















Quantitative Image Informatics for Cancer Research (QIICR)

Pls: Andrey Fedorov, PhD, and Ron Kikinis, MD

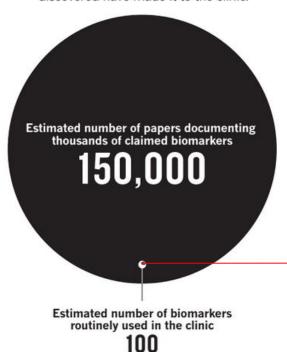
Brigham and Women's Hospital/Harvard Medical School

ITCR Face-to-face meeting 13 June 2016



A DROP IN THE OCEAN

Few of the numerous biomarkers so far discovered have made it to the clinic.



Actual number of FDA qualified imaging biomarkers

1





"The way the system now is set up, researchers are not incentivized to share their data. [...] every expert I've spoken to says we need to share this data in order to move the process more rapidly."

"I made a commitment that I will [...] eliminate the barriers that get in your way, get in the way of science, the research and development."

Joe Biden, Vice president of the United States of America

"When we have the opportunity to share data, knowledge, tools, or techniques, we must do it."

Warren Kibbe, PhD, acting NCI Deputy Director, Director of the Center for Biomedical Informatics and Information Technology (CBIIT)





"We need that big data to be accessible. It's not enough to say that we are in a big data era for cancer. We also need to be in a big data access era."

Francis Collins, PhD, MD, director of the NIH

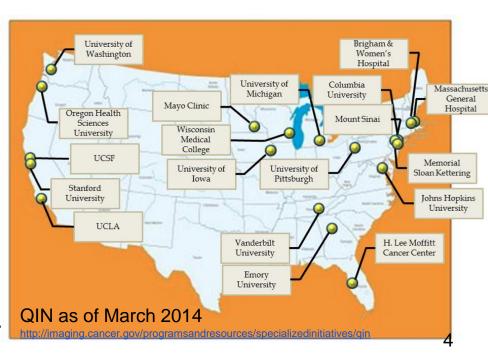


NCI Quantitative Imaging Network (QIN)

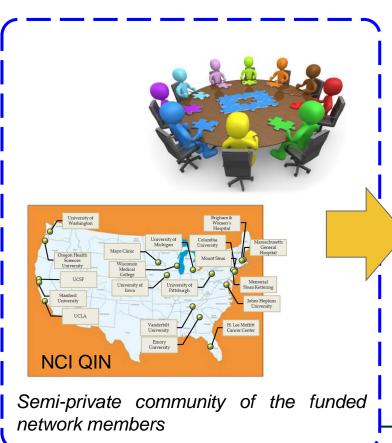
"The network is designed to promote research and development of quantitative imaging methods for the measurement of tumor response to therapies in clinical trial settings, with the overall goal of facilitating clinical decision making."

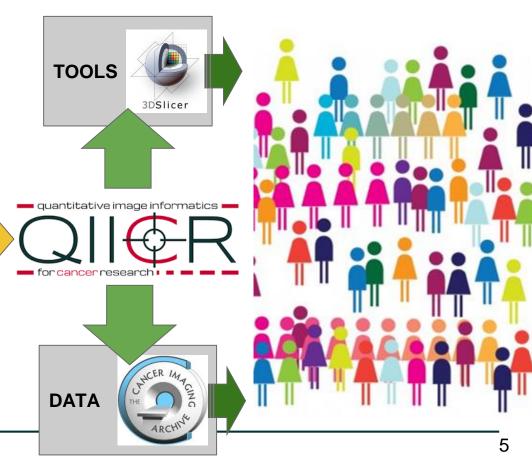
- Consensus building
- Best practices
- Performance measurement
- Collaborative activities
- Working Groups
- Tool comparison
- Image data sharing





Quantitative Image Informatics for Cancer Research (QIICR)





QIICR goals

Technology and tools for:

standards-based sharing reproducible calculation

of (candidate) imaging biomarkers from clinical image data



QIICR guiding principles

Driven by the needs of clinical research projects Head and neck, brain and prostate cancers "No-strings-attached" open source Tools for developers and non-developers Standardized data format imaging and non-imaging Outreach, sample data, tutorials

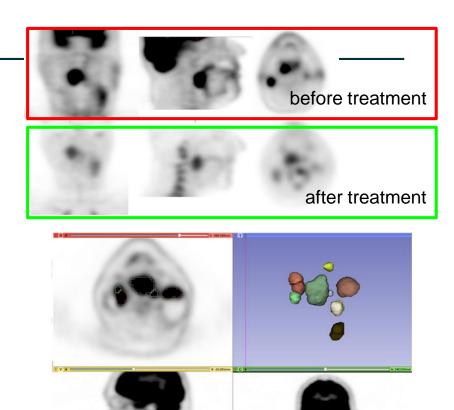


Tumor metabolic activity

Imaging surrogate: *Positron Emission Tomography*Quantitative image analysis:
localization of the uptake regions
quantification of metabolic activity

