

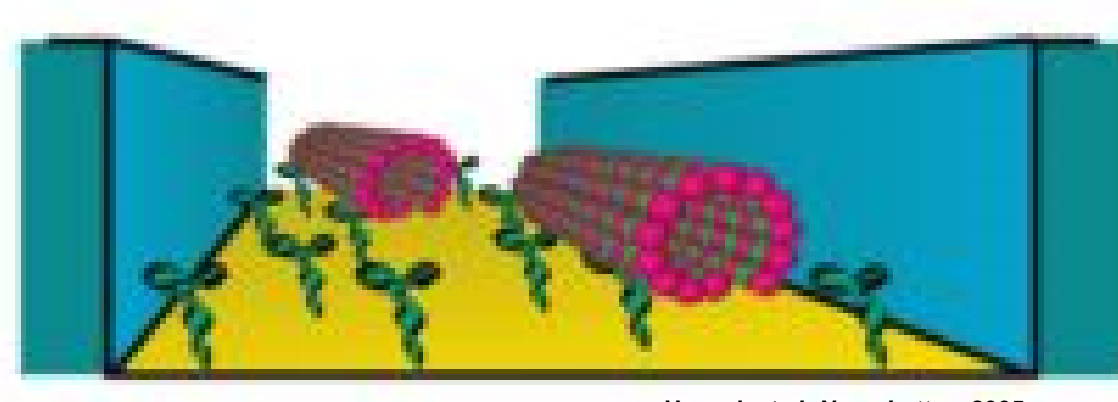
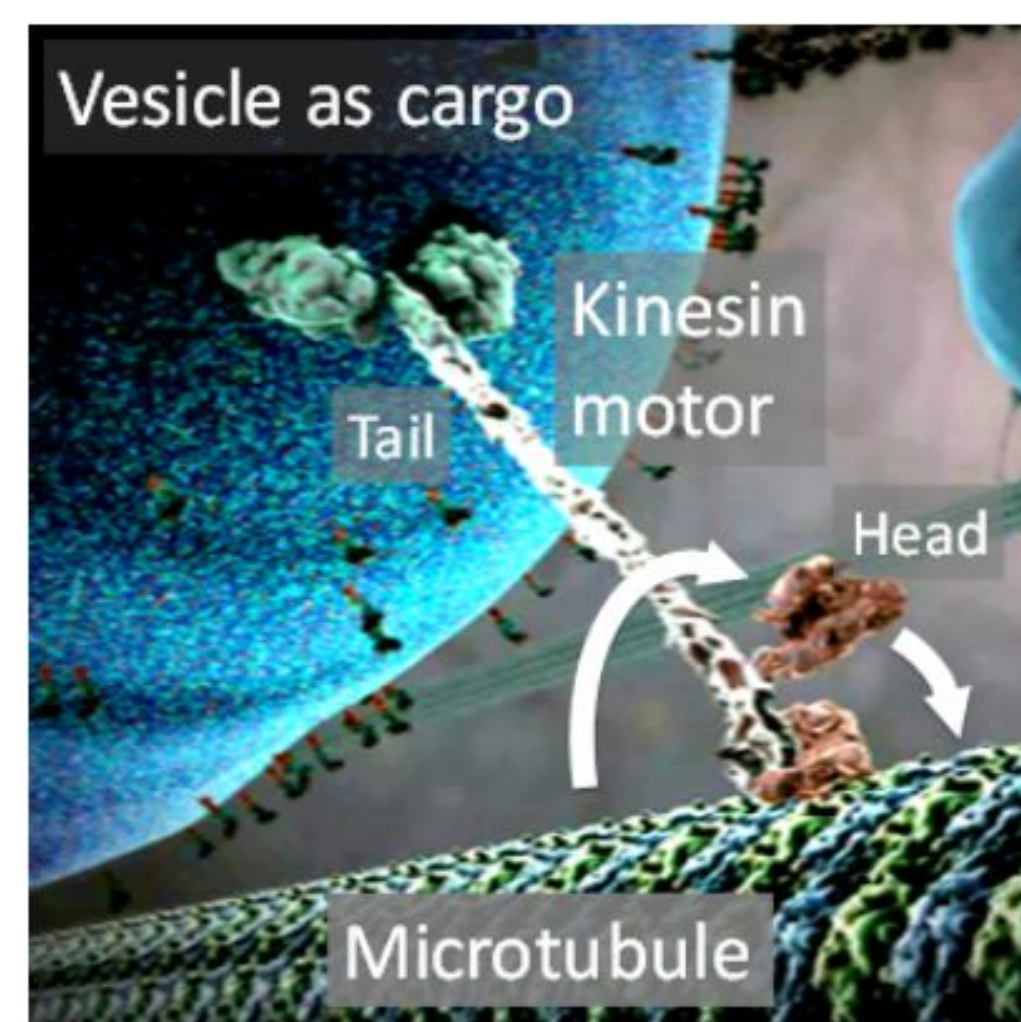
Microtubule Transport on 3D Biocompatible Nanostructures

