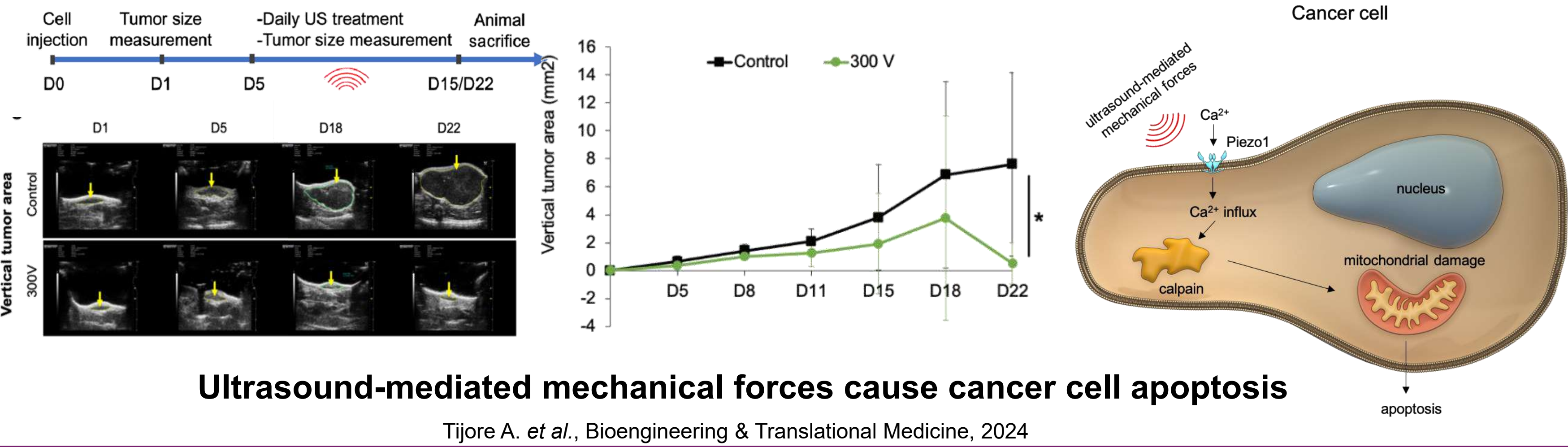


Biomechanical Vulnerability of Oral Cancer Cells Enables Low-Frequency Ultrasound Treatment

Rashmita Luha¹, Gomathi Sankar¹, Akshay Kumar², Alka Kumari¹, Ketan Kulkarni¹, Rudra Pratap², Aravind Kapali³, Ajay Tijore¹

¹Department of Bioengineering, Indian Institute of Science, Bangalore, India, ²Centre for Nano Science and Engineering, Indian Institute of Science, ³Department of Surgical Oncology, M. S. Ramaiah Medical College & Hospitals
Email: rashmitaluha@iisc.ac.in

