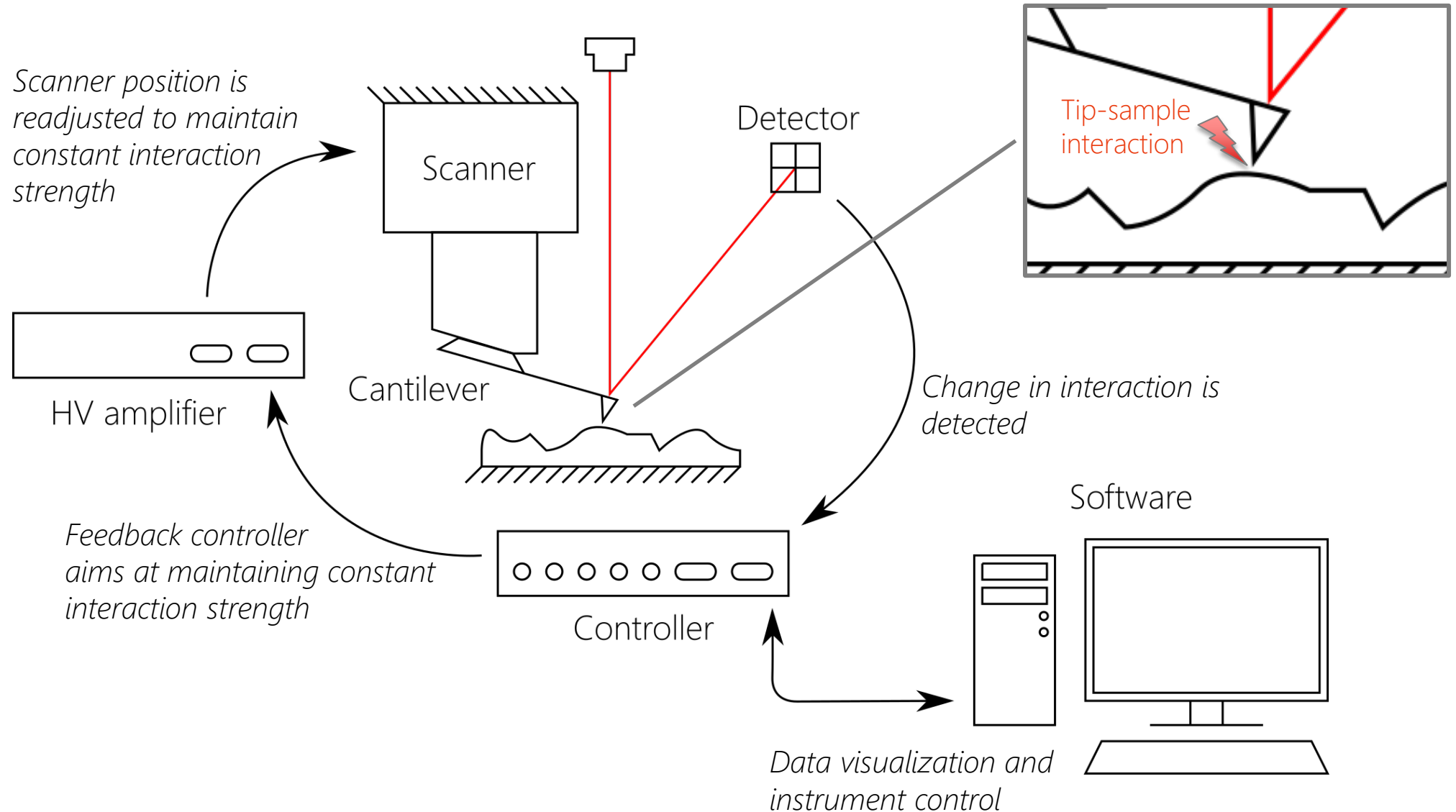


# Mass and viscoelasticity of single cells

Dr. Marco Portalupi, Nanosurf AG

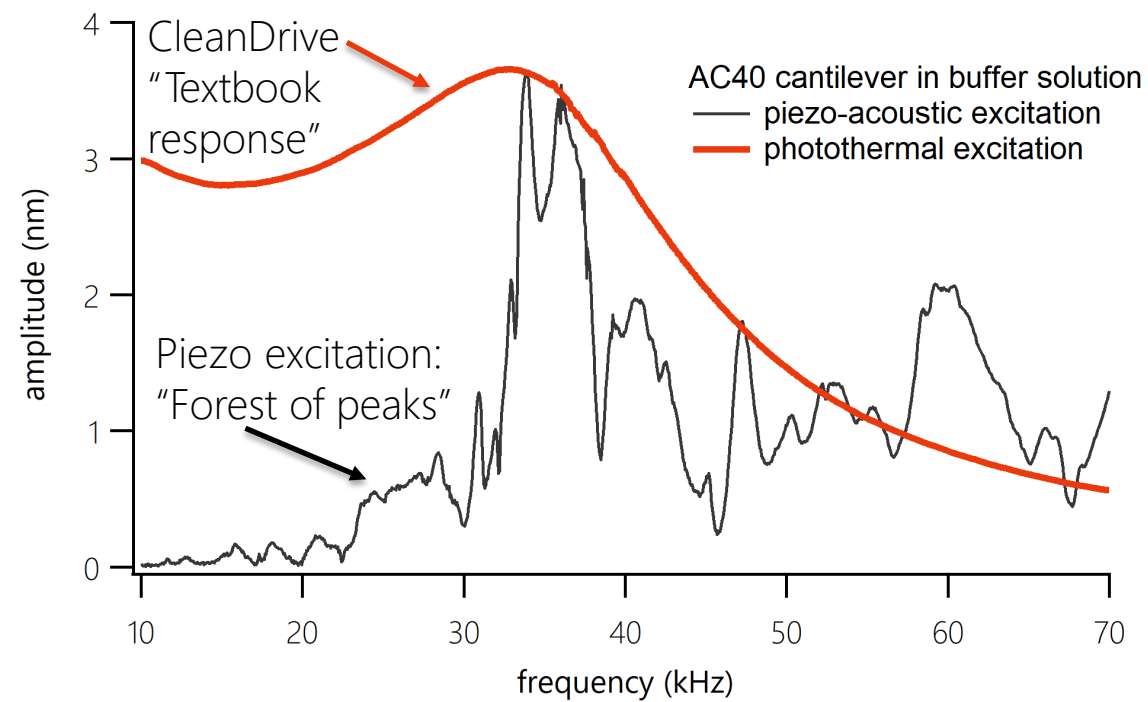
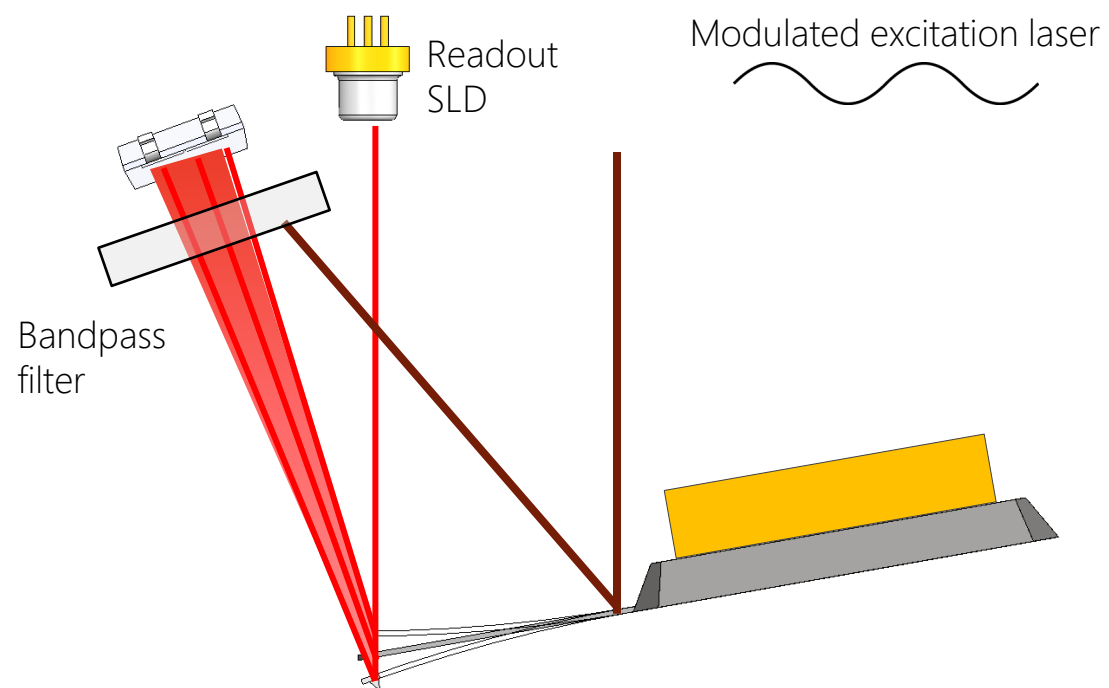


