

Image of snake removed due to copyright restrictions.  
Source: Jain, Rakesh K. "An indirect way to tame cancer."  
Scientific American 310, no. 2 (2014): 46-53.

MEDICINE

# An Indirect Way to Tame Cancer

Cells and a stiff material called the matrix can squeeze blood vessels in tumors and thus block delivery of cancer-fighting drugs to malignant cells. Now researchers may have a way to reopen vessels and restore the drugs' power

*By Rakesh K. Jain*

Rakesh K. Jain  
*Scientific American*  
2014

Illustration by Brian Douglas

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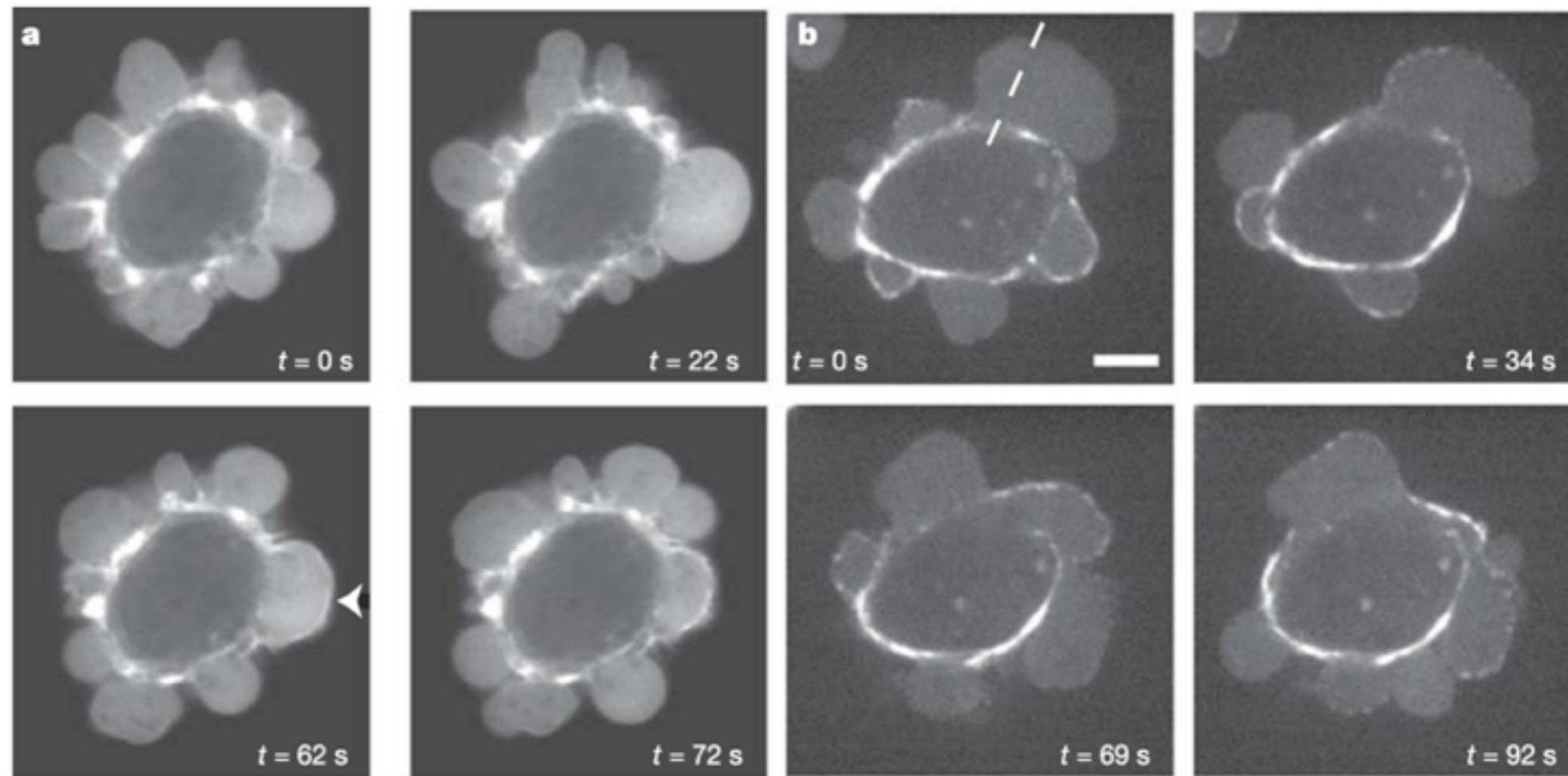
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Scientific American 310, no. 2 (2014): 46-53.

