

The dynamics of Life

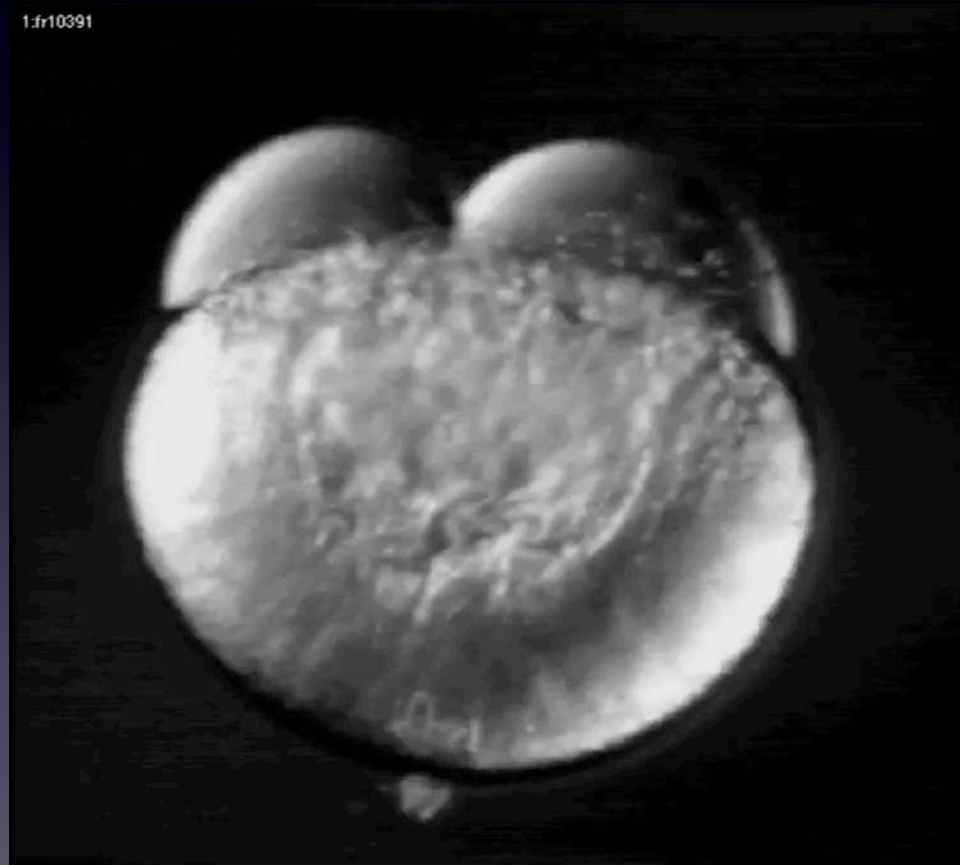
Karsten Kruse



**UNIVERSITÉ
DE GENÈVE**

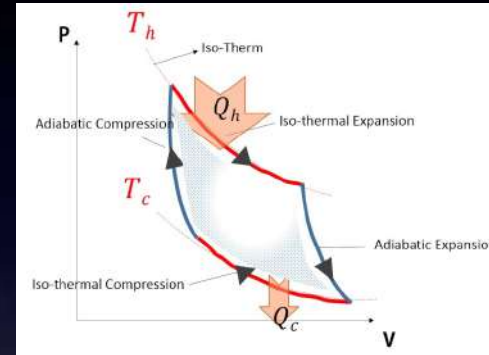
FACULTÉ DES SCIENCES

The dynamics of living beings is fascinating

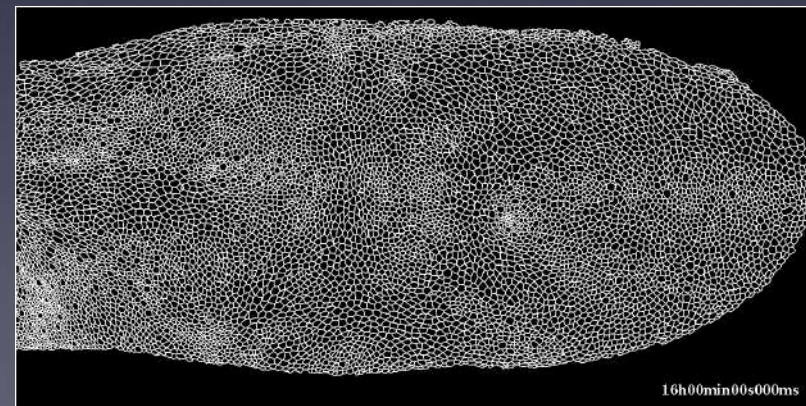
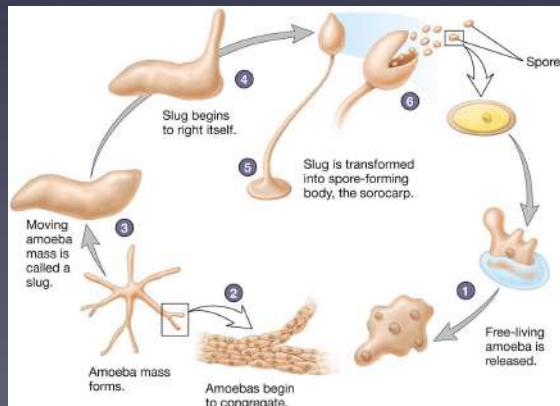


Developing zebrafish

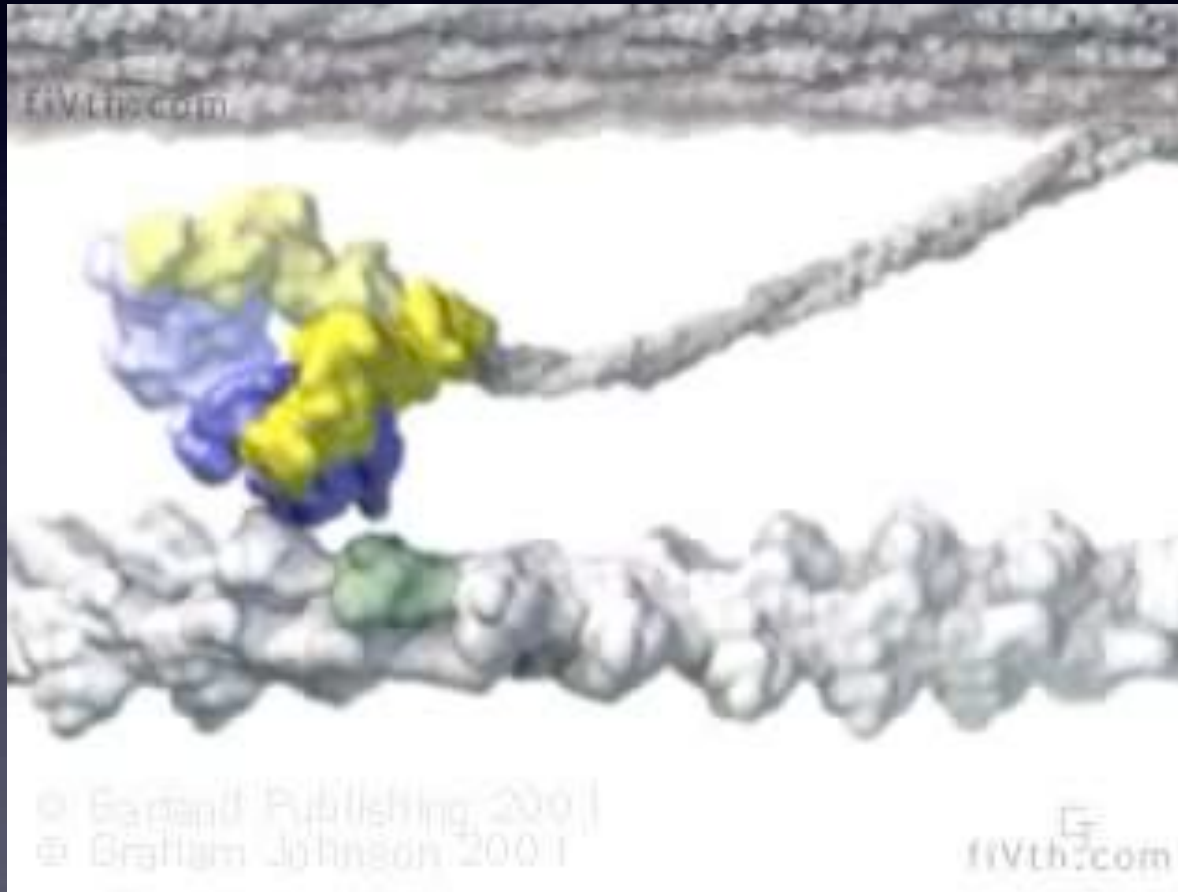
Thermodynamics



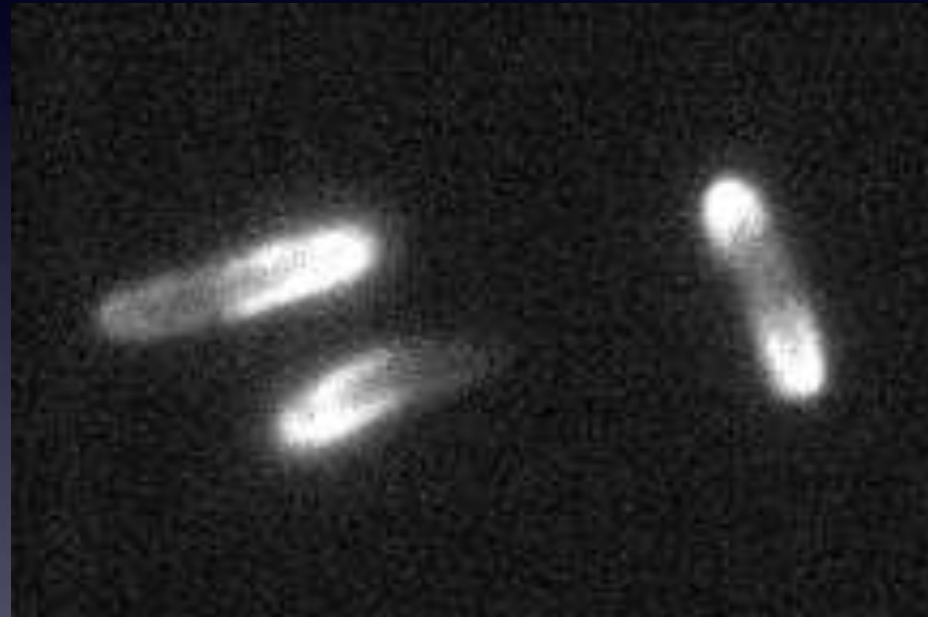
Vividynamics



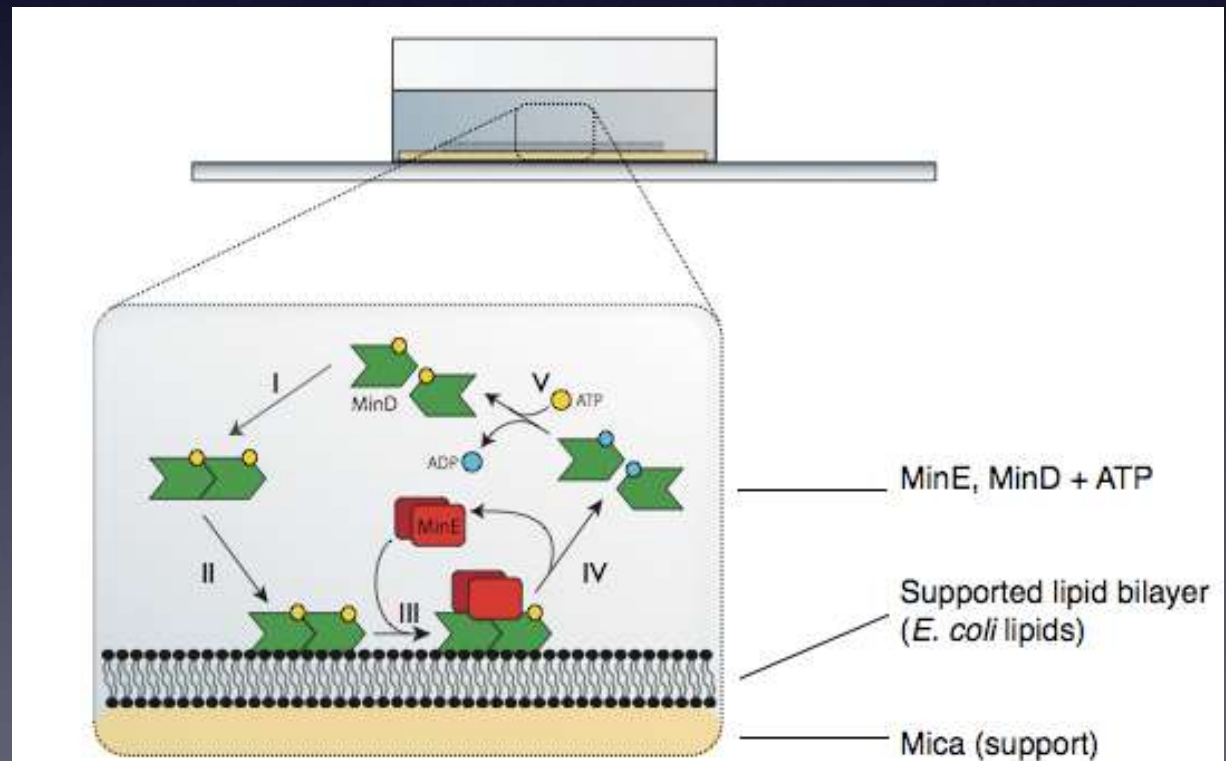
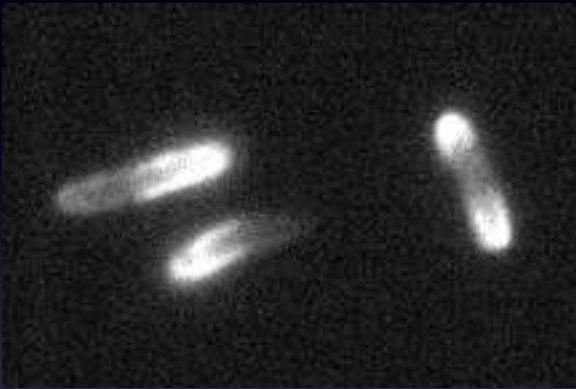
There are no living biomolecules



