0 (0%)	19 (19.4%)	1 (1%)
Minimal uptake	3 (0.67%)		
Mild uptake	36 (8%)	79 (80.6%)	97 (99%)
Moderate uptake	84 (18.67%)		
Intense uptake	327 (72.67%)		
Total	450	98	98



FES Uptake Did Not Predict PFS



FDG Uptake Strongly Predicted PFS



Lessons Learned:

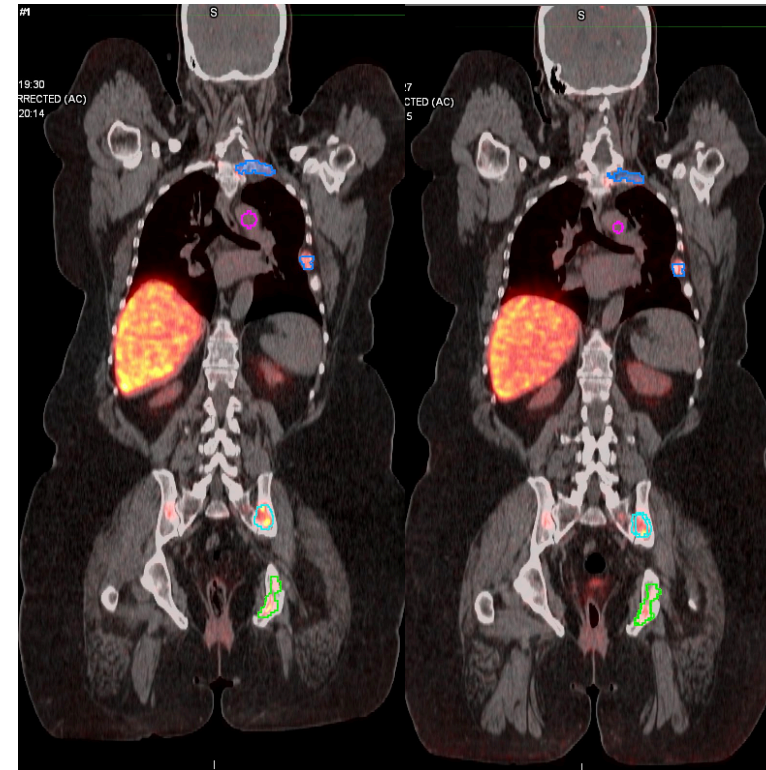
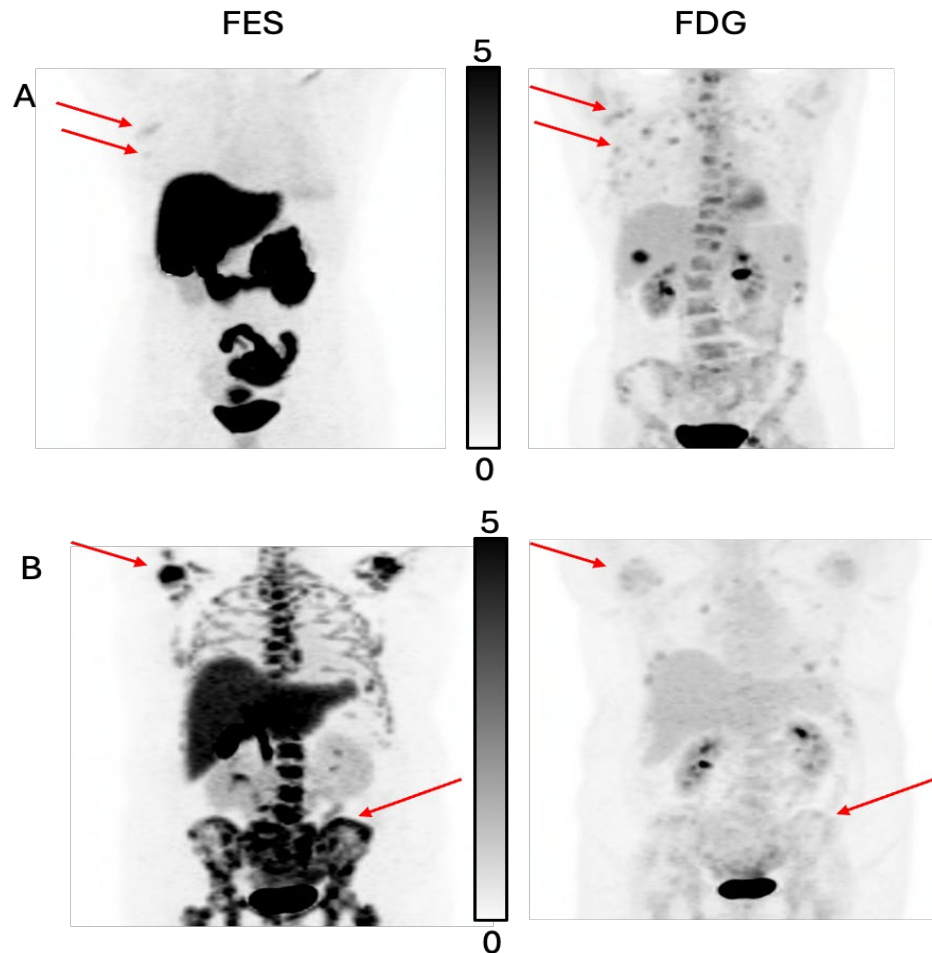
- Prospective, multi-center trials are important!
- Predictive value of imaging biomarkers depends upon patient presentation and treatment stage
- Consistent retention of ER expression in untreated MBC support a role for FES PET/CT in staging