# AFM – how does it work?

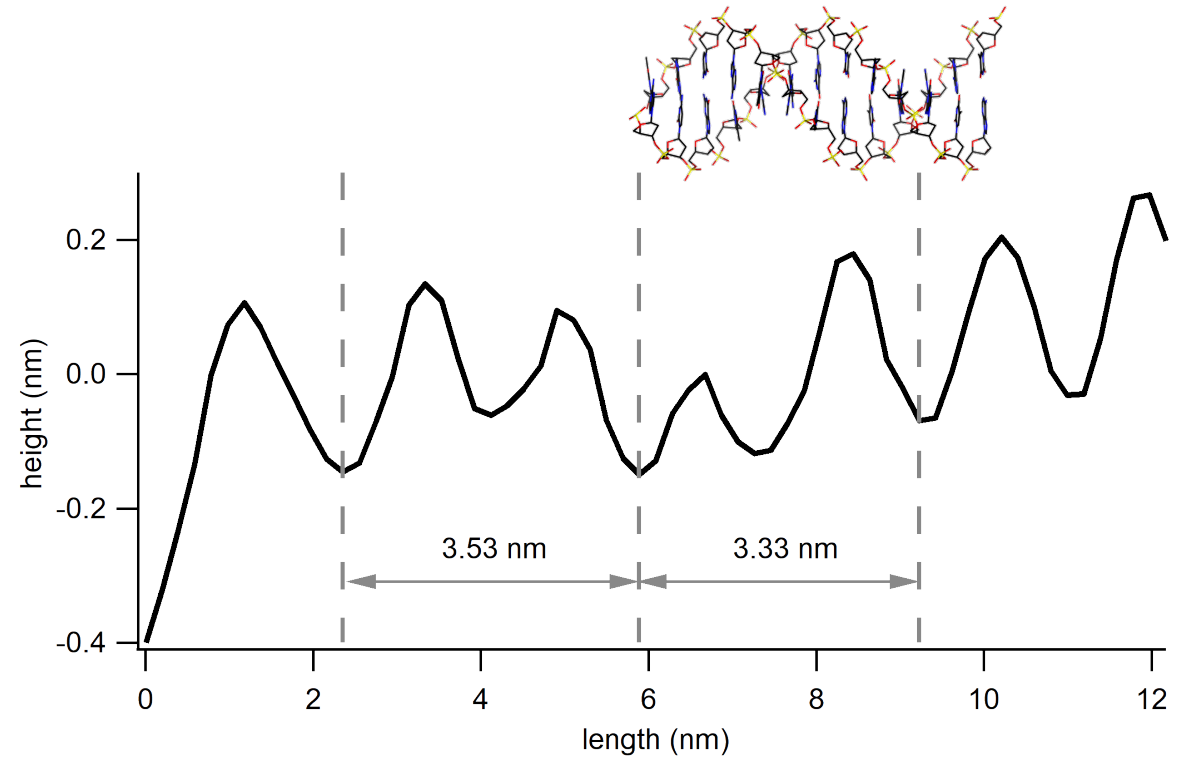
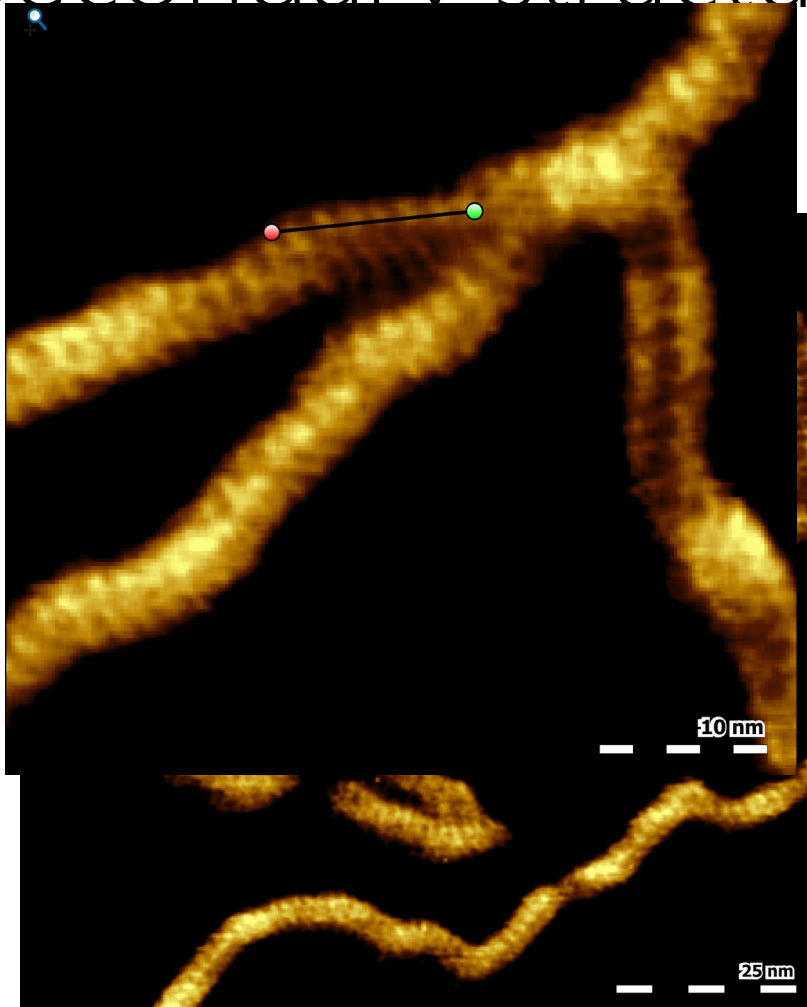