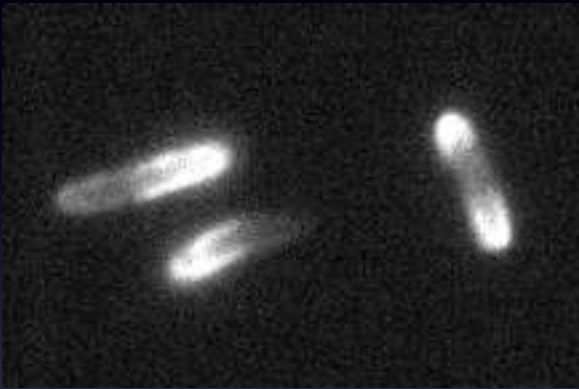
The Min-proteins form a standing wave in *Escherichia coli*



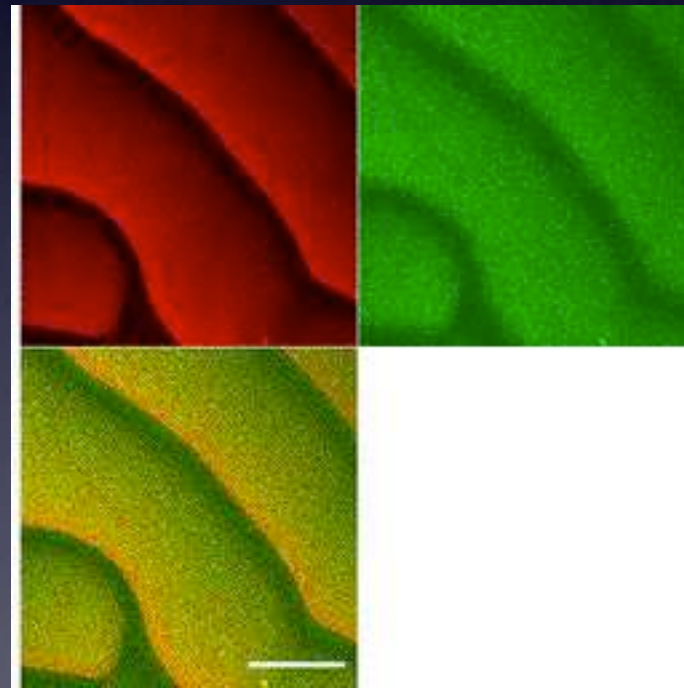
The Min-proteins dynamics can be reconstituted *in vitro*



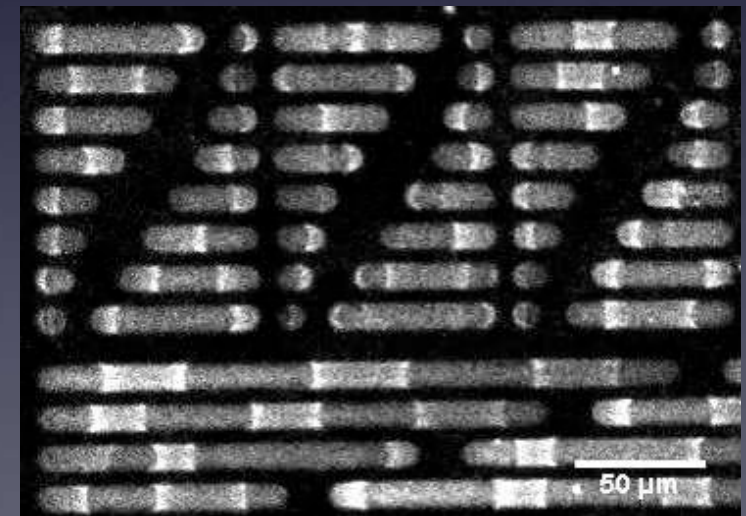
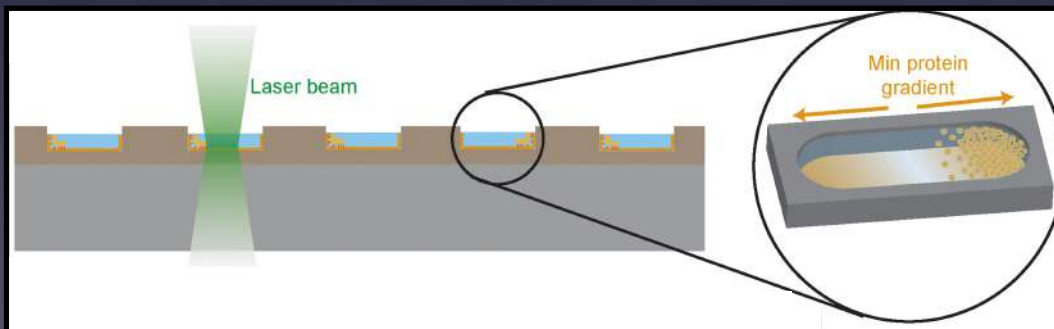
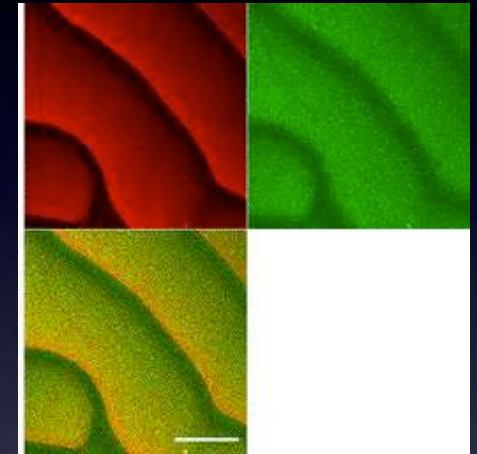
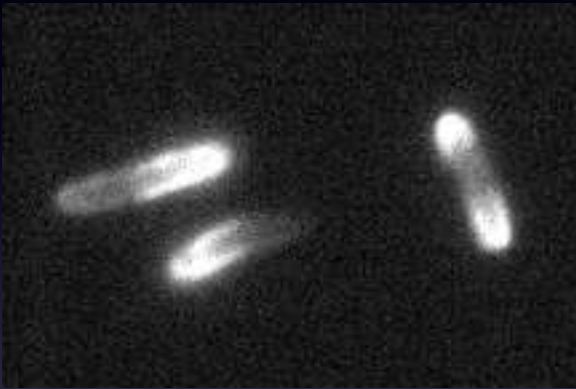
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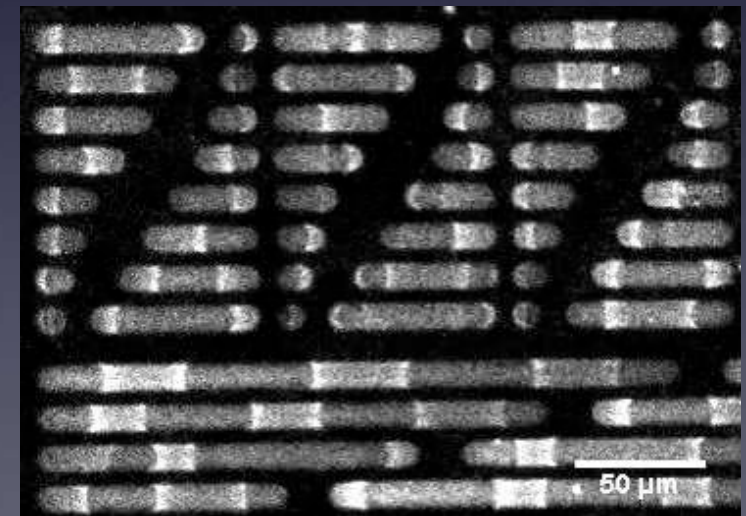
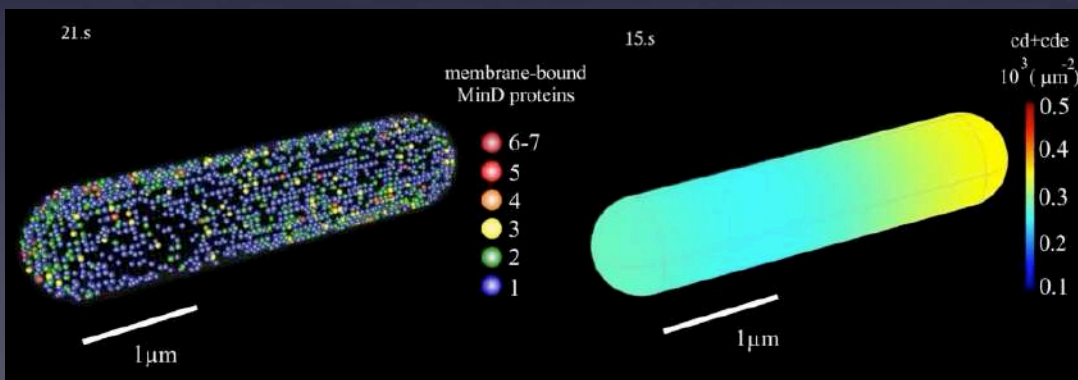
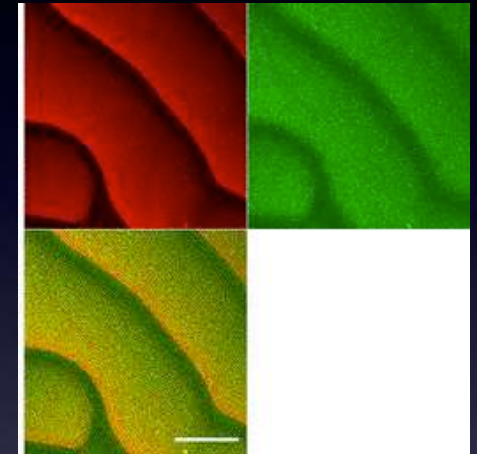
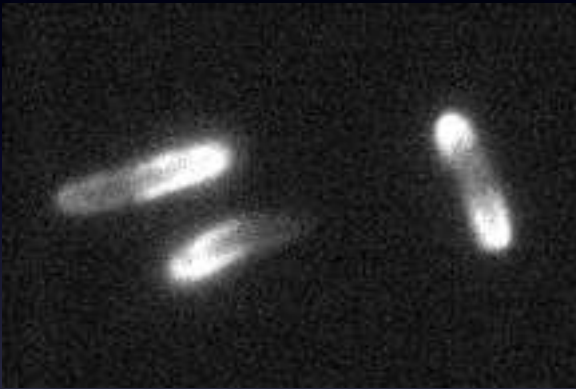
scale bar: 50 μ m
velocity: 0.5 μ m/s



