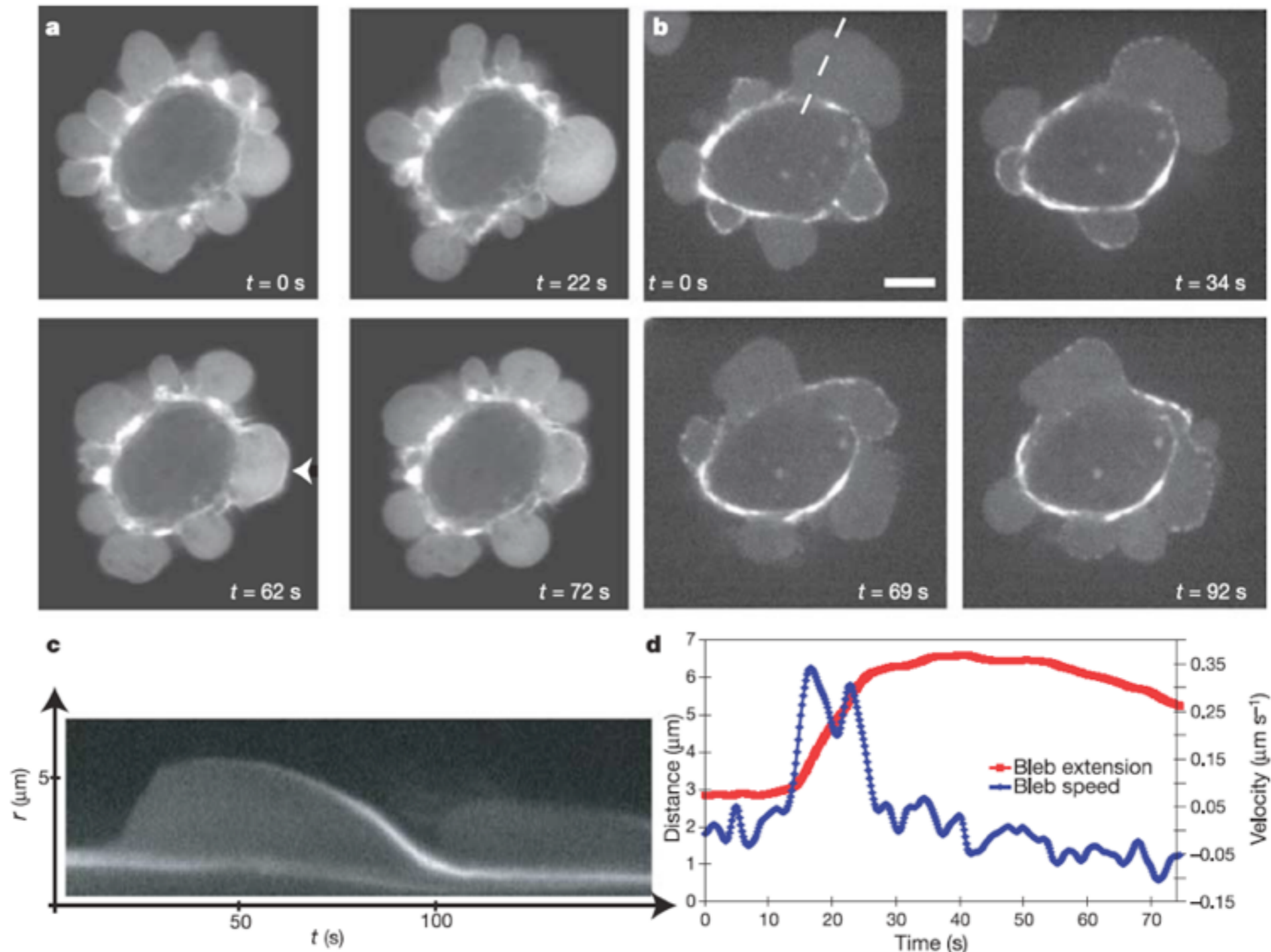


Non-equilibration of hydrostatic pressure in blebbing cells

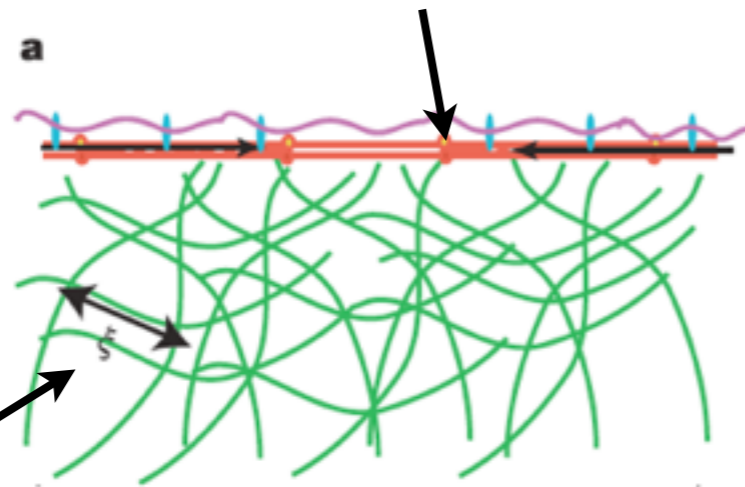
Guillaume T. Charras¹, Justin C. Yarrow¹, Mike A. Horton², L. Mahadevan^{1,3,4} & T. J. Mitchison¹

Vol 435|19 May 2005|doi:10.1038/nature03550

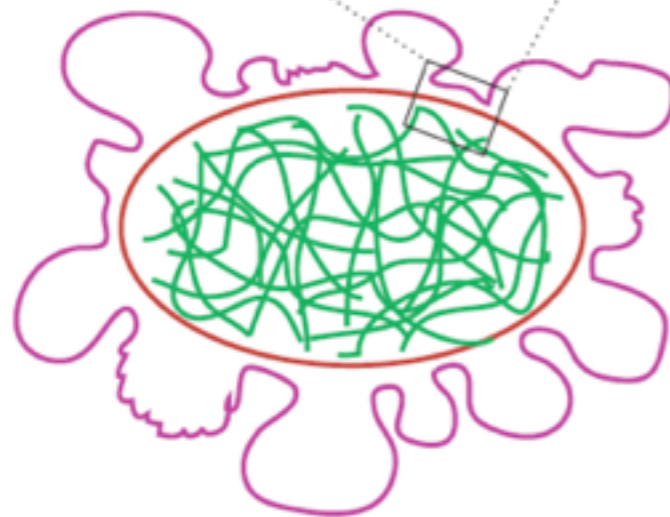


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Source: Charras, Guillaume T. et al. "Non-equilibration of hydrostatic pressure in blebbing cells." Nature 435, no. 7040 (2005): 365-369.