EAI142 Planned Secondary Analysis Papers

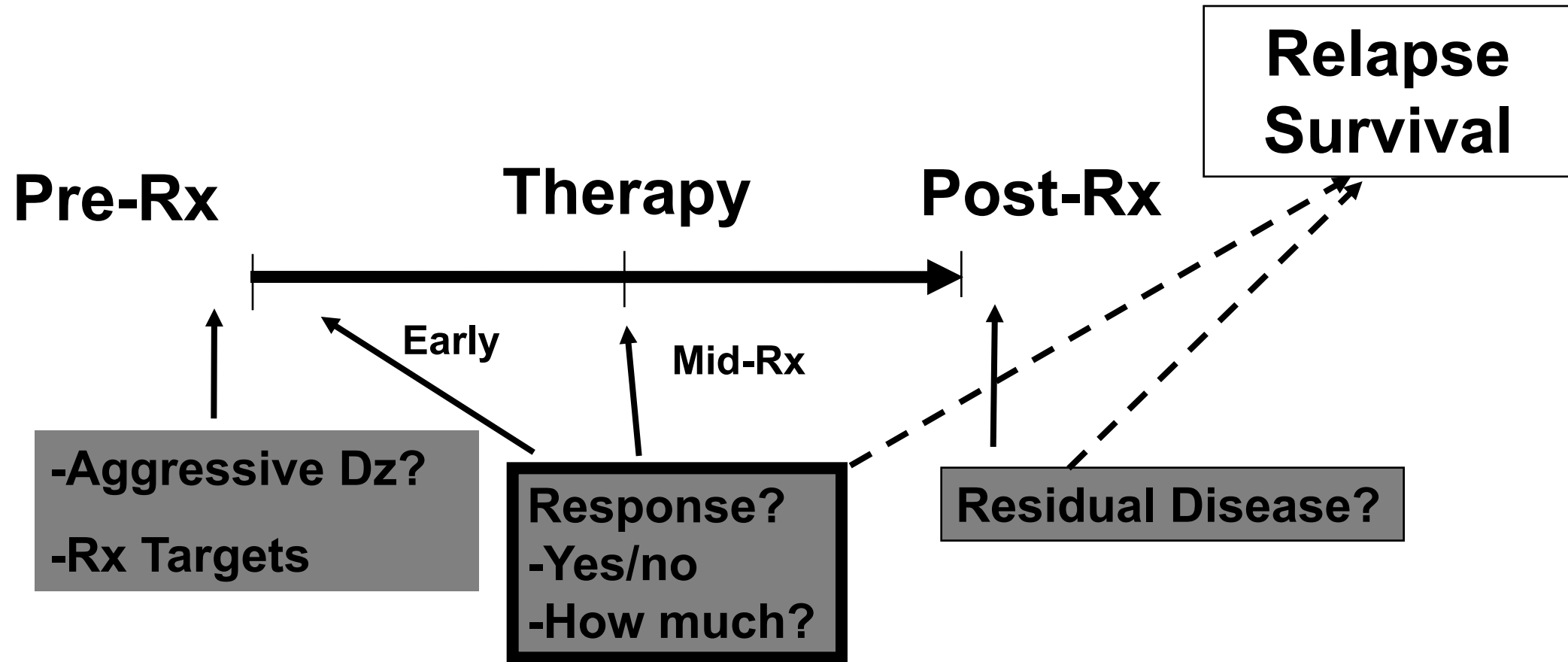


**Comparative Staging of FES vs FDG PET/CT:
Is FES > FDG for ILC?**

**Methodology for Automated, Clinically
Practical Comparative FDG and FES Analysis**

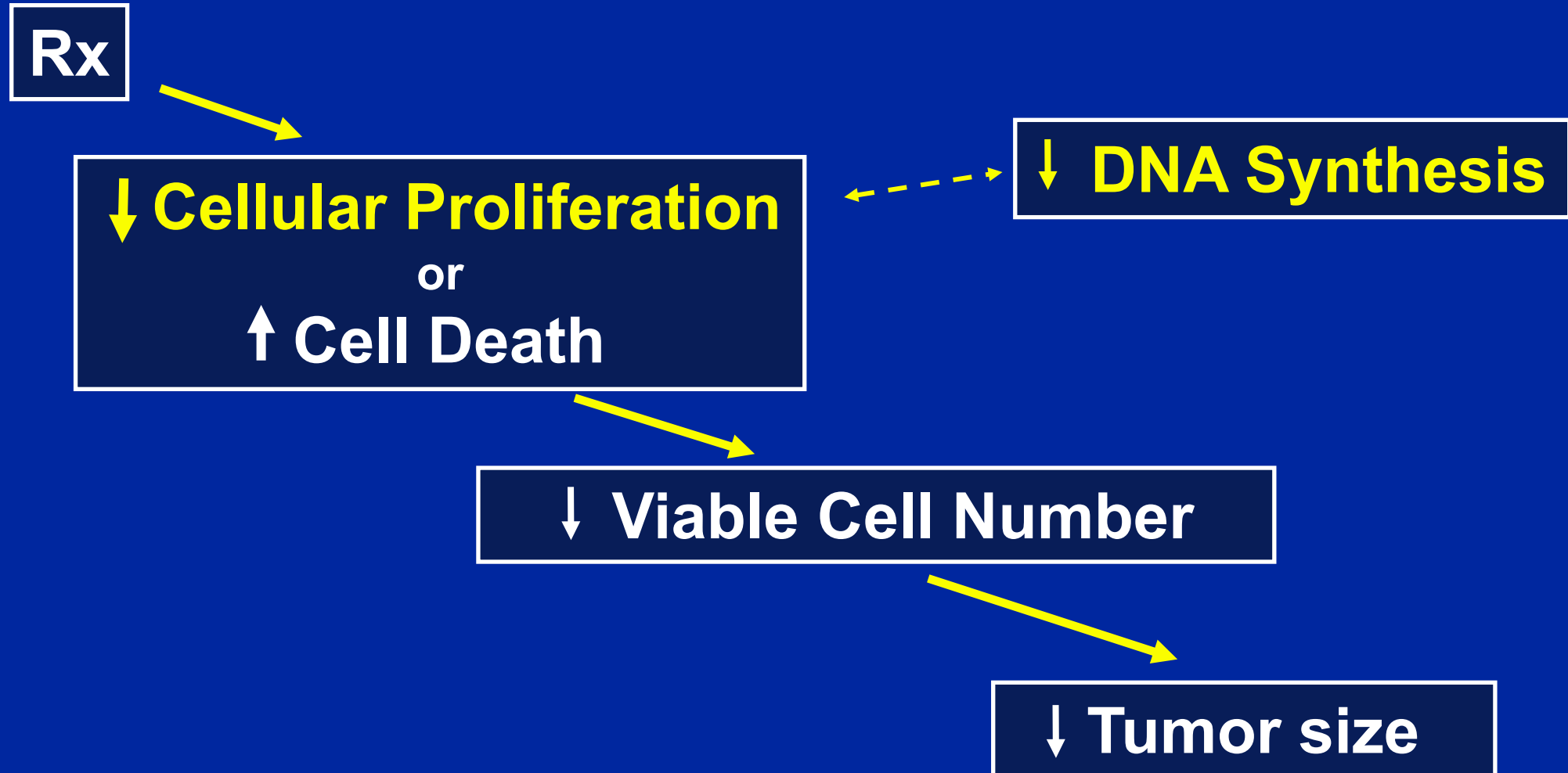


Imaging to Guide Cancer Therapy: Clinical Needs

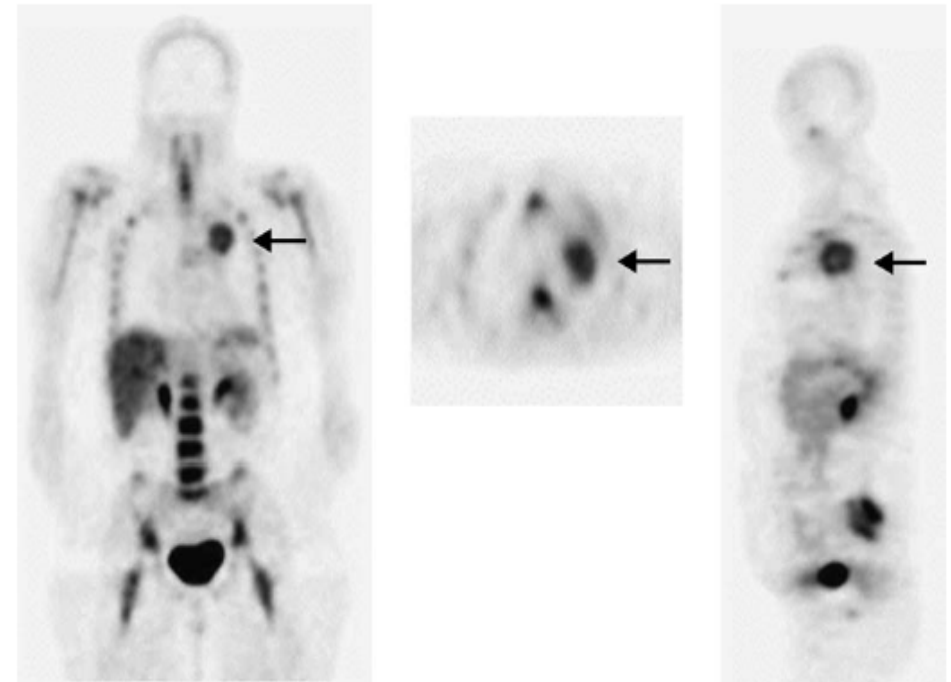
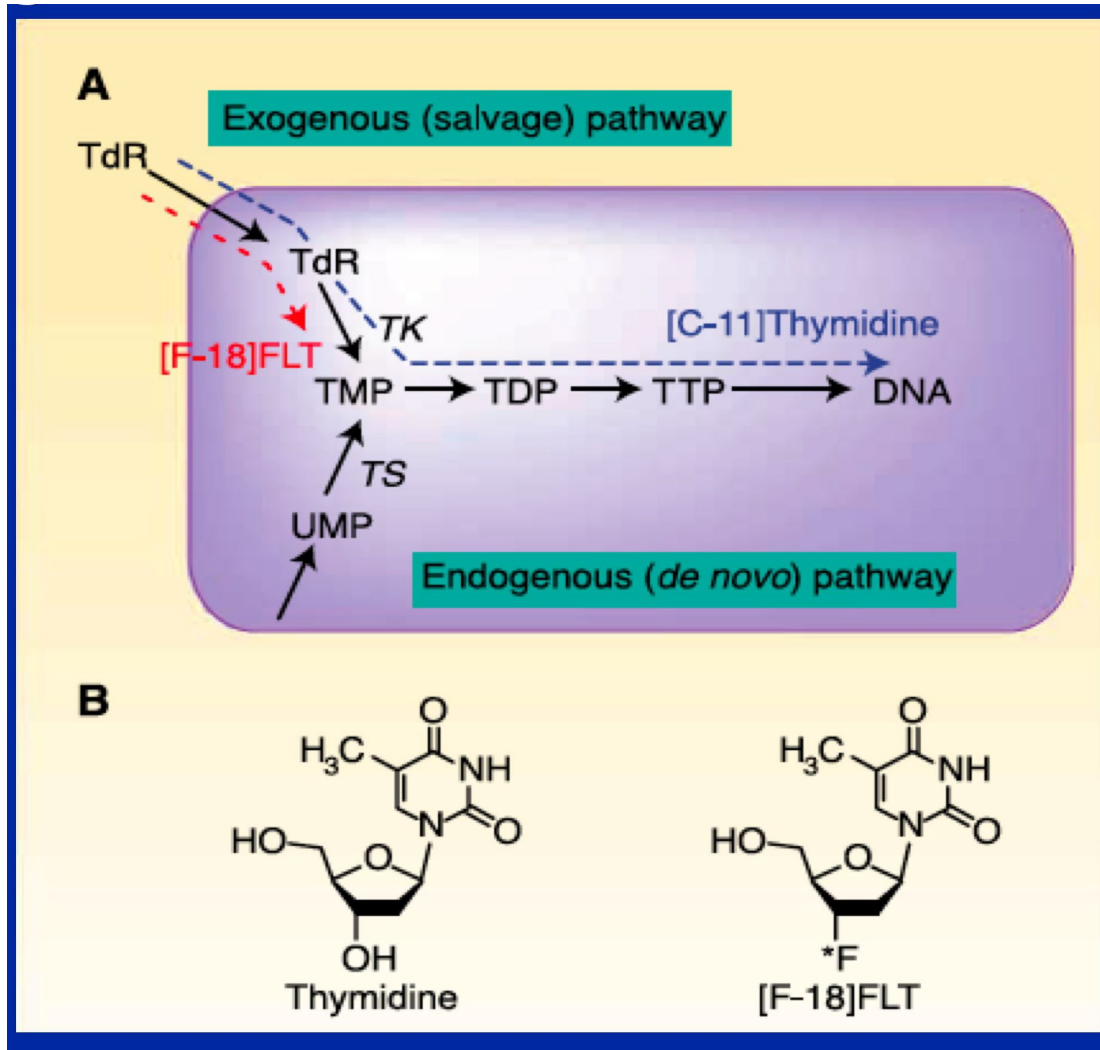
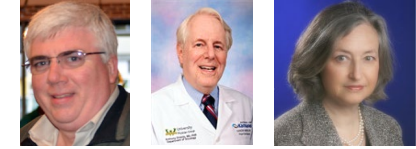


Biologic Events in Early Response to Successful Cancer Therapy

Rationale for Measuring Early Response by Cell Proliferation Imaging



Imaging Cellular Proliferation: [^{18}F]fluorothymidine (FLT)