Introduction



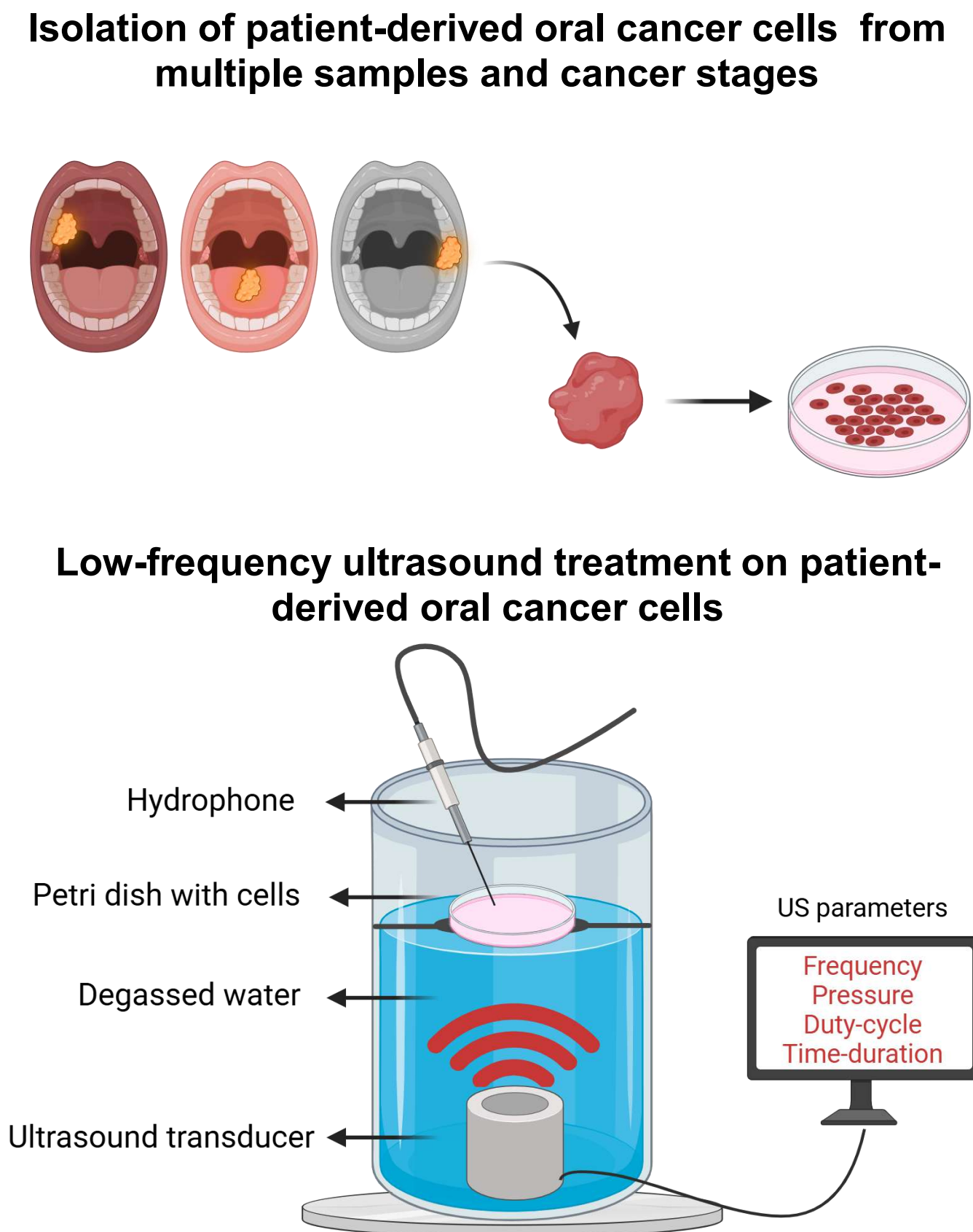
Key features of low-frequency ultrasound for cancer therapy

- Personalised
- Selectively induces apoptosis in cancer cells without damaging normal cells
- No thermal ablation