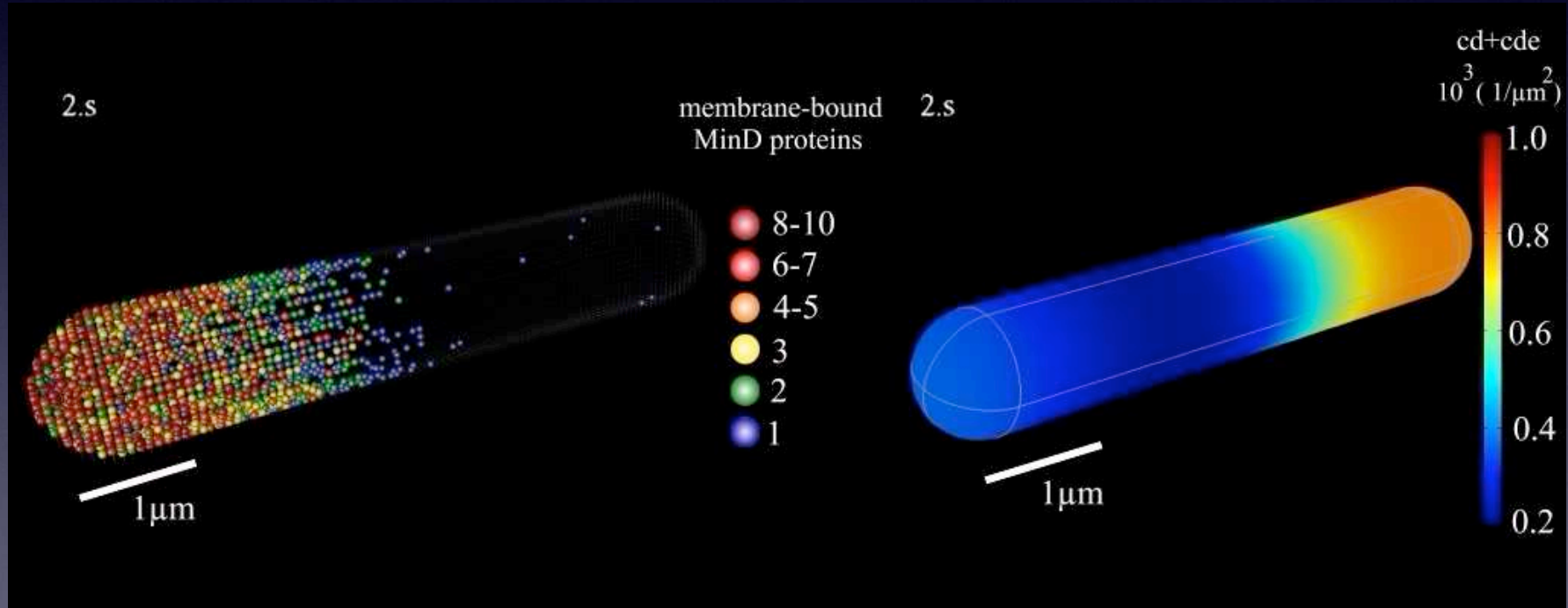
The Min-proteins dynamics can be reconstituted *in vitro*



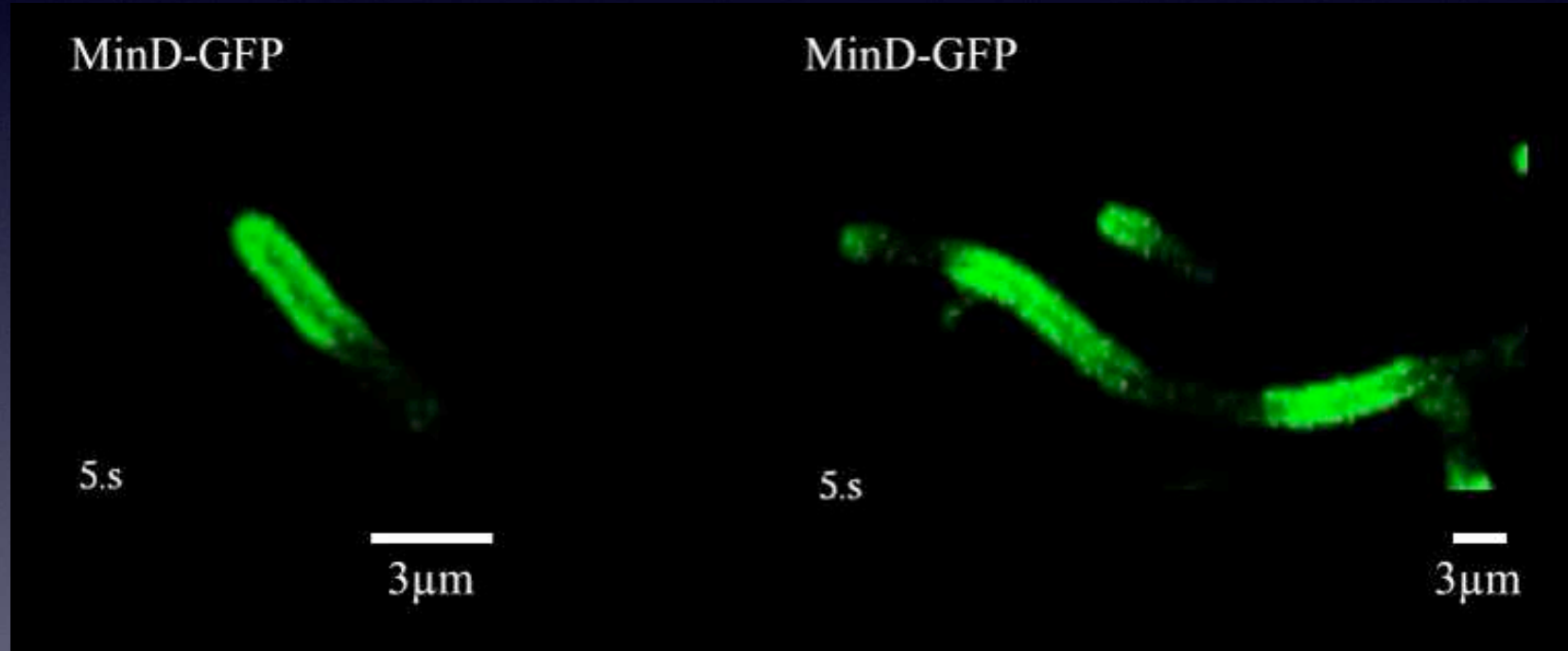
The Min-protein patterns result from reactions and diffusion



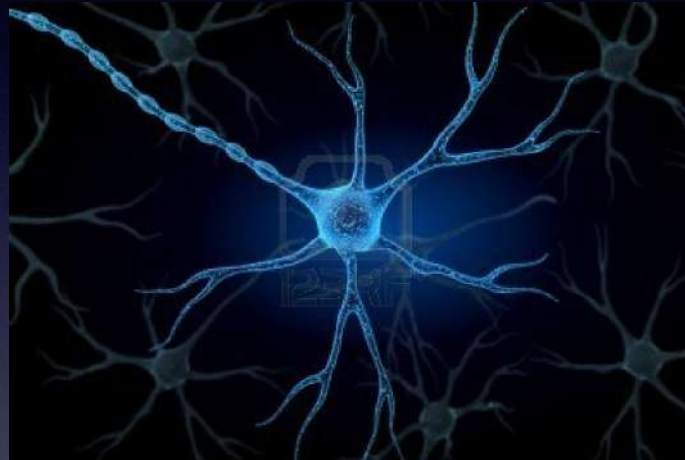
Computations predict traveling waves if MinD/E are over-expressed



Traveling waves can be observed in *E. coli*



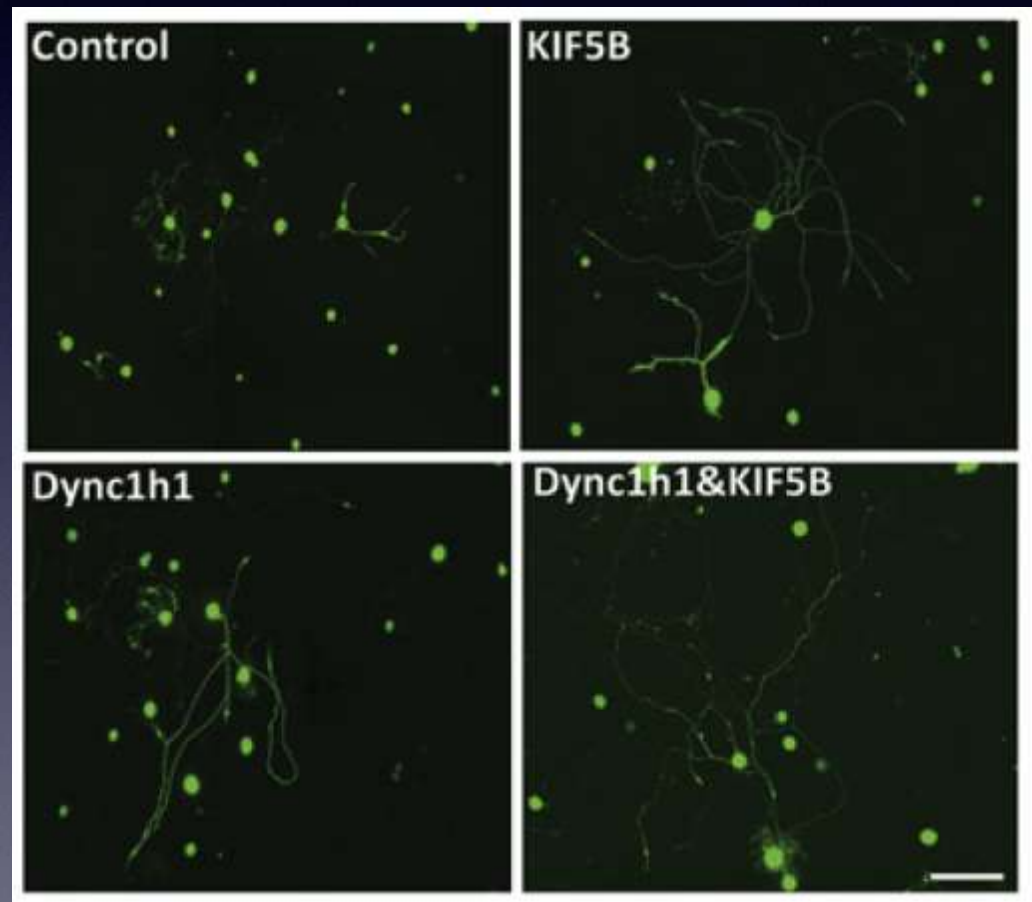
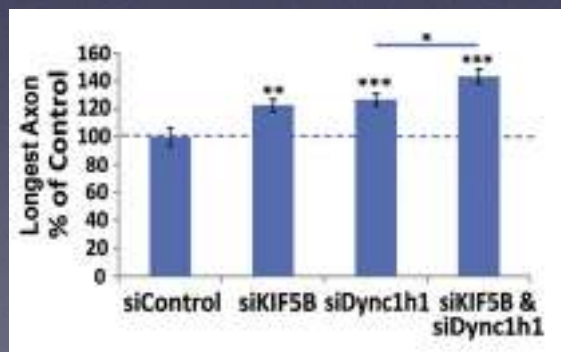
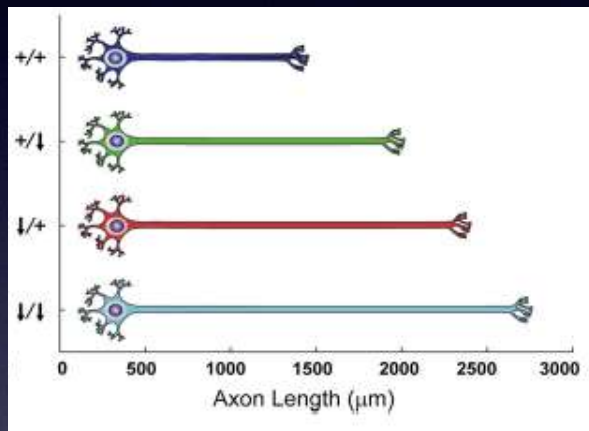
Neurons sense the length of their axons



