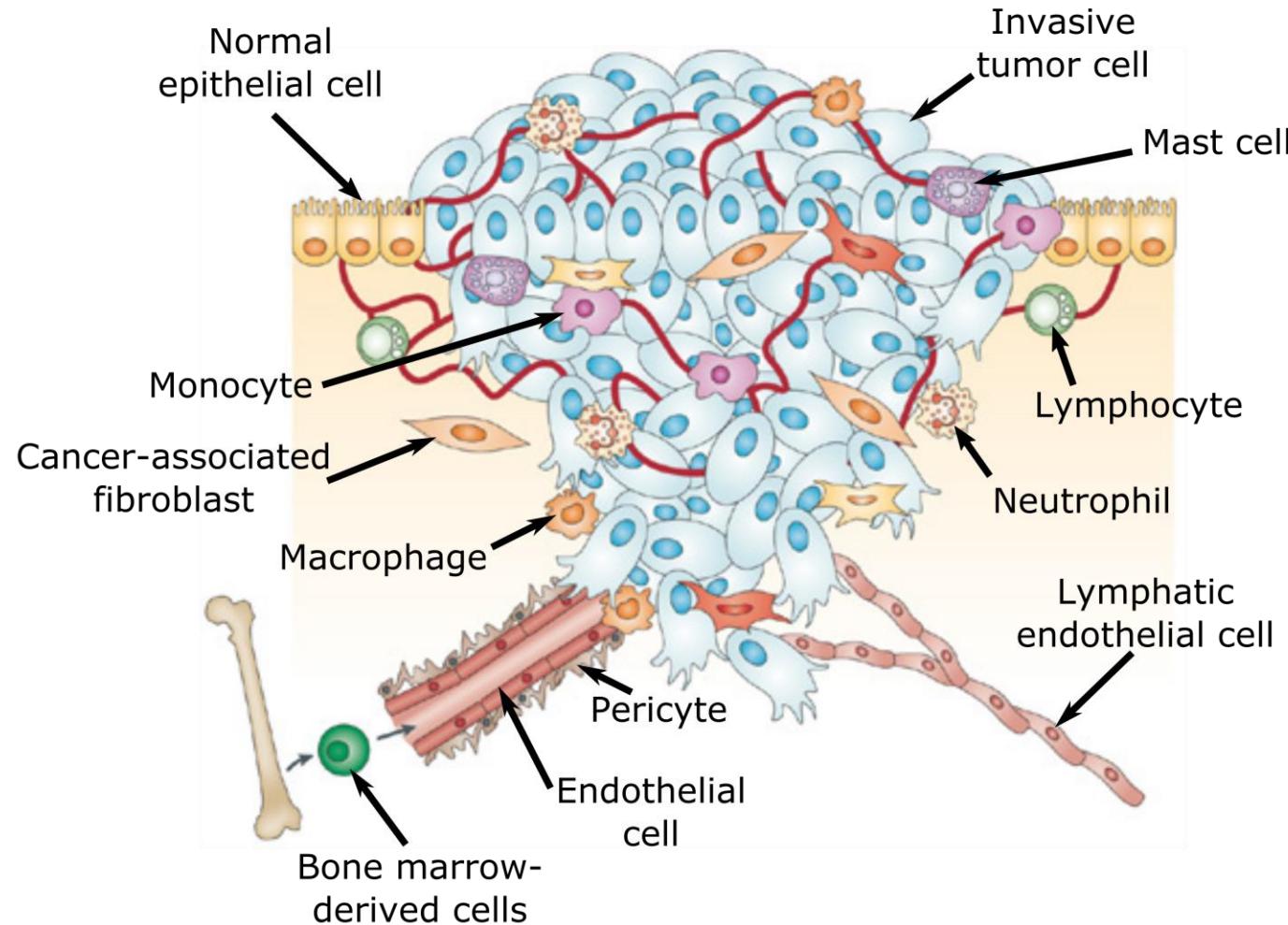


# Informatics Tools for Spatial Intra-Tumor Heterogeneity in Multiplexed Fluorescence Images

*s. chakra chennubhotla*

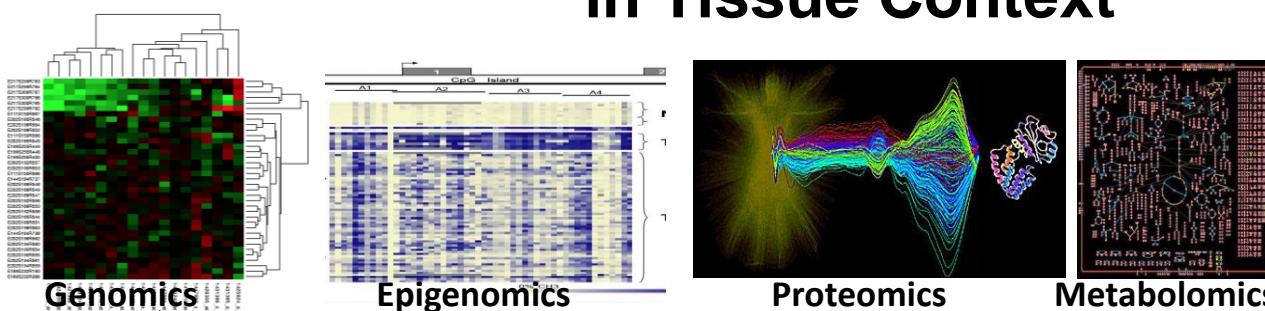
department of computational & systems biology  
drug discovery institute  
university of pittsburgh