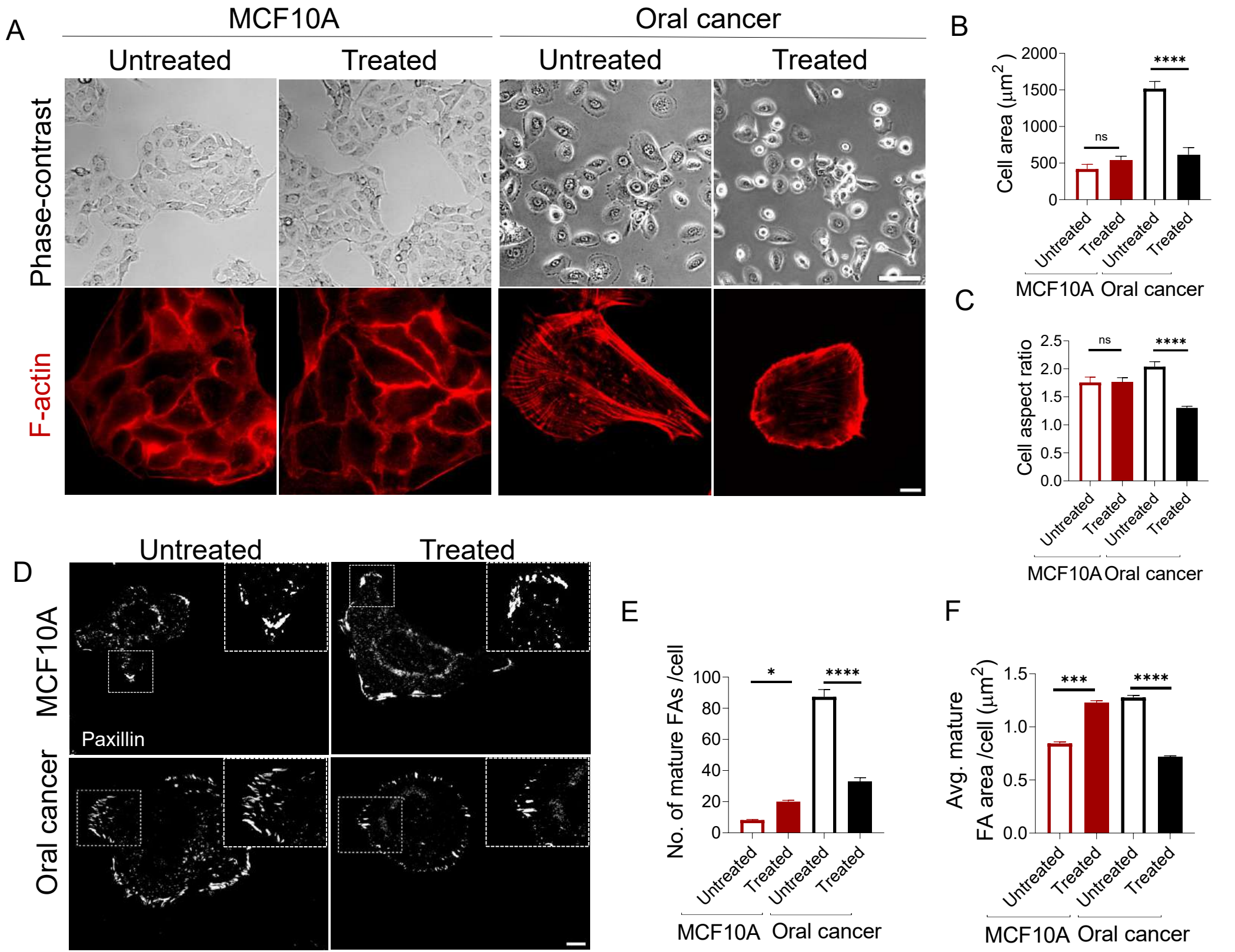
Research Aims

1. Investigate the effect of low-frequency ultrasound (LFU) on patient-derived oral cancer cell survivability and invasion.
2. Develop a biomaterial-engineered 3D coculture platform to evaluate the effect of LFU.