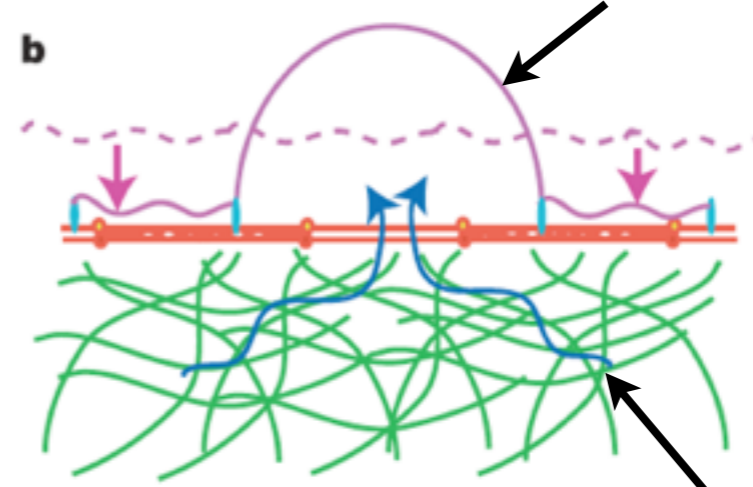
cortical acto-myosin contraction



pore-size or mesh-size



bleb forms (rupture event)



fluid flows through pores due to hydrostatic pressure

Courtesy of Macmillan Publishers Limited. Used with permission.
Source: Charras, Guillaume T. et al. "Non-equilibration of hydrostatic pressure in blebbing cells." Nature 435, no. 7040 (2005): 365-369.