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Introduction

- Microtubules (MTs) are cytoskeletal protein filaments that provide mechanical support for the cell, and serve as “tracks” for motor proteins to transport organelles
- Kinesin is a microtubule-based motor protein that “walks” along MTs by dissipating chemical energy, with a force of $\sim 40\text{pN}$ nm and efficiency of $\sim 50\%$ ¹
- Kinesin-MT transport system has been used in many nanotechnological applications including biosensing², cargo transportation³, and assembly of ring nanocomposites⁴

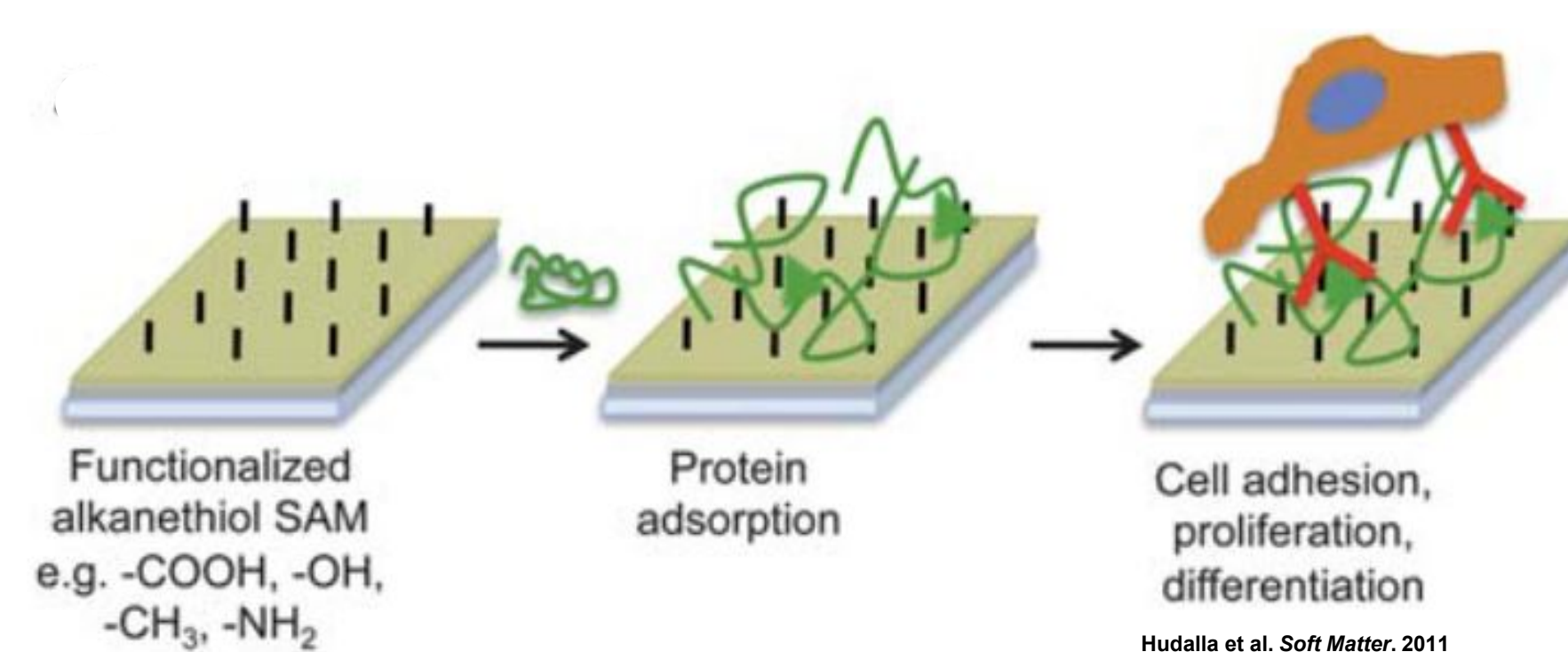


- MT guiding using lithographically nanostructured surfaces hinder MT motility and lead to MT loss⁵