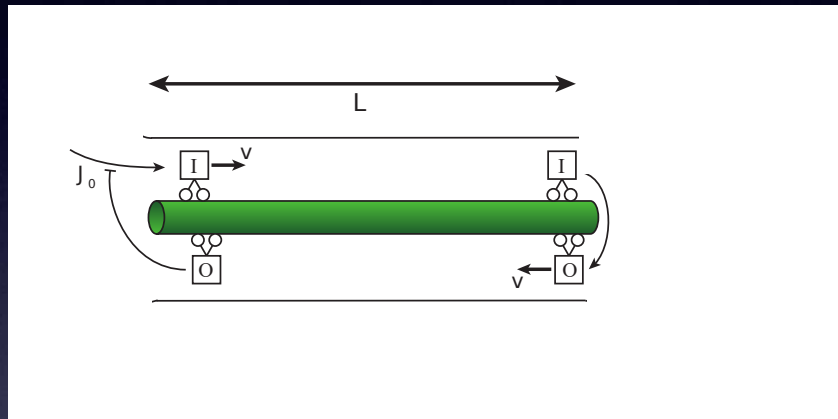
Kane Thomas

- Axons vary dramatically in length (μm to m)
- Neurons adapt (protein) metabolism to maintain homeostasis
- There is evidence for intrinsic length sensors
- Diffusion gradients only for lengths $<100\mu\text{m}$
- (Microtubule-based) rulers?
- There is evidence for intrinsic axonal length control (Albus et al, Trends Cell Biol 23, 305 (2013))

Molecular motors are involved in length sensing and control

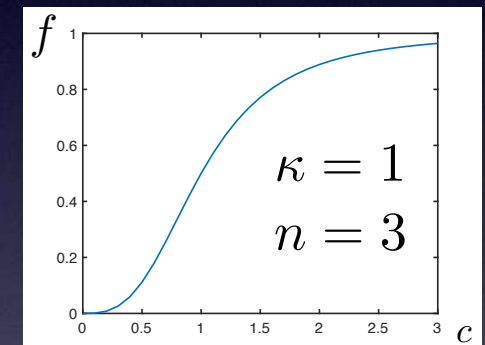


Oscillation with length-dependent period



Sigmoidal dose response

$$f_{\kappa}(c) = \frac{c^n}{\kappa^n + c^n}$$

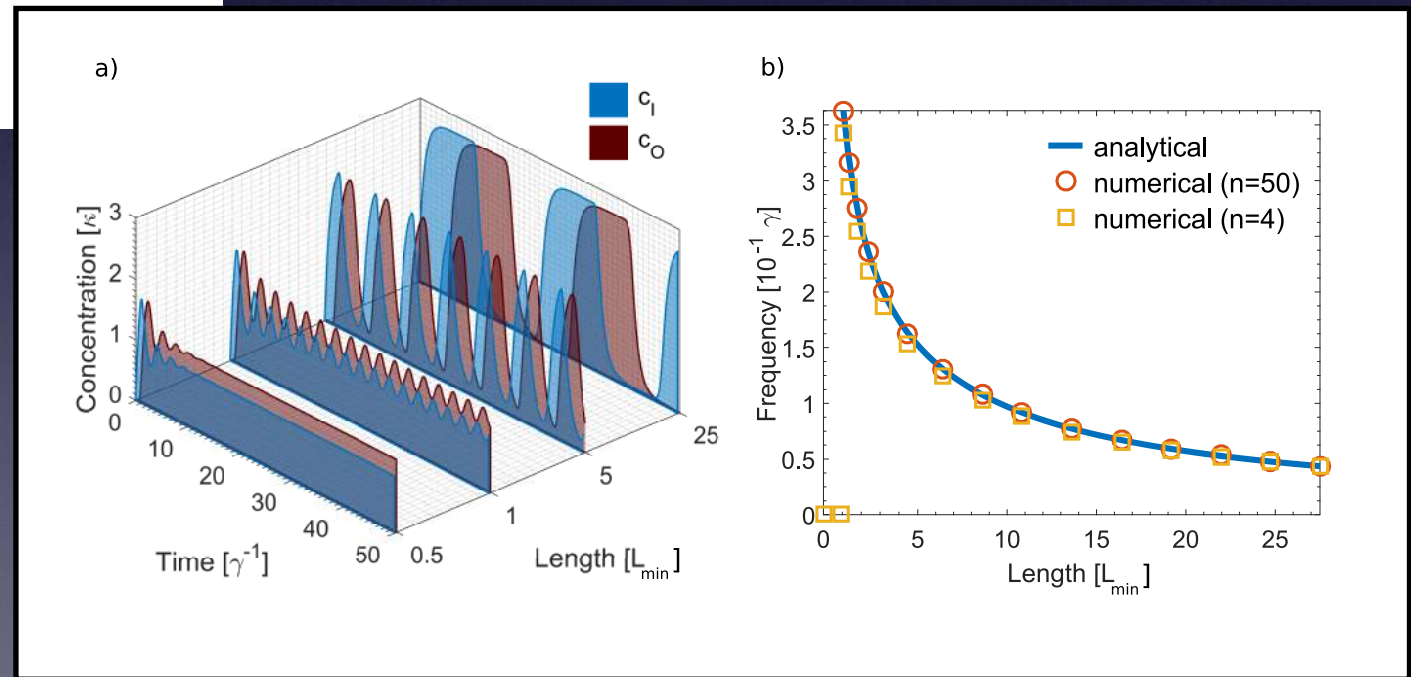
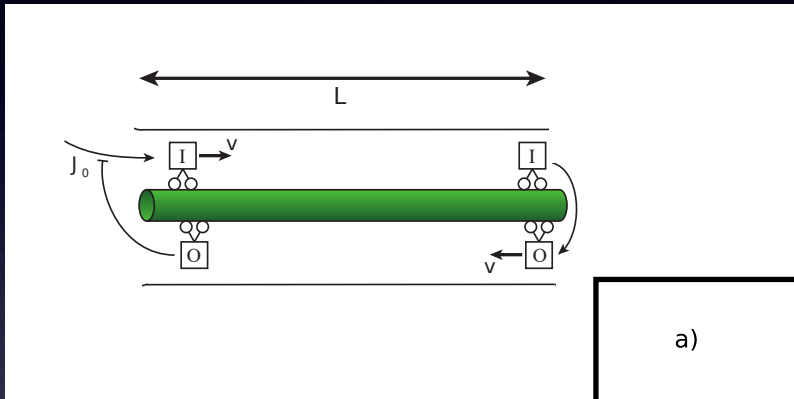


$$\dot{c}_I(t) = J_0 - J_I f_{\kappa}(c_O(t - \tau)) - \gamma_I c_I(t)$$

$$\dot{c}_O(t) = J_O f_{\kappa}(c_I(t - \tau)) - \gamma_O c_O(t)$$

$$\tau = L/v$$

Oscillation with length-dependent period



How to read out length-dependent oscillations?

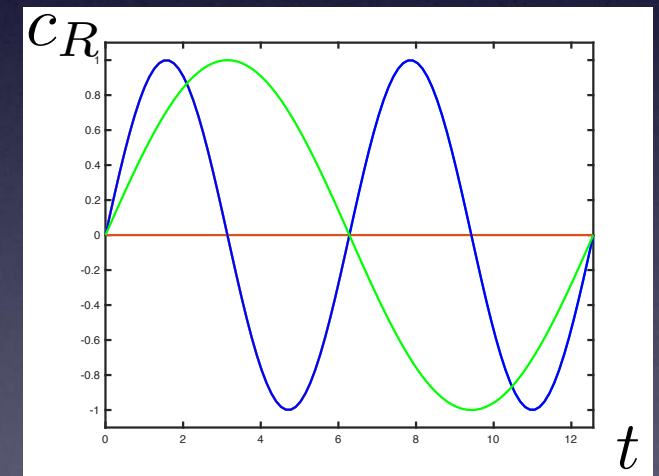
Incoming signal

$$c_I(t + T) = c_I(t)$$

Average signal

$$\langle c_I \rangle \equiv \frac{1}{T} \int_0^T c_I(t) dt$$

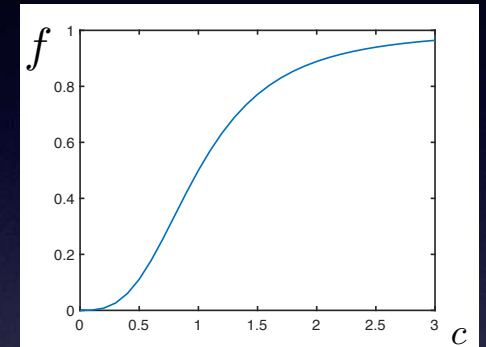
independent of T



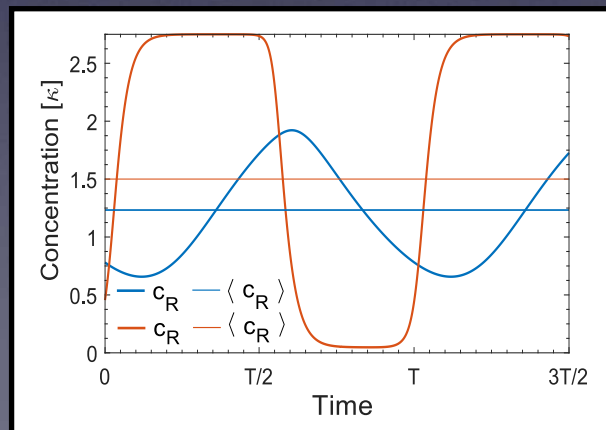
How to read out length-dependent oscillations?

Response

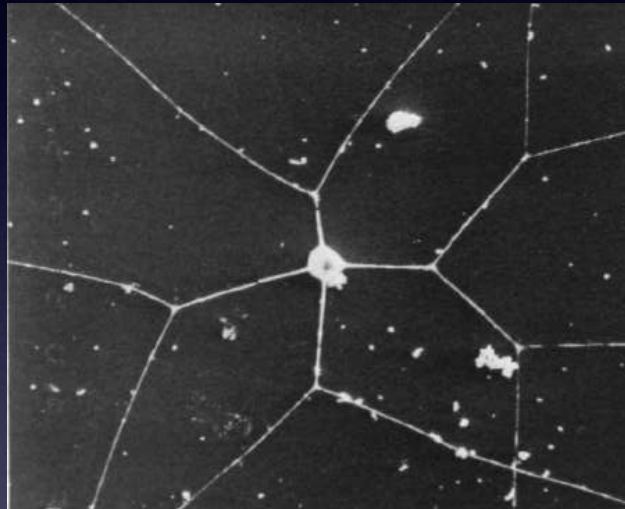
$$\dot{c}_R = J_R(1 - f_\kappa(c_I)) - \gamma_R c_R$$



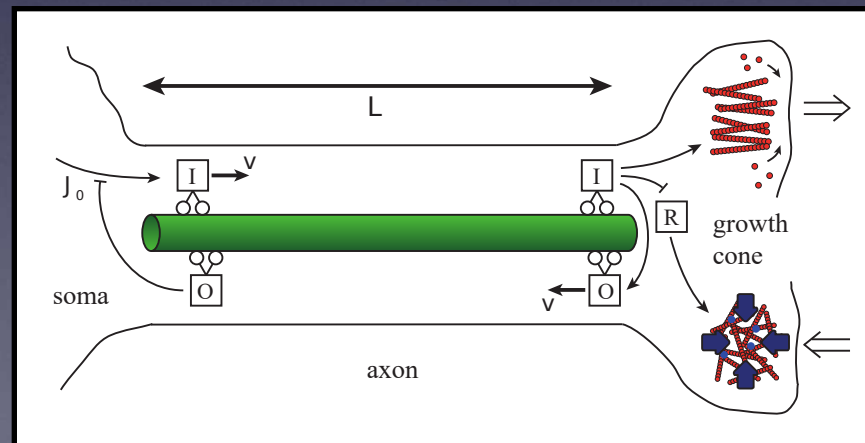
If the shape of c_R depends on T then so does $\langle c_R \rangle$



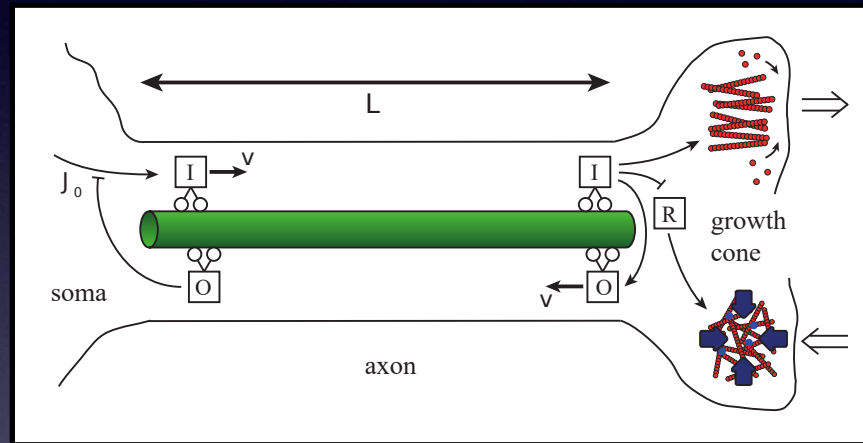
Mechanics is important for axonal growth