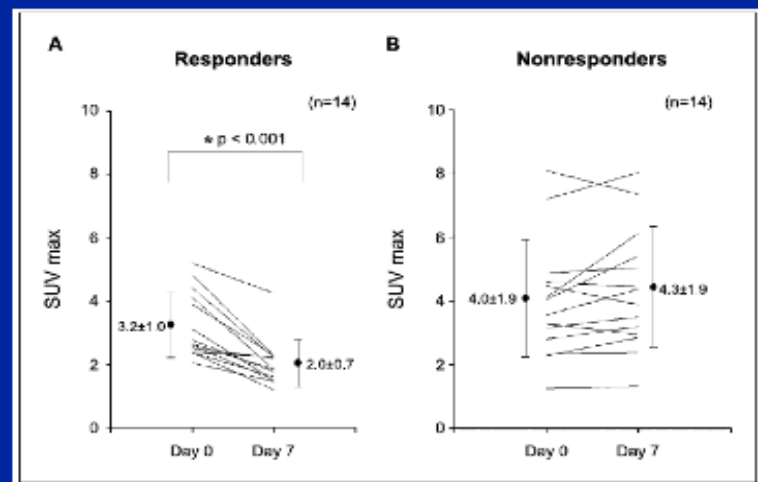
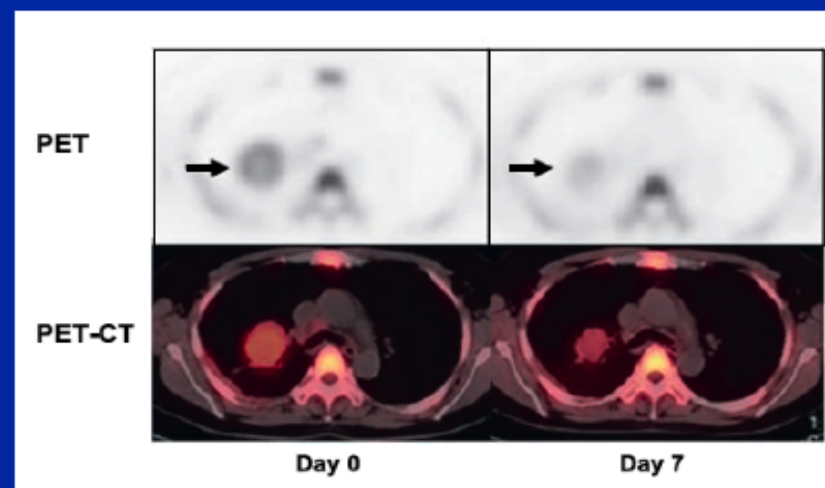
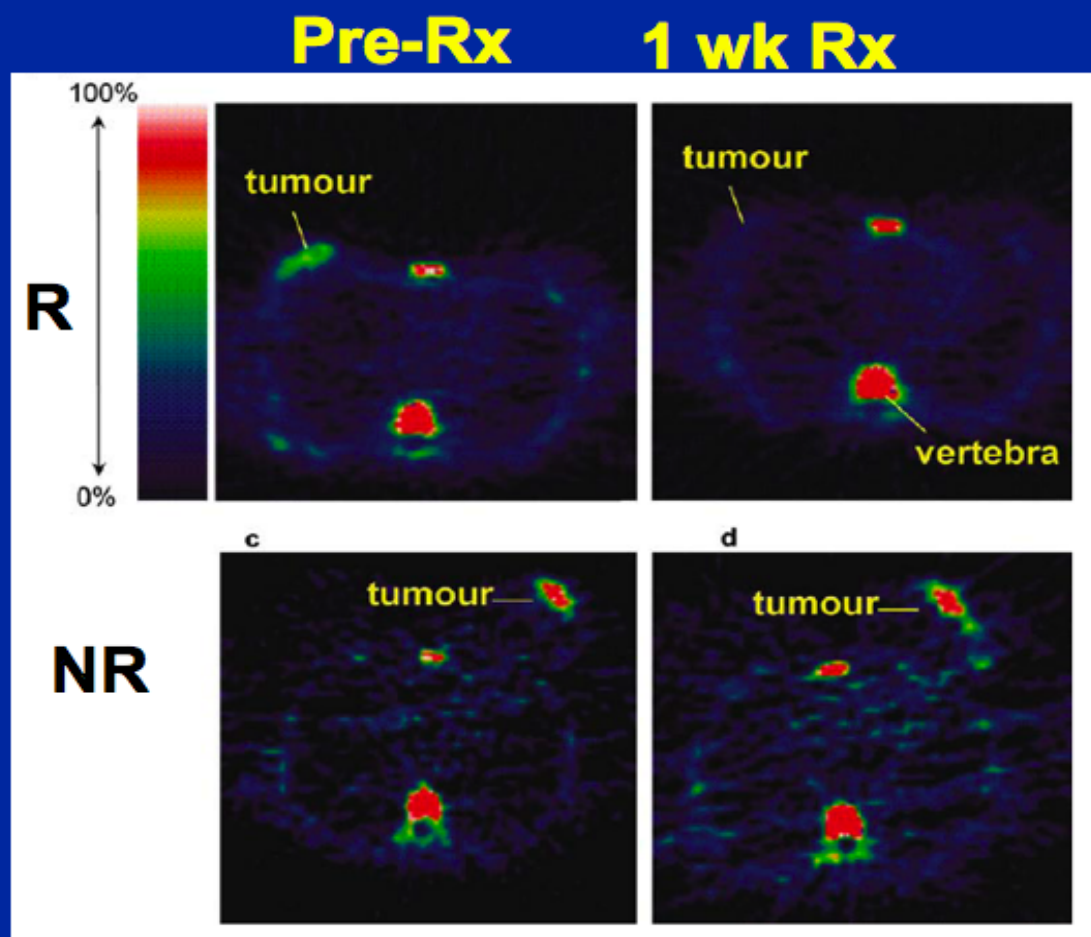
(Shields AF, Nature Med 4:1334, 1998)

(Grierson, Nucl Med Biol 27:143, 2000;
Mankoff and Eary, Clin Cancer Res 14:7519, 2008)

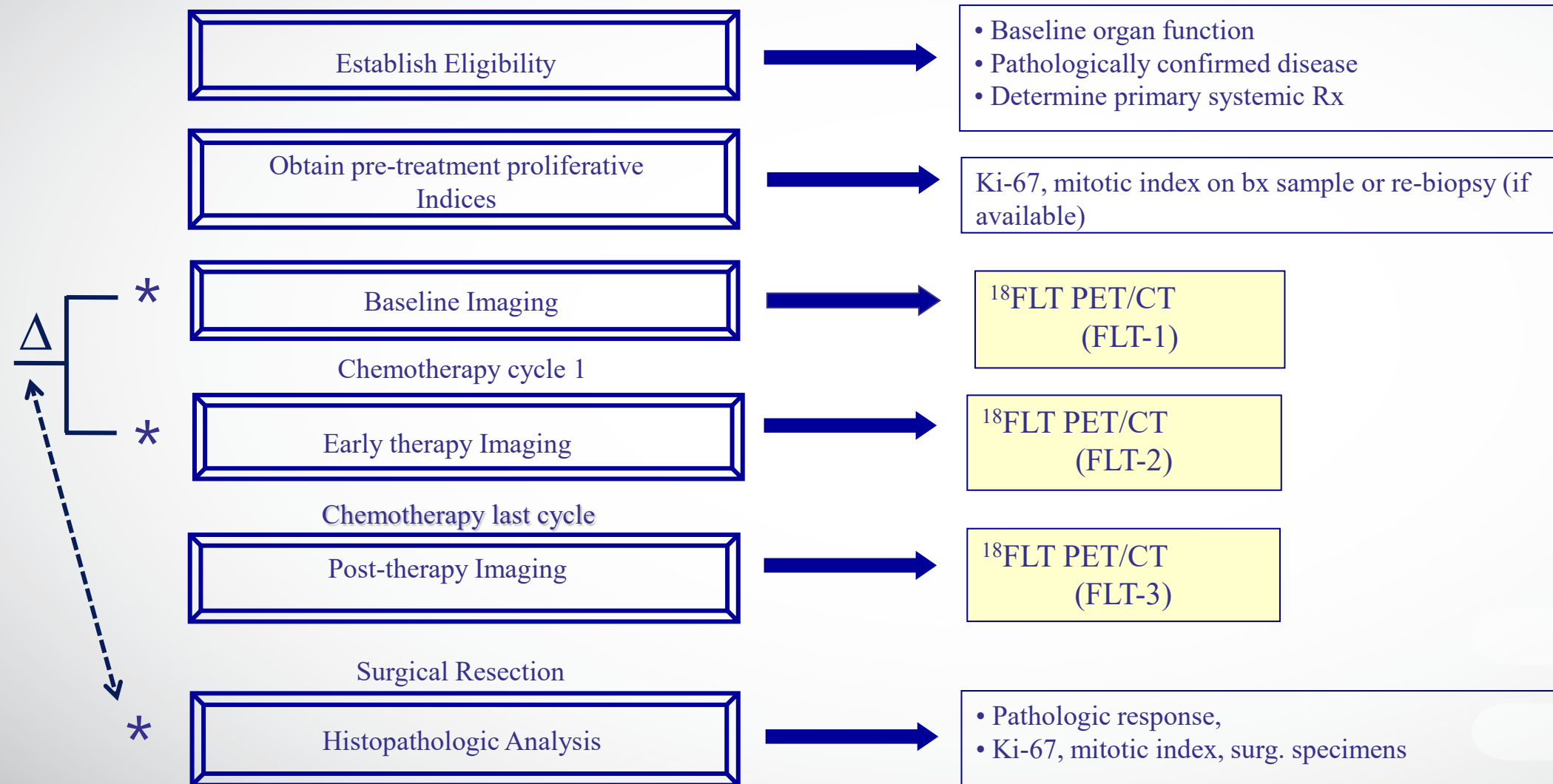
Early Response Measured by ^{18}F -fluorothymidine (FLT) PET

Breast CA, ChemoRx
(Kenny, EJNMMI 34:1339, 2007)

Lung CA, Genfitinib Rx
(Sohn, Clin Cancer Res 14: 7423, 2008)



ACRIN 6688: Phase II Study of FLT-PET in Invasive Breast Cancer



ACRIN 6688: FLT PET to Measure Early Breast Cancer Response

(PI: Lale Kostakoglu)

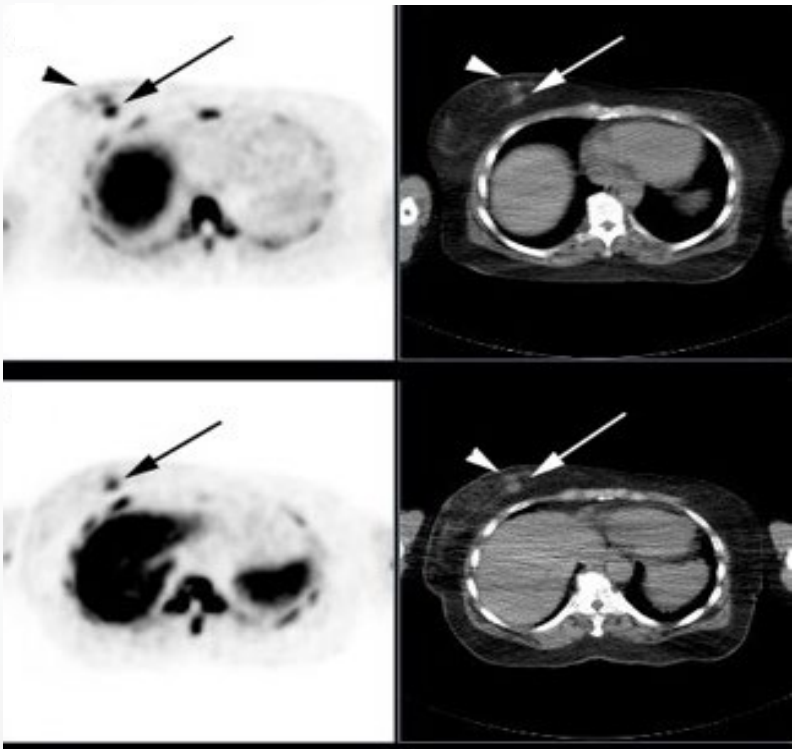


Lale Kostakoglu, PI
Jen Specht, UW Site PI

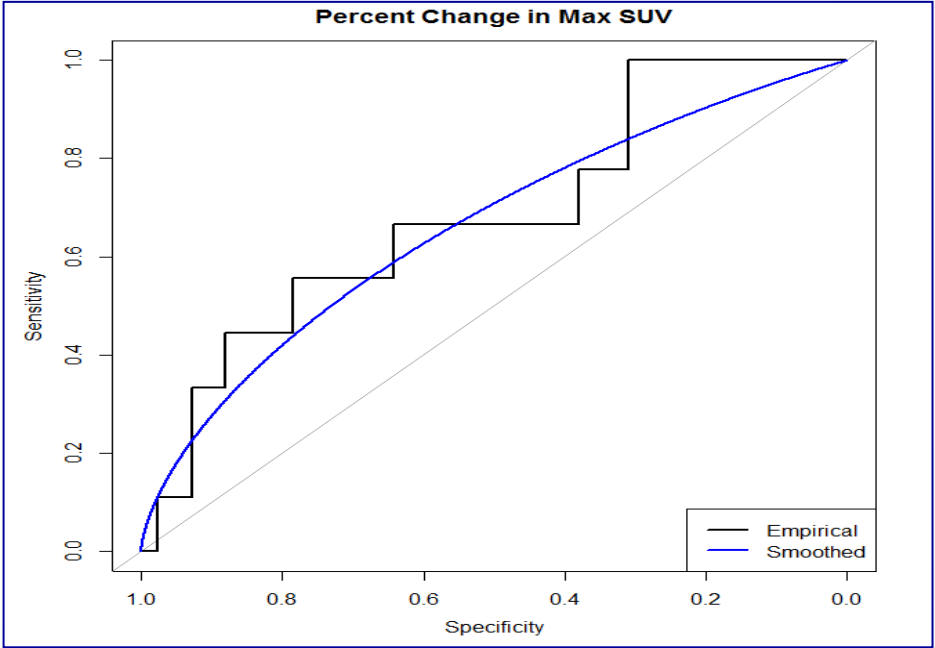
A Phase II Study of 3'-Deoxy-3'-¹⁸F-Fluorothymidine PET in the Assessment of Early Response of Breast Cancer to Neoadjuvant Chemotherapy: Results from ACRIN 6688

