The NCI CIRP Program

Quantitative Imaging Research Resources At Preclinical and Clinical Setting

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CIRP Objectives

- Establish co-clinical quantitative imaging protocols via revere-translation from clinical imaging setting to pre-clinical imaging setting, with emphasis on optimization of preclinical imaging,
- Use available, credential GEMMs or PDXs to re-capitulate disease biological
- features and to address important cancer issues for adult or pediatric diseases,
- Use known therapeutic or prevention interventions as testing bed,
- Implement optimized imaging protocols to co-clinical interventions to collect coclinical images,
- Deliver a Web-Resource populated with all methods, protocols, tools and results by the 3rd quarter of yr-5.



https://ncihub.org/groups/cirphub

CIRP Web-Resource

Basic Required Content

Protocols

- Animal protocols
- Biology/Pathology/histopathology protocols
- Imaging protocols: QA/QC, PET, CT, MRI, etc.
- □ Methods
- □ Software
 - Imaging process
 - Image reconstruction
 - Data analysis
- Representative Data that are correlated to each other (preclinical & clinical)
 - Biology data
 - Pathology Data
 - Imaging data

Results

<u>Workflow</u>





CIRP Teams: Project Overview

			Patient Tumor Stage Group		Mouse Therapy Model		Quantitative Imaging								
Cancer Type	Diseases	Team	Adult	Pediatric	Primary	Metastatic	PDXs	GEMMs	Therapy	Combined	Anatomic	Metabolic	Cellular	Perfusion	Molecular
Hematology	Myelofibrosis	U Michigan	Y		Y			Y	Target		Y		Y		Y
Bone	Osteosarcoma	Stanford		Y	Y		Y		Immuno- therapy			Y			Y
Breast	Breast Cancer, TNBC	WUSTL	Y		Y		Y		Chemo- therapy		Y	Y	Y	Y	
	Breast Cancer, TNBC	BCM/UT/Stanford	Y		Y		Y		Chemo- therapy				Y	Y	
Colon	Colorectal Cancer	MDACC	Y		Y		Y		Target	Immuno- therapy		Y			Y
Lung	Lung Cancer, NSCLC	UW/FHCRC/BWH	Y		Y			Y	Immuno- therapy	Target		Y			
Muscle	Soft Tissue Sarcoma	Duke	Y		Y	Lung		Y	Immunot herapy	Radiation	Y		Y		
Pancreas	Pancreatic Cancer, PDA	UPENN	Y		Y			Y	Target				Y	Y	
Prostate	Prostate Metastatic cancer, SCNC	UCSF	Y		Y	Bone Liver	Y		Chemo- therapy		Y		Y	Y	Y

Cancer Issue: Therapeutic response, efficacy, failure and prediction, personalized treatment,

Disease: 8 cancer types (6 primary, 2 metastasis), 7 Adult diseases, 1 pediatric diseases,

Animal Model: 5 sites use PDXs, 4 sites use GEMMs,

Clinical Trial: 4 institutes on-going, 2 SPOREs, 1 CTEP, 1 R01, 1 Consortium; 8 Co-Clinical prospective, 1 matched pre-clinical prospective,

Therapy: 3 Chemotherapy, 2 target therapy, 2 target/immunotherapy, 1 immunotherapy, 1 immunotherapy/radiation therapy,

Imaging Methods: 1 CT, 2 PET, 5 MRI, 2 PET/MRI, anatomic, molecular, metabolic, cellular, perfusion, extendable to other diseases & therapies.

CIRP: Imaging Methods



In vivo quantitative imaging of anatomy, function, metabolism, molecules, vascularity, cellularity.

Cancers and Orthotopic Mouse Models Studied by CIRP



Potential of CIRP as Publicly Available Web-Resources

- Methodology to use animal models (PDXs or GEMMs) to recapitulate biological features of diseases,
- Protocols to detect dynamic spatio-temporal changes in tumor burdens at molecular, cellular, functional, physiological, anatomic level,
- Tools to generate quantitative, non-invasive, real time, longitudinal assessment for large cohort study,
- Strategies to inform clinical trials with preclinical study non-invasively and to bridge preclinical-clinical translational divide efficiently.

Web-resources Demonstrations: 7 teams





Team, Awarded Time

D Anderson

PREDICT

WUSTL	May 2017
Duke	Sept 2017
MDACC	Sept 2018
UPENN	Sept 2018
UMich	Sept 2019
Baylor	Sept 2019
UCSF	Sept 2020



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The Duke Preclinical Research Resources for Quantitative Imaging Biomarkers

Duke Radiology



CIRP Resource Example: WUSTL https://c2ir2.wustl.edu/

Washington University School of Medicine in St. Louis

Co-Clinical Imaging Research Resource (C2IR2)



Resources

- Animal Models
- Preclinical Phantoms for QA/QC
- **SOPs for Animal Modeling**
- SOPs for Scanner QA/QC
- Software/Code
- Datasets
- CIRP Resource Sharing Policy

