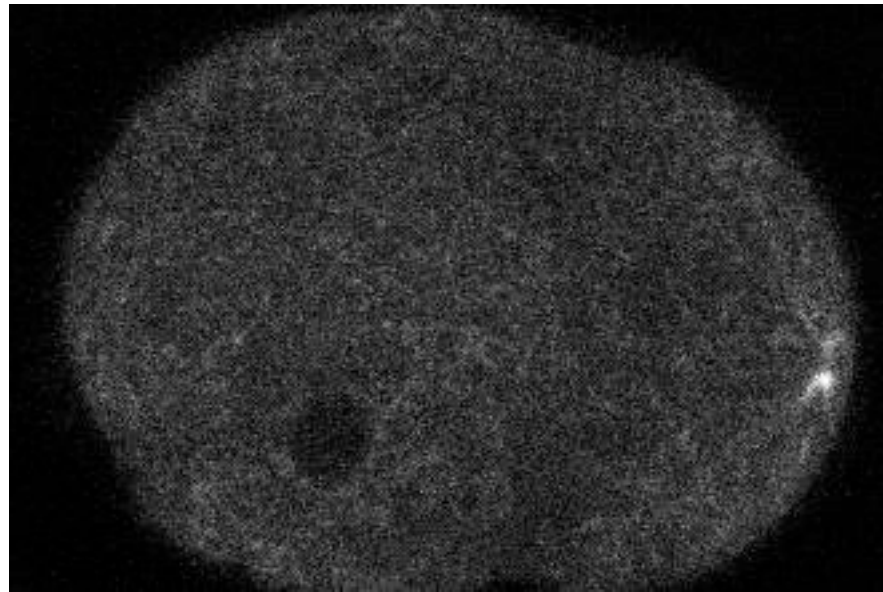


# Cytoskeleton-based positioning during development, cell division

---



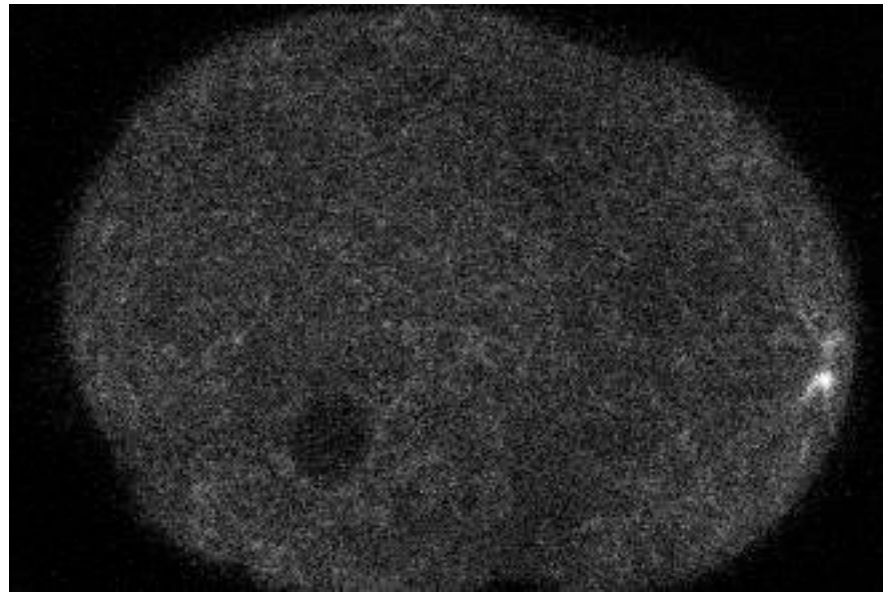
C elegans fertilization - pronuclei  
migration and reorientation  
Ed Munro and Chris Schoff (Center for  
Cell Dynamics)

C elegans fertilization - chromosome alignment  
and segregation  
George von Dassow (Center for Cell Dynamics)



# Cytoskeleton-based positioning during development, cell division

---



C elegans fertilization - pronuclei  
migration and reorientation  
Ed Munro and Chris Schoff (Center for  
Cell Dynamics)