Bray, J Cell Sci **37**, 391 (1979)

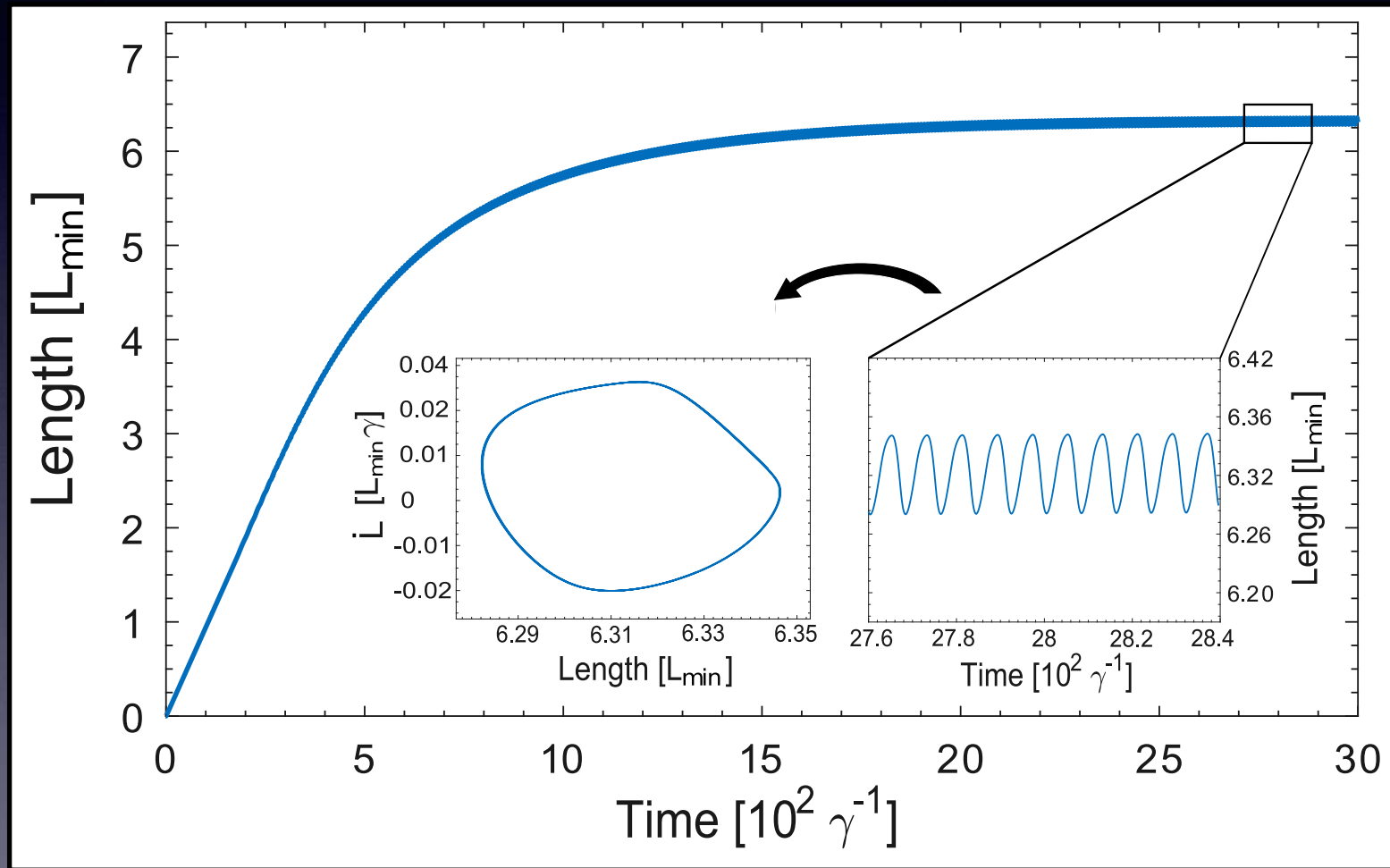


The signals regulate growth cone extension and retraction

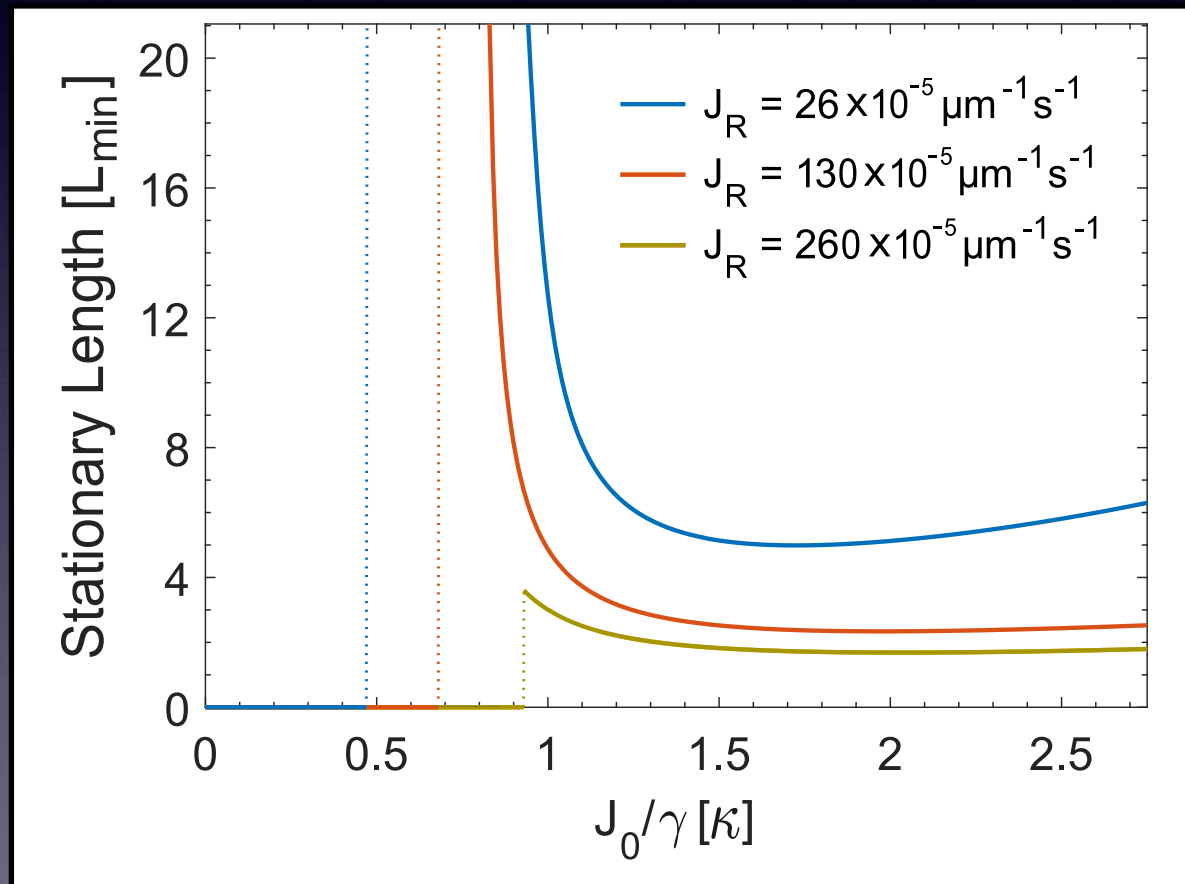


$$\dot{L} = v_g c_I - v_s c_R$$

The oscillatory signal can set the axon length

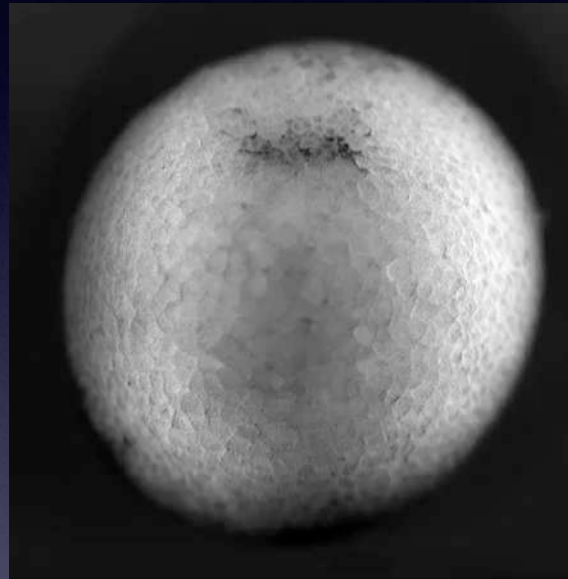


Length can increase with decreasing motor concentration



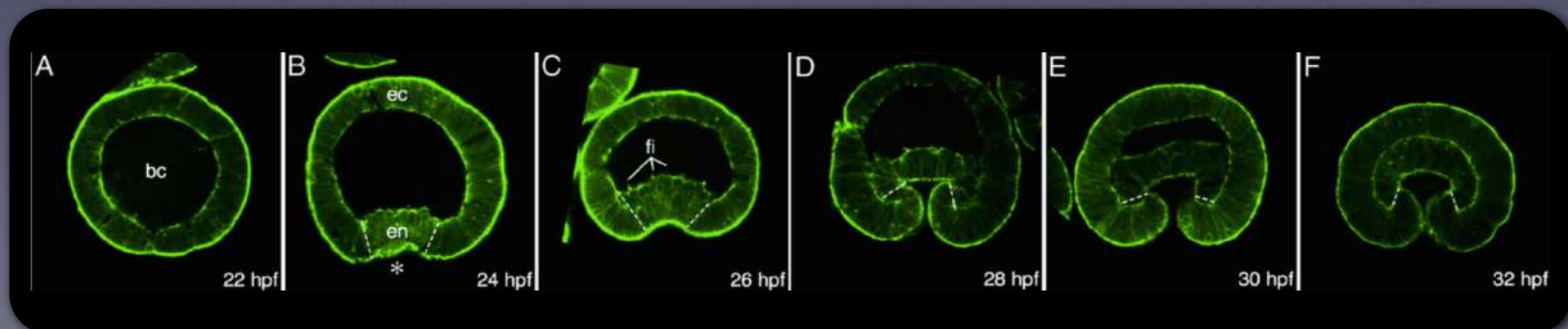
A single-cell layer invaginates during gastrulation