Lale Kostakoglu¹, Fenghai Duan², Michael O. Idowu³, Paul R. Jolles³, Harry D. Bear^{3,4}, Mark Muzi⁵, Jean Cormack², John P. Muzi⁵, Daniel A. Pryma⁶, Jennifer M. Specht⁵, Linda Hovanessian-Larsen⁷, John Miliziano⁸, Sharon Mallett⁹, Anthony F. Shields¹⁰, and David A. Mankoff⁶, on behalf of the ACRIN 668 Investigative Team

Pre-Rx



7 d Post-Rx



Best $\Delta\text{SUV}_{\text{max}}$ cut-off for predicting pCR = -51% (sensitivity 56%; specificity 79%).

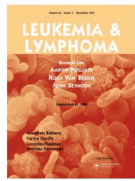
(Kostakoglu, J Nucl Med, 56: 1681, 2015)

ACRIN 6688 Secondary Impact & Opportunity

Does Kinetic Image Analysis improve Response Prediction?

Paved the Way for an AML FLT Trial (EAI141)

Kinetic Analysis (Methodology)



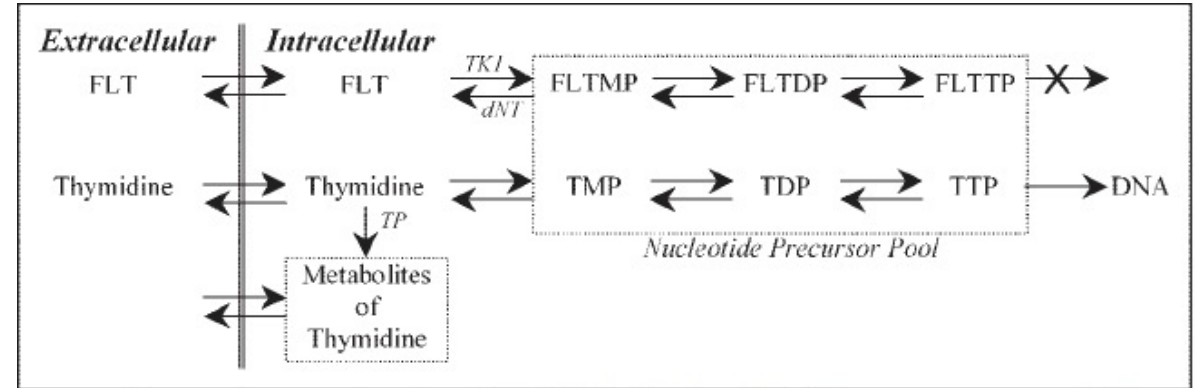
Leukemia & Lymphoma



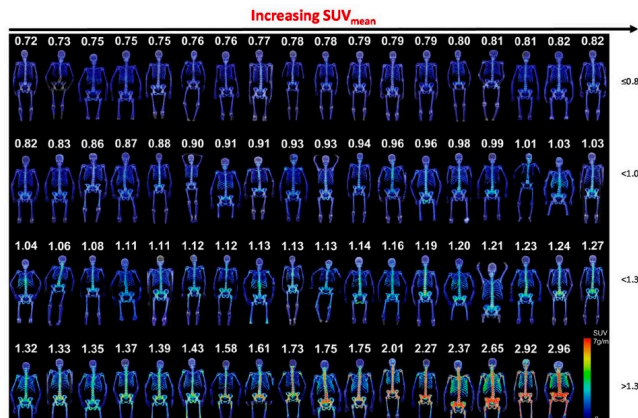
ISSN: 1042-8194 (Print) 1029-2403 (Online) Journal homepage: www.tandfonline.com/journals/lal20

Early assessment of treatment response in AML using FLT PET/cT: a trial of the ECOG-ACRIN Cancer Research Group (EAI141)

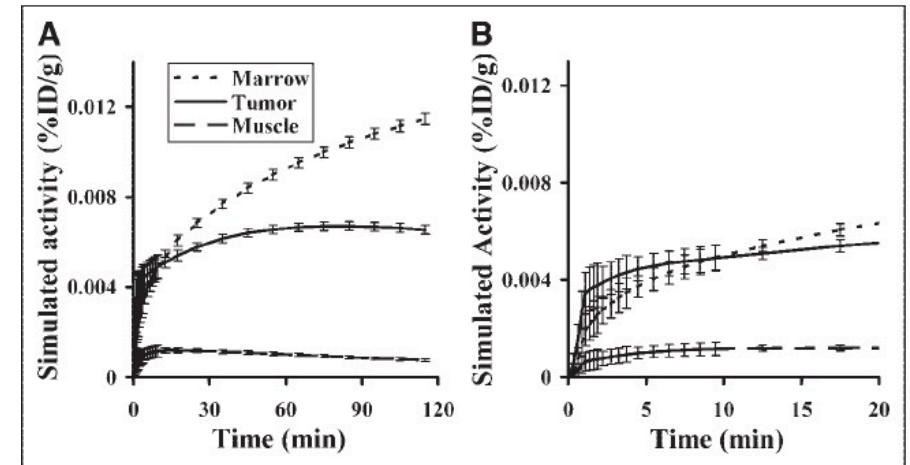
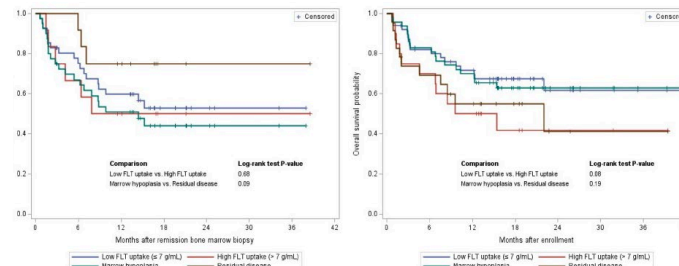
Robert Jeraj, Fenghai Duan, Ryan J. Mattison, Justin Romanoff, Lale Kostakoglu, Daniel A. Arber, Elisabeth M. Paletta, Jennifer L. Holter-Chakrabarty, Charles D. Arnold, Stephen A. Strickland, Jennifer G. Whisenant, Barry A. Siegel, Geoffrey L. Uy, Kebede H. Begna, Gregory A. Wiseman, Aric C. Hall, Scott B. Perlman, Mark R. Litzow & David A. Mankoff



WB FLT PET Methodology



Predictive Value for AML Induction



(Muzi, J Nucl Med 46:371, 2005)

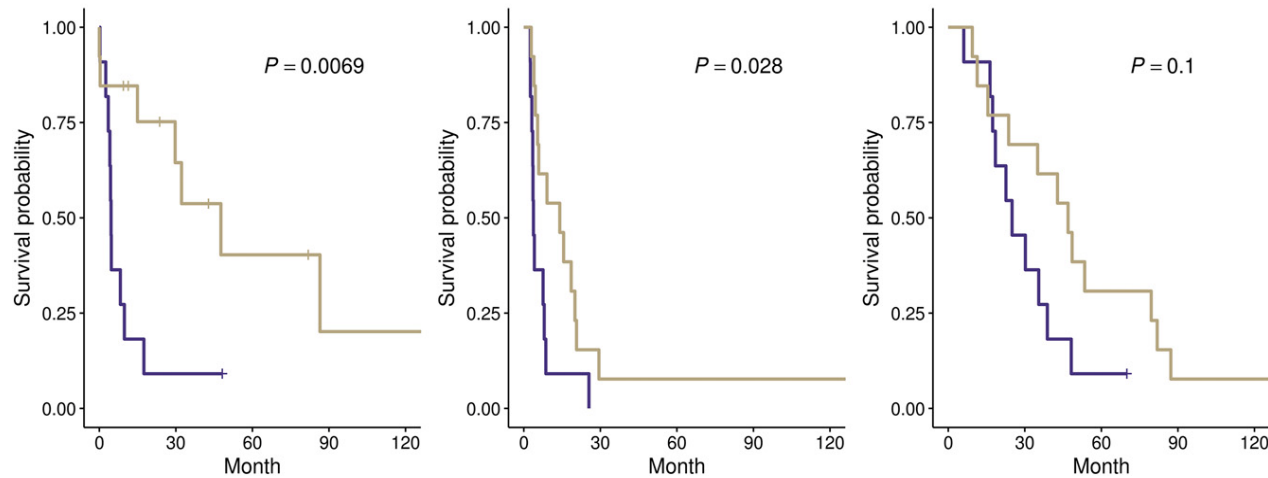
(Jeraj, Leuk Lymphoma. 66: 2765, 2025)

FDG PET Measures Bone Metastasis Therapeutic Response

Progression-Free Survival

Time-to- Skeletal Event

Overall Survival



Modified PERCIST Criteria: Metabolic Progression versus Other

(Peterson, J Nucl Med 2018; 59:1823)