C elegans fertilization - chromosome alignment  
and segregation  
George von Dassow (Center for Cell Dynamics)



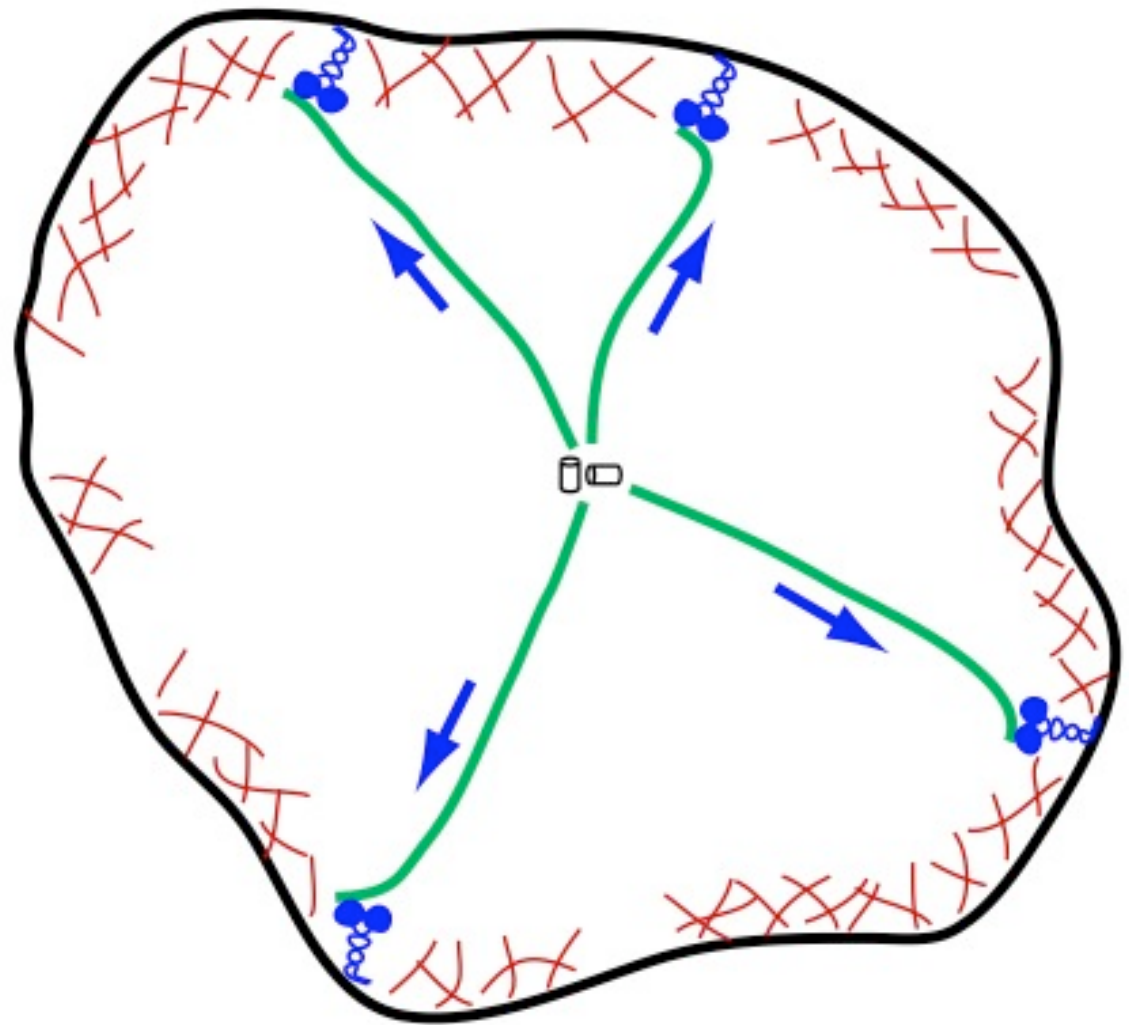
# What forces are involved?