Xenopus laevis



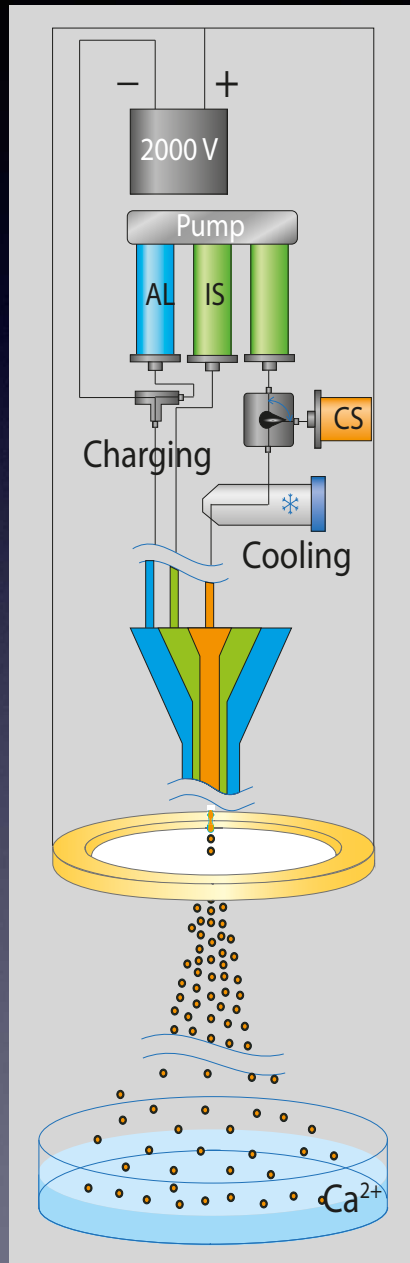
Total time: 15 hours

D. Shook from: Gastrulation: From Cells to Embryo, CSHL Press (2004)

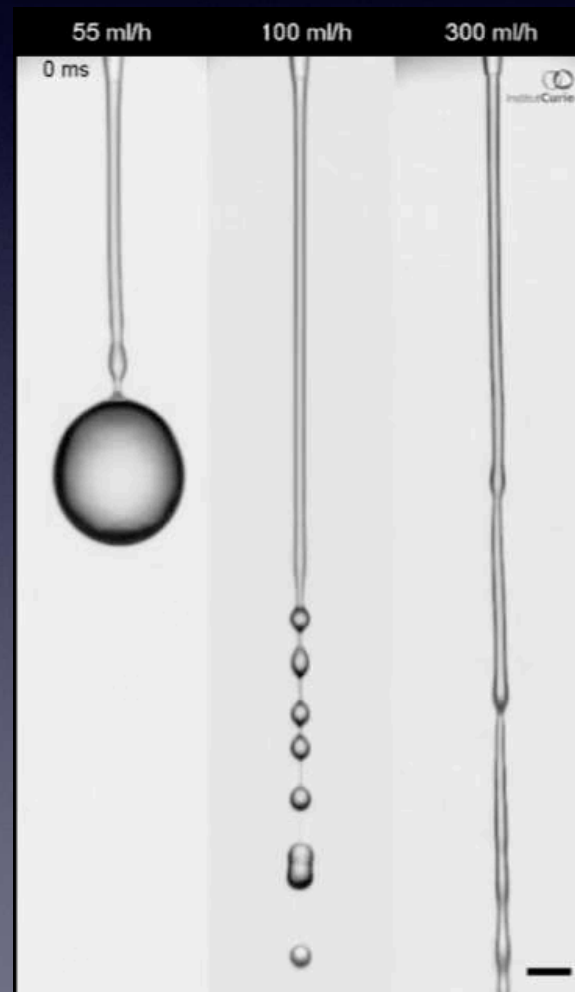
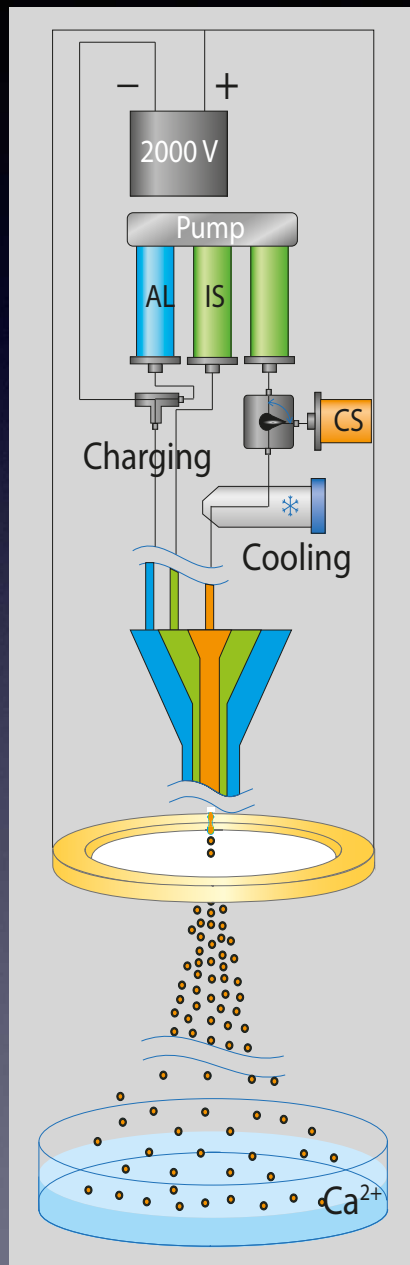


Tamulonis et al, Dev Biol (2011)

Epithelial cells can be confined in alginate capsules

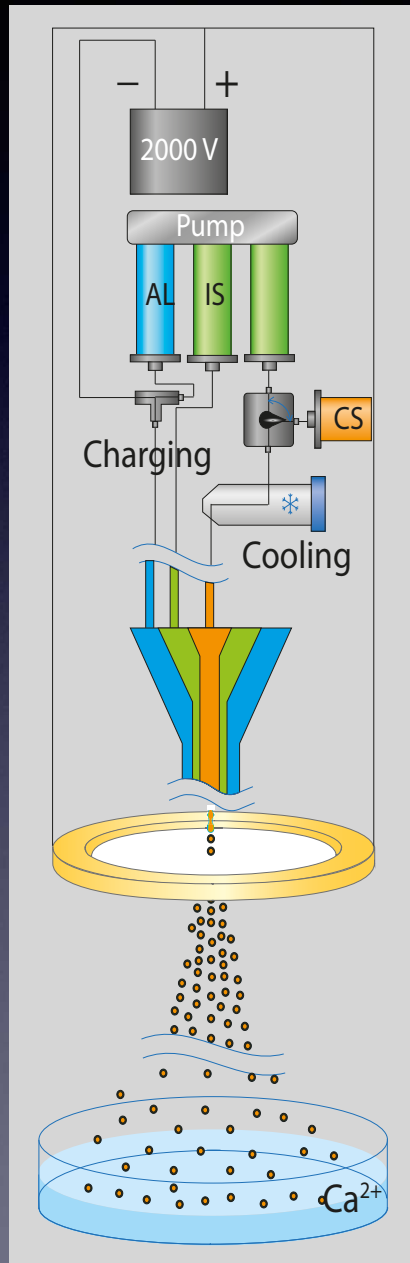


Epithelial cells can be confined in alginate capsules

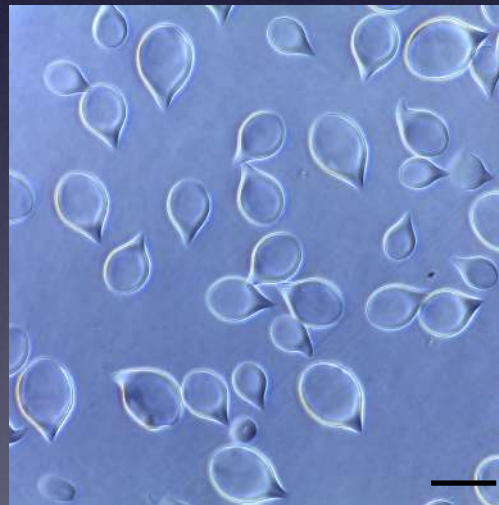


Drop radius:
 $\sim 100\text{-}200 \mu\text{m}$

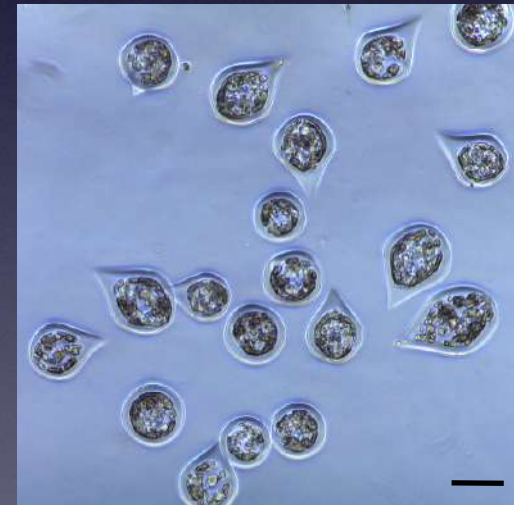
Epithelial cells can be confined in alginate capsules



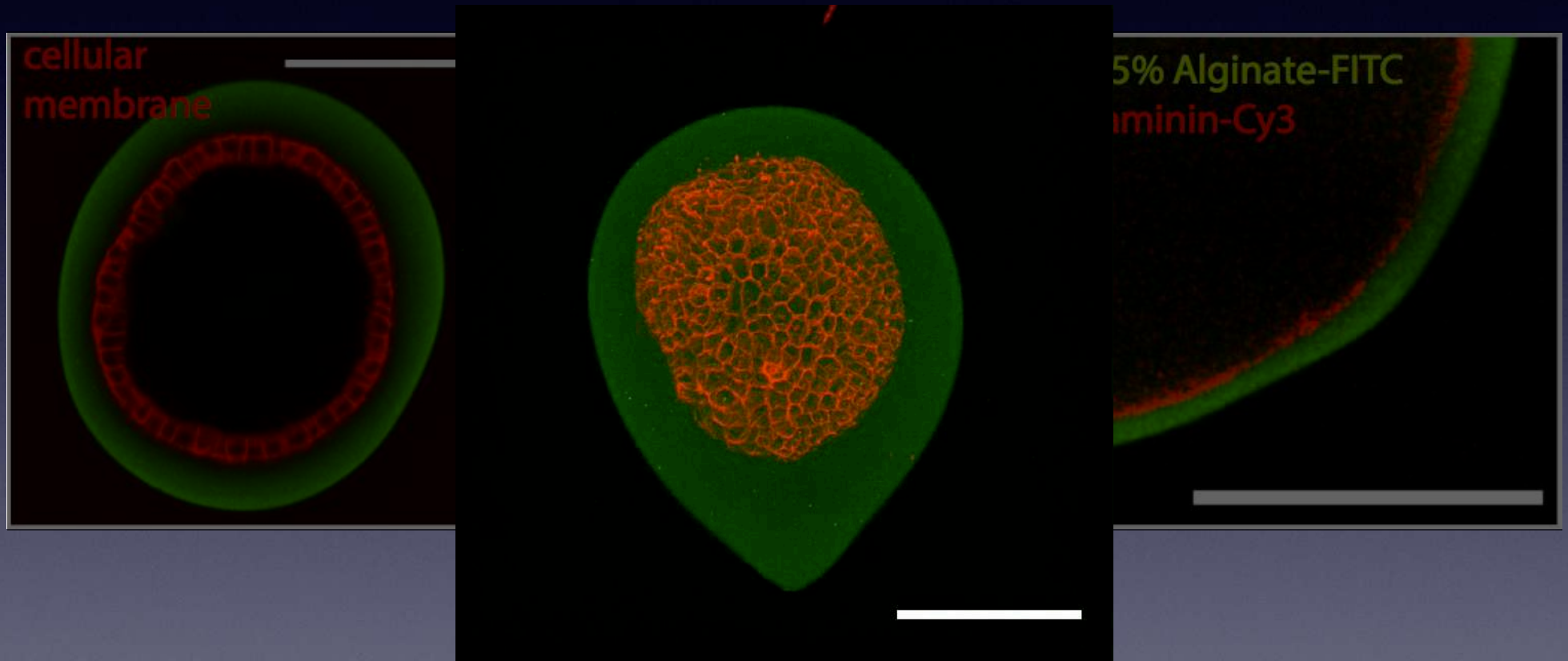
w/o cells



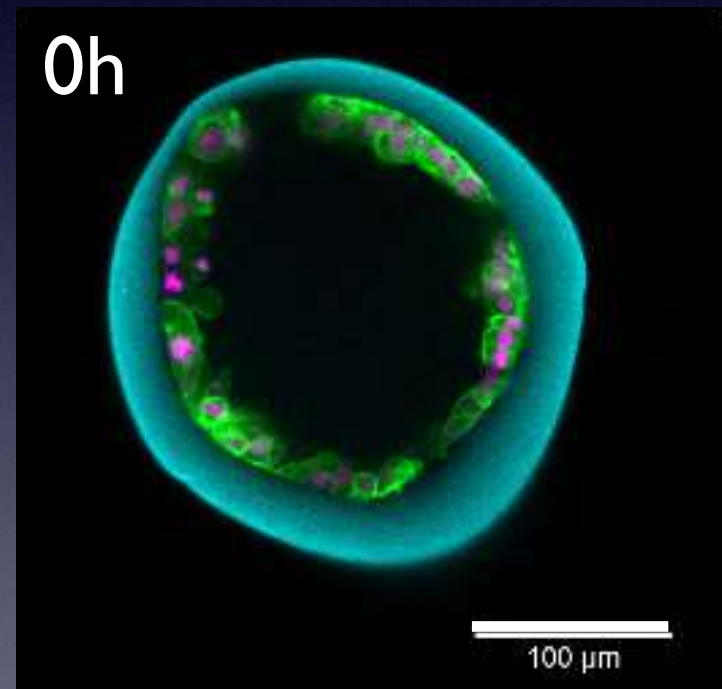
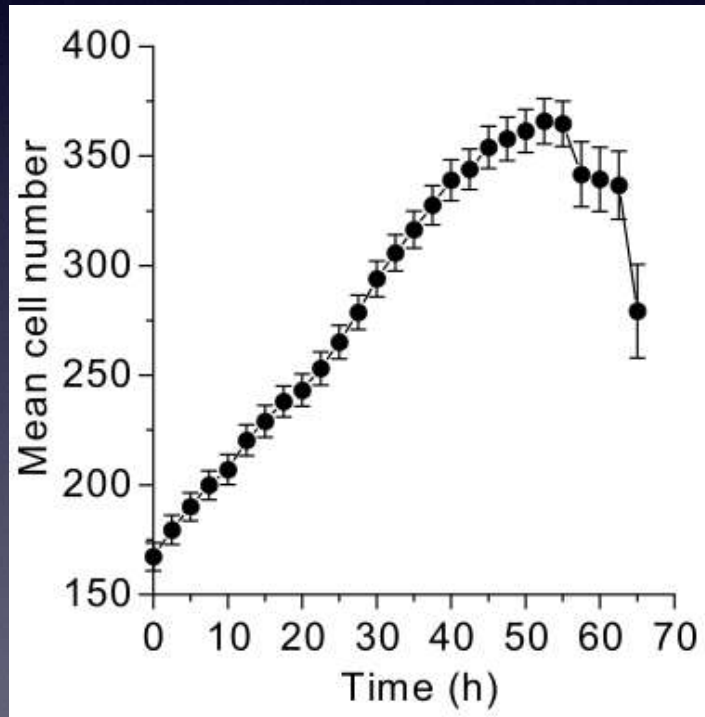
with cells