# Non-equilibration of hydrostatic pressure in blebbing cells

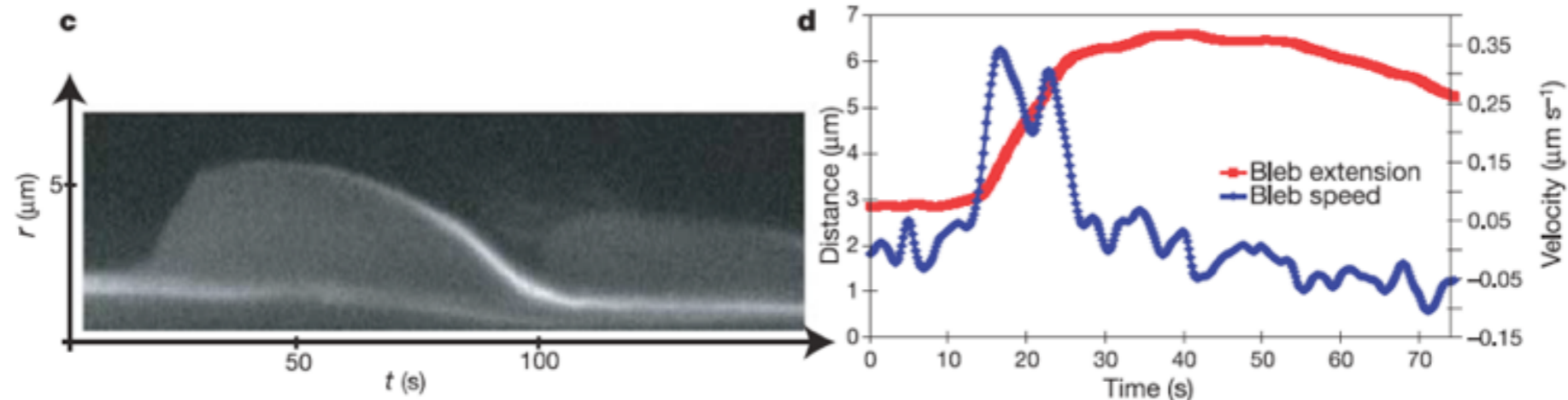
Guillaume T. Charras<sup>1</sup>, Justin C. Yarrow<sup>1</sup>, Mike A. Horton<sup>2</sup>, L. Mahadevan<sup>1,3,4</sup> & T. J. Mitchison<sup>1</sup>