together with drug studies discussed below, support the following qualitative model for bleb dynamics¹⁹. Cortical acto-myosin contracts (Supplementary Videos 10 and 11), generating hydrostatic pressure that causes a patch of plasma membrane to tear free from its attachment to the cortical cytoskeleton. This patch of cytoskeleton-free membrane rapidly inflates as cytosol flows in, with its base enlarging by further tearing (Supplementary Fig. 3). Later, inflation slows and a mesh of actin and myosin II assembles in the bleb to form a contractile cortex attached to the plasma membrane (Fig. 1b). Finally, contraction of this cortical mesh causes the bleb to shrink, driving the extruded cytosol back into the cell body.

Figures 7.14 and 7.15 removed due to copyright restrictions.
Source: Grodzinsky, Alan. Field, Forces and Flows in Biological
Systems. Garland Science, 2011.

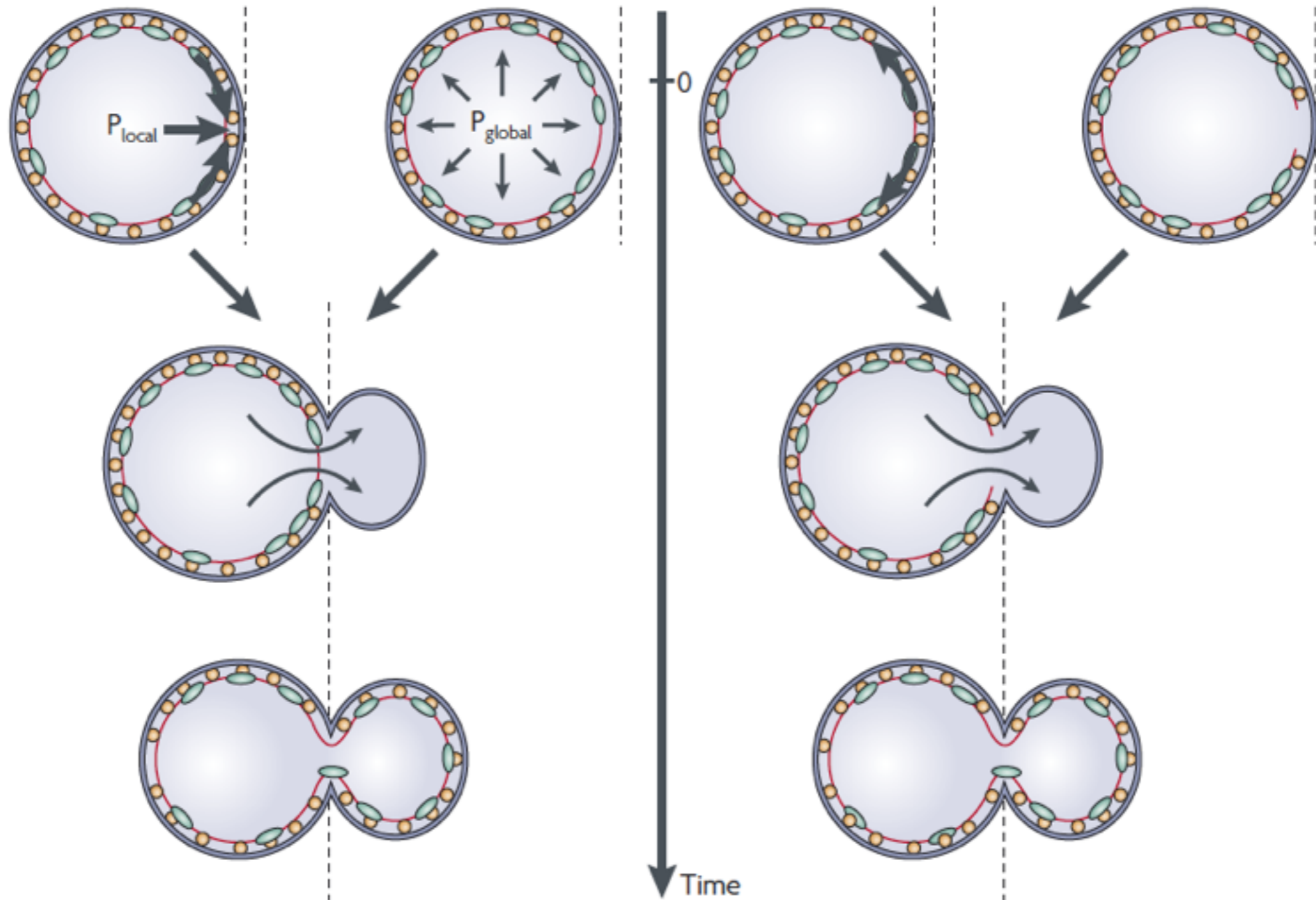
Fields, Forces, & Flows, Figures 7.14 & 7.15