Disease: TNBC Breast Cancer

Selected publications

- X. Ge, et al., Test–Retest Performance of a 1-Hour Multiparametric MR Image Acquisition Pipeline with Orthotopic Triple-Negative Breast Cancer Patient-Derived Tumor Xenografts, Tomography, 2019, 5, 320-331,
- S. Roy, et al., Optimal co-clinical radiomics: Sensitivity of radiomic features to tumour volume, image noise and resolution in co-clinical T1-weighted and T2weighted magnetic resonance imaging, EBioMedicine 59 (2020) 102963,
- M. A. Savaikar, et al., Preclinical PERCIST and 25% of SUVmax Threshold: Precision Imaging of Response to Therapy in Co-clinical 18F-FDG PET Imaging of Triple-Negative Breast Cancer Patient–Derived Tumor Xenografts, J Nucl Med 2020; 61:842–849,
- K. I. Shoghi, et al., Co-Clinical Imaging Resource Program (CIRP): Bridging the Translational Divide to Advance Precision Medicine, Tomography, 2020, 6, 273-287,
- K. Dutta, et al., Deep Learning Segmentation of Triple-Negative Breast Cancer (TNBC) Patient Derived Tumor Xenograft (PDX) and Sensitivity of Radiomic Pipeline to Tumor Probability Boundary, Cancers 2021, 13, 3795.
- S. Roy, et al., Co-clinical FDG-PET radiomic signature in predicting response to neoadjuvant chemotherapy in triple-negative breast cancer, European Journal of Nuclear Medicine and Molecular Imaging, 2021, <u>https://doi.org/10.1007/s00259-021-05489-8</u>.

CIRP Resource Example: Duke https://sites.duke.edu/pcqiba/

The Duke Preclinical Research Resources for Quantitative Imaging Biomarkers

Duke Radiology



Resources

MRI

Micro-CT

Protocols

Code

Data

Disease: Soft Tissue Sarcoma with Lung Metastasis

Selected publications

- □ S. J. Blocker, et al., Bridging the translational gap: Implementation of multimodal small animal imaging strategies for tumor burden assessment in a co-clinical trial, PLoS ONE 14(4): e0207555, 2019
- □ S. J. Blocker, et al., *The impact of respiratory gating on improving volume measurement of murine lung tumors in micro-CT imaging*, PLoS ONE 15(2): e0225019, 2020
- □ M. D. Holbrook, et al., *MRI-Based Deep Learning Segmentation and Radiomics of Sarcoma in Mice,* Tomography, 2020, 6, 23–33,
- □ A. J. Wisdom, et al., Single cell analysis reveals distinct immune landscapes in transplant and primary sarcomas that determine response or resistance to immunotherapy, NATURE COMMUNICATIONS, 6410 (2020),
- M. D. HolBrook, et al., Detection of Lung Nodules in Micro-CT Imaging Using Deep Learning, Tomography 2021, 7, 358–372.
- Blocker, S. et al., Ex Vivo MR Histology and Cytometric Feature Mapping Connect Three-dimensional in Vivo MR Images to Two dimensional Histopathologic Images of Murine Sarcomas, Radiology: Imaging Cancer 2021; 3(3): e200103.

CIRPHub: Central Hub links to Individual Resources https://nciphub.org/groups/cirphub

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poweredby NCIP Hub	Login Register Request Membership
Co-Clinical Imaging Research Resources Program Netwo	ork (CIRP) [cirphub]
Overview Members Resources Forum Projects Calendar Announcements Collections Activity	
About CIRP	
The Co-Clinical Imaging Research Resources Program Network (CIRP) is based on the trans-NCI initiative, currently, PAR-18-841. This FOA imaging methods are optimized to improve the quality of imaging results for co-clinical trials. Projects include optimization of pre-clinical que the data, methods, workflow documentation, and results collected from cancer therapeutic or prevention co-clinical investigations. To achie human investigations, imaging platforms, quantitative imaging methods, decision support software and informatics to populate the research	invites Cooperative Agreement applications to develop research resources that encourage a consensus on how quantitative uantitative imaging methods, implementation in co-clinical trials, and creating a web-accessible research resource that contains all eve the goals of the CIRP, applicants are encouraged to organize multi-disciplinary teams with experience in mouse models research, the resource. Each resource contains four essential elements: animal models, co-clinical trials, quantitative imaging, and informatics.
Four essential elements	
Animal Models Co-Clinical Trials Corection Co-Clinical Trials	Informatics

- **Teams:** abstracts of CIRP projects
- **Team Websites:** links to individual CIRP web-resource
- ❑ Activities: meeting agenda and program slides
- Selected publications: CIRP papers

CIRP Working Groups

- □ Animal Model & Co-Clinical Trial (AMCT) WG:
 - **Cancer issues and Unmet Needs**
- Imaging Acquisition & Data Process (IADP) WG: Standardization & Metrology
- □ Informatics & Outreach (IMOR) WG:
 - Metadata & Interoperability

CIRP Network: A special issue is coming

https://www.mdpi.com/journal/tomography/special_issues/AQCCIR



CIRP Outreach & Dissemination



CIRP Annual Meeting: Open to public,

- Outside speakers invited by the CIRP working groups,
- CIRP organizes sessions at scientific meetings or conferences,
- CIRP investigators present at scientific meetings,
- CIRP invites outside investigators as associate members,
- CIRP Hub as central hub (<u>https://nciphub.org/groups/cirphub</u>),
- □ Individual CIRP Web-resources.

Questions?

This set of slides will be available at https://ncihub.org/groups/cirphub



www.cancer.gov/espanol

www.cancer.gov