---

# What forces are involved?

---

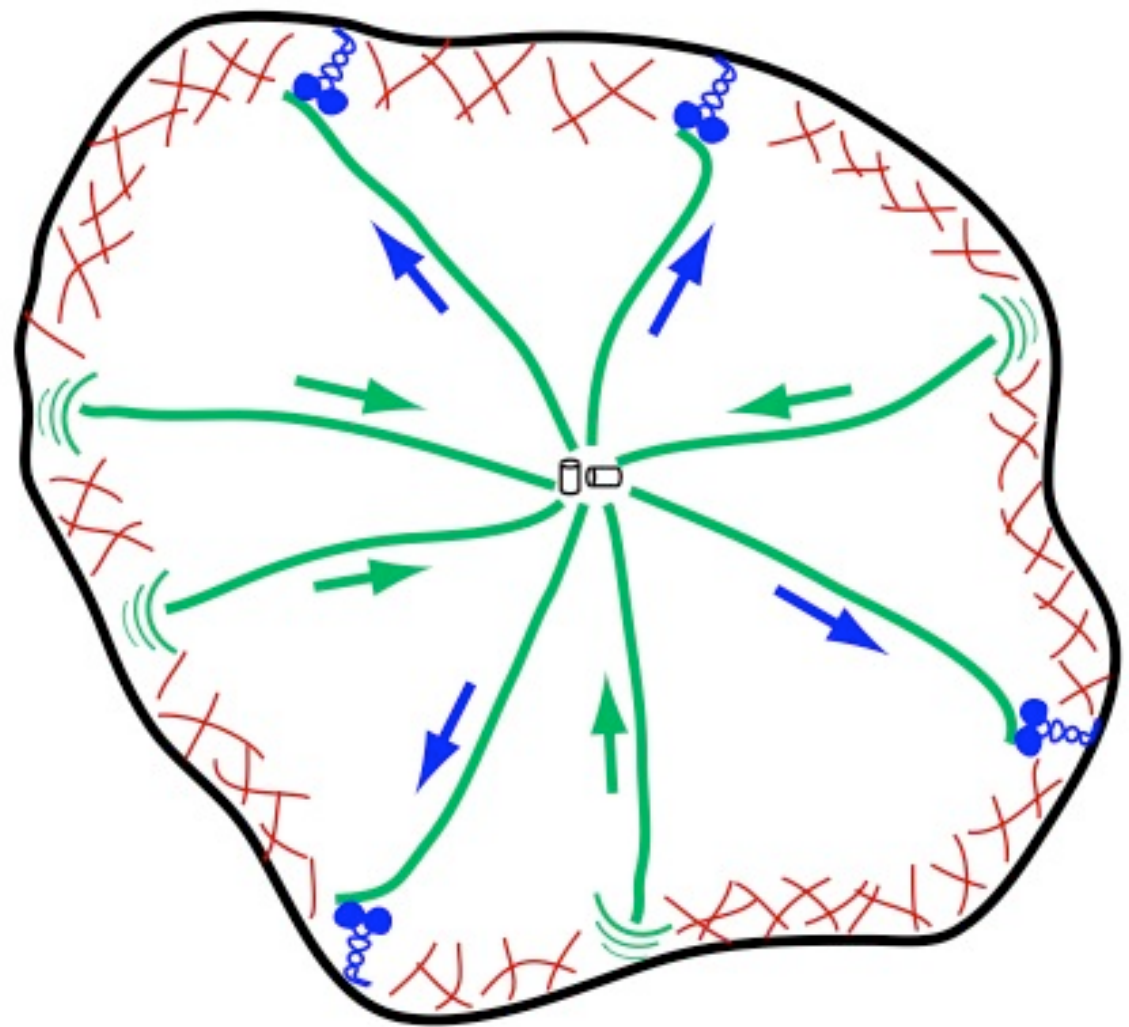
- Cortical dynein pulls on aster.