Figure 7.13 and Equations 7.45 through 7.51 removed due to copyright restrictions.
Source: Grodzinsky, Alan. *Field, Forces and Flows in Biological Systems*. Garland Science, 2011.

Cortical Actomyosin: "Surface Tension"

a Bleb initiation by membrane detachment

b Bleb initiation by cortex rupture



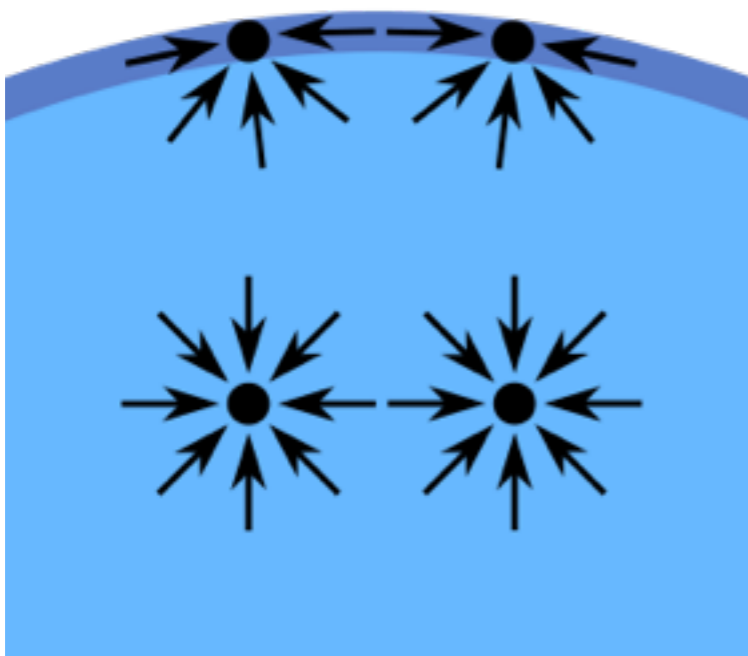
Charras & Paluch,
NatRev MCB 2008

Courtesy of Macmillan Publishers Limited. Used with permission.
Source: Charras, Guillaume and Ewa Paluch. "Blebs lead the way: how to migrate without lamellipodia." *Nature reviews Molecular cell biology* 9, no. 9 (2008): 730-736.