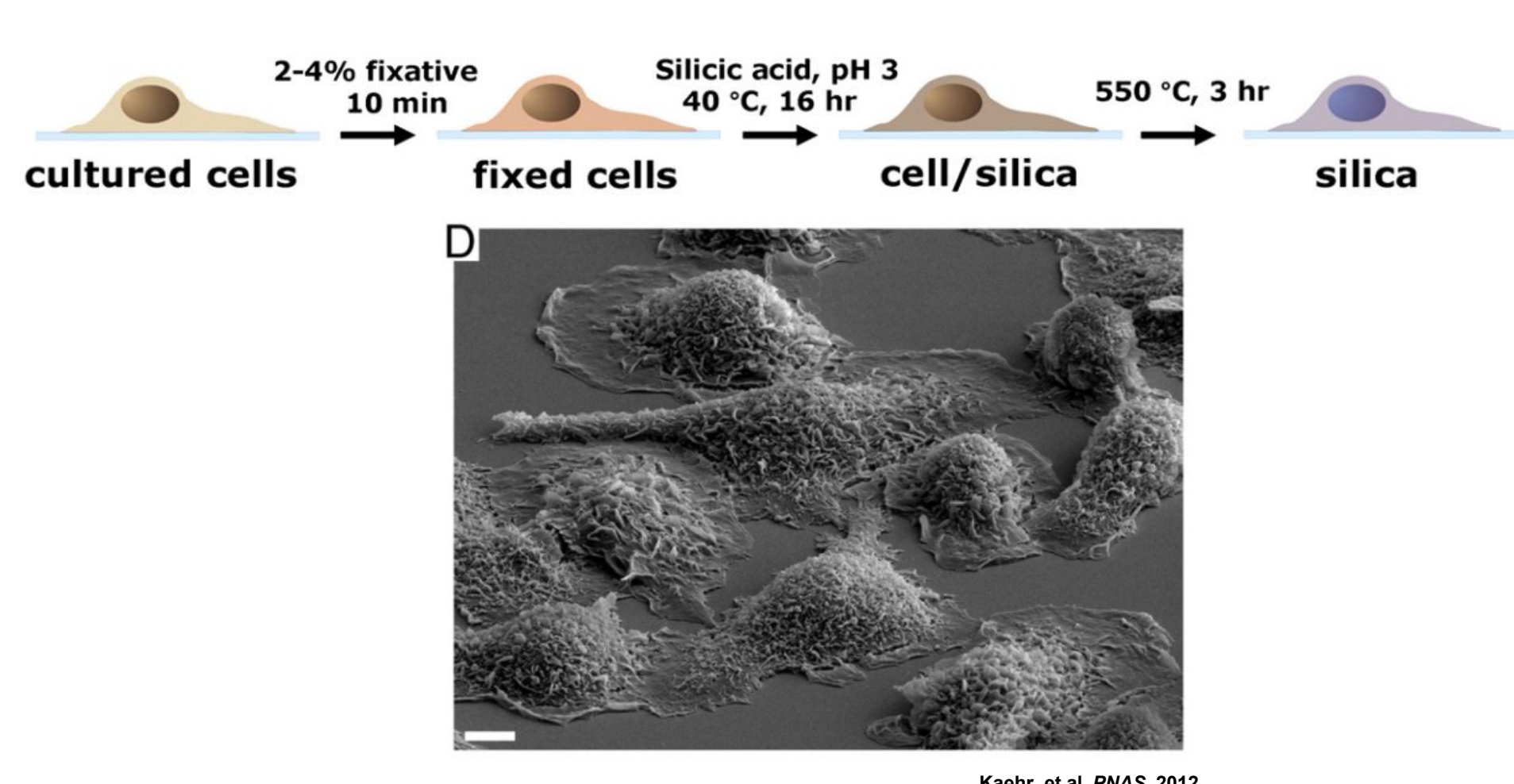
Exploring alternative nanostructures is essential for reliable MT guiding