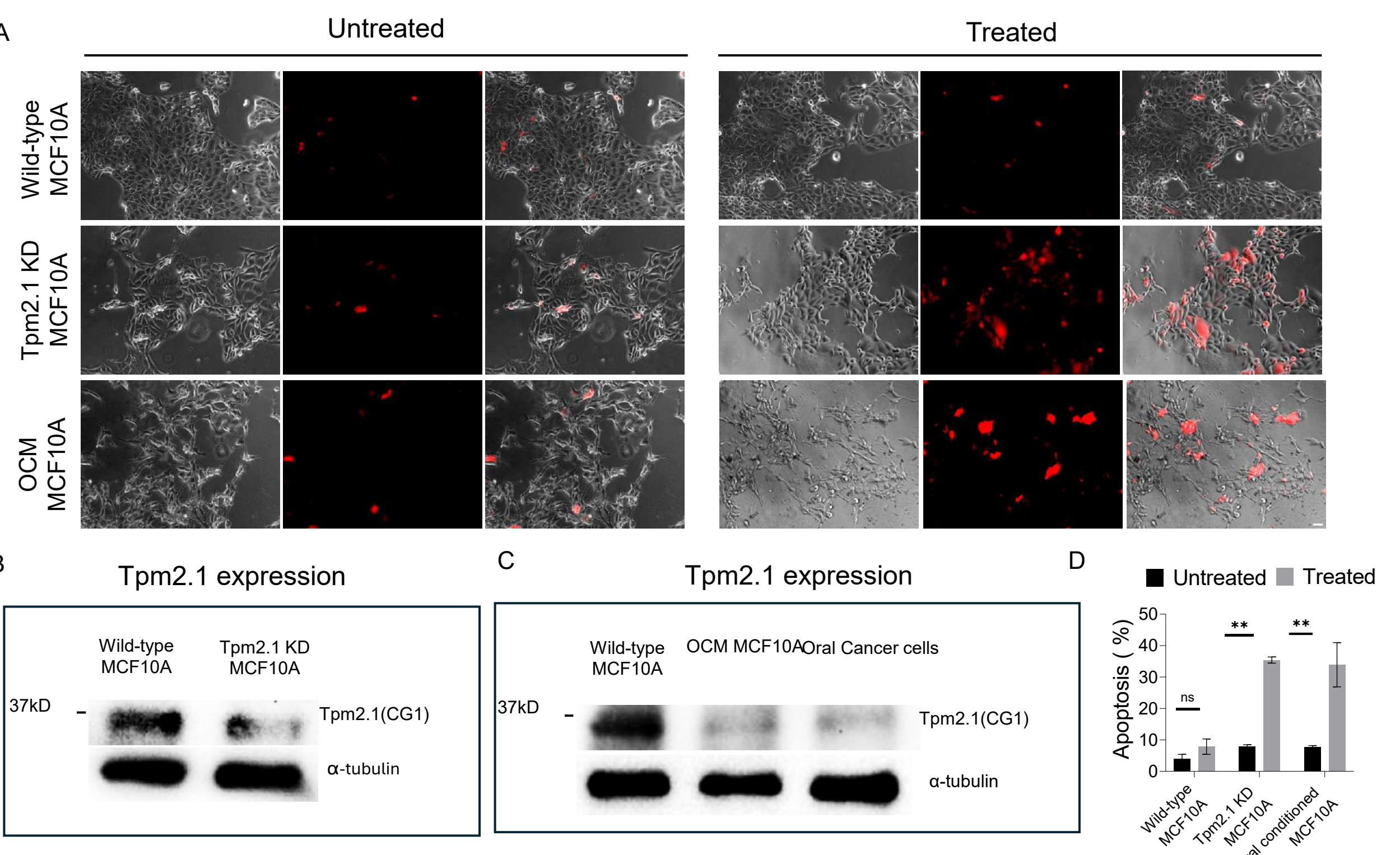
Methodology



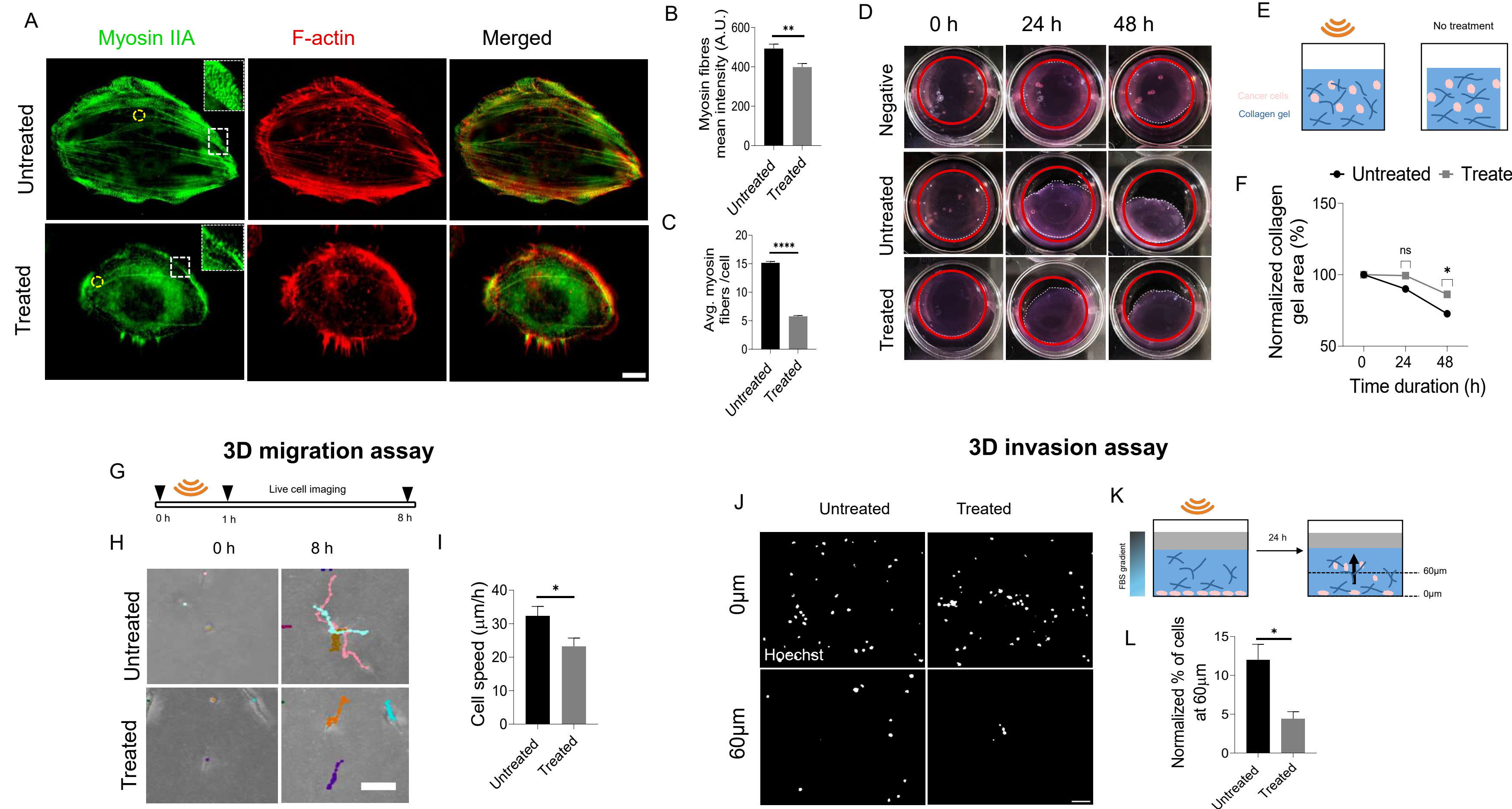
1. Low-frequency ultrasound reduces cancer cell spreading by disassembling focal adhesions