# What forces are involved?

---

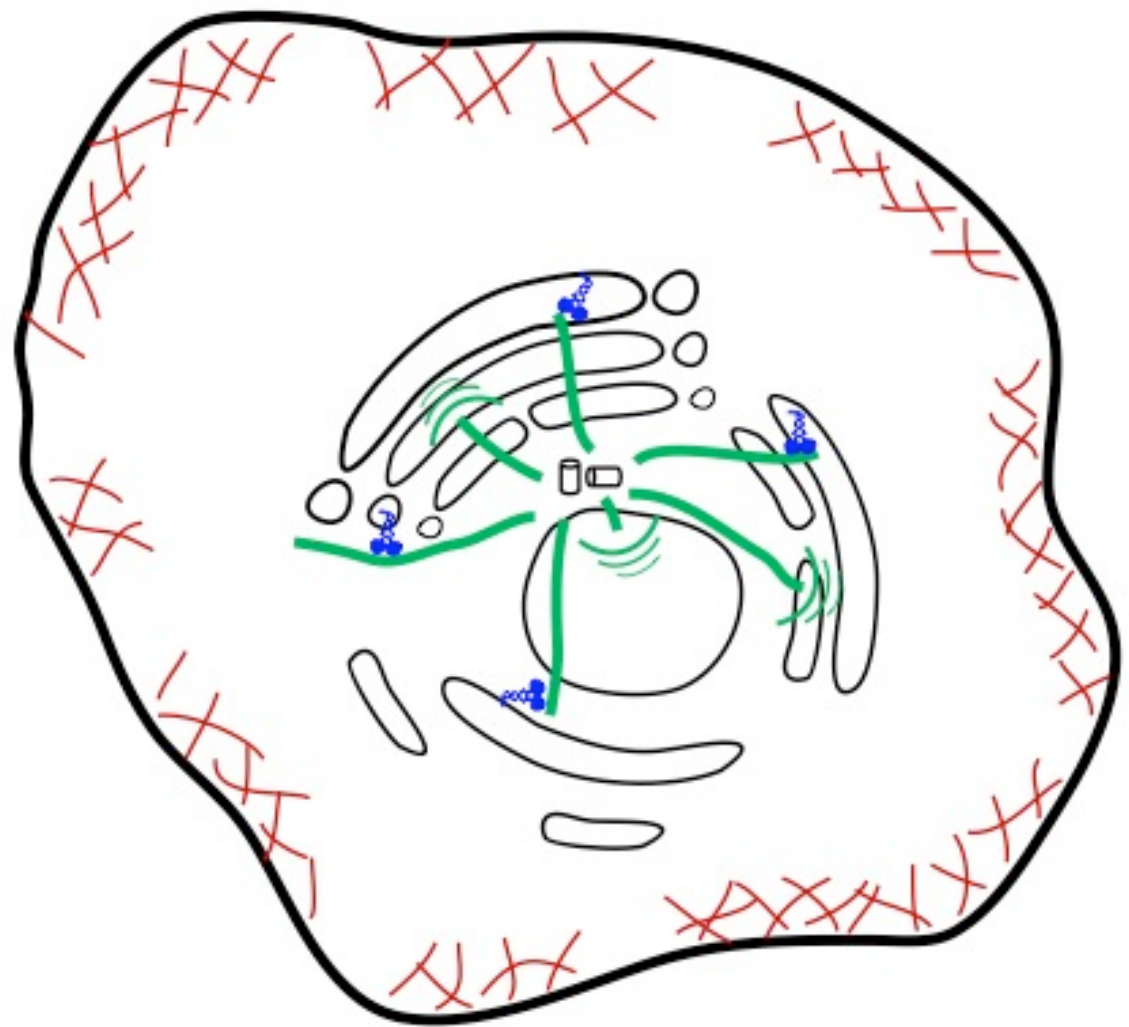
- Cortical dynein pulls on aster.
- MTs push against cortex by polymerizing.



# What forces are involved?

---

- Cortical dynein pulls on aster.
- MTs push against cortex by polymerizing.
- Motors push/pull against organelles.