Cells form a monolayer in the capsule

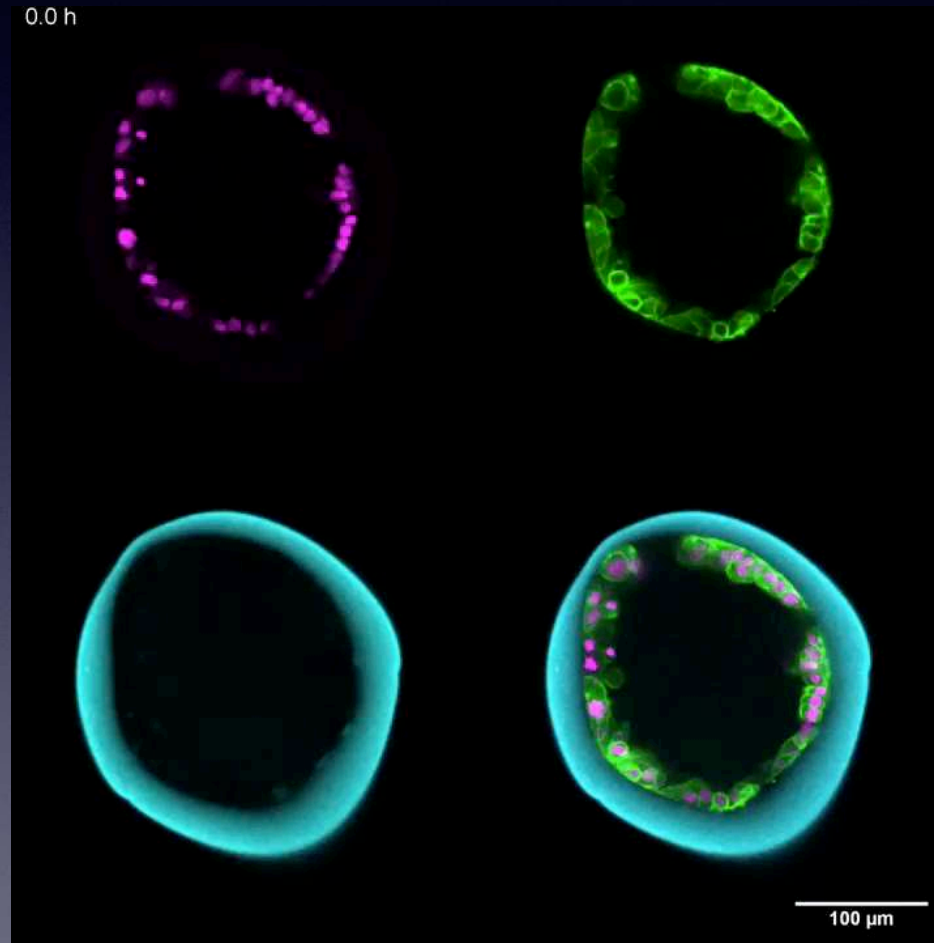


Cell number increases by proliferation



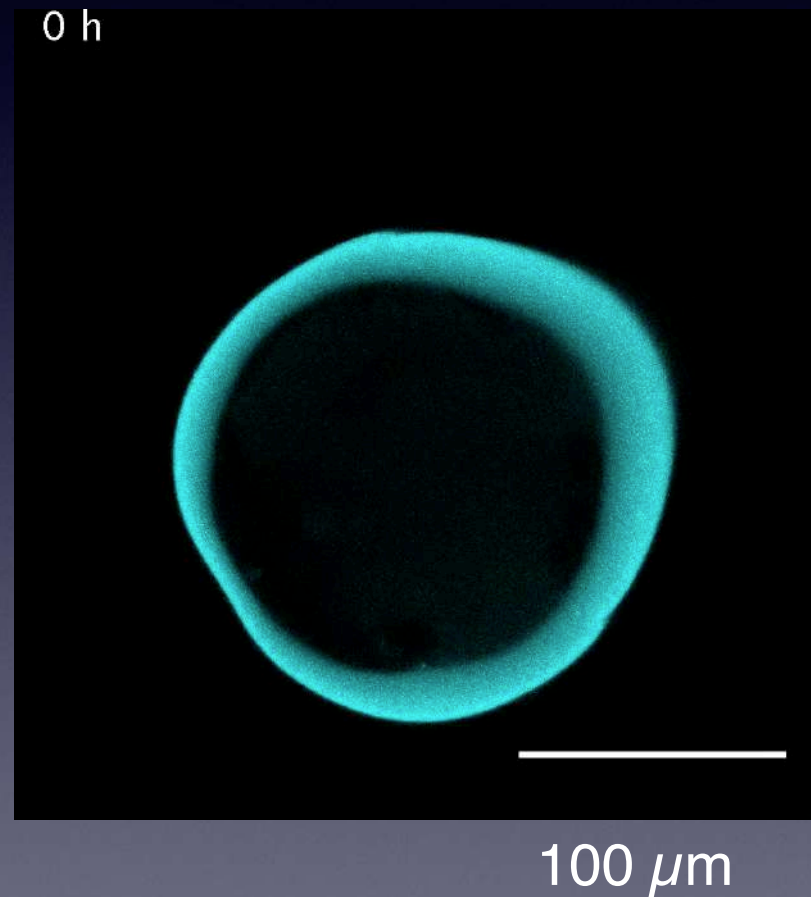
100 μm

Cell layers growing in capsules can fold

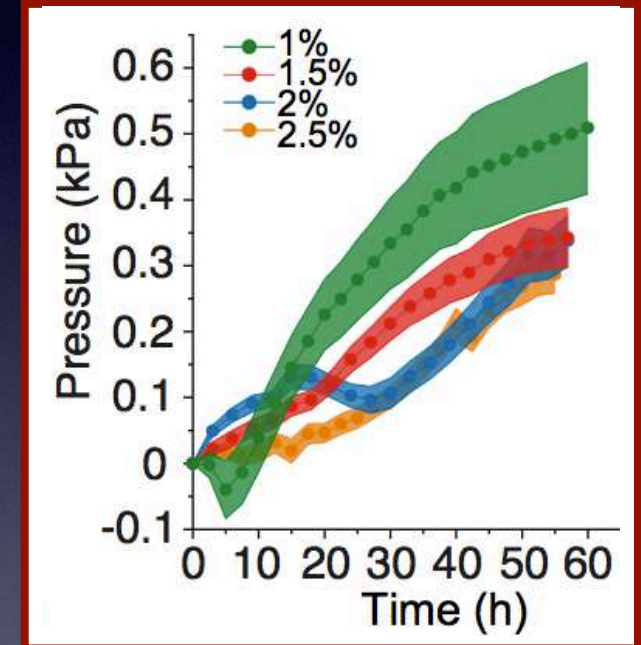
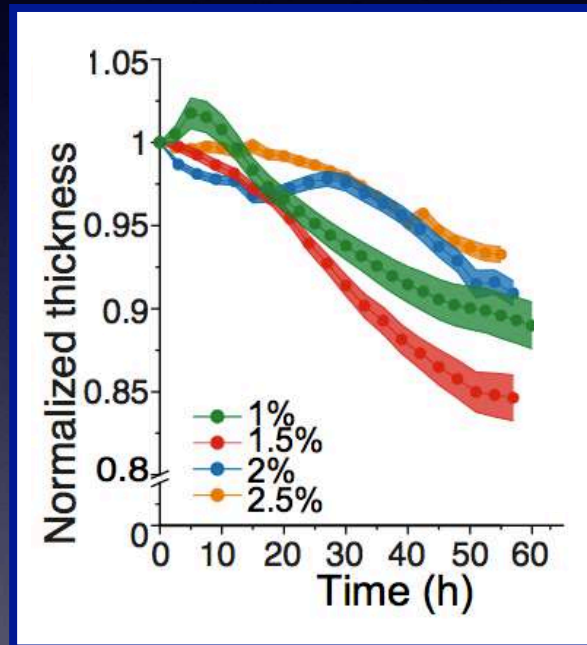
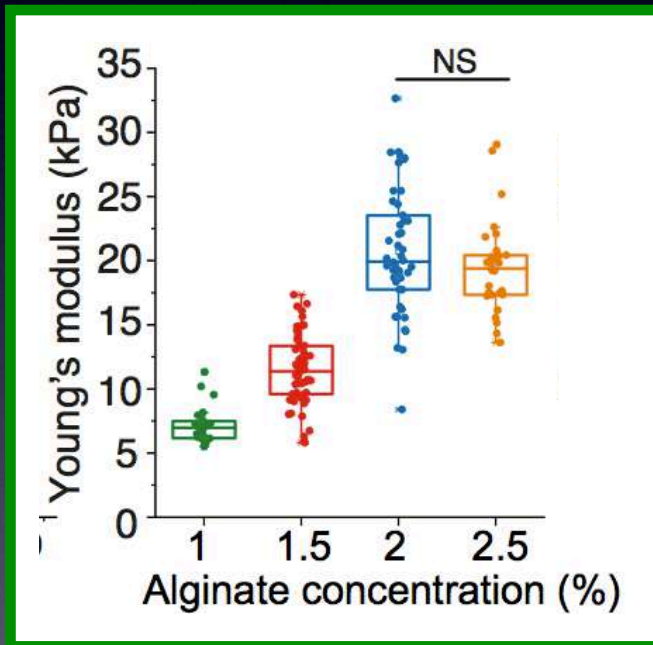


100 μm

Capsule thickness changes with time



Capsule deformations allow to measure pressure



$$\frac{2E}{1 - \nu} \frac{h(0)h(t)}{R^2} = P$$

We describe the monolayer as an elastic material

$$\mathcal{E} = \boxed{K} \int_L \kappa^2 ds + \boxed{\lambda} \int_{L_0} \epsilon^2 ds_0 + \boxed{k} \int_L (r(s) - R)^2 ds$$