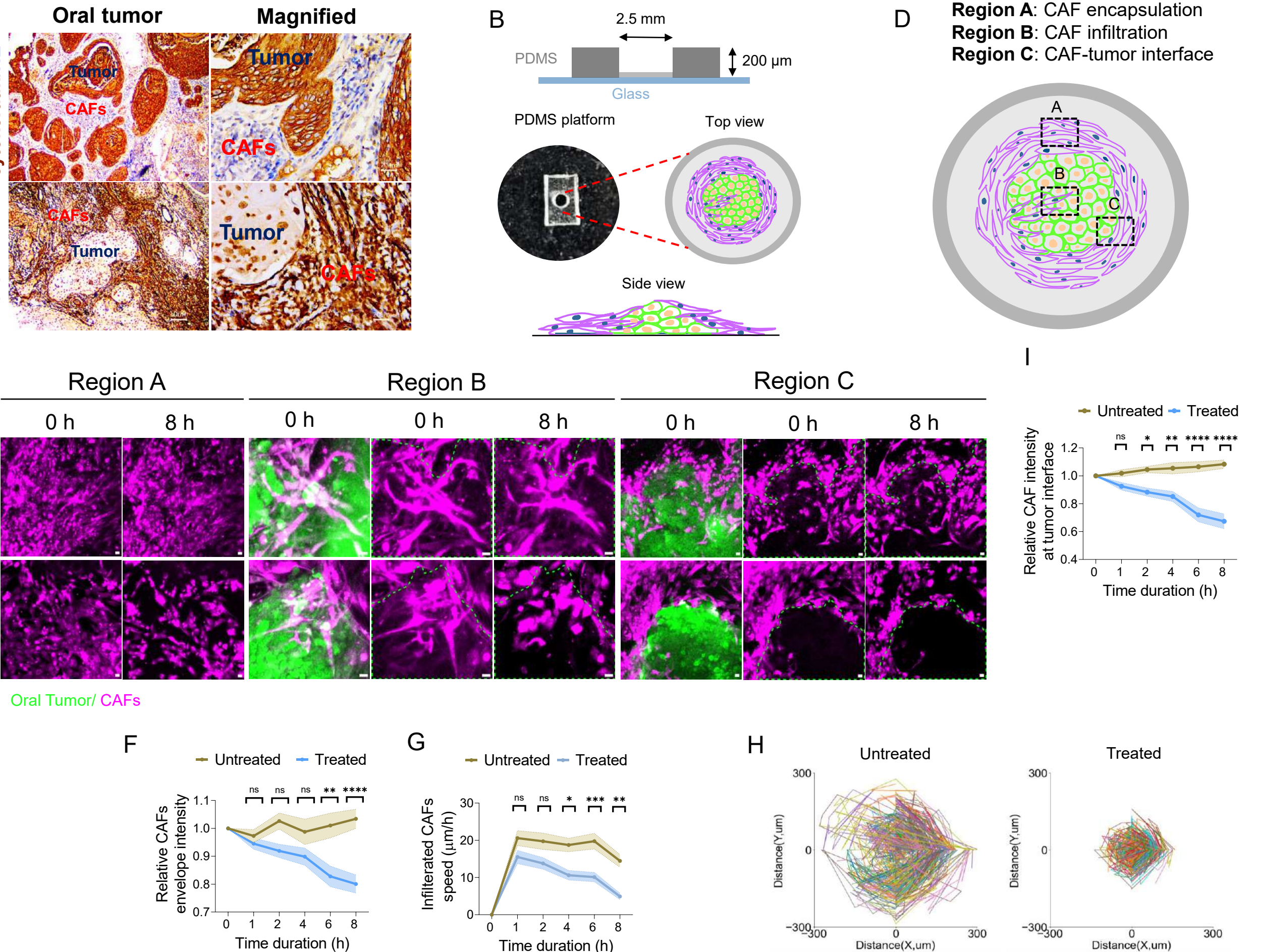
4. The biomechanical vulnerability of oral cancer cells to US treatment is due to the mechanosensory protein Tropomyosin 2.1 (Tpm2.1)