# CleanDrive photothermal excitation

Bio-compatible 785nm NIR laser source  
No "forest of peaks" in any environment  
Reliable automatic tuning  
Ultra stable excitation



# Secondary structure of dsDNA





# DriveAFM as a table-top system...

Stage mounted on Isostage 300 active vibration isolation



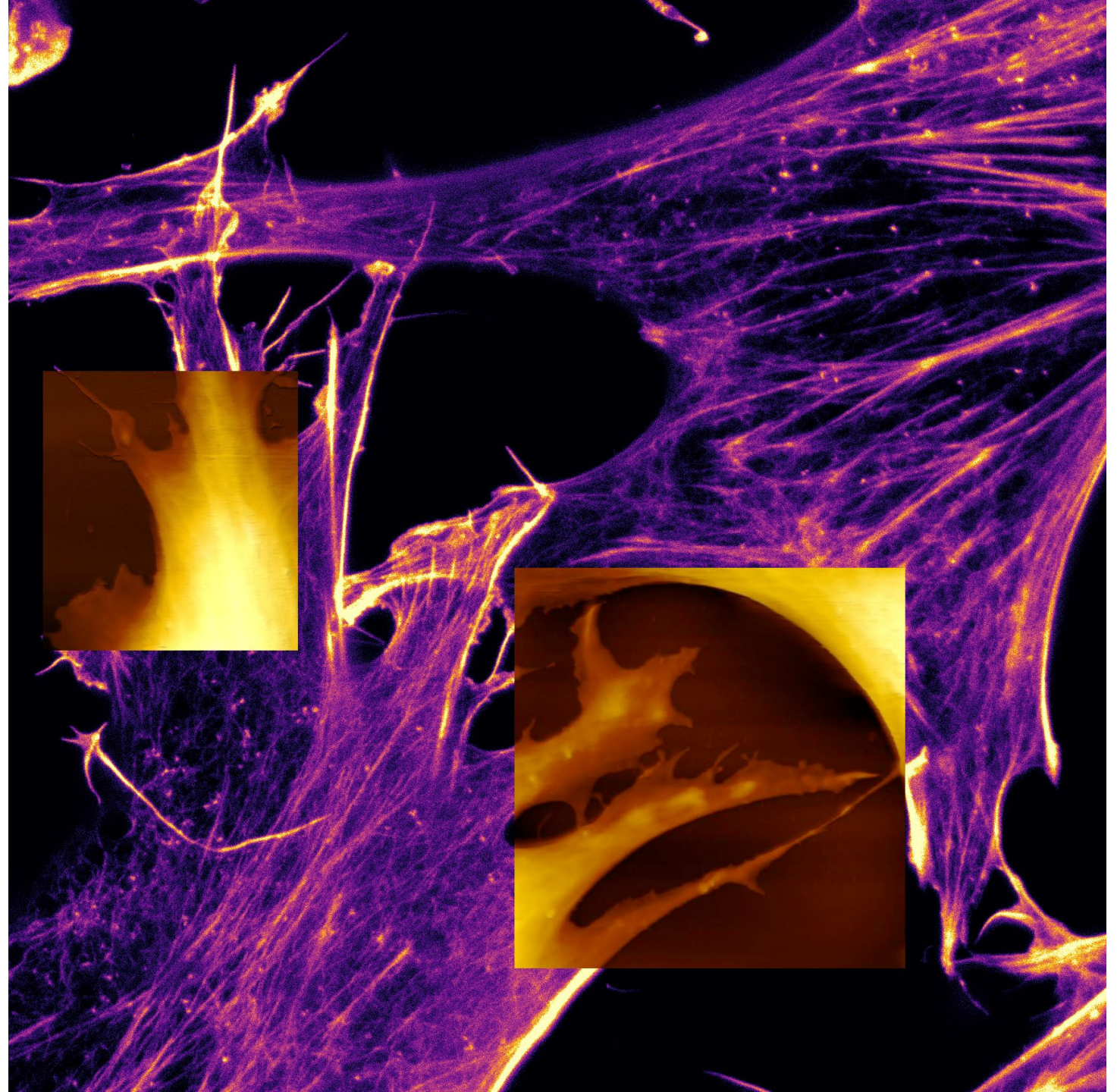
## ...or on an inverted optical microscope

A variety of stages for different inverted optical microscope brands is available.