Bending rigidity

Compressibility modulus

Capsule stiffness

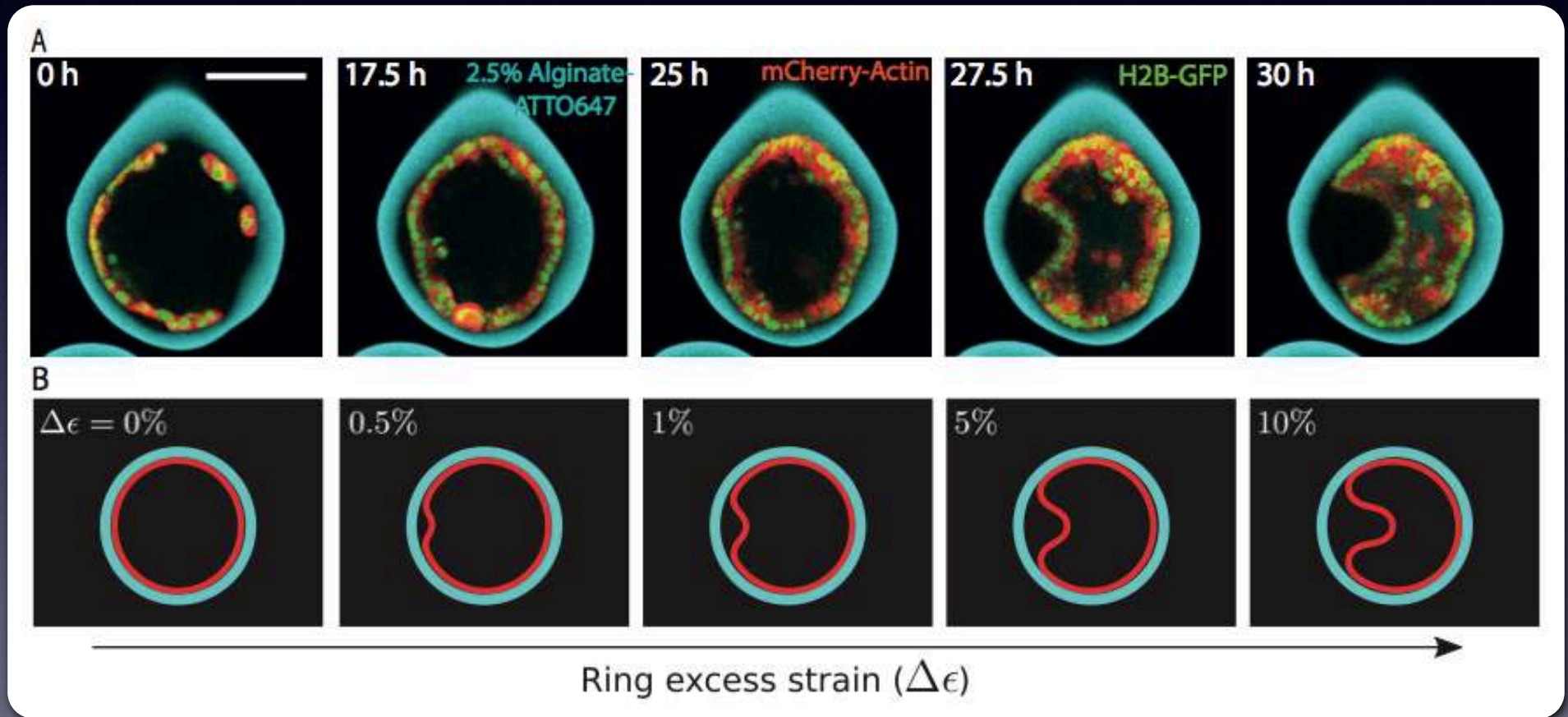
Dimensionless parameters:

$$\frac{\lambda R^2}{K} \quad \frac{\lambda}{kR^2} \ll 1 \quad \Delta\epsilon = \frac{L - 2\pi R}{2\pi R}$$

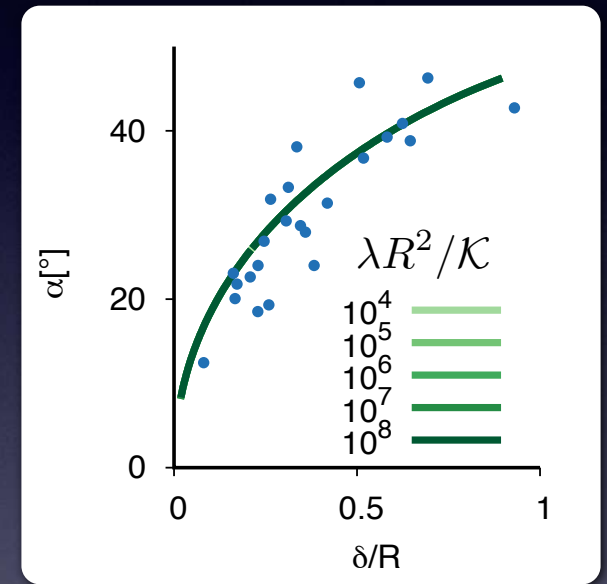
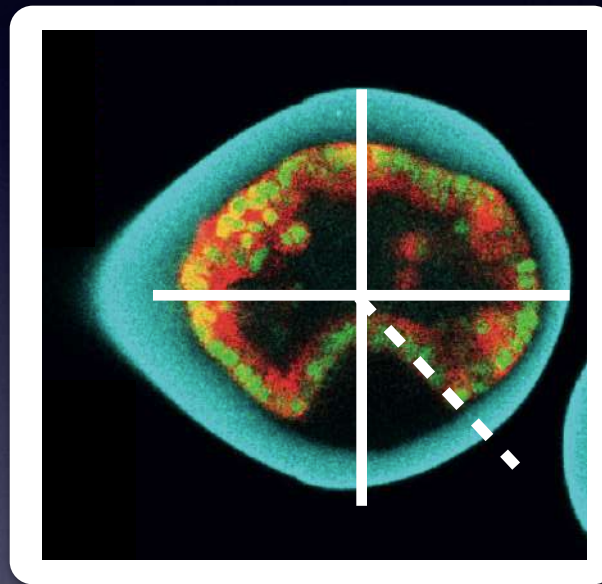
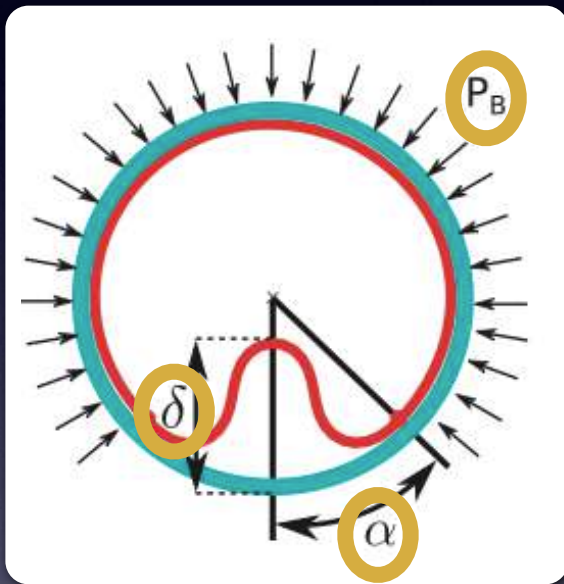


C. Blanch Mercader

The folded shapes of elastic rings and monolayers are similar



The similarity can be quantified



α : opening angle

δ : invagination depth

Our parameter values are consistent with the measured capsule stiffness

Parameter values

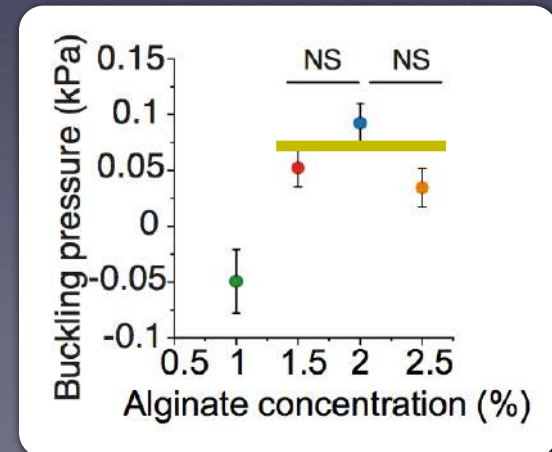
$$\lambda \approx 0.1 \frac{\mu\text{N}}{\mu\text{m}}$$

$$K \approx 1 \mu\text{N } \mu\text{m}$$

$$R \approx 100 \mu\text{m}$$

Pressure at buckling

$$P_{\text{buck}} \sim \frac{\lambda^{2/5} K^{3/5}}{R^{11/5}} \\ \approx 0.1 \text{ kPa}$$



Das Leben ist tot.

Das Leben ist tot.

Und wir haben es getötet.

Das Leben ist tot.

Und wir haben es getötet.

Life is dead.

And we have killed it.

Alors on peut bien parier que la
vie s'effacerait, comme à la
limite de la mer un visage de
sable.

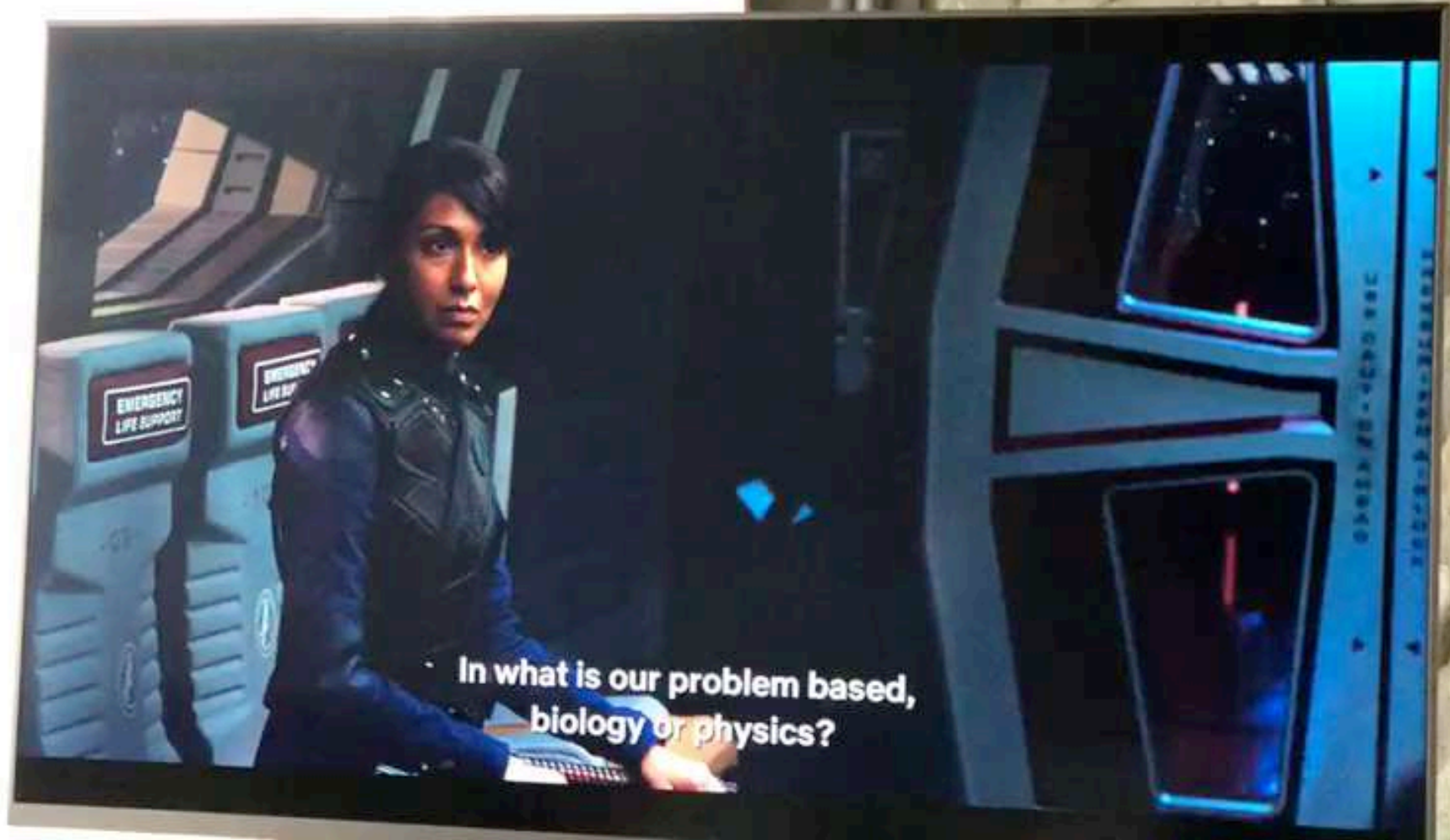
Alors on peut bien parier que la
vie s'effacerait, comme à la
limite de la mer un visage de
sable.

So we can surely bet that Life would
erase itself, like a face in the sand at the
border of the sea.

At the quantum level, there is no difference between physics and biology.

No difference at all. (...) Physics and biology? No; physics as biology.

Paul Stamets, Astromycologist, 2256



Paul Stamets, astromycologist, 2256

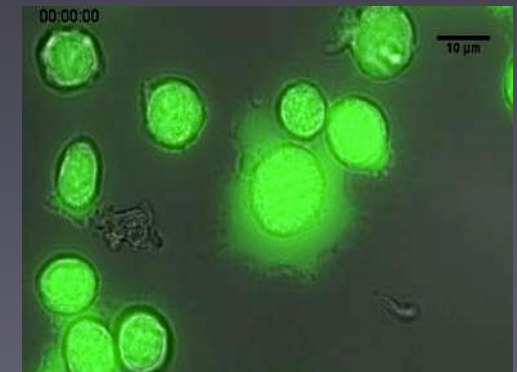
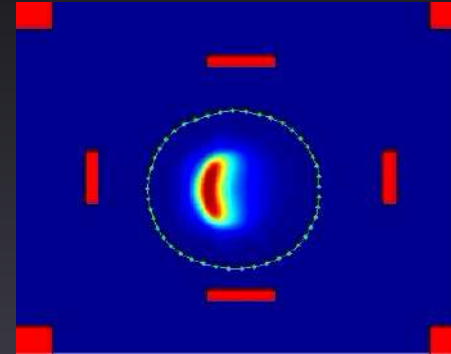
Acknowledgements

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