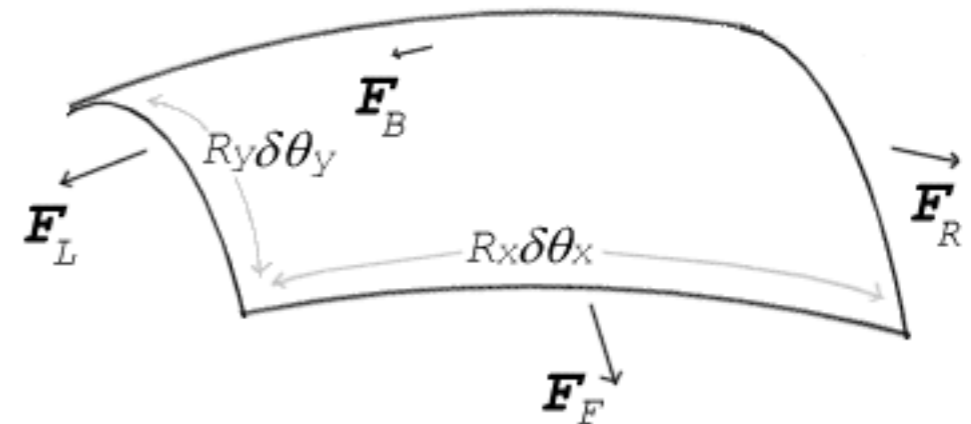
Cortical Actomyosin: “Surface Tension”



$$\Delta p = \gamma \left(\frac{1}{R_x} + \frac{1}{R_y} \right)$$



$$\gamma = \left(\frac{\partial G}{\partial A} \right)_{T,P,n}$$



$$\gamma = 1 \frac{\text{dyn}}{\text{cm}} = 1 \frac{\text{erg}}{\text{cm}^2} = 0.001 \frac{\text{N}}{\text{m}} = 0.001 \frac{\text{J}}{\text{m}^2}$$

Neutrophils behave like Newtonian fluids with surface tension:

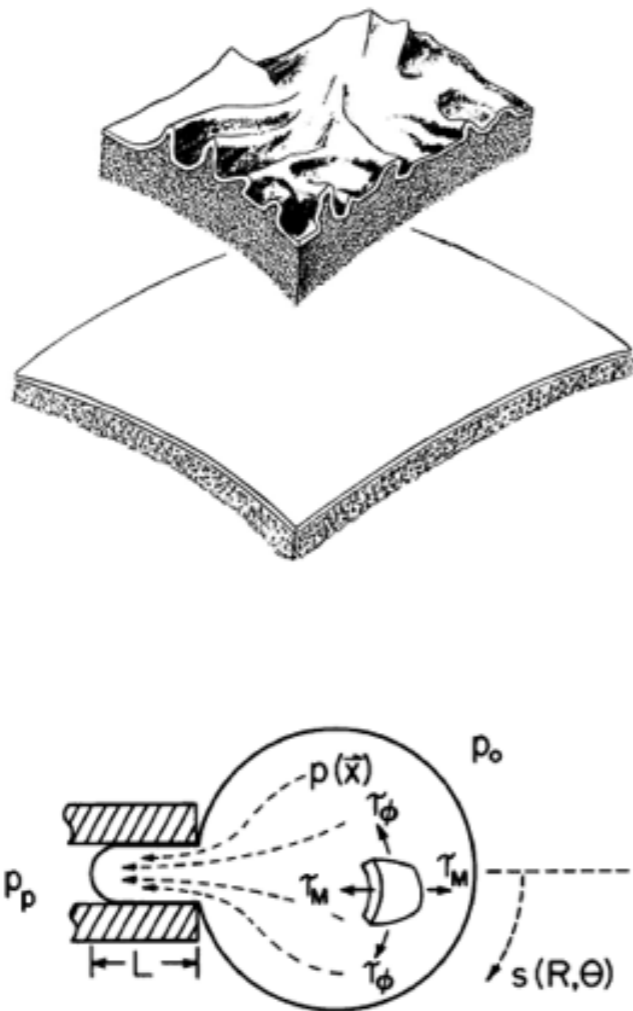


FIGURE 3 Schematic of the convergent flow into the pipet and the in-plane stress resultants supported by the cortical shell.

Evans & Yeung *Biophys J* 1989



FIGURE 1 Sequence of video micrographs of a blood granulocyte aspirated into a $3.5 \mu\text{m}$ caliber pipet: (a) first, formation of a hemispherical cap at the threshold pressure with no further flow; (b) next, the extent of flow after the suction was elevated above the threshold for a fixed time period then returned to the threshold value; and (c) recovery of the cell to its initial spherical form after release from the pipet.