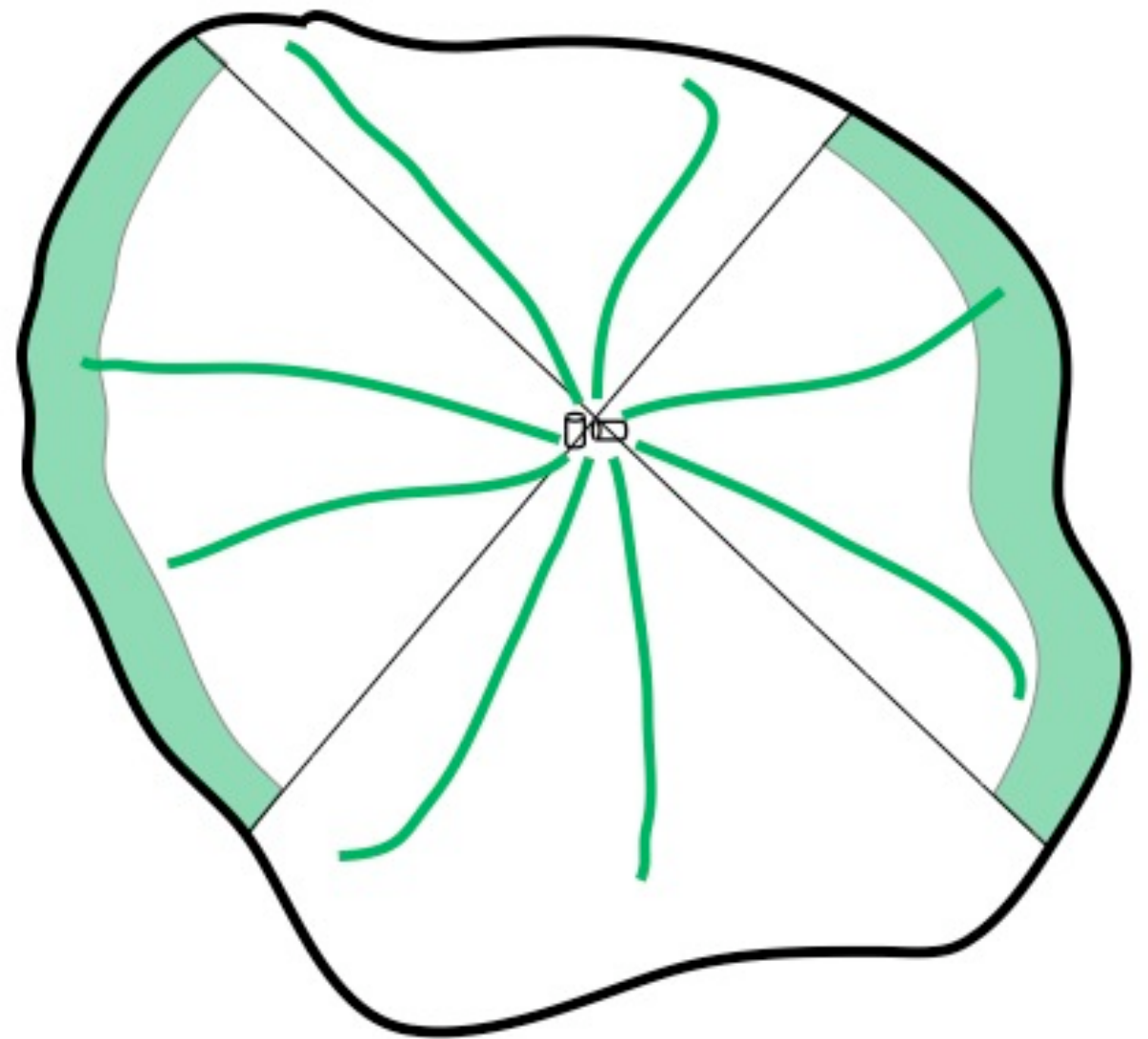
# Forces, steady states and their stability

---

# Forces, steady states and their stability

---

- Cortical pulling - destabilizing.