Cell patterning and preservation

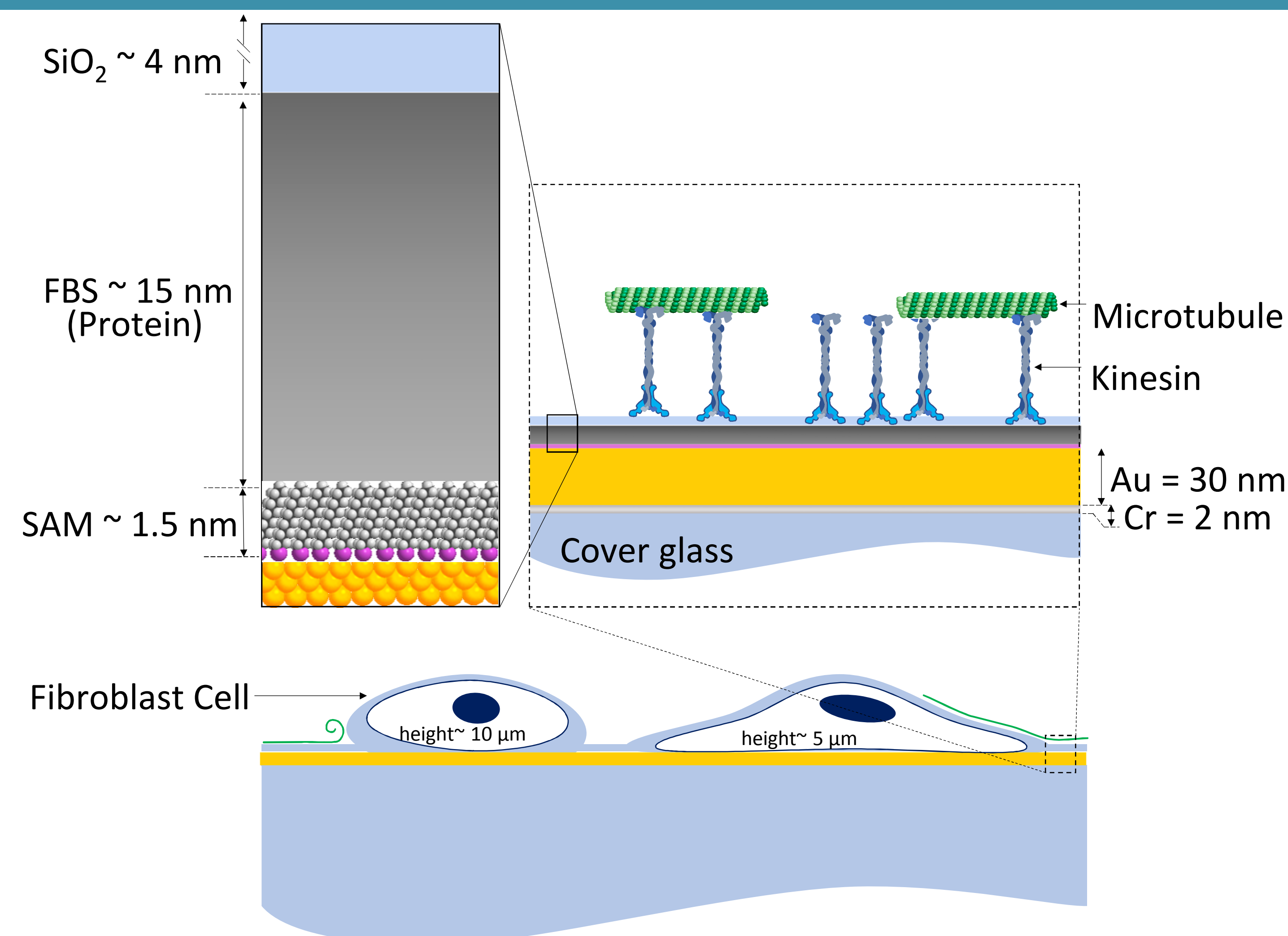
- Self-assembled monolayers (SAMs) previously used to modulate cell adhesion and spreading, allowing for size and shape control of cell patterns
- Limitation:** environmental conditions render cells unstable for long-term applications



- Preservation of cellular architecture through silicification process
- Preserve user-defined 3D features
- Provides simple alternative to specimen preparation and preservation (no expertise or specialized equipment needed)
- Tolerate extreme environments

