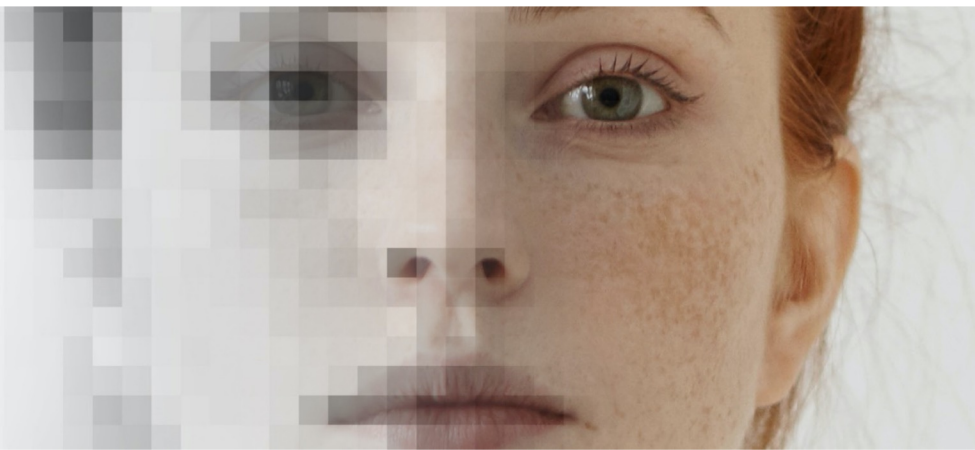
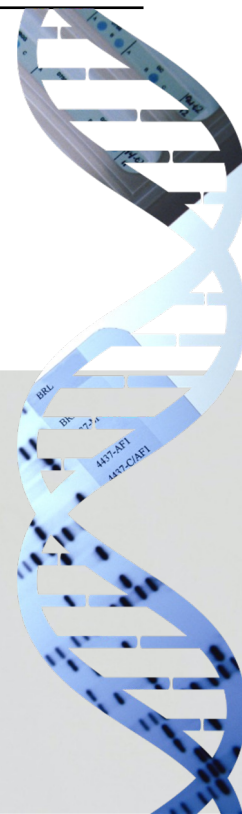


Automated Tumor Burden Assessment from CT-scans

Thomas Bengtsson

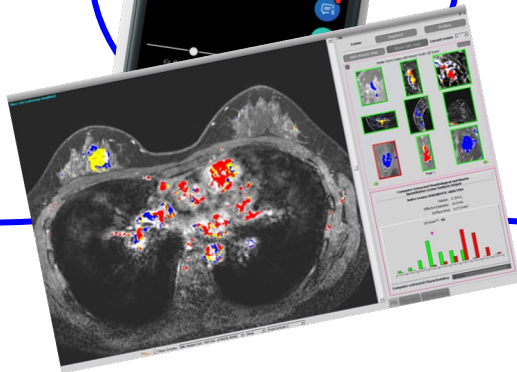
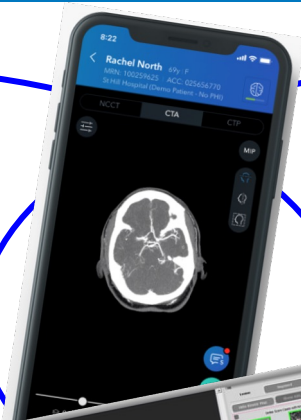
Skander Jemaa, Rick Carano, Jill Fredrickson, Shelby Wyatt,
Tina Nielsen, Alex de Crespigny
Genentech/Roche

BBSW, Nov 8, 2019



A Revolution at the Interface of Radiology and AI

Diagnostic Radiology



Deep Learning

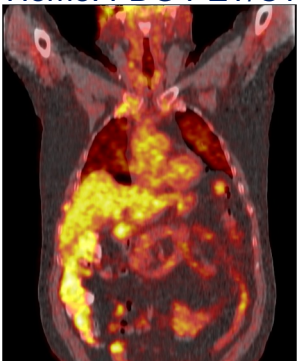


<http://www.image-net.org/>

What does all this mean for drug development?

Radiographic endpoints are a standard part of many of our trials, especially in oncology..