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

Cortical  
actin  
assembly

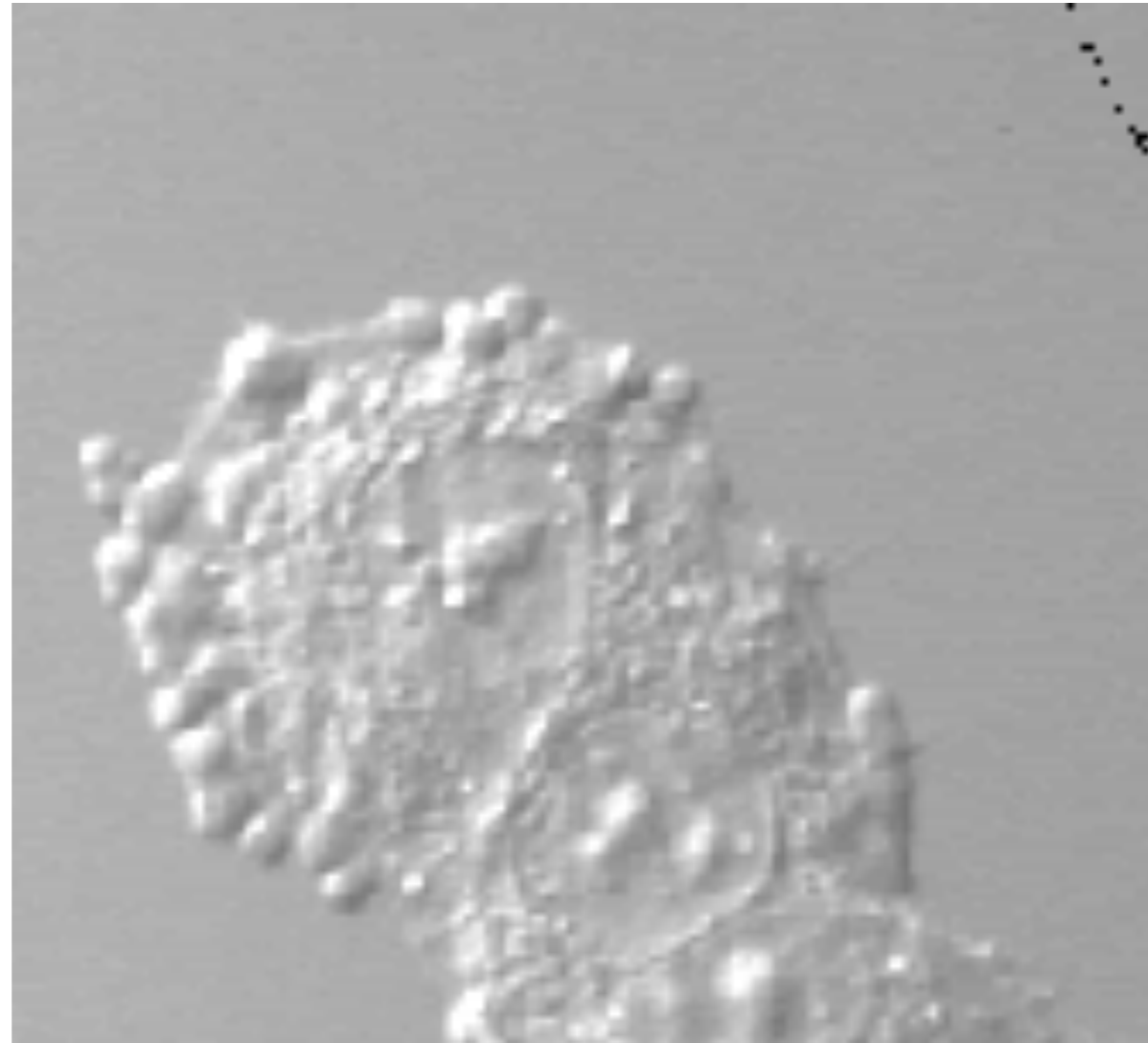


Cortical  
myosin  
assembly



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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.