Evaluation in H&N cancer

Collaboration with U.of Iowa QIN site



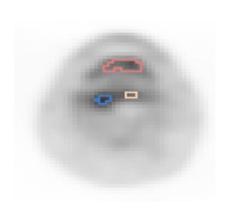


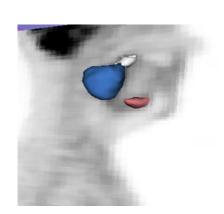
PET normalization

Standardized Uptake Value (SUV)

$$SUV(t) = rac{c_{img}(t)}{ID/BW}$$
 c_{img} - measured activity, \emph{ID} - injected dose, \emph{BW} - body weight

Reference regions - SUV Ratio (SUVR)











Quantitative imaging biomarkers: A review of statistical methods for technical performance assessment

Statistical Methods in Medical Research 2015, Vol. 24(1) 27–67
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DOI: 10.1177/0962280214537344
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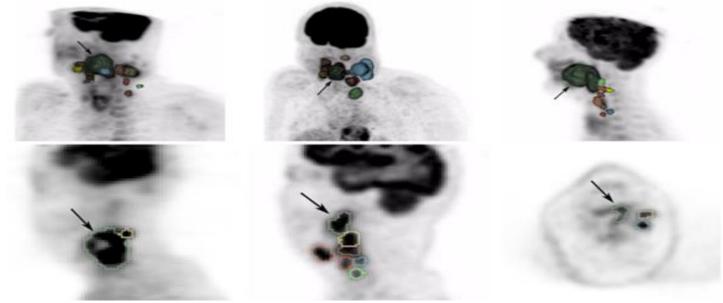
SSAGE

David L Raunig, ¹ Lisa M McShane, ²
Gene Pennello, ³ Constantine Gatsonis, ⁴ Paul L Carson, ⁵
James T Voyvodic, ⁶ Richard L Wahl, ⁷ Brenda F Kurland, ⁸
Adam J Schwarz, ⁹ Mithat Gönen, ¹⁰ Gudrun Zahlmann, ¹¹
Marina V Kondratovich, ³ Kevin O'Donnell, ¹² Nicholas Petrick, ³
Patricia E Cole, ¹³ Brian Garra, ³ Daniel C Sullivan ¹⁴ and
QIBA Technical Performance Working Group

Anecdotally, reproducibility of structural measurements is reader- and algorithm-dependent and is amenable to analysis using these summary measures, ⁵⁹ but the greater source of error for functional measurements such as the standardized uptake value (SUV) from fluorodeoxyglucose (FDG) PET imaging is outliers due to mistaken transfer of information such as patient weight and injection dose. ⁶⁰ Specification of the random effects model is critical to the design, analysis, and interpretation of these studies. Additionally, note that published results of studies involving rater agreement should adhere to Guidelines for Reporting Reliability and Agreement Studies. ⁶¹



PET quantitation: Region of Interest (ROI)



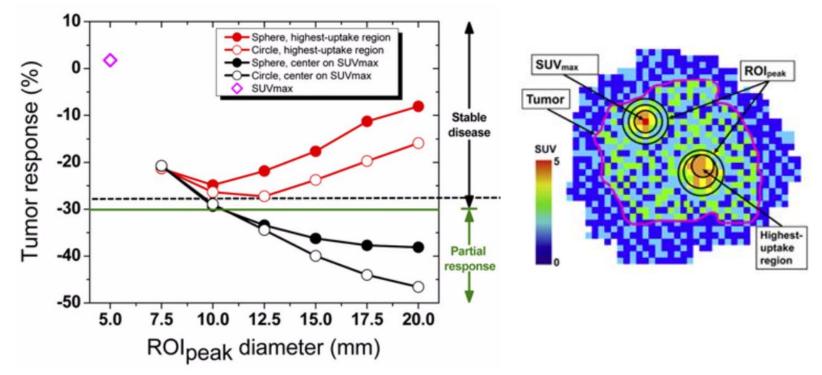
Primary tumor *vs* affected lymph nodes Splitting lesions Varying levels of uptake