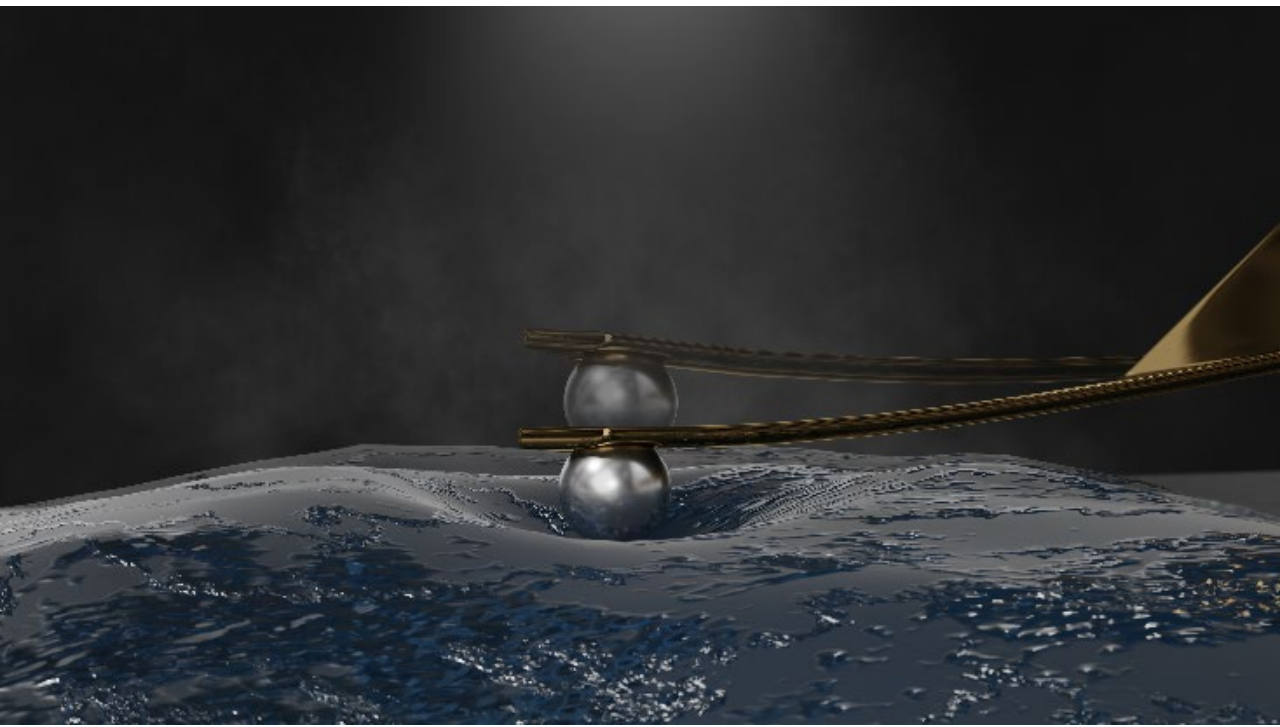
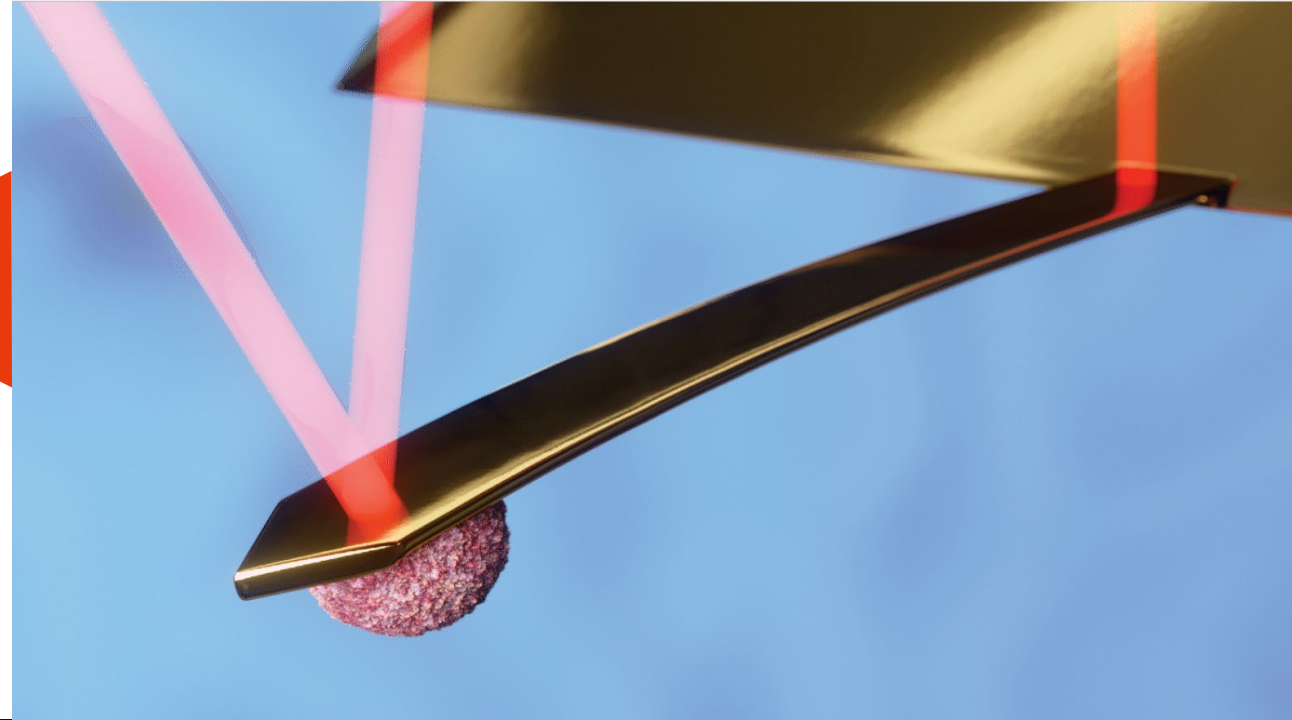
# AFM + optical microscopy

Super-resolution techniques  
challenge AFM in the biosphere

Mechanical interaction → other  
insights than just structure



Mass measurements

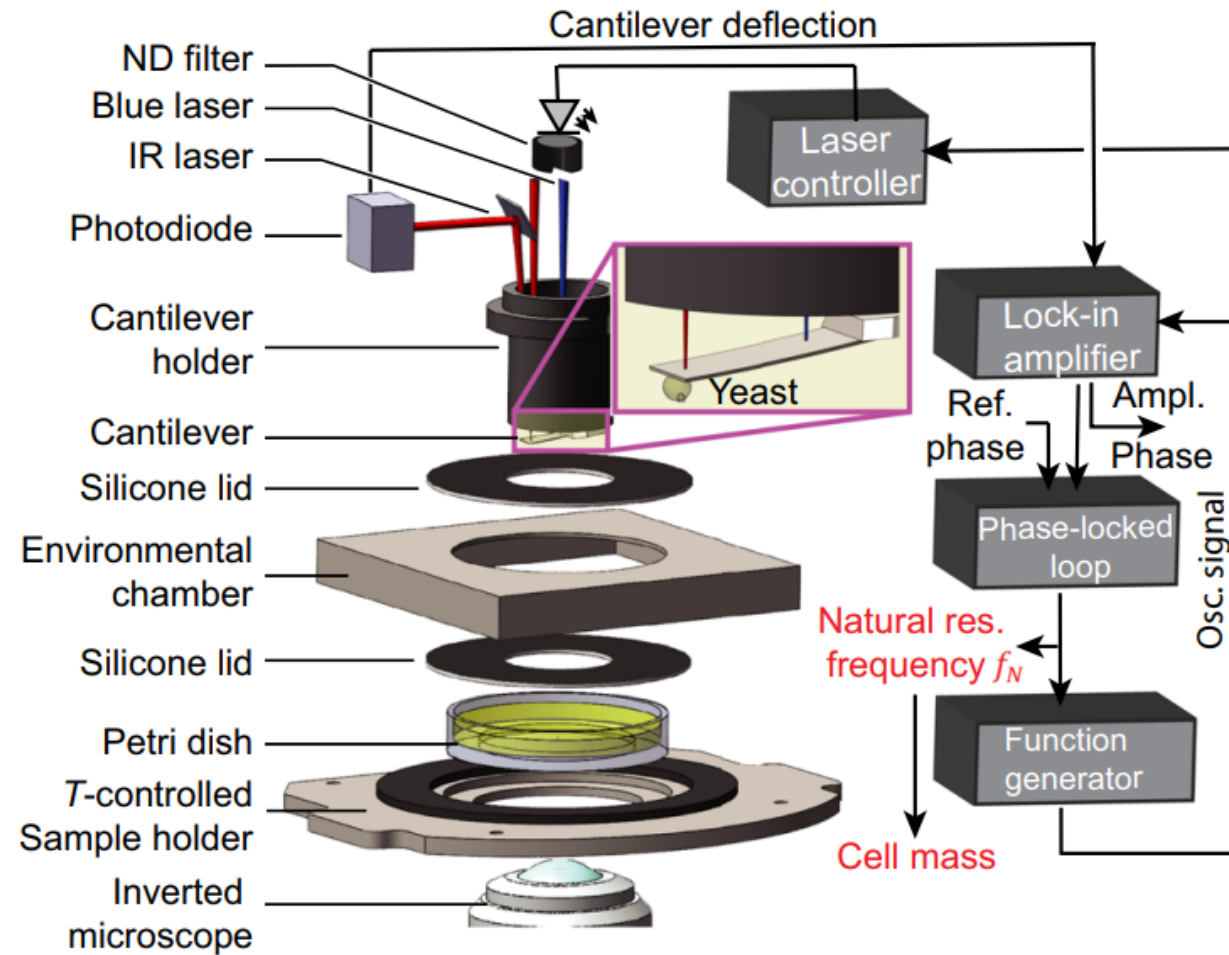


Viscoelasticity measurements

# Mass measurements of single yeast cells

Advancing the original technology  
(Martinez-Martin & Fläschner et. al. Nature 2017)