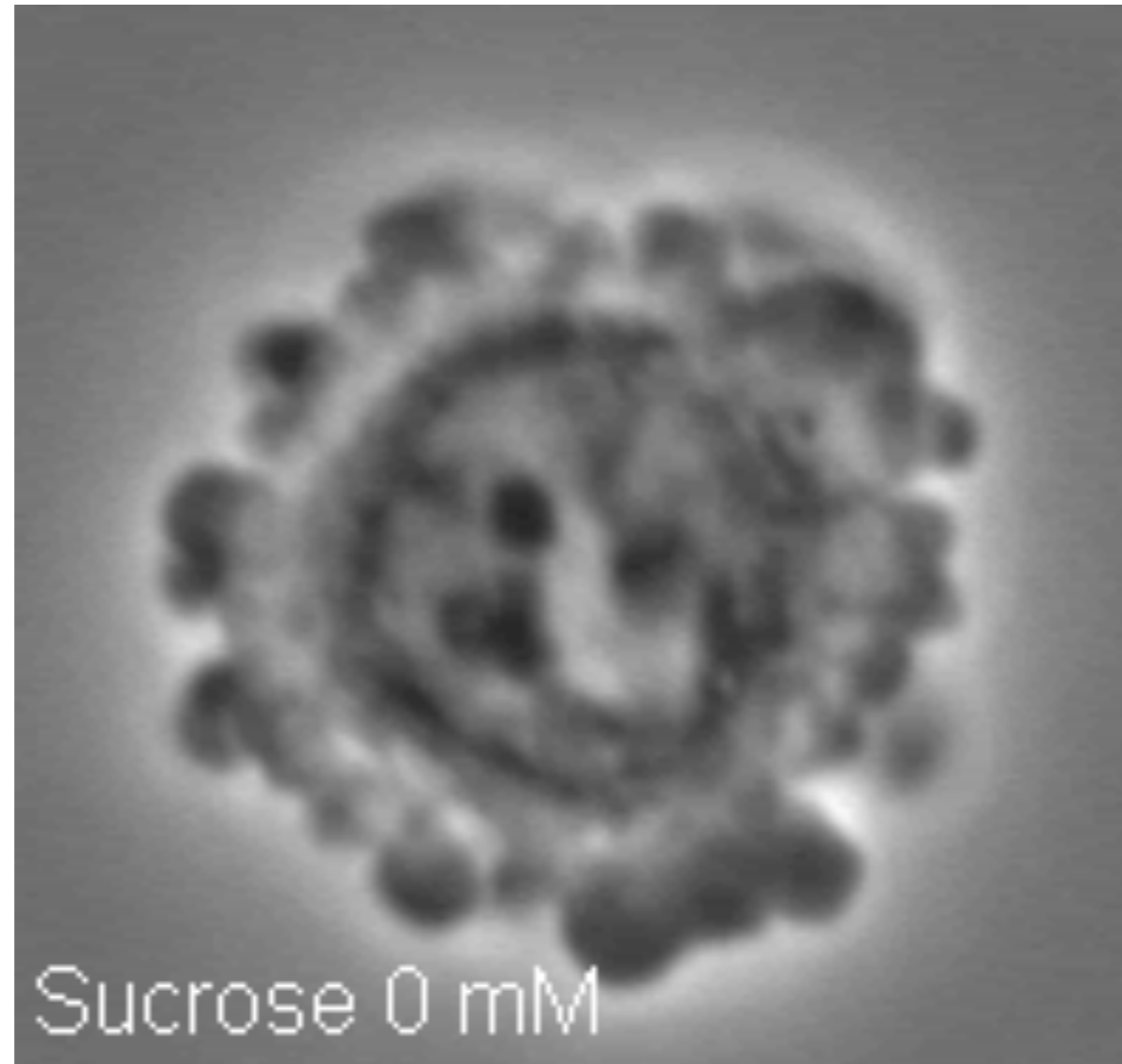
# Microplast (cell fragment) volume remains constant during blebbing



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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.

We first confirmed that the cell volume stays approximately constant during blebbing (Supplementary Data, Supplementary Video 12 and Supplementary Fig. 4)<sup>18</sup>, implying that blebbing is driven primarily by flow of fluid within the cell rather than water crossing the plasma membrane. We then confirmed that bleb