Pre-Rx

4 wks Rx



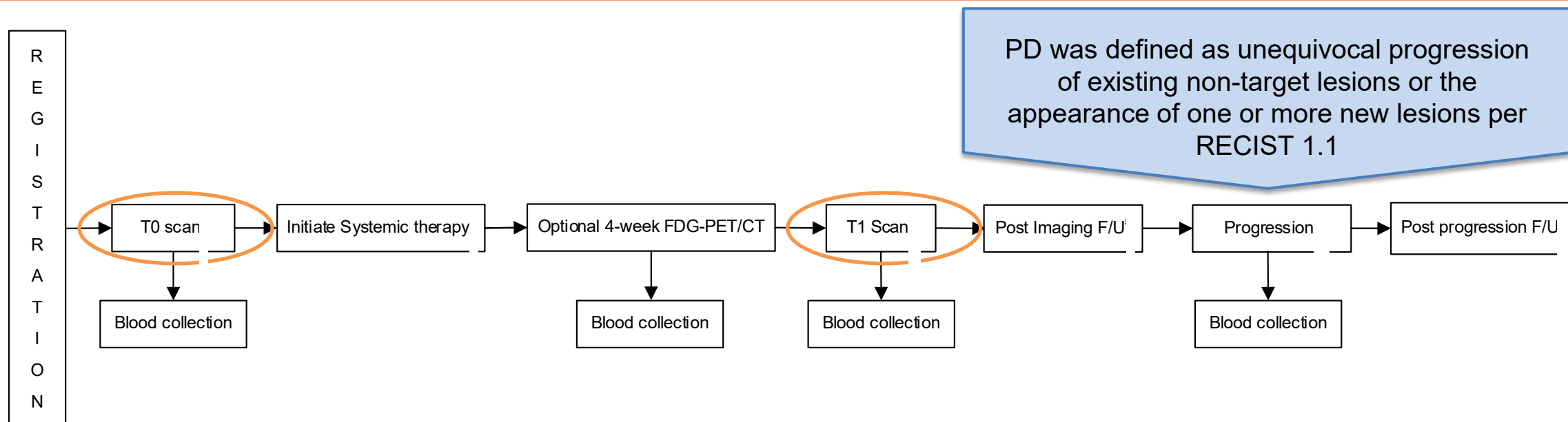
(Mahlkin, Radiol Imaging Cancer. 2022 Nov;4(6):e220032)



Penn Medicine
Abramson Cancer Center



EA1183 Schema



- Radiographically confirmed MBC with histologic confirmation of either metastatic or primary tumor biopsy; **hormone receptor positive with known HER2 status**.
- **BO/BD disease**
- RECIST 1.1 measurable lesions in viscera, active CNS, LMD, or pleural or peritoneal disease were excluded.
- Plan to receive **1st, 2nd, or 3rd line endocrine therapy, or chemotherapy or antibody drug conjugates or HER2-targeted therapies**. Patients who have received >3 lines of cytotoxic chemotherapy were excluded.

FDG-PET to Assess Therapeutic Response in Patients with Bone-dominant Metastatic Breast Cancer, **FEATURE** ECOG-ACRIN EA1183



Jennifer M. Specht¹, Fenghai Duan², Heather Jacene³, Eric Q. Konnick⁴, Frank Brescia⁵, Austin R. Pantel⁶, Samuel J. Galgano⁷, Justin Romanoff², Lanell M. Peterson¹, Mark Muzi⁴, Nancy U. Lin³, Ahmed K. Abou Hussein⁸, Wajeeha Razaq⁹, William R. Gwin¹, Ami N. Shah¹⁰, Mathew A. Cherian¹¹, Blanche H. Mavromatis¹², Chadwick L. Wright¹³, Daniel G. Stover¹¹, Amy M. Fowler¹⁴, Hannah M. Linden¹, Angie M. DeMichele⁶, Jonathan E. McConathy⁷, Christopher Comstock¹⁵, Antonio C. Wolff¹⁶, David A. Mankoff⁶

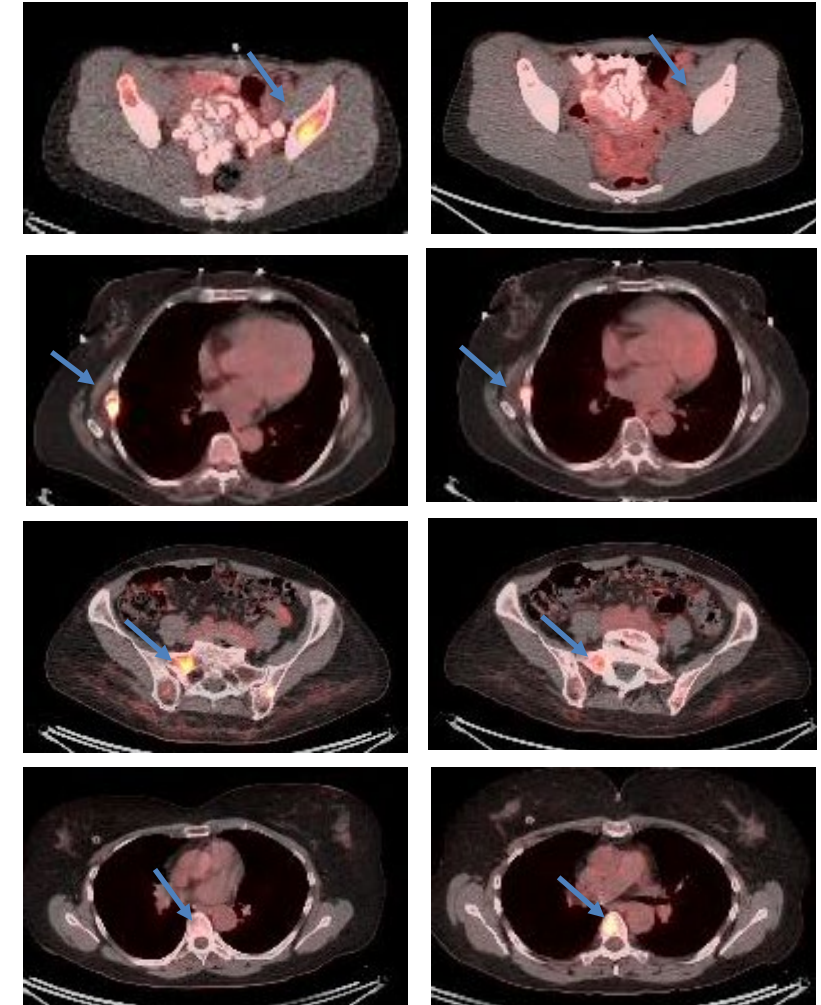
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Modified PERCIST Adapted for Bone

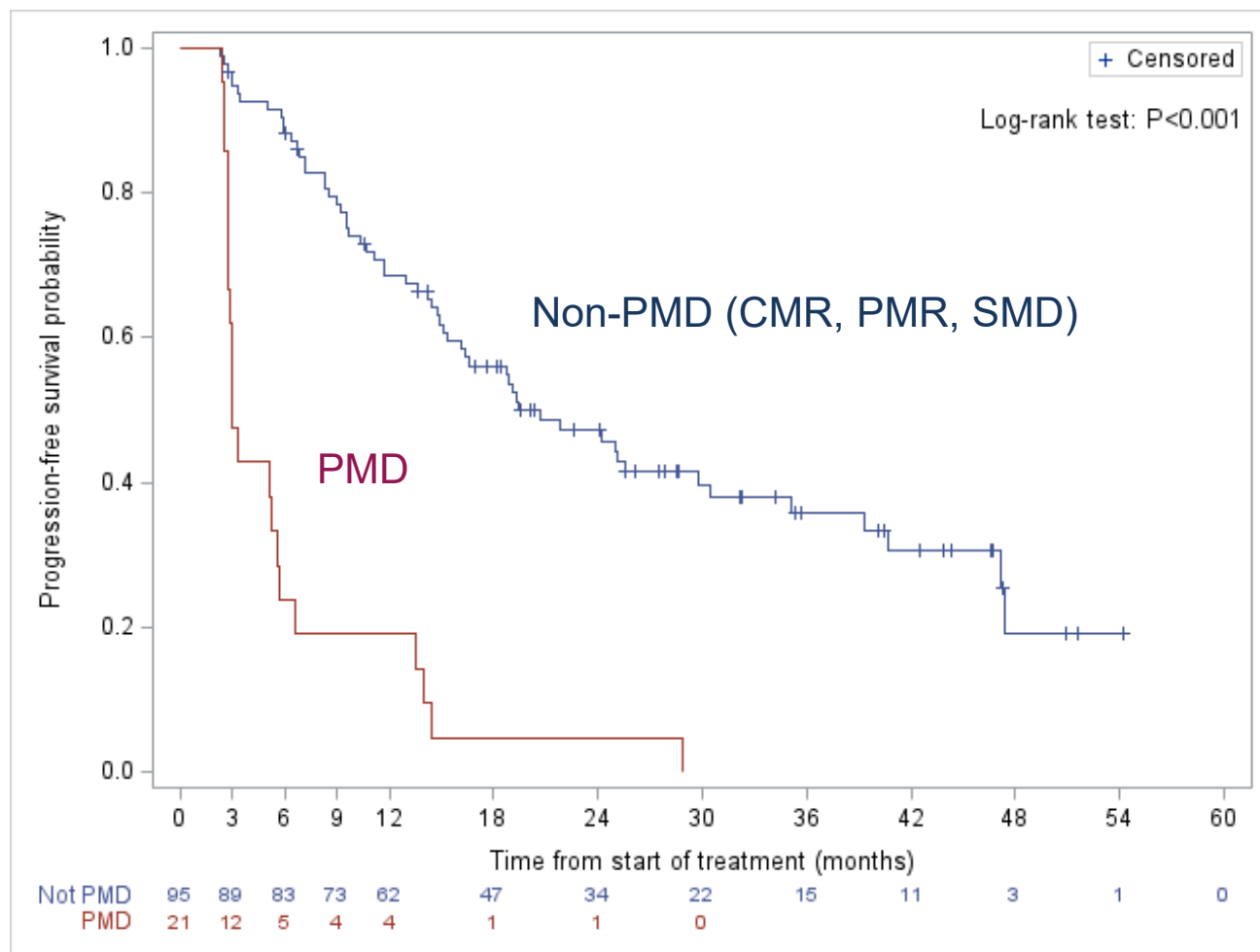


Measurable lesion: $SUL_{peak} \geq 1.5 \times \text{liver } SUL_{mean}$ (3 cm spherical ROI in normal right lobe of liver)

Complete metabolic response (CMR)	Complete resolution of FDG uptake in all active tumor to the level of background; No new FDG-avid lesions
Partial metabolic response (PMR)	$\geq 30\%$ and absolute 0.8 SUL unit decline in SUL_{peak} from single-hottest lesion at baseline to single hottest lesion at follow up; No increase $>30\%$ in SUL_{peak} or size of target or nontarget lesions; No new FDG-avid lesions.
Stable metabolic disease (SMD)	Not complete, partial metabolic response or progressive metabolic disease.
Progressive metabolic disease (PMD)	$\geq 30\%$ and absolute 0.8 SUL unit increase in SUL_{peak} from single-hottest lesion at baseline to single hottest lesion at follow up; or visible increase in extent of FDG uptake (75% in TLG volume with no decline in SUL); or new FDG-avid lesions in a pattern typical of cancer



