Unraveling Breast Cancer Heterogeneity: Microfluidic Sorting and Bioassay-Based Functional Analysis

Esra Yilmaz^{1,*}, Zhimeng Fan², Jason P. Beech¹, Vinay Swaminathan² and Jonas O. Tegenfeldt¹
*esra.yilmaz@ftf.lth.se





OncobioMech Team

¹Division of Solid<mark>-State</mark> Physics, Physics Department and NanoLund, Lund University

¹²Department of Clin<mark>ical</mark> Sciences, Faculty of Medicine and NanoLund, Lund University

Current problems

Metastasis causes 90% of cancer deaths.

Alteration in the cancer cell mechanics correlate with disease state.

Solution

Breast cancer cells can be sorted into sub-populations as **small** and **large** based on size.



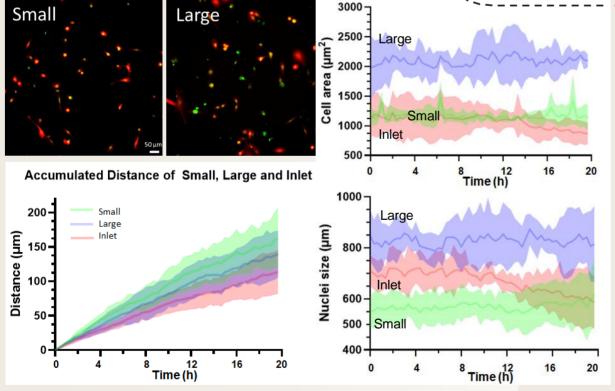
Method: Deterministic lateral displacement (DLD)

Buffer inlet Bu

№
 №
 №
 №
 №
 №
 №
 №

Findings II

Small Cells Exhibit Enhanced Migration on Basement Membrane Extract (BME)



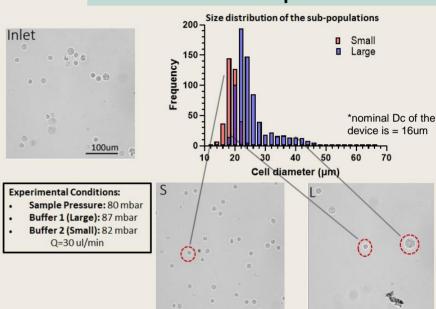
- Small breast cancer cells showed significantly greater migration on BME over 20 hours compared to large cells.
- Nuclei size and cell area confirmed persistent morphological of differences between subpopulations as well as inlet.

Our research goal

- •Characterizing cancer cells by their mechanical properties, which often relate to disease fate.
- •Exploring subpopulation functionality and their role in cancer metastasis.

Finding I

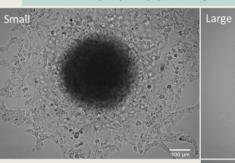
Size-Based Sorting Reveals Distinct Cancer Cell Populations

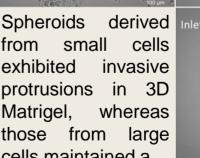


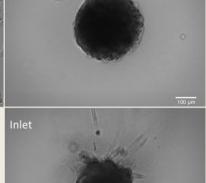
•Small cells display a narrow, uniform size range whereas large cells show broader heterogeneity, including clusters.

Findings III

Small-Cell Spheroids Show Invasive Behaviour in 3D Matrigel







compact and organized structure, highlighting functional differences in invasiveness.

This suggests higher invasive potential in the small-cell subpopulation.



Future work will examine patient-derived circulating tumor cell deformability under varying pressures and their long-term behavior related to proliferation and motility.



The analyses will comprise qRT-PCR, antibody assay, and long-term time-lapse imaging to deepen our understanding of the cancer cascade.

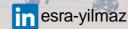
Acknowledgments



This research received funding from the Swedish Research Council (grant number 2019-02355) the European Union's Horizon 2020 research and innovation programme under the Marie Skłodowska-Curie grant agreement. All device fabrication was done in Lund NanoLab.

Esra Yilmaz

Doctoral Researcher Tegenfeldt Research Group Nanolund, Lund University, Sweden





SMILS 16-17 June 2025, Lund, Sweden