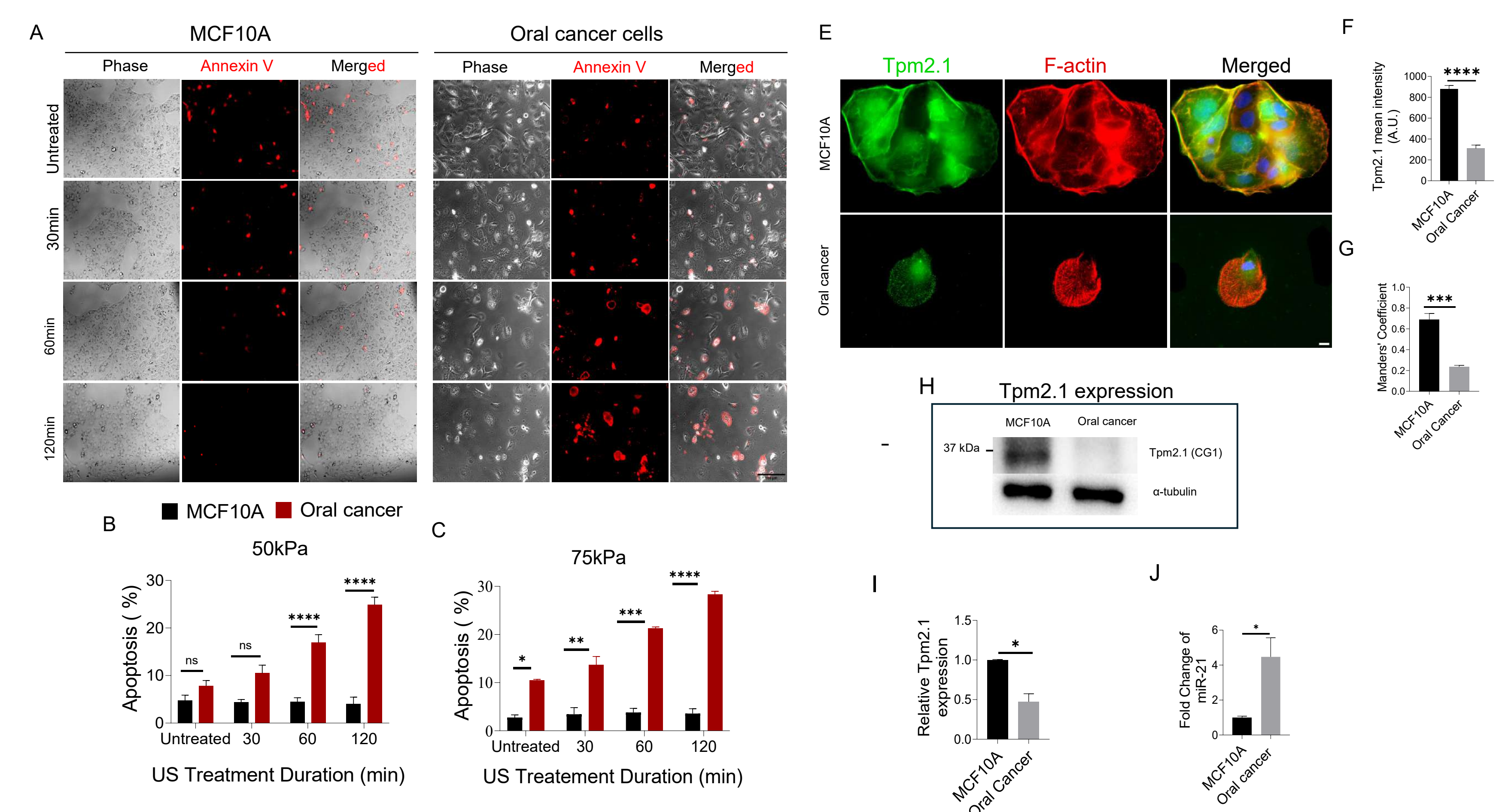
2. Low-frequency ultrasound suppresses oral cancer cell migration and invasion by disrupting contractility