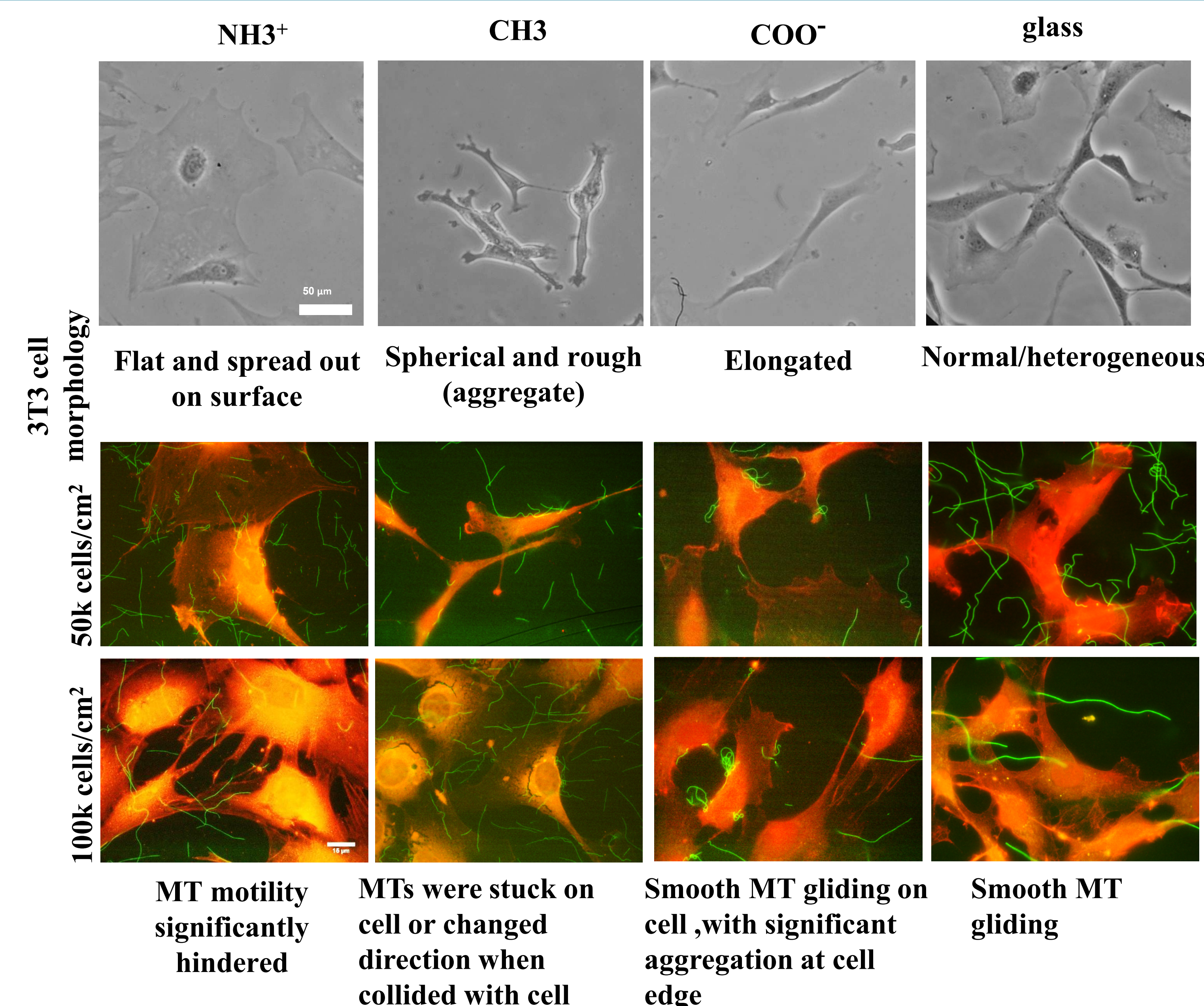


Approach

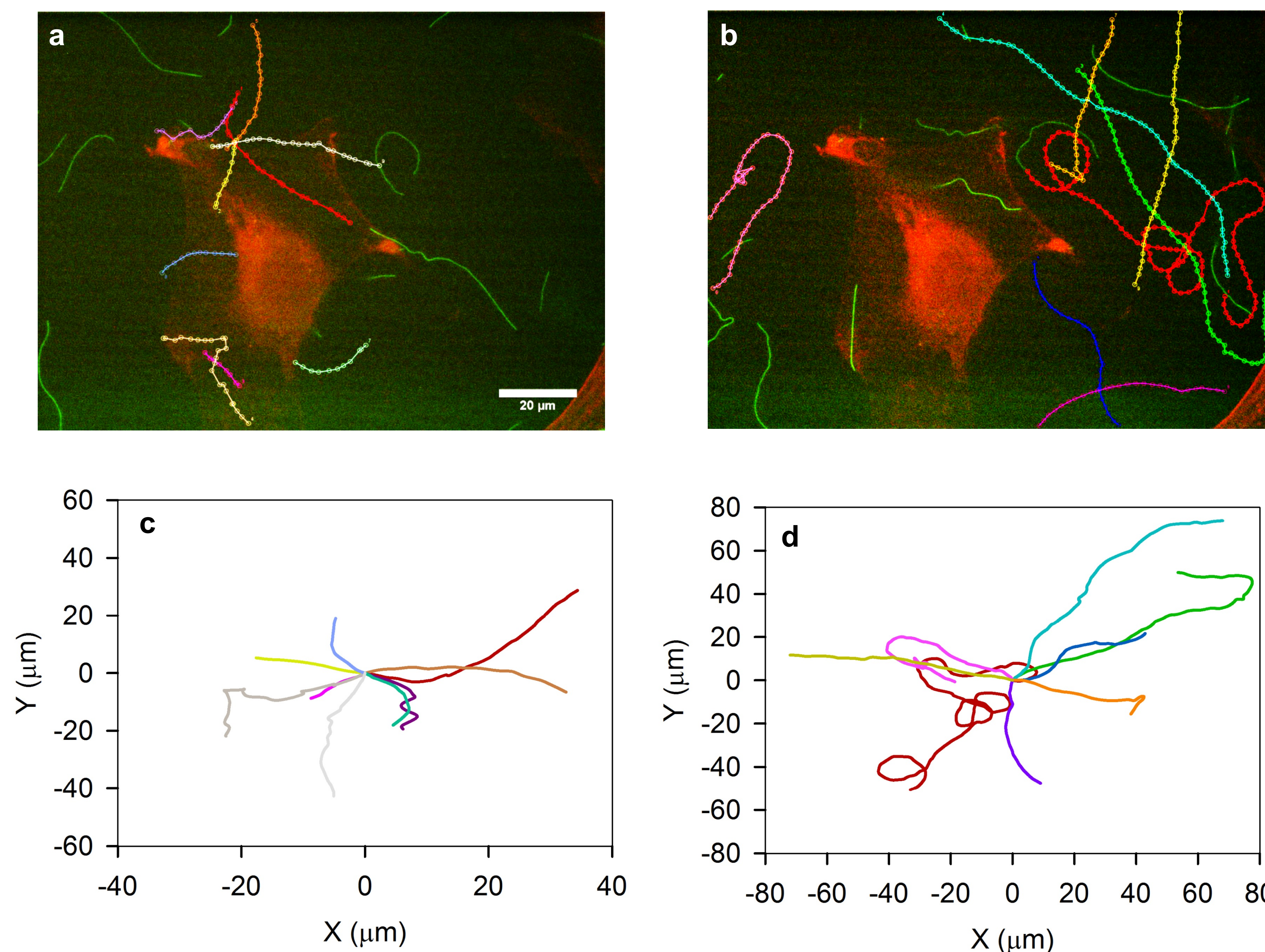


- Self-assembled monolayers (SAMs) with various functional groups are used to alter mammalian cell morphology
- Cells are preserved through silicification process, and used as 3D nanostructures to explore the behavior of kinesin-MT system

Effect of SAMs and cell confluency on morphology and MTs



Microtubule trajectories and velocity



- MT gliding trajectories were evaluated using carboxyl (COOH) terminated SAMs and low cell count (50k cells/cm²)
- MT trajectories were linear on silicified cells (a,c), while curved trajectories were observed on silicified SAMs surface (b,d)
- Gliding velocity remained constant with an average of 1 μm/s, independent of surface

CONCLUSIONS

- We established a unique “bottom-up approach” by combining well-established techniques to generate preserved, 3D biocompatible structures dictated by SAMs and cell confluency
- Preliminary experiments show promising results of surface topographies influencing MT translocation
- Future experiments will provide insight into the development of applications involving Kinesin-MT transport on complex nanostructures

References:

- [1] Liu et al., (2008). *Adv. Mater.* **20**, 4476-4481
- [2] Fischer et al., (2009). *Nat. Nanotechnol.* **4**, 162-166
- [3] Brunner et al., (2007). *Lab Chip*, **7**, 1263-1271
- [4] Lam et al., (2014). *Soft Matter*, **10**, 8731-8736
- [5] Korten et al., (2016). *IEEE Trans. Nanobioscience*, **15**, 62-69

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