Increasing mass resolution,  
improving optical microscopy



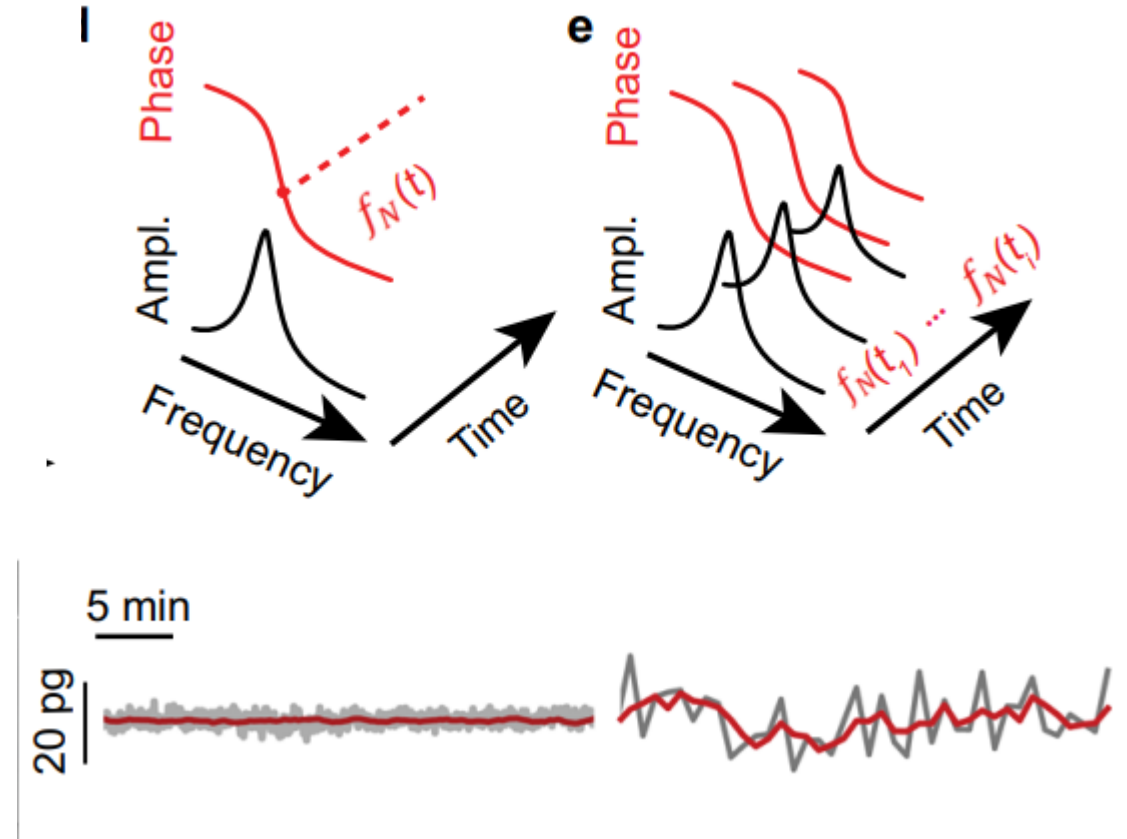
Cuny, Sapra, Martinez-Martin, Fläschner *et al.*,  
Nat. Commun. 2022



# Mass measurements of single yeast cells

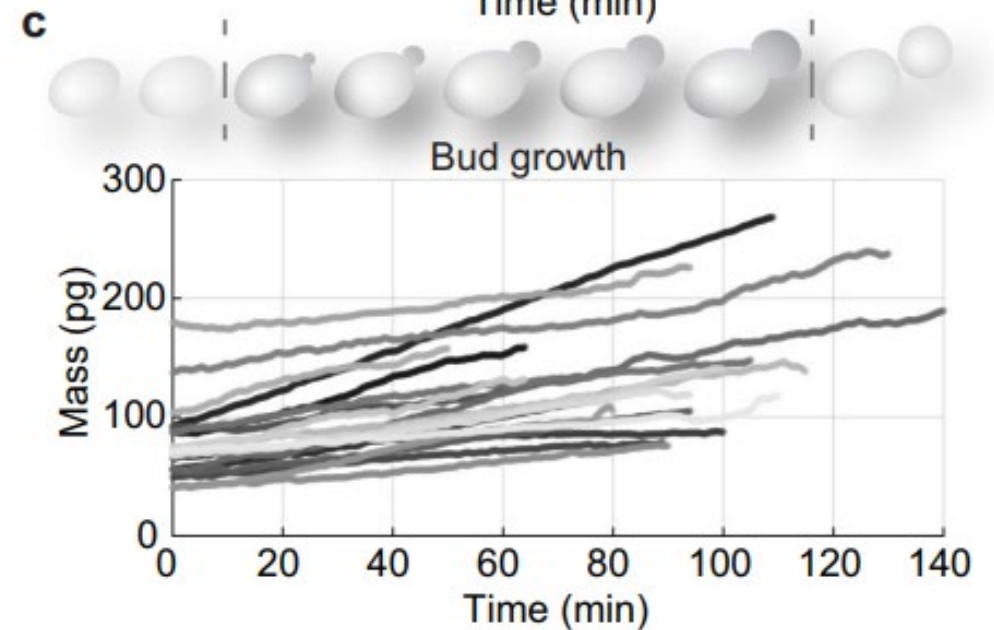
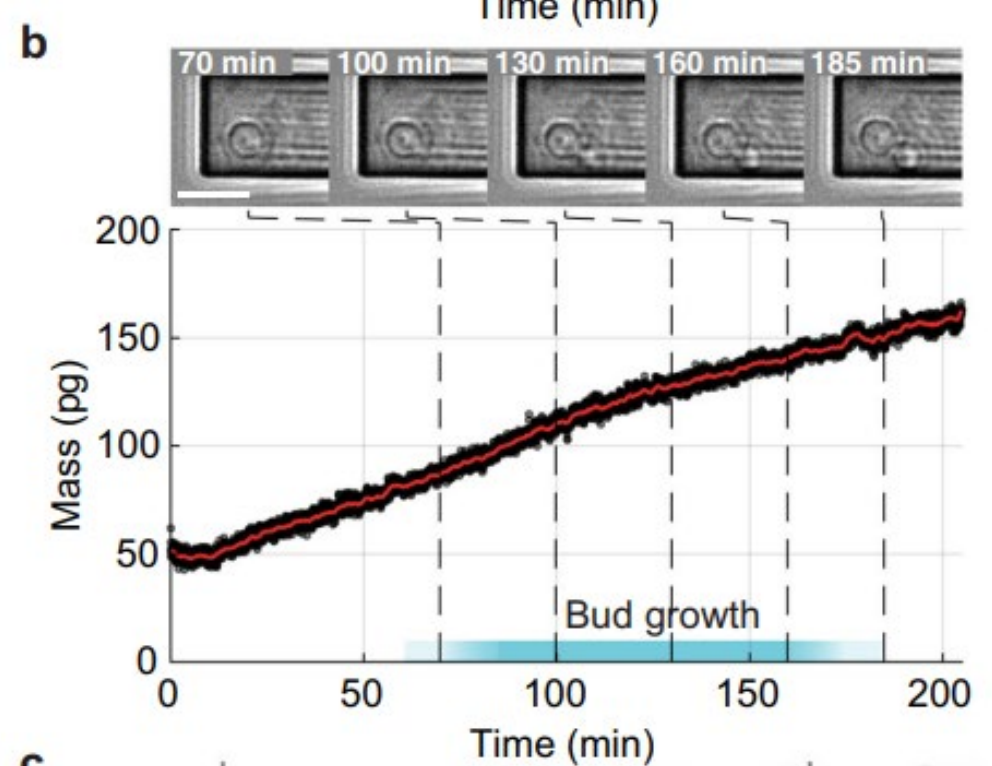
Two measurement modes:

- Continuous mode
- Sweep mode



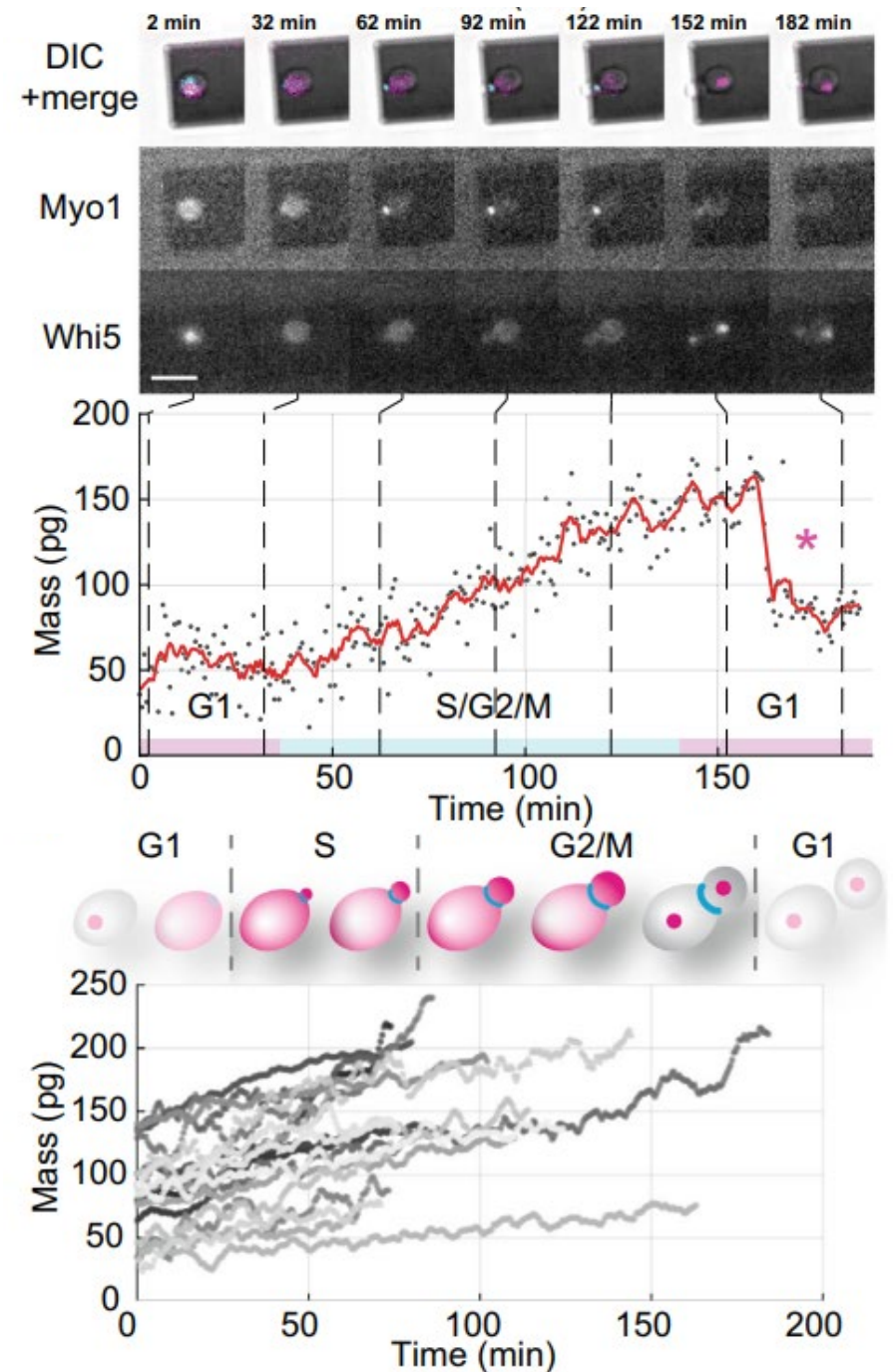
# Mass measurements of single yeast cells

Increasing mass resolution



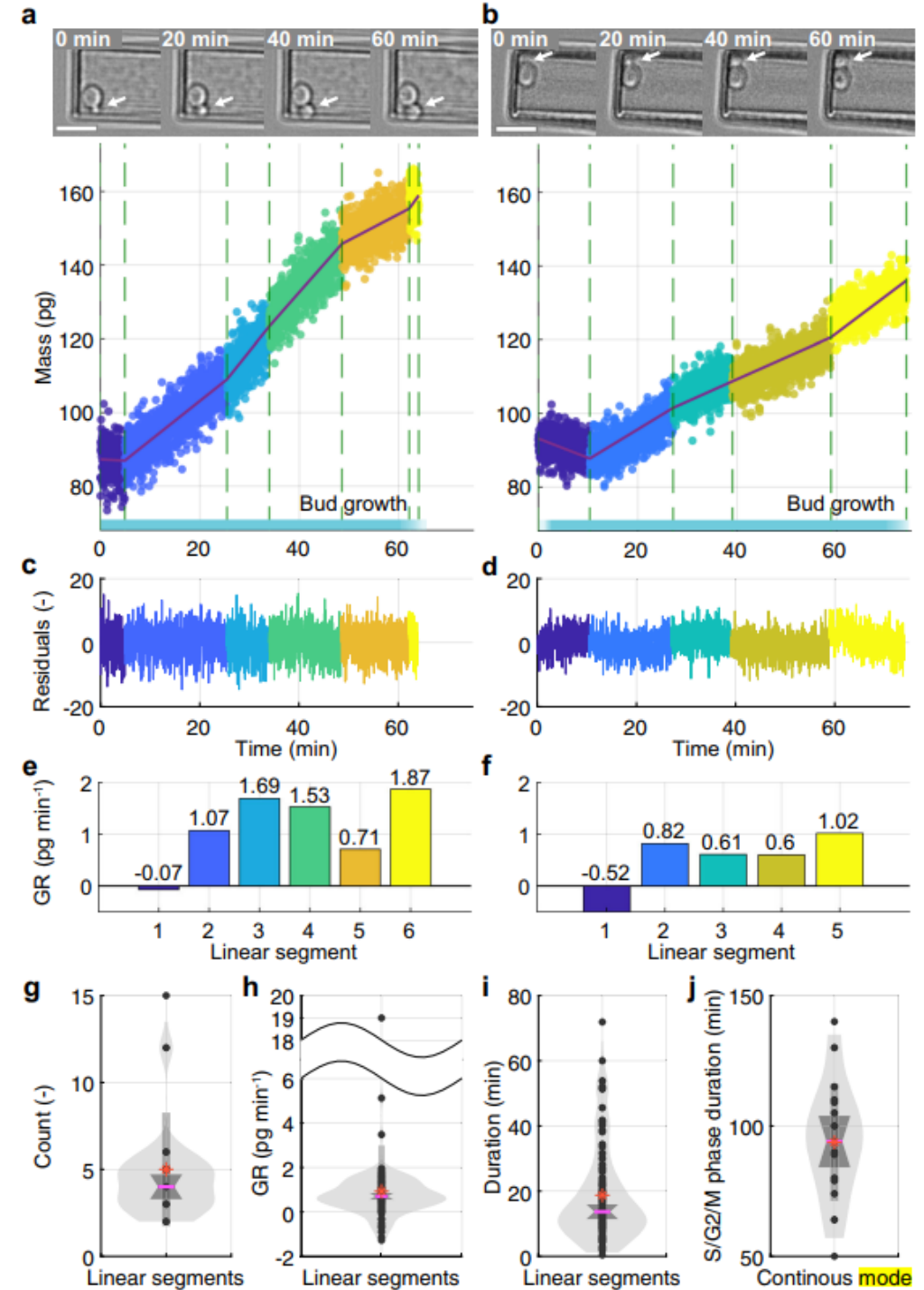
# Mass measurements of single yeast cells