Heme: FDG PET/CT



Metabolic Response
by **Cheson** or
Lugano criteria

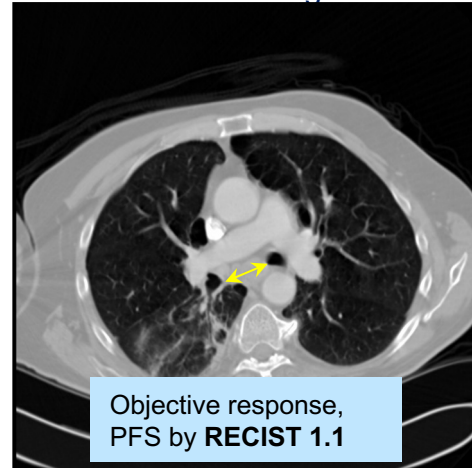
Current paradigm is central image review, with **adjudication rates 10--60%**

Automation: faster,
cheaper, more reliable

**Better endpoints,
patient selection**

**New approvable
endpoints, real-world
impact**

Solid tumors: Diagnostic CT



Objective response,
PFS by **RECIST 1.1**

Auto CT Project Overview

- **Background:**

- Computed Tomography - Response Evaluation Criteria In Solid Tumors (CT-RECIST)
 - Radiological standard method to quantify treatment response in solid tumors
 - Tumor Quantification: **Identify up to 5 target lesions (most 2 per organ)**, Identify up to 10 non-target lesions, quantify longest diameter for target lesions
 - Clinical Assessment: Define response (Ex. CR, PR, SD, PD) based on **changes in target and non-target lesion btw time points**
- CT-RECIST is an FDA accepted clinical trial endpoint for evaluating investigational therapies

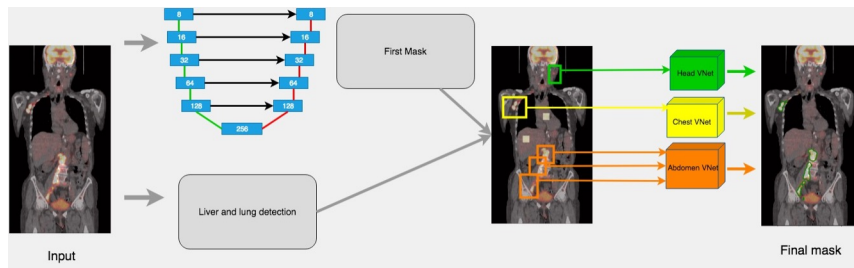
- **Goal: Develop a fully-automated image analysis CT-RECIST algorithm**

What is possible?

- More informative and robust endpoints for trials
- Faster readouts → Replace IRFs,
- Tools to help clinicians to select optimal therapies

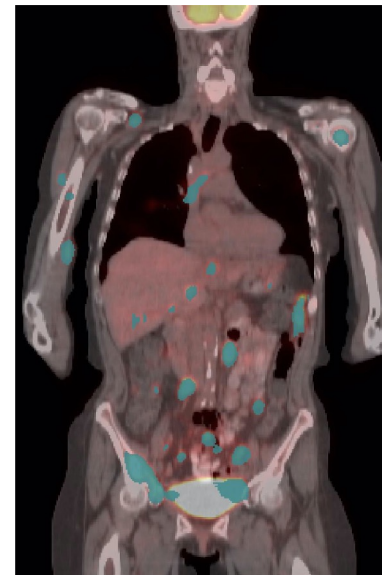
Automatic tumor burden assessment on FDG-PET

- Sparsity of disease (signal <1% body vol.) and lack of specificity of PET uptake relative disease requires multiscale residual learning DL architecture based on pyramidal dilations



- Spatial agreement* (trained on Goya, validated on Gallium)
 - DICE = 0.89
 - Corr. between reader- and predicted TMTV = 0.98
- Independent of IPI* (in Goya), a prognostic risk score based on $\{TMTV > 330\text{ml} \cap \text{Bulky disease} \cap \text{Nr. Lesions} > 12\}$ was an independent predictor of PFS (HR=2.01; CI 1.42-2.86)

NSCLC pat. w/ ~90 lesions

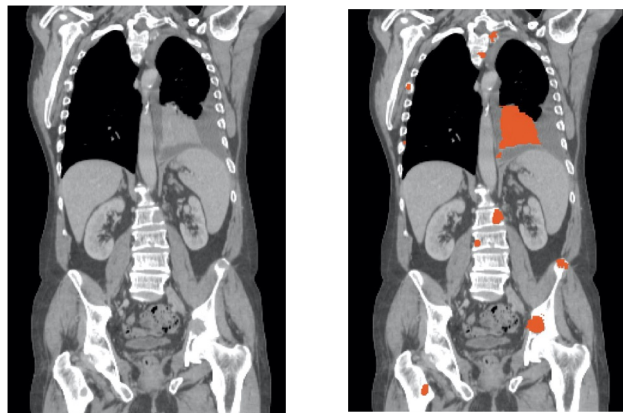


The model can process 1000 scans in < 1d
(cf. radiologist ~100 days)

*Ref: Jemaa et al. (SIIM 2019, ASH 2019)