# Intra-Tumor Spatial Heterogeneity Complicates Accurate Diagnosis/Prognosis

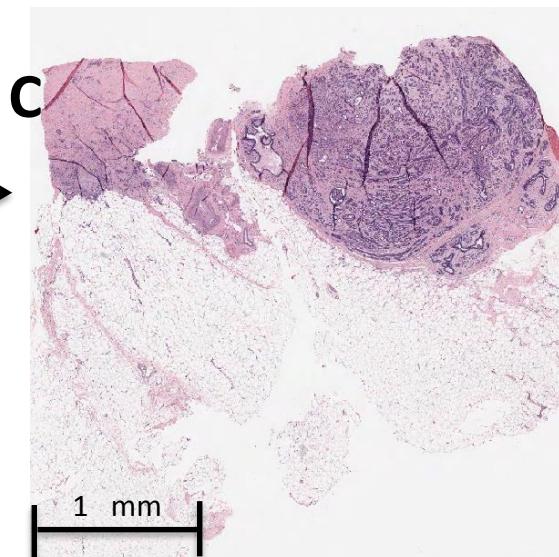
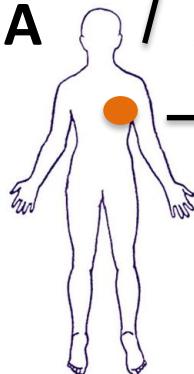


# Continuum of H&E, IHC and Multiplexed/Hyperplexed IF Computational Pathology Tools to Address Heterogeneity in Tissue Context

B



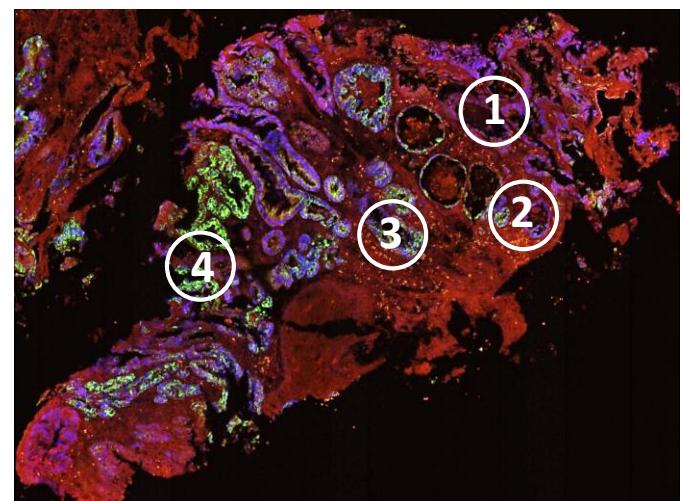
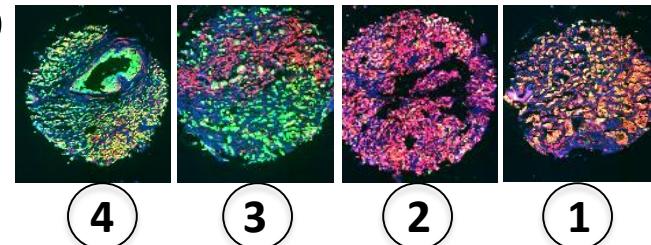
A



H&E stained whole tissue section from FFPE tumor sample

C

D

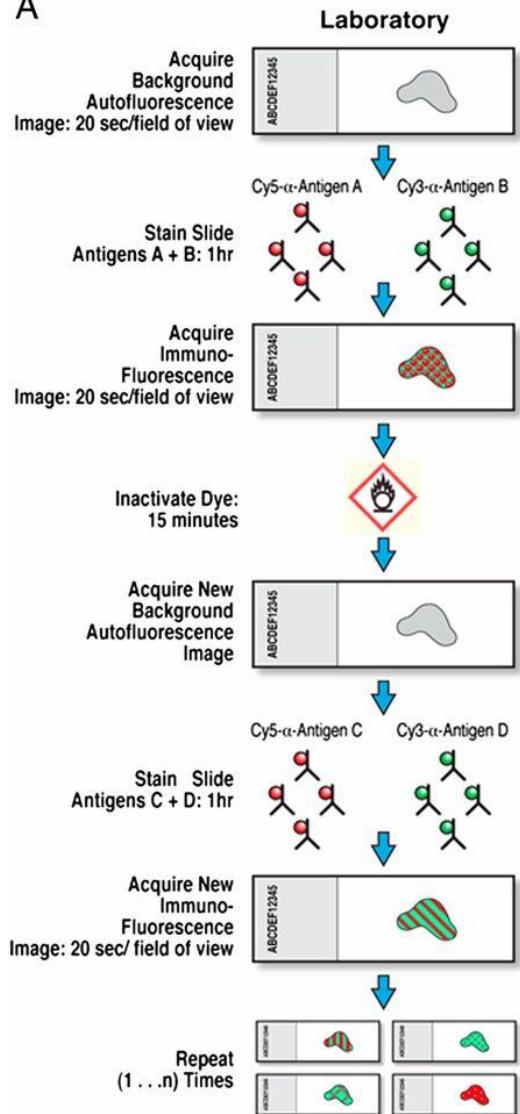


Multi to hyperplexed fluorescence imaging of whole<sup>3</sup> section for higher spatial resolution and tissue context