Increasing mass resolution,  
improving optical microscopy



Single budding yeast cells (S/G2/M phase) increase total mass in multiple linear segments sequentially

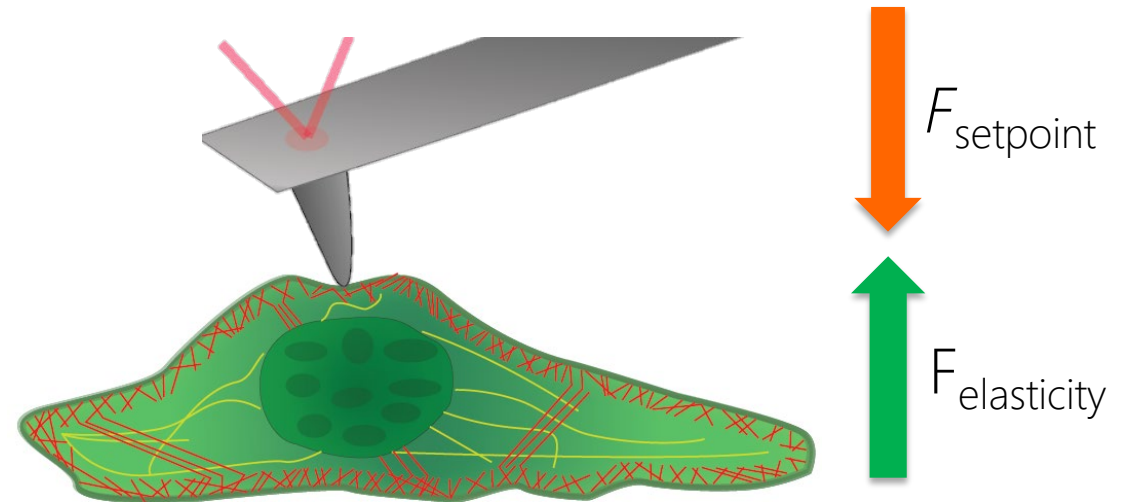
Cuny, Sapra, Martinez-Martin, Fläschner *et al.*,  
Nat. Commun. 2022



# Viscoelastic measurements

Materials exhibit only elasticity

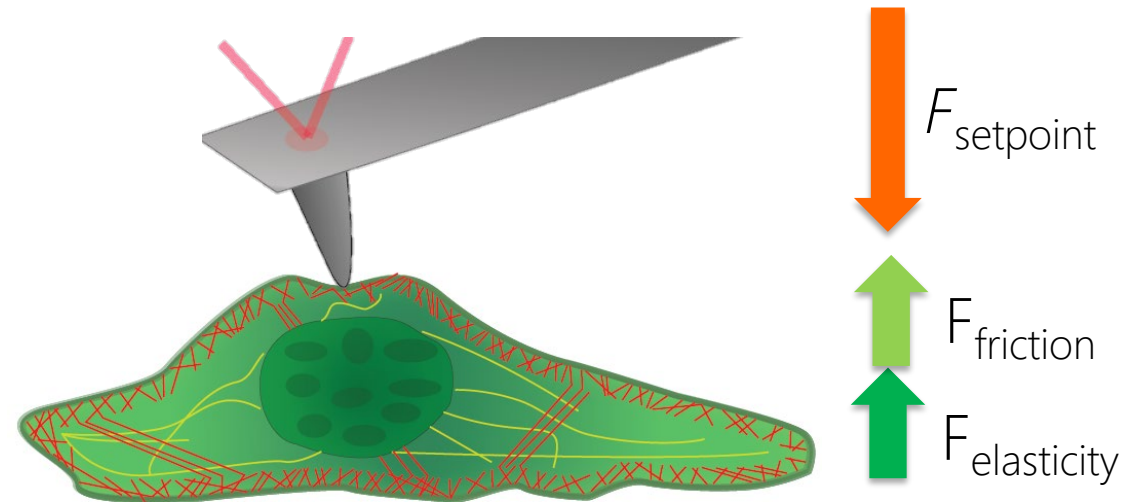
- Reactive force:  $F_{\text{elasticity}}$
- All compression energy is stored
- From  $F_{\text{elasticity}}$  and probe geometry:  $E_{\text{Youngs}}$



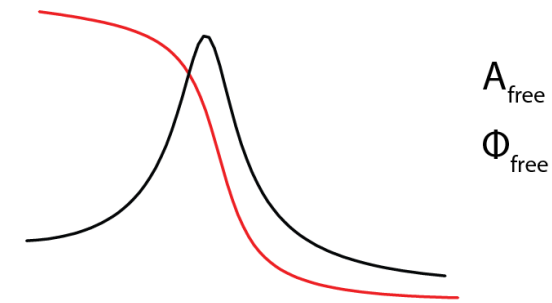
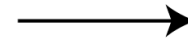
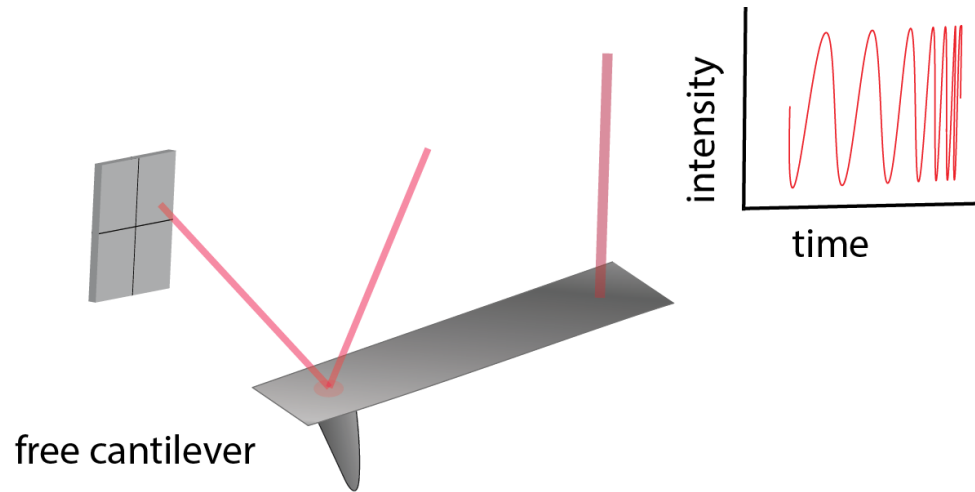
# Viscoelastic measurements