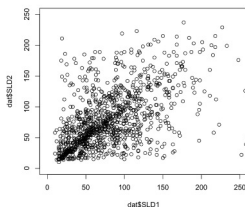
Automatic tumor burden assessment on CT

- Automatic lesion detection & segmentation on CT is a v. difficult problem, especially for advanced stage patients

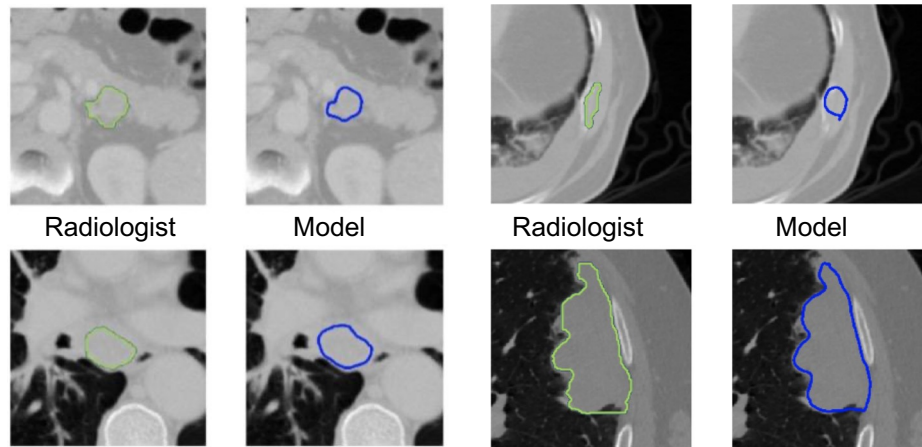


- High number of lesions: up to 250
- Substantial fraction of small lesions

Tumor selection problem leads to low inter-reader agreement



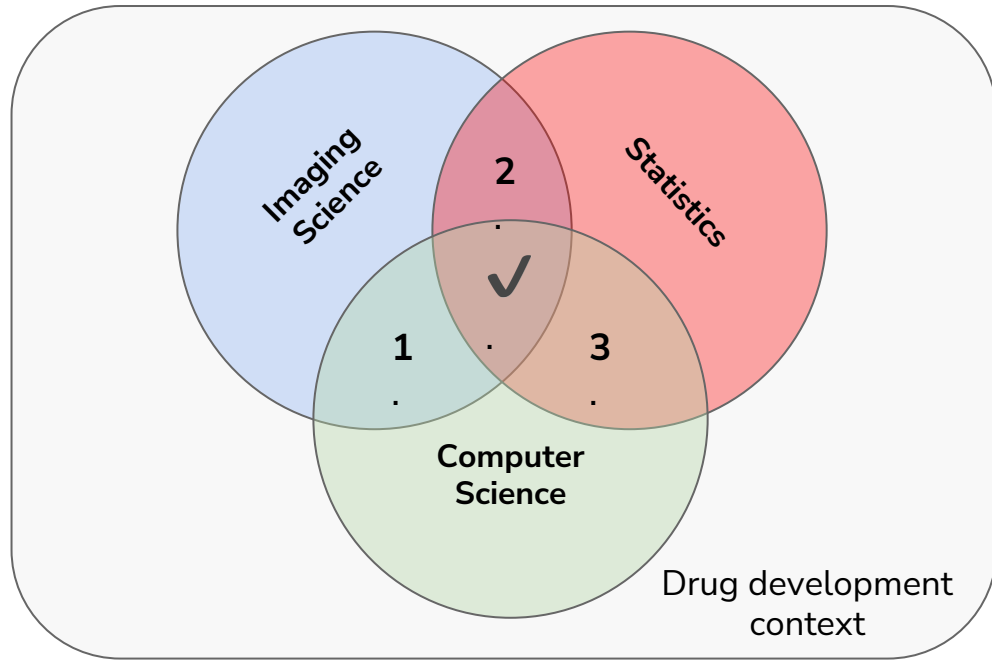
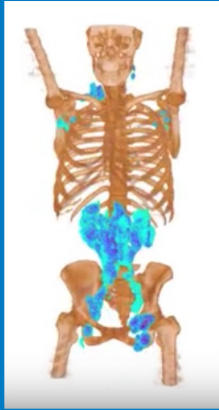
- A cascaded & memory efficient DL model has been designed to delineate tumors in Whole Body CT scans.



Results:

- For CT-R identified lesions (by radiologist), our **full-body model** has a voxel-wise sensitivity of 0.89 and a lesion level sensitivity of 0.94
- The segmentation achieves a 0.82 Dice Similarity Coefficient.

Cross-functional collaboration on AI and medical imaging



1. Lots of loops...
 2. Traditional analysis of tabular data...
 3. Analysis of imaging artifacts...
- ✓ required blend of imaging, CS, & Stats