Neutrophils behave like Newtonian fluids with surface tension:

Critical pressure for hemispherical cap:

$$P_{cr} = 2\bar{\tau}_o(1/R_p - 1/R_c)$$

Varying pipette radius:

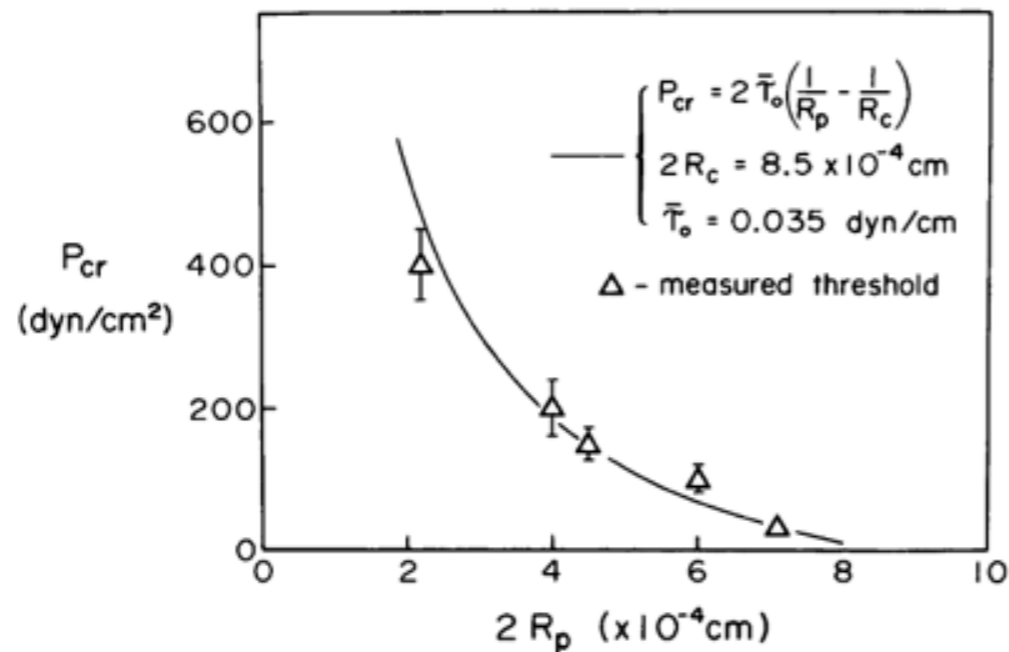


FIGURE 2. Threshold pressures required to form initial hemispherical projections inside pipets without further flow versus pipet radii. The solid curve is the theoretical correlation based on the presence of a persistent tension $\bar{\tau}_o = 0.035$ dyn/cm in the cell cortex. The average cell diameter was measured to be $8.5 \mu\text{m}$.