Materials exhibit elasticity and internal friction

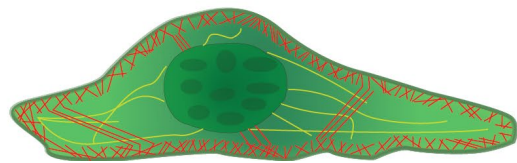
- Some energy is stored, some lost
- Two force components:  $F_{\text{elasticity}}$  and  $F_{\text{friction}}$
- Analogous to  $E_{\text{Youngs}}$ :  $E_{\text{storage}}$  and  $E_{\text{loss}}$



# Calibration

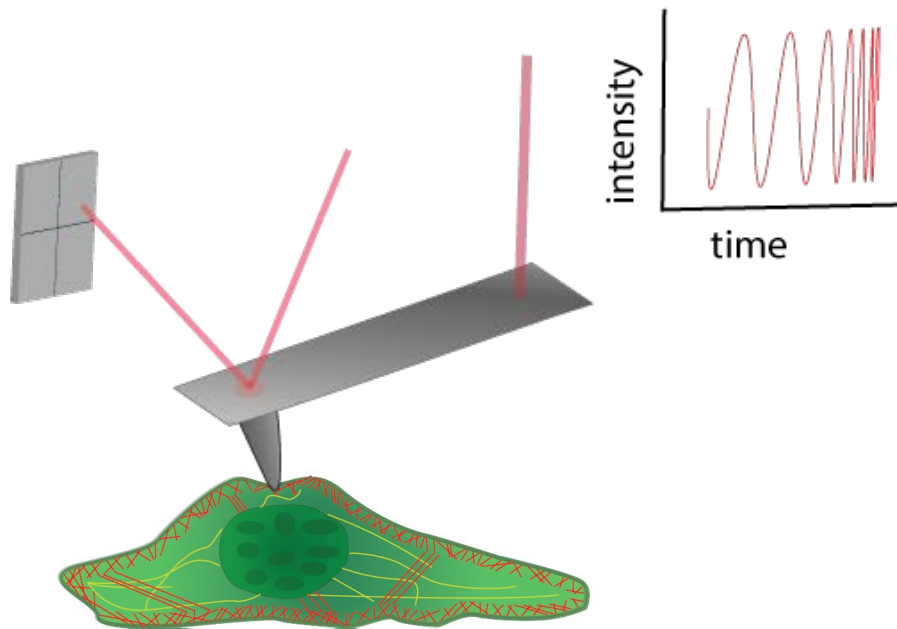


Amplitude  $A_{\text{free}}$ , and phase  $\Phi_{\text{free}}$  characterize the cantilever behavior in absence of conservative and dissipative forces of the sample (i.e. stiffness and viscosity of the sample)

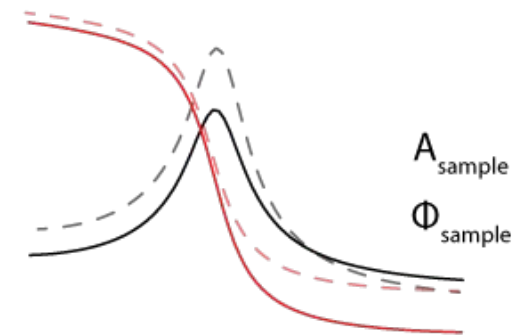


# Characterization

Go in contact. Stay there. Modulate.



Using the same “driving” of the cantilever, the oscillation of the cantilever changes, characterized by its new amplitude  $A_{\text{sample}}$ , and phase  $\phi_{\text{sample}}$





# Calculation

Dynamic sample stiffness:

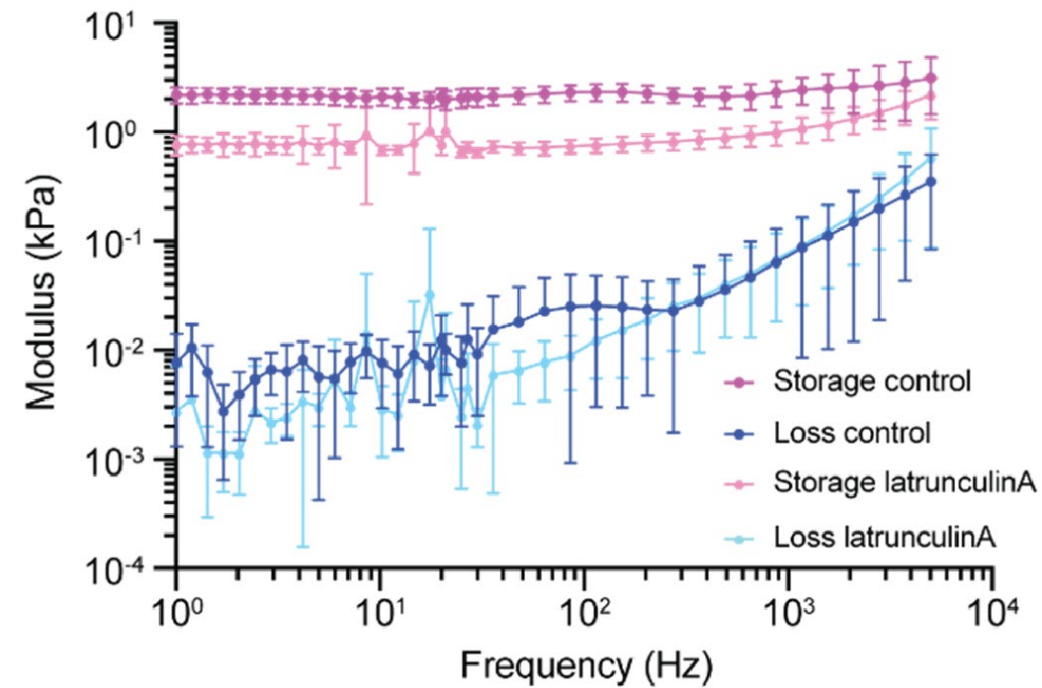
$$k_{\text{sample}} = k_{\text{cantilever}} \left( \frac{A_{\text{free}}}{A_{\text{sample}}} - 1 \right)$$

Loss tangent:

$$\tan(\delta) = \tan(\phi_{\text{sample}} - \phi_{\text{free}})$$

\*L.M. Rebelo et al. Soft Matter 10 (2014) 2141

- Use contact models to extract  $E_{\text{storage}}$  from  $k_{\text{sample}}$
- With  $E_{\text{storage}}$  and  $\tan(\delta)$  get  $E_{\text{loss}}$



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# Thank you for your attention

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