Effect of ROI on the measurement up to 55%



PET quantitation: Feature extraction

Variety of quantitative descriptors





Candidate biomarker evaluation

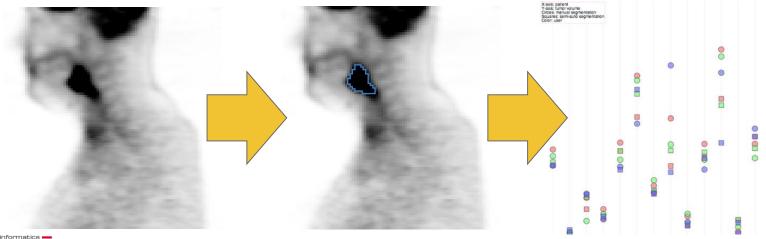
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treatment response assessment (QIN mission)
validate against clinical endpoint
    "a characteristic or variable that reflects how a patient feels,
       functions, or survives"
must consider relevant clinical descriptors
    Demographics
    Environment
    Therapy
    Complications
```



QIICR tools for PET/CT quantitative imaging

End-to-end automated processing tools

From loading a PET image to candidate imaging biomarker measurements



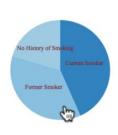


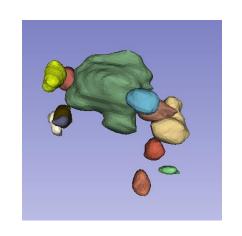
QIICR data for PET/CT quantitative imaging

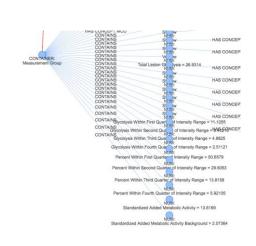
DICOM encoding of biomarker data

Segmentations, image normalization, semantics,
provenance, clinical data, outcomes, treatment, ...

Public on The Cancer Imaging Archive (TCIA)









buccal mucosa (2%) floor of mouth (11%) glottis (13%)

hypopharynx (5%)

lower alveolar ridge (2%)

maxillary sinus (1%)

nasal cavity (2%) nasopharynx (4%)

oral cavity (7%) oropharynx (7%)

palatine uvula (1%) paranasal sinus (1%)

pyriform sinus (5%)

supraglottis (18%)

retromolar trigone (8%) salivary gland (1%)

pharyngeal tonsil (adenoid) (1%)

larvnx (11%)

lip (0%)

PeerJ

✓ PEER-REVIEWED

DICOM for quantitative imaging biomarker development: a standards based approach to sharing clinical data and structured PET/CT analysis results in head and neck cancer research

Bioinformatics Clinical Trials Oncology Radiology and Medical Imaging Andriy Fedorov^{1,2}, David Clunie³, Ethan Ulrich^{4,5}, Christian Bauer^{4,5}, Andreas Wahle^{4,5}, Bartley Brown ⁶, Michael Onken ⁷, Jörg Riesmeier ⁸, Steve Pieper ⁹, Ron Kikinis ^{1,2,10,11}, John Buatti 12, Reinhard R. Beichel 4, 5, 13 Published May 24, 2016 Note that a Preprint of this article also exists, first published November 26, 2015. PubMed 27257542 Author and article information Abstract



Graphical abstract



Meta

Peer Review history

Citations in Google Scholar

Questions

Links

Visitors 349

Views 514

Downloads 99



Broader utility contributions

Contributions to the DICOM standard process 1 supplement and 18 correction proposals Implementation of the relevant DICOM components in DCMTK library Simplify creation of relevant DICOM objects Tools to simplify interaction with DICOM Web search index Editor plugin



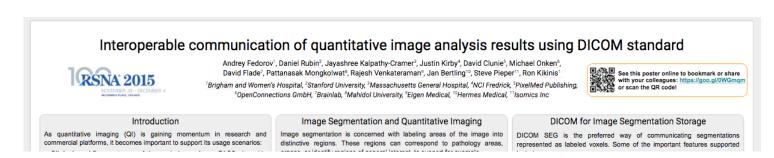
Collaborations - ITCR

- Standardized encoding of radiomics features (U24 CA194354, PI Aerts)
- Development of a 3D Slicer SlicerPathology extension for digital pathology image annotation (U24 CA180924, PI Saltz)
- Standardized exchange of annotations; integration of webbased (client) and 3D Slicer (server) tools to support quantitative imaging research (U24 CA199460, PI Harris)



Collaborations - beyond ITCR

- DICOM interoperability community effort:
 - o connectathon, sample data, demonstrations
 - RSNA activities (academic and industry participation: MEVIS, OHIF, ePAD, MITK, Brainlab, 3D Slicer, etc.)





Community building: NA-MIC Project Week

Twice a year, hands-on, open invite, academic/industry

welcome

January 9-13

MICCAI-endorsed event

Last event: 77 attendees

Join us for the next project week!

June 20-25, 2016: Heidelberg, Germany (CARS/IPCAI/DKFZ)





Demo video





https://youtu.be/wK2TGyVQjzs