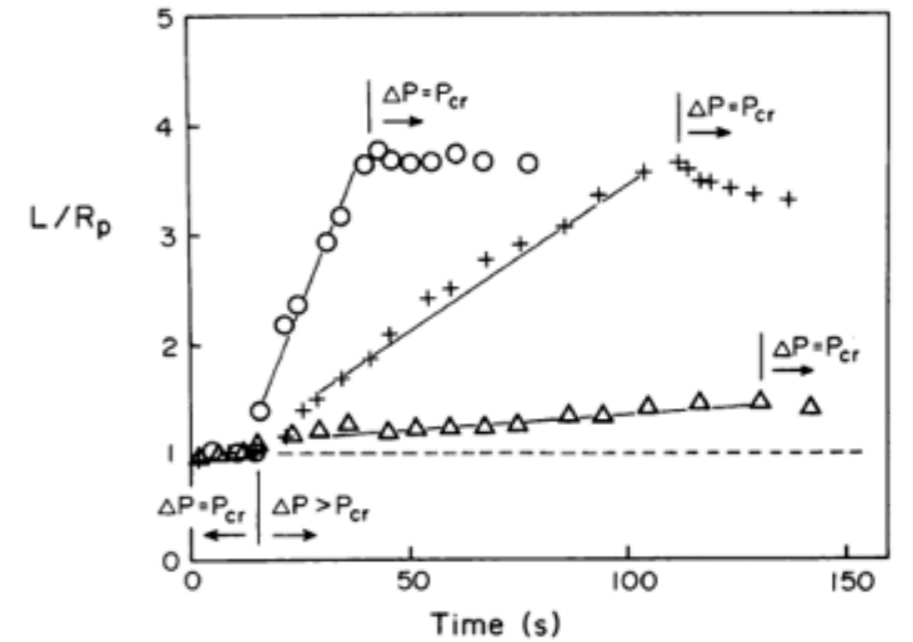
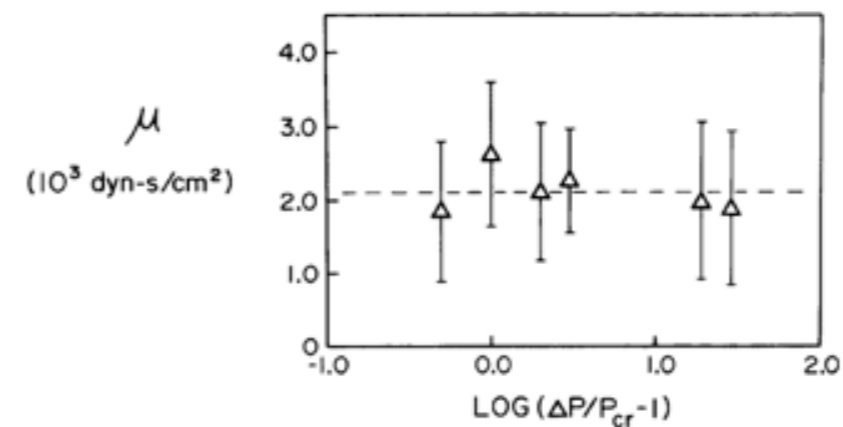


FIGURE 4. Measurements of projection length versus time for a single granulocyte subjected to three aspiration tests with a medium-size pipet where suction pressure was varied between 1.5–4.3 times the threshold value. This figure illustrates the proportional increase in entry flow rate with excess pressure above the threshold value.



Chondrocytes are more like elastic spheres

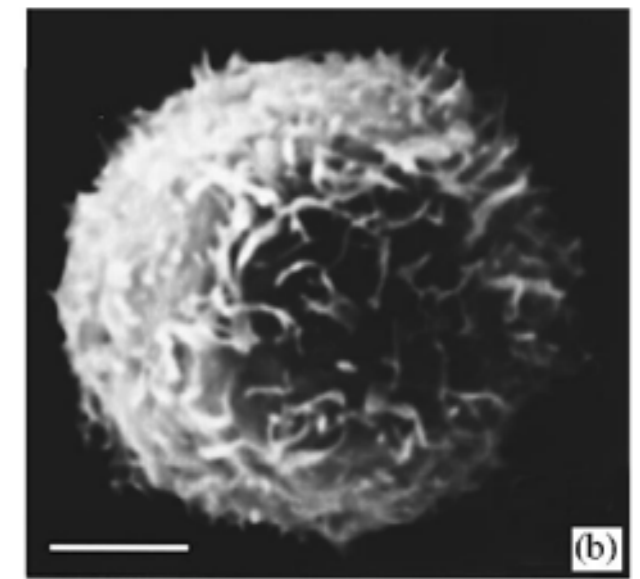
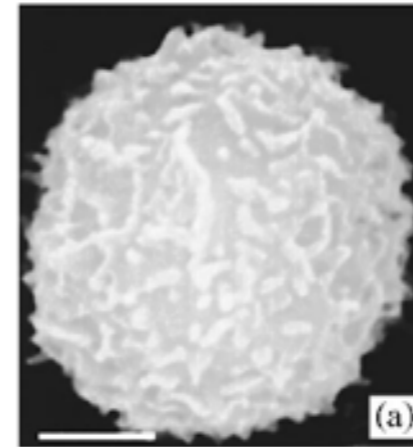