# Increasing extracellular osmolarity makes blebs smaller

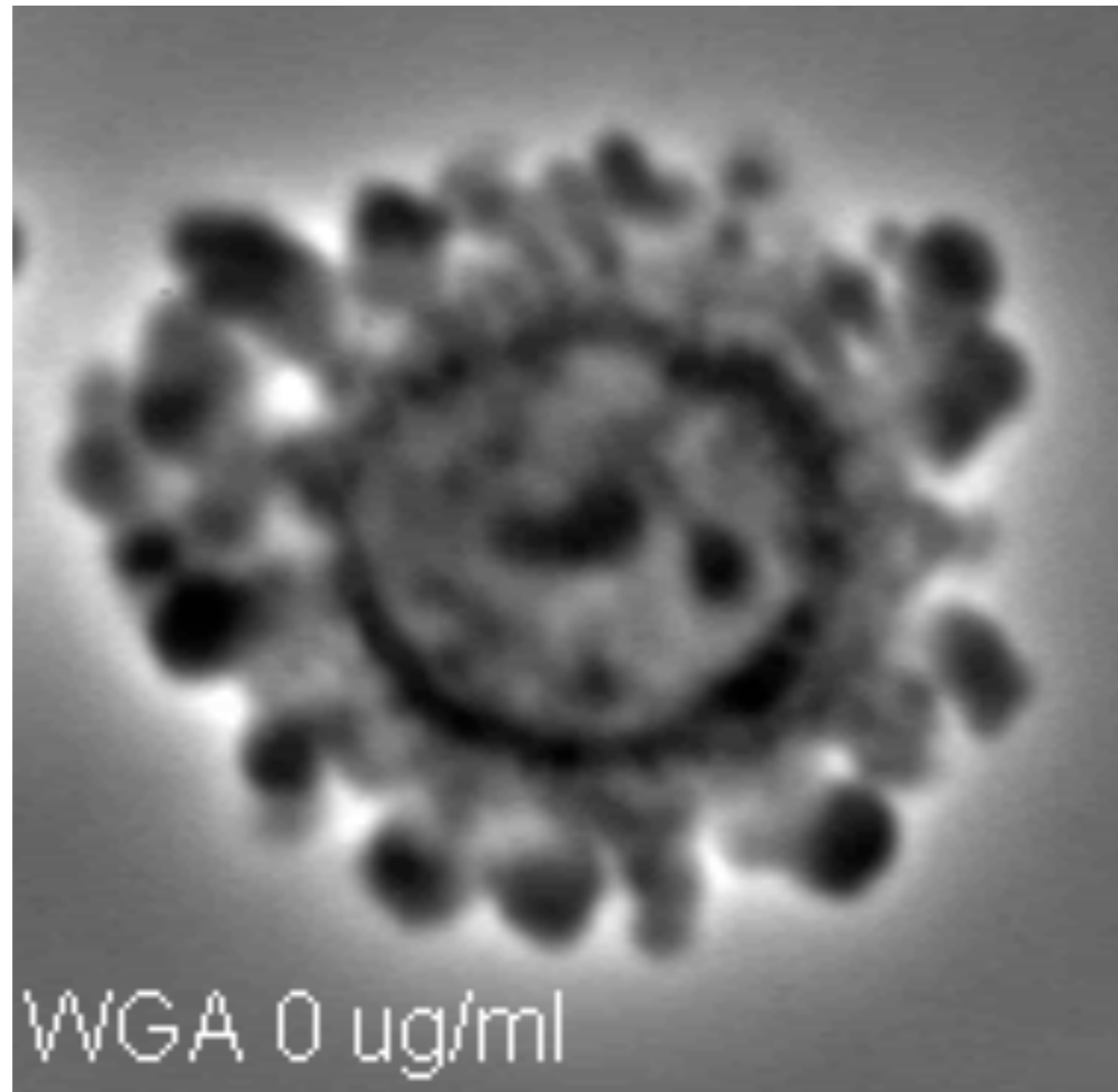


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Source: Charras, Guillaume T. et al. "Non-equilibration of hydrostatic pressure in blebbing cells." *Nature* 435, no. 7040 (2005): 365-369.

contraction. Increasing extracellular osmolarity made blebs smaller, whereas decreasing it made them larger<sup>22</sup> (Fig. 1; see also Supplementary Video 1). Increasing membrane rigidity by crosslinking the