Results: PFS by mPERCIST at T1 (12-wks)



- Data cut from 1 Sept 2025; median follow up 14.4 months; 21 pts with PMD and 95 without PMD
- Median PFS was **3.0 mos** (95% CI: 2.8 to 5.6 mos) for the PMD group and **19.4 mos** (95% CI: 15.1 to 30.5 mos) for the non-PMD group (log-rank test $P < 0.001$).
- Significantly lower risk of progression in pts with non-PMD compared to PMD pts; Univariate Cox HR **0.16** (95% CI: 0.10 to 0.28; $P < 0.001$)

Kaplan-Meier curves for PFS for participants with positive (not PMD) and negative (PMD) mPERCIST responses

Benefit to Investigators

R01 for Exploratory Endpoints (Early PET, ctDNA); Specht PI, Konnick Co-I



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SUMMARY STATEMENT
(Privileged Communication)

Release Date: 11/06/2020
Revised Date:

Principal Investigator
ODWYER, PETER J

Application Number: 1 R01 CA251803-01A1
Formerly: 1R01CA251803-01

Applicant Organization: ECOG-ACRIN MEDICAL RESEARCH FOUNDATION

Review Group: ZRG1 CTIS-C (07)
Center for Scientific Review Special Emphasis Panel
Clinical Translational Imaging Science

Meeting Date: 10/08/2020
Council: JAN 2021
Requested Start: 04/01/2021

RFA/PA: PAR19-363
PCC: Y3DI

Project Title: Role of ctDNA change as a response measure in the EA1183 patient population and how ctDNA changes correlate with metabolic response by serial FDG PET/CT

SRG Action: Impact Score:23 Percentile:6 &
Next Steps: Visit https://grants.nih.gov/grants/next_steps.htm

Human Subjects: 30-Human subjects involved - Certified, no SRG concerns
Animal Subjects: 10-No live vertebrate animals involved for competing appl.
Gender: 2A-Only women, scientifically acceptable
Minority: 1A-Minorities and non-minorities, scientifically acceptable
Age: 3A-No children included, scientifically acceptable

Project Year	Direct Costs Requested	Estimated Total Cost
1	340,667	493,748
2	331,845	480,962
3	217,688	315,508
4	194,725	282,226
5	185,637	269,054
TOTAL	1,270,562	1,841,498

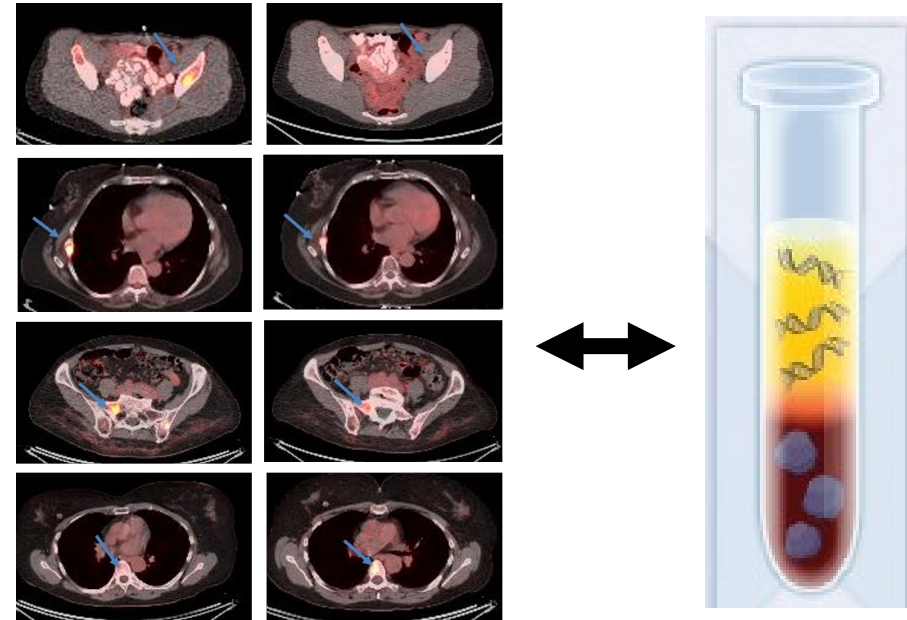
EAI142 Secondary Analysis Papers Planned

Early Response by FDG PET/CT



- 4-week FDG PET/CT response vs 12-week response
- Both vs PSF

ctDNA Response & Correlation with PET Results



ctDNA Data vs FDG PET/CT vs PFS

ACRIN 6687



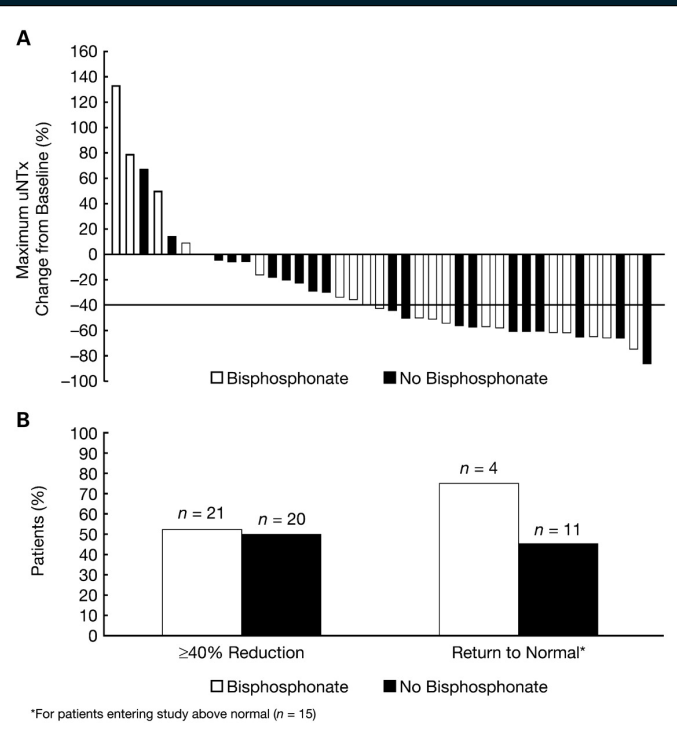
**A Phase 2, Multicenter Evaluation of ^{18}F -Fluoride PET
as a Pharmacodynamic Biomarker
for Dasatinib, a Src Kinase Inhibitor, in Men With
Castration-Resistant
Prostate Cancer and Bone Metastases**

***Principal Investigators: Evan Y. Yu, MD and
David A. Mankoff, MD, PhD***

Background for ACRIN 6687

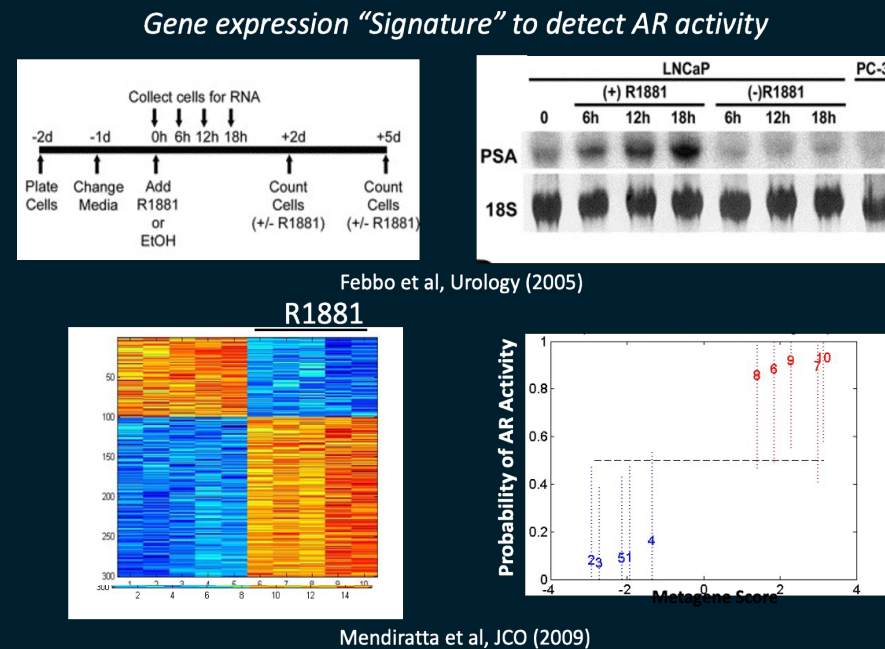


Dasatinib Leads to a Decrease in Bone Turnover Markers Is This Effect Metastasis-Specific?

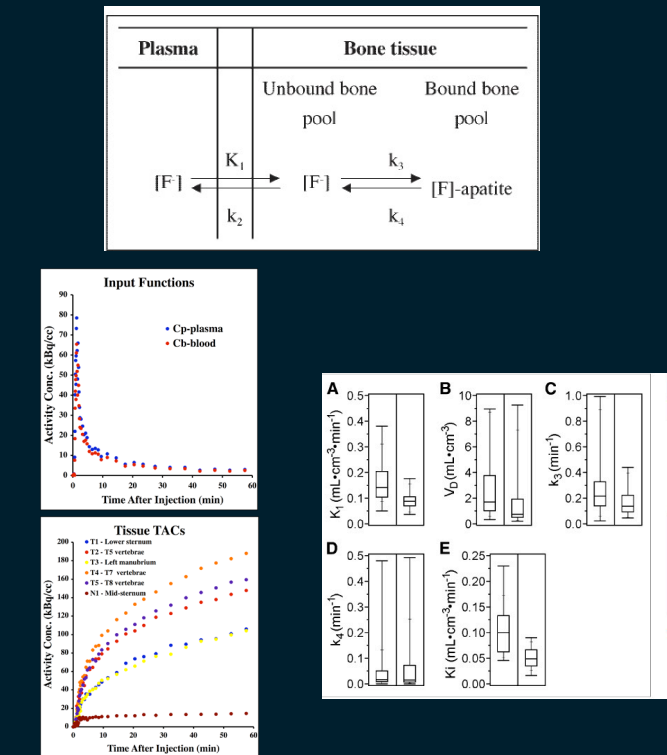


Yu EY, Clin Cancer Res 2009;15:7421)

Gene Expression to Predicts AR-Targeted Therapy Response Biomarker-Guide Crossover: Nilutamide vs Dasatinib



