Intra-tumor Heterogeneity and Data-Guided Modeling of Cancer

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Introduction

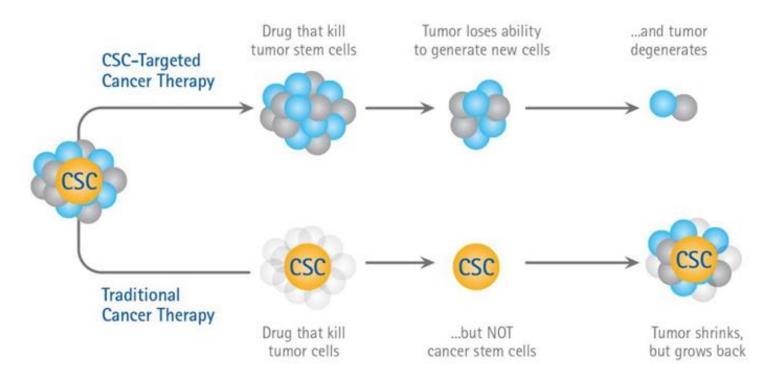
Intratumor heterogeneity

- ▶ Within a tumor, all cells originate from one mutated cell
- Cells within a tumor are not identical; they show genetic and nongenetic variability
- Sources of tumor heterogeneity:
 - Genetic: accumulation of mutations and evolution
 - ► Nongenetic: epigenetic, transcriptomic variations → affect cell state
 - Environmental: dynamic microenvironment influences tumor progression and clinical outcomes

Cancer Stem Cells

- CSC model: certain tumor cells have stem cell characteristics
 - Can self-renew or differentiate into cancer cells
- Where do CSCs originate?
 - Arise from mutated regular (tissue) stem cells
 - Cell state transition of tumor cells (dedifferentiation)
 - Epithelial-mesenchymal transition during metastasis
- Possible that stemness is a dynamic characteristic

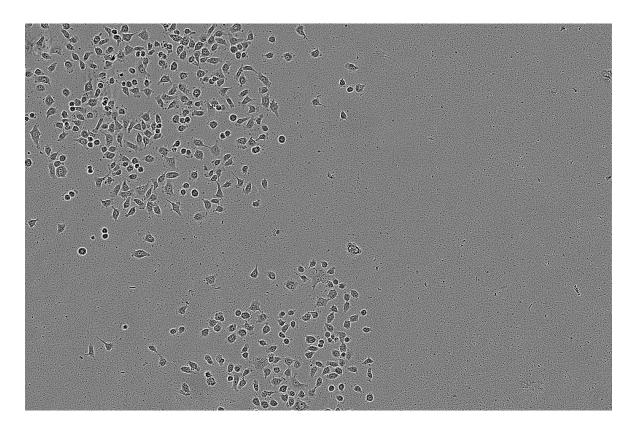
Cancer Treatment



Deeper understanding of CSCs could guide cancer treatment strategies

Research Goals

Investigate phenotypes and "stemness" of cancer cells during tumor growth, and possible changes over time



Acknowledgements

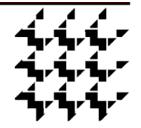
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Sources

- "Cancer Stem Cells: New Targets for Cancer Therapy." Sigma-Aldrich, www.sigmaaldrich.com/technical-documents/articles/biofiles/thecancer-stem-cell.html#ref.
- Biswas A, De S. Drivers of dynamic intratumor heterogeneity and phenotypic plasticity. Am J Physiol Cell Physiol 320: C750-C760, 2021. doi: 10.1152/ajpcell.00575.2020
- Gross, Emilie T.E. "Are Cancer Stem Cells a Prime Target for Therapy?" CrownBio Blog, 22 Nov. 2018, <u>https://blog.crownbio.com/cancer-stem-cell-models</u>.