Fig. 3. Comparison of a human neutrophil (a) to a chondrocyte (b). The neutrophil has a diameter of about 8 μm while the majority of chondrocytes have diameters between about 12 and 16 μm . The scale bars indicate 2 μm , but note that the significant shrinkage of the cell has occurred during the preparation of the cells for scanning electron microscopy.

Table 1
“Natural” SI units at the level of the cell

	“Micro SI”	Application
Distance (m)	1 μm (10^{-6} m)	All
Force (N)	1 pN (10^{-12} N) 1 nN (10^{-9} N)	Molecular bonds “soft” cells Stiff cells
Pressure, stress (Pa)	1 pN/ μm^2 (1 Pa) 1 nN/ μm^2 (1 kPa)	Soft cells (blood cells) Stiff cells
Tension (mN/m)	1 pN/ μm (10^{-3} mN/m) 1 nN/ μm (1 mN/m)	Cortical elasticity of soft cells Elasticity of lipid bilayer

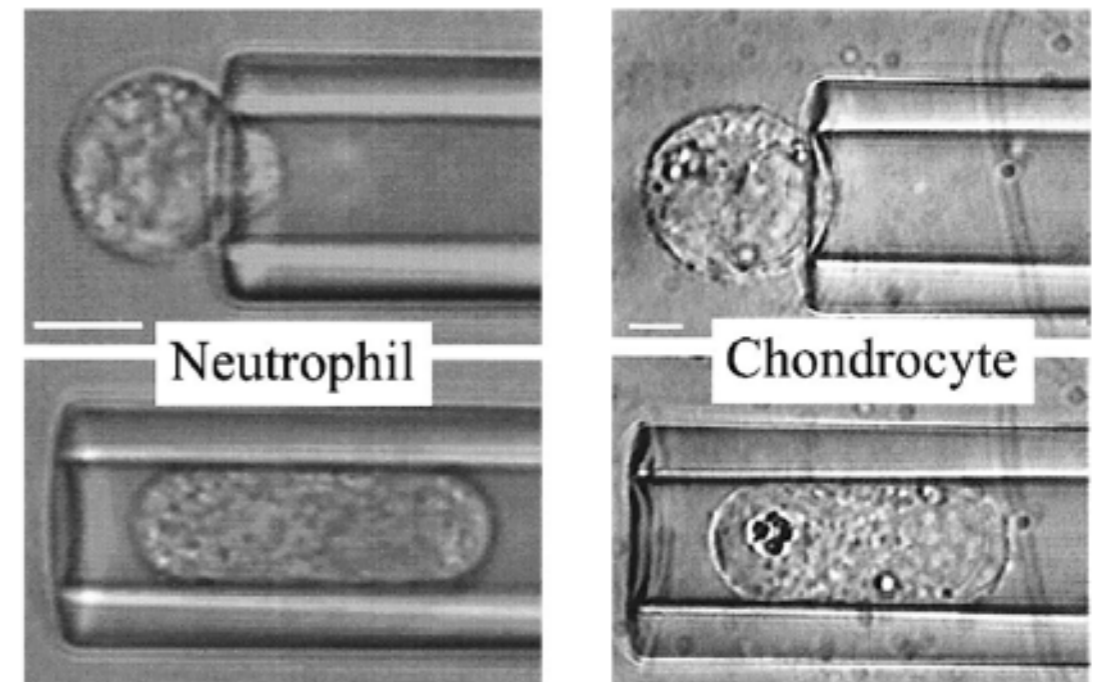


Fig. 4. A neutrophil and a chondrocyte each being aspirated into a micropipette. The photomicrographs of the chondrocyte are adapted from Jones et al. (1999). The scale bars indicate 5 μm .

Hochmuth *J Biomech* 2000

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