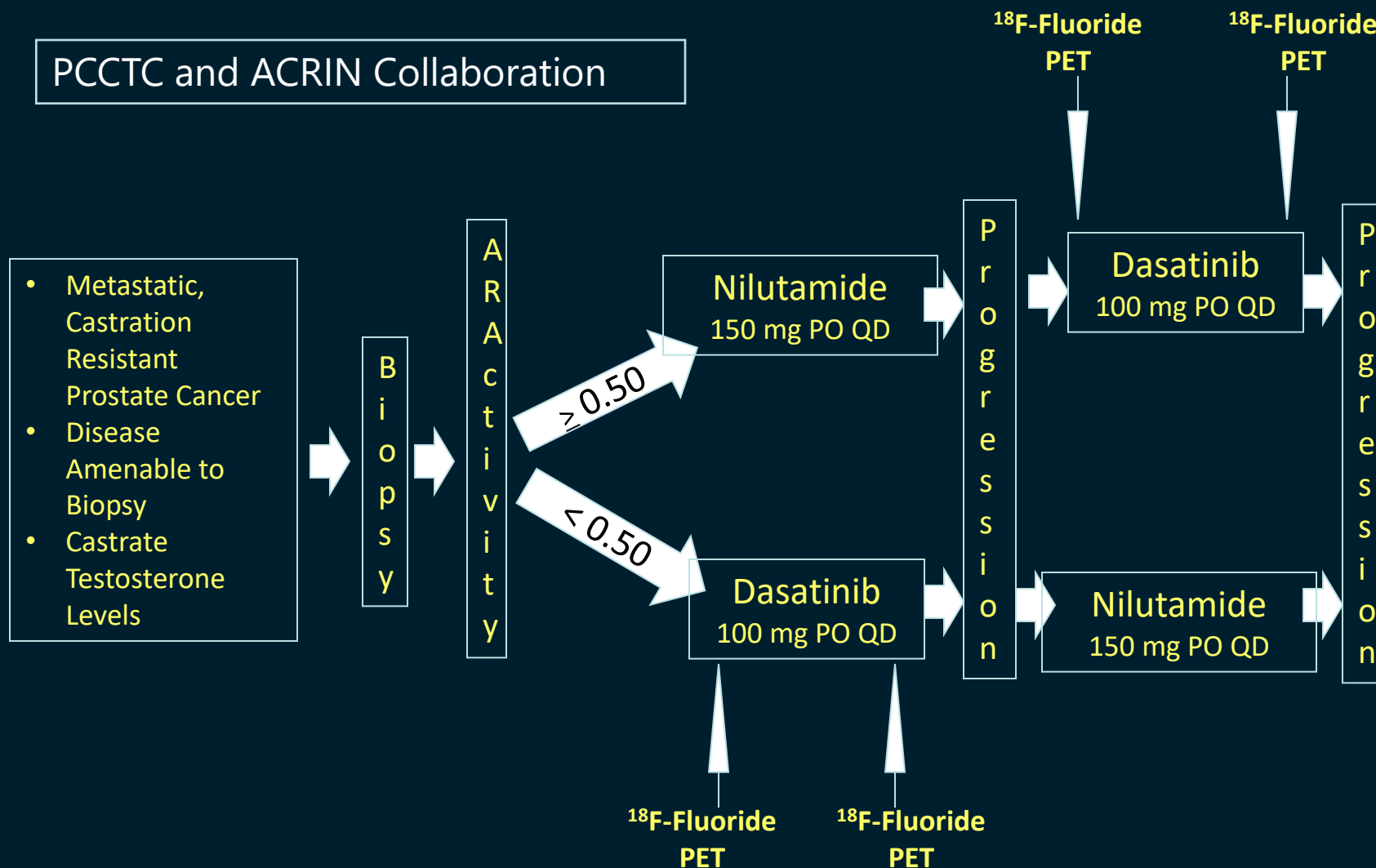
Dynamic [¹⁸F]NaF PET to Assess Regional Bone Turnover Kinetic Model Developed for Saroma & Br Ca



(Brenner, J Nucl Med 45:1493, 2004)
(Doot, J Nucl Med 51:521, 2010)

Study Schema

PCCTC and ACRIN Collaboration





ACRIN 6687: Image Analysis Tools

(Mark Muzi)



Forms and Scans

P2 ACRIN 6687
Evaluation of ¹⁸F-Fluoride PET for
Dysplasia
Visit 2: PET/CT Imaging Form

ACRIN Study 6687
PLACE LABEL HERE

Participant Initials: _____ Case No.: _____

7. Image Quality Interpretability?
a. Adequate present to get: ☐ 1
b. Inadequate present to get: ☐ 2
7a. Reason (mark all that apply):
☐ 1 Motion ☐ 2 Patient motion ☐ 3 Low contrast ☐ 4 Poor technique ☐ 5 Low image quality ☐ 6 Poor technique ☐ 7 Incomplete coverage

8. Name of Reader: _____

9. Reader ID#: _____

10. Expected Dosimetry: _____

11. Total number of tumor regions to report: _____

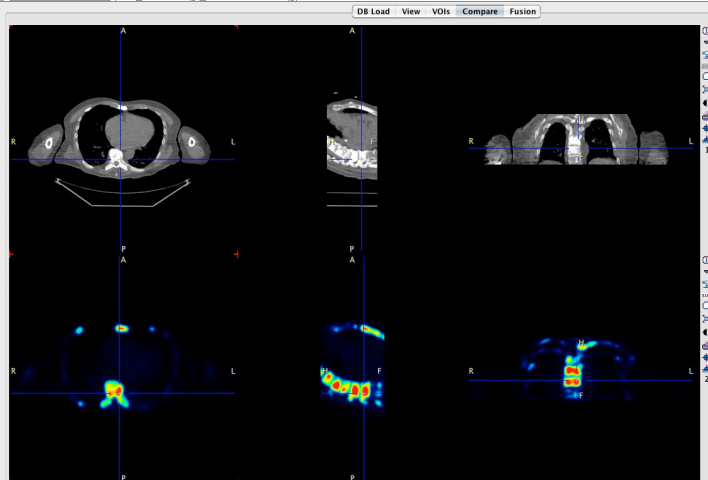
12. List up to the 5 most prominent regions to report (1-5) and identify anatomic site (1-5) in column 2.

13. List identifying normal regions to report (1-5) in column 1. Identify anatomic site and description of the region. Indicate the slice on which the region is located corresponding to the slice.

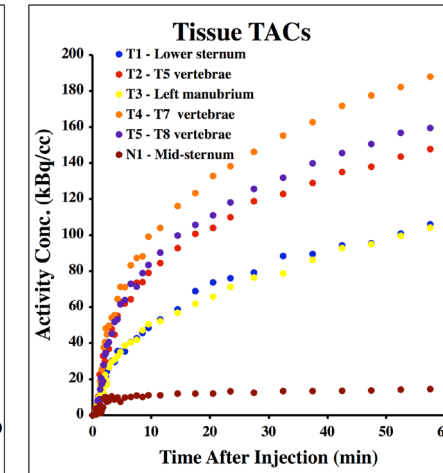
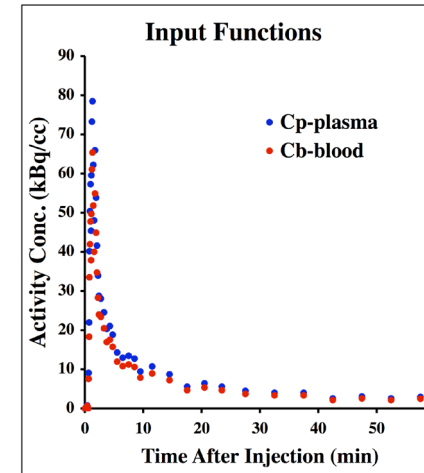
14. Initials of person completing form: _____

15. Date form completed: _____

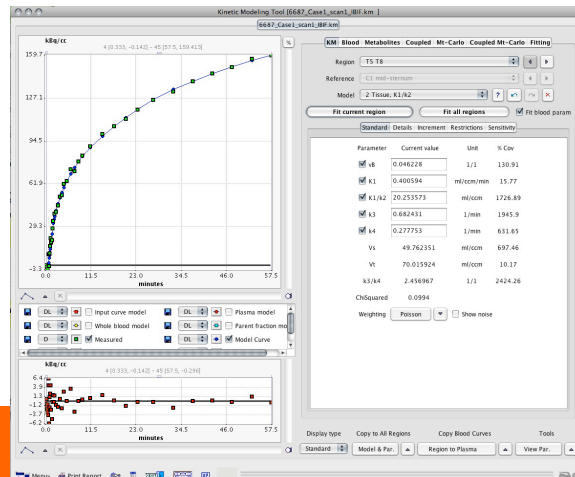
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Blood and Tissue Curves



Modeling, Parameter Optimization



UW PET Lab Data sent to ACRIN

ACRIN 6687 Dynamic Core Lab Data													
ACRIN Site	PET Study	PET Scan	ACRIN Case	Patient	Core	Core	PET Image	Core					
ID	Date	Visit Number	Number	Initials	Interpretation	Lab Data	Size	Analysis Sheet					
(###)	(mm-dd-yyyy)	(2 or 4)	(###)	(F/M)	(#)	(mm-dd-yyyy)	(cc)	(1, 2, 3, or ...)					
4221	03-06-2010	2	1	RWB	08-27-2010	5L	0.0504	1					
FROM THE P2 and P4 FORMS													
The Most Prominent Tumors	Local Tumor ID#	Local Tumor Site Description	Local Tumor SUVmax (g/mL)	Core Tumor ID#	Core Tumor Site Description	Core Tumor SUVmax (g/mL)	Core Tumor SUVmean (g/mL)	Core Tumor SUVpeak (g/mL)	Core Tumor Patlak Flux (mL/min/mL)	Core Tumor Transport K1 (mL/min/mL)	Core Tumor Flux Ki (mL/min/mL)	Core Tumor Flux Ki (mL/min/mL)	Identification Agree between Local and Core (Yes/No)
1	1	Lower sternum	36.7	T1	Lower sternum	27	43.62	36.99	0.190	0.280	0.241	Yes	
2	2	T5	43.3	T2	T5	27	50.81	41.78	0.260	0.412	0.273	No	
3	3	Left manubrium	32.9	T3	Left manubrium	27	36.40	28.49	0.194	0.245	0.216	No	
4	4	T7	49.8	T4	T7	27	60.23	53.14	0.337	0.591	0.373	No	
5	5	T8	46.7	T5	T8	27	55.26	45.27	0.302	0.397	0.262	No	
Tumor-Matched Normal Bone													
Local Normal Bone ID#	Local Normal Bone Site Description	Local Normal Bone SUVmax (g/mL)	Core Normal Bone ID#	Core Normal Bone Site Description	Core Normal Bone SUVmax (g/mL)	Core Normal Bone SUVmean (g/mL)	Core Normal Bone SUVpeak (g/mL)	Core Normal Bone Patlak Flux (mL/min/mL)	Core Normal Bone Transport K1 (mL/min/mL)	Core Normal Bone Flux Ki (mL/min/mL)	Core Normal Bone Flux Ki (mL/min/mL)	Identification Agree between Local and Core (Yes/No)	
1	Mid-sternum	10	N1	Mid-sternum	18	4.31	2.40	2.41	0.013	0.042	0.015	No	