## Increasing membrane rigidity inhibits blebbing

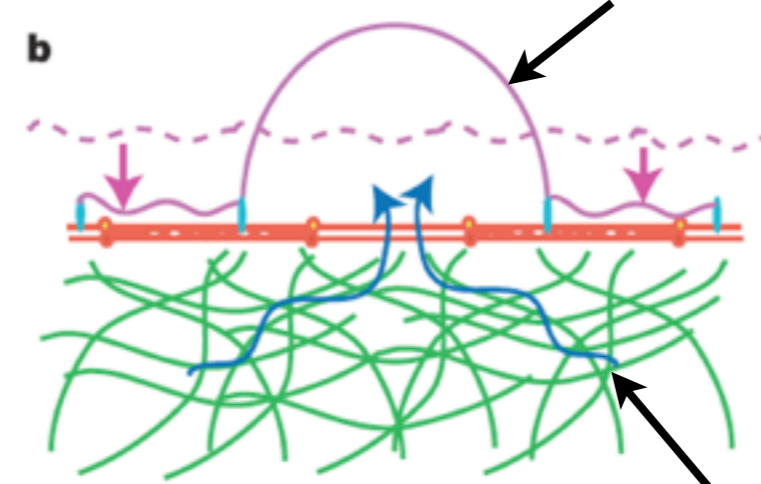
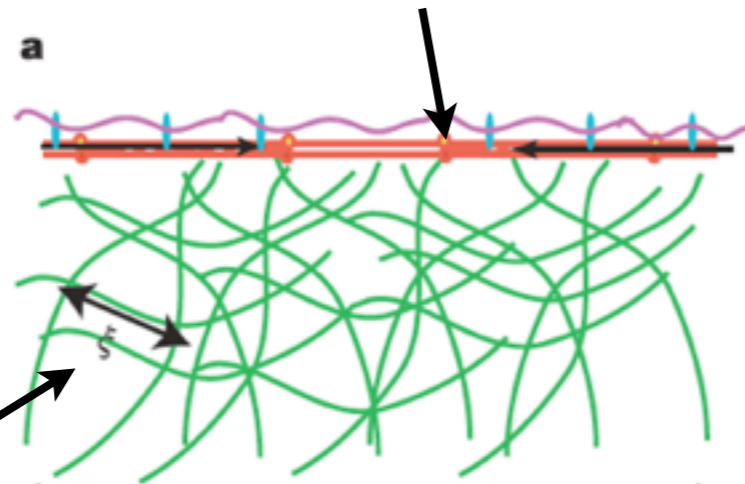


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Source: Charras, Guillaume T. et al. "[Non-equilibration of hydrostatic pressure in blebbing cells.](#)" Nature 435, no. 7040 (2005): 365-369.

tary Video 1). **Increasing membrane rigidity by crosslinking the glyocalix polysaccharides with wheat germ agglutinin (WGA)<sup>23</sup> inhibits blebbing (Supplementary Video 2).** These observations,

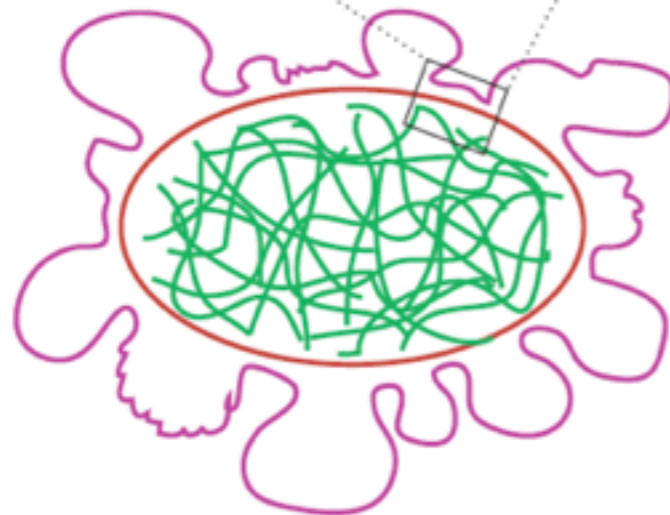
cortical acto-myosin contraction

bleb forms (rupture event)



pore-size or mesh-size

fluid flows through pores due to hydrostatic pressure



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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.

together with drug studies discussed below, support the following qualitative model for bleb dynamics<sup>19</sup>. 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

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