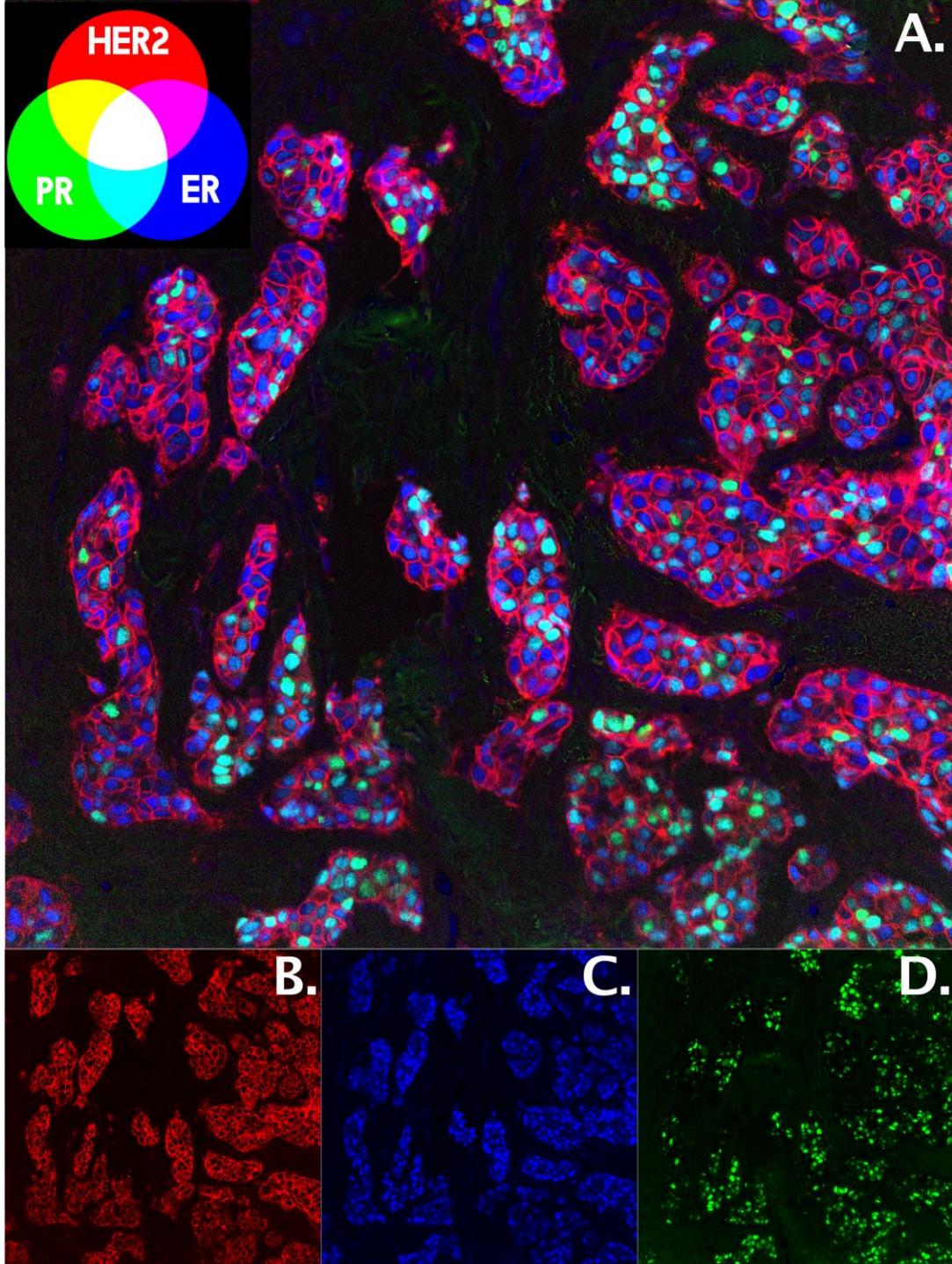
# Multiplexed IF

- **9 biomarkers**
- **Up to 60Ab**
- **Multiple FISH**

A



Gerdes et al. PNAS 2013;110:11982-11987



# Spatial Intra-Tumor Heterogeneity

1. Quantitate spatial intra-tumoral heterogeneity (ITH)
  - Quadratic entropy, Shannon index, Pointwise Mutual Information and others
2. Test spatial ITH metrics for correlation with tumor stage, grade and clinical outcome
3. Interaction with multiplexed/hyperplexed IF images and visualize spatial ITH

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