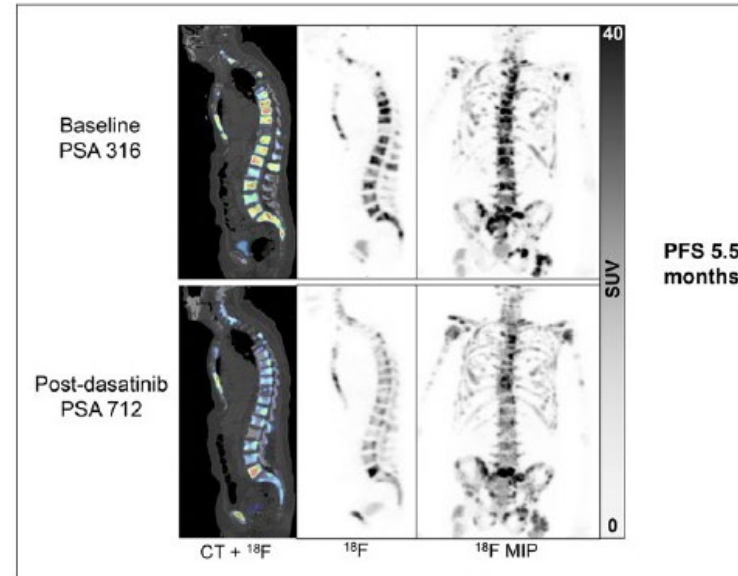
ACRIN 6687 Primary Paper

Yu, J Nucl Med 56: 354, 2015



Castration-Resistant Prostate Cancer Bone Metastasis Response Measured by ^{18}F -Fluoride PET After Treatment with Dasatinib and Correlation with Progression-Free Survival: Results from American College of Radiology Imaging Network 6687

Evan Y. Yu¹, Fenghai Duan², Mark Muzi¹, Xuan Deng², Bennett B. Chin³, Joshi J. Alumkal⁴, Mary-Ellen Taplin⁵, Jina M. Taub¹, Ben Herman², Celestia S. Higano¹, Robert K. Doot⁶, Donna Hartfeil⁷, Philip G. Febbo⁸, and David A. Mankoff⁶

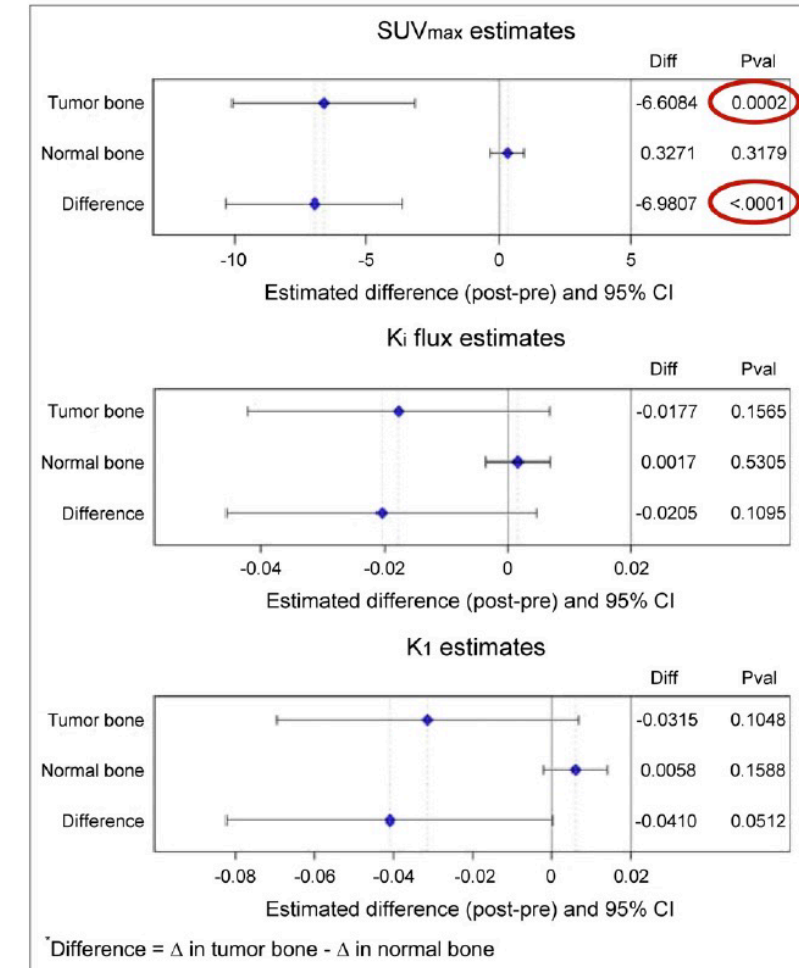


Conclusions:

- Effect of dasatinib on bone turnover – metastases > normal bone
- Borderline associate between NaF PET tumor changes and PFS
- Response can be heterogenous from site-to-site

Next Steps:

- Are other NaF PET image analysis methods more predictive?
- Would other imaging agents work better?



Secondary Papers for ACRIN 6687

Planned Secondary Aim

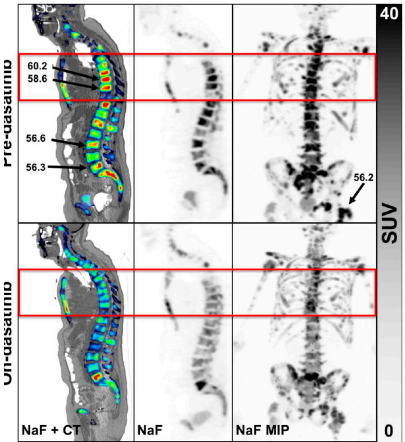


Article

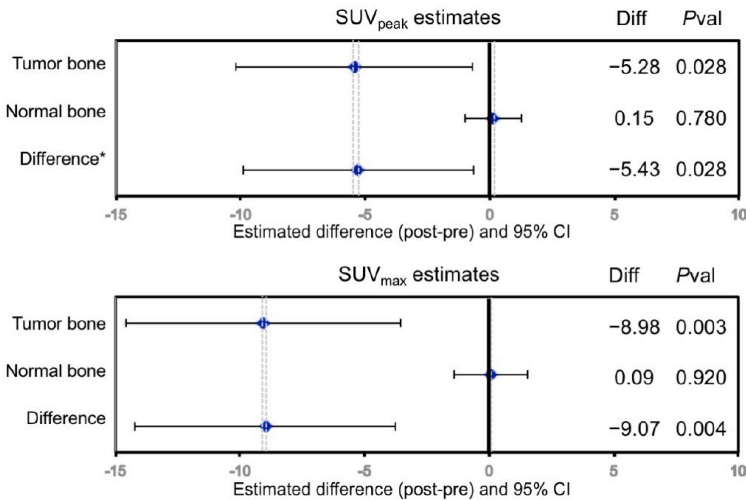
Whole-Body [18F]-Fluoride PET SUV Imaging to Monitor Response to Dasatinib Therapy in Castration-Resistant Prostate Cancer Bone Metastases: Secondary Results from ACRIN 6687



WB NaF PET Methodology



Improved Assessment of Dasatinib Impact on Metastasis Bone Turnover



(Muzi, Tomography, 7: 139, 2021)

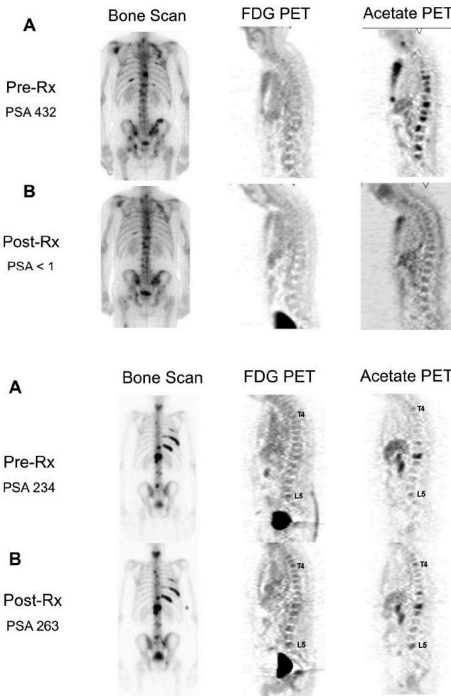
Pilot Trial Motivated by Trial Results

ORIGINAL ARTICLE

C11-Acetate and F-18 FDG PET for Men With Prostate Cancer Bone Metastases

Relative Findings and Response to Therapy

Evan Y. Yu, MD,* Mark Muzi, PhD,† Joy A. Hackenbracht, MA,* Brian B. Rezvani, MD,* Jeanne M. Link, PhD,† Robert Bruce Montgomery, MD,* Celestia S. Higano, MD,* Janet F. Eary, MD,† and David A. Mankoff, MD, PhD†



(Yu, Clin Nucl Med 36:192, 2011)

Cancer Imaging-Focused Trials in the NCTN

- My background and path NCTN participation and leadership
- How Does Imaging Fit into the NCTN Portfolio?
- Types of Imaging trials in the NCTN
- Examples of ECOG-ACRIN Imaging Trials and Opportunities for Early Investigators
- **Guidance and Future Directions**

NCTN Imaging Studies: Opportunities for Early Investigators

- **Opportunities**

- For non-imagers

- Leverage expertise and NCTN leadership in imaging at UW/FH to become an interface between therapy trial design and imaging tools for the trial
 - Write secondary papers for trial secondary, image-based endpoints
 - Be part of grants for exploratory imaging endpoints (e.g. EA1183 early PET/ctDNA R01)

- For imagers

- Provide expertise for therapy trials – leadership role in trial
 - Be PI or co-PI for a primary imaging trial
 - Be PI or co-I image-focused grants
 - Write secondary image analysis papers

- **Future Directions**

- New trials will increasingly require both imaging and tissue companion diagnostic
 - RP theranostics trials: by necessity, need oncology-NM coPIs, especially for combination Rx

Acknowledgements

- **Organizations**

- ACRIN/ECOG-ACRIN, NCI CIP, NCI QIN

- **Individuals**

- Charle Apgar, Amy Clark, Angie DeMichele, Janet Eary, Fegnhai Duan, Diana Ewen, Amy Fowler, Constantine Gatsonis, Elizabeth Gerstner, Donna Hartfeil, Heather Jacene, Robert Jeraj, Paul Kinahan, Eric Konnick, Lale Kostakoglu, Hannah Linden, Mark Muzi, Peter O'Dwyer, Lanell Peterson, Eva Ratai, Mitch Schnall, Erin Schubert, Jennifer Specht, Evan Yu