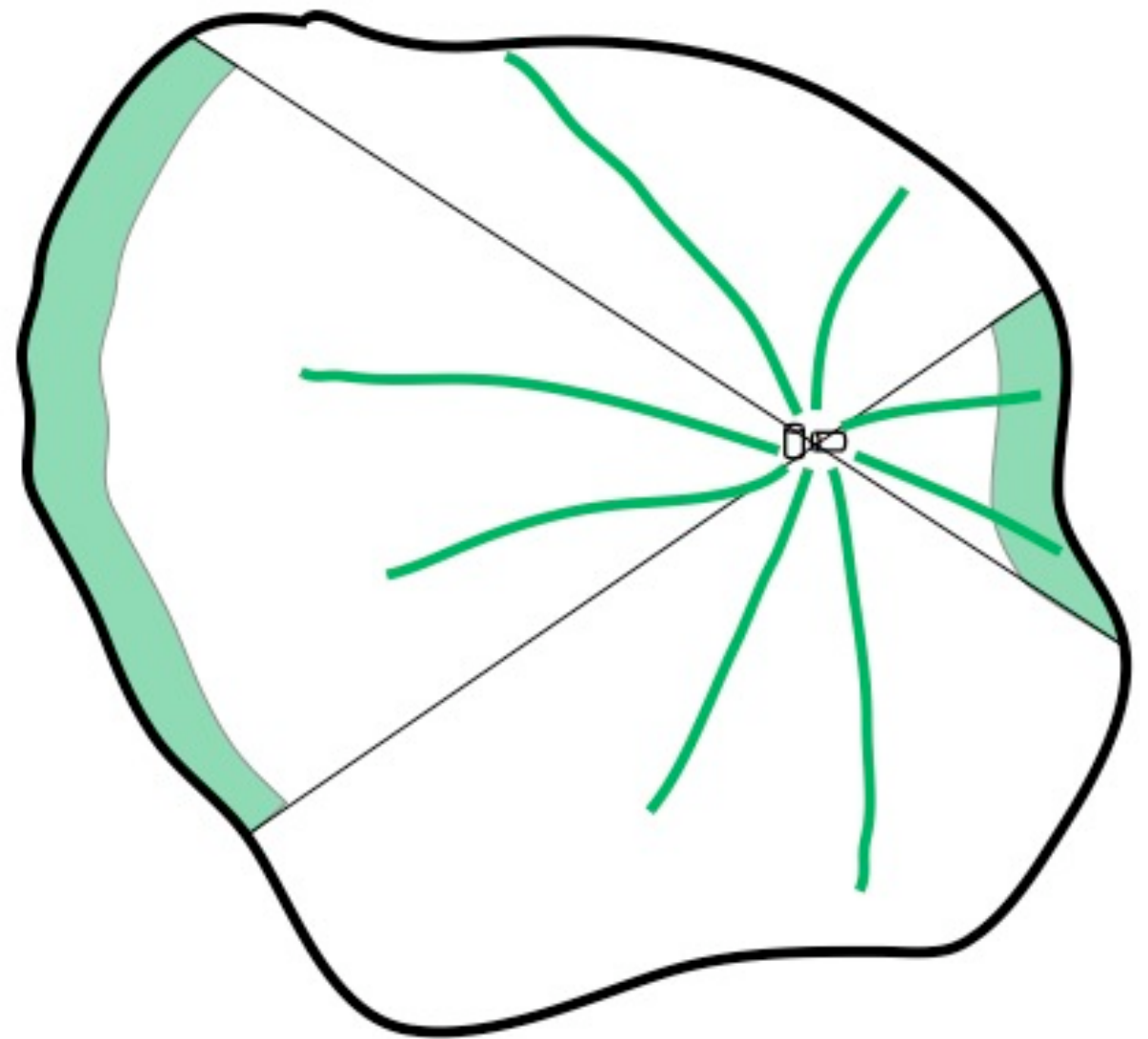




# Forces, steady states and their stability

---

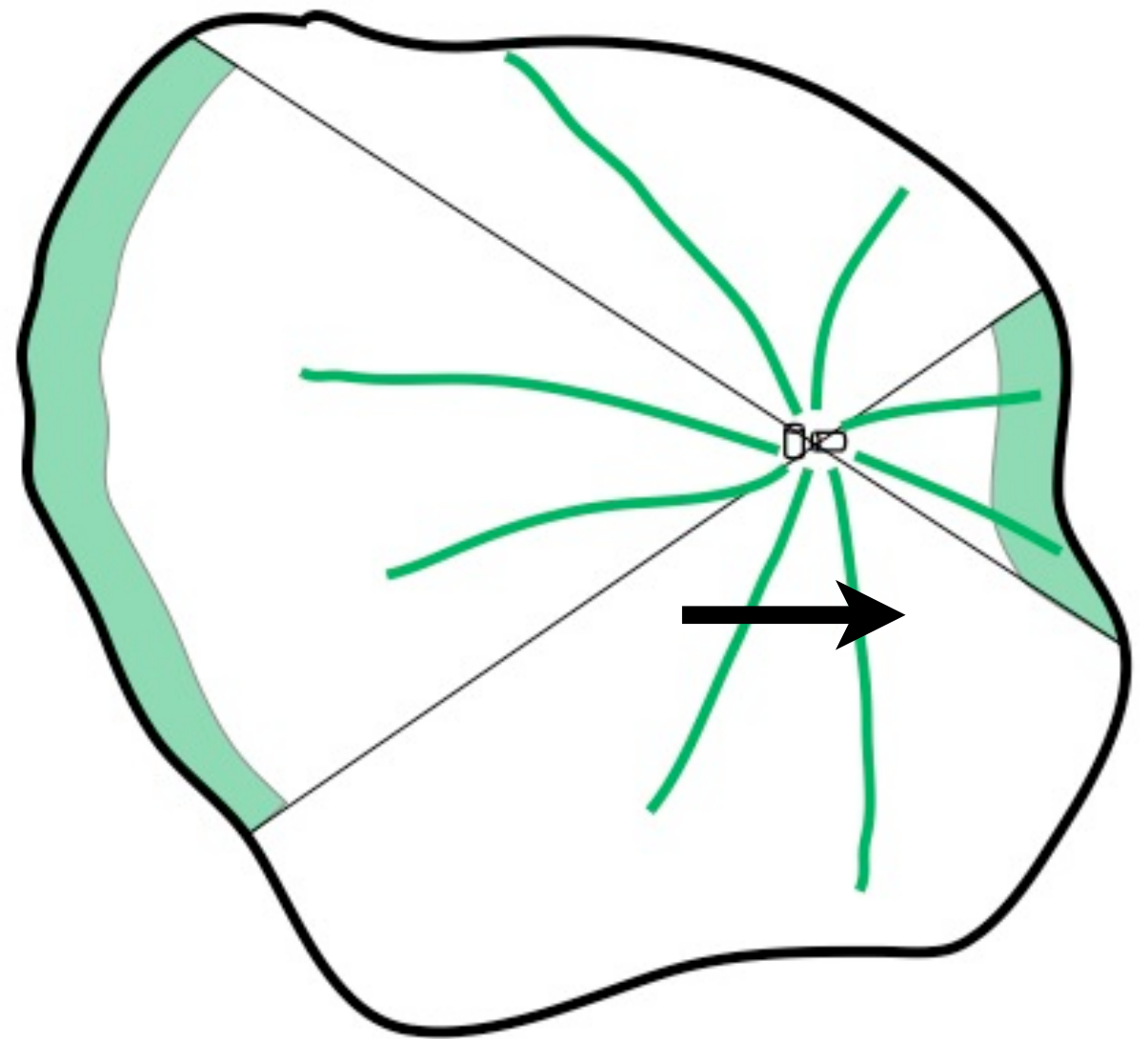
- Cortical pulling - destabilizing.



# Forces, steady states and their stability

---