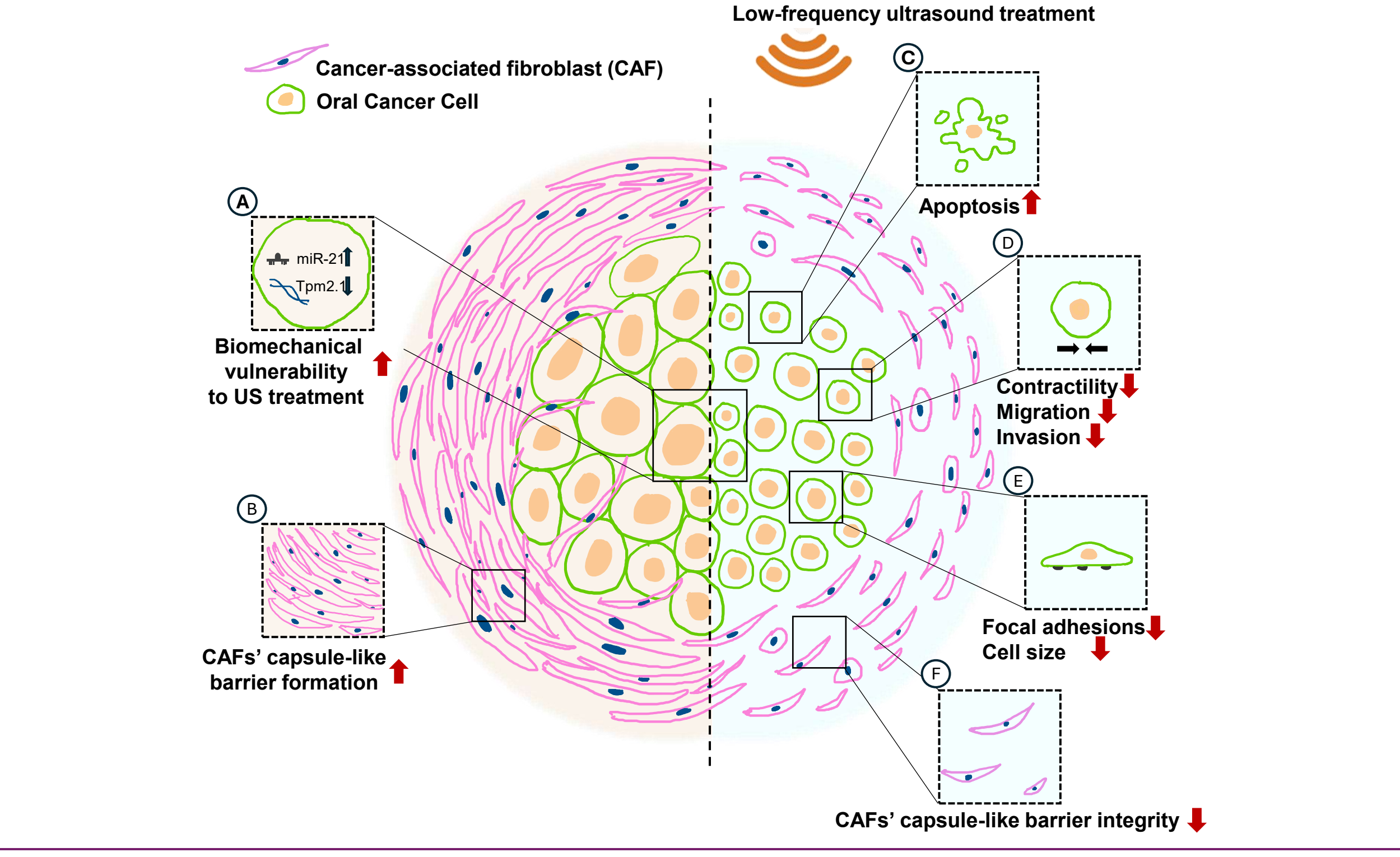
5. Ultrasound treatment disrupts *in vivo* mimicking oral cancer cell-CAF 3D organization



3. Low-frequency ultrasound causes mechanoptosis in patient-derived oral cancer cells without damaging normal cells.



Summary



Significance: The selective killing of cancer cells without damaging normal cells, along with the disruption of the stromal barrier surrounding the tumor core, compromises the CAF's capsule-like barrier integrity, highlight the potential of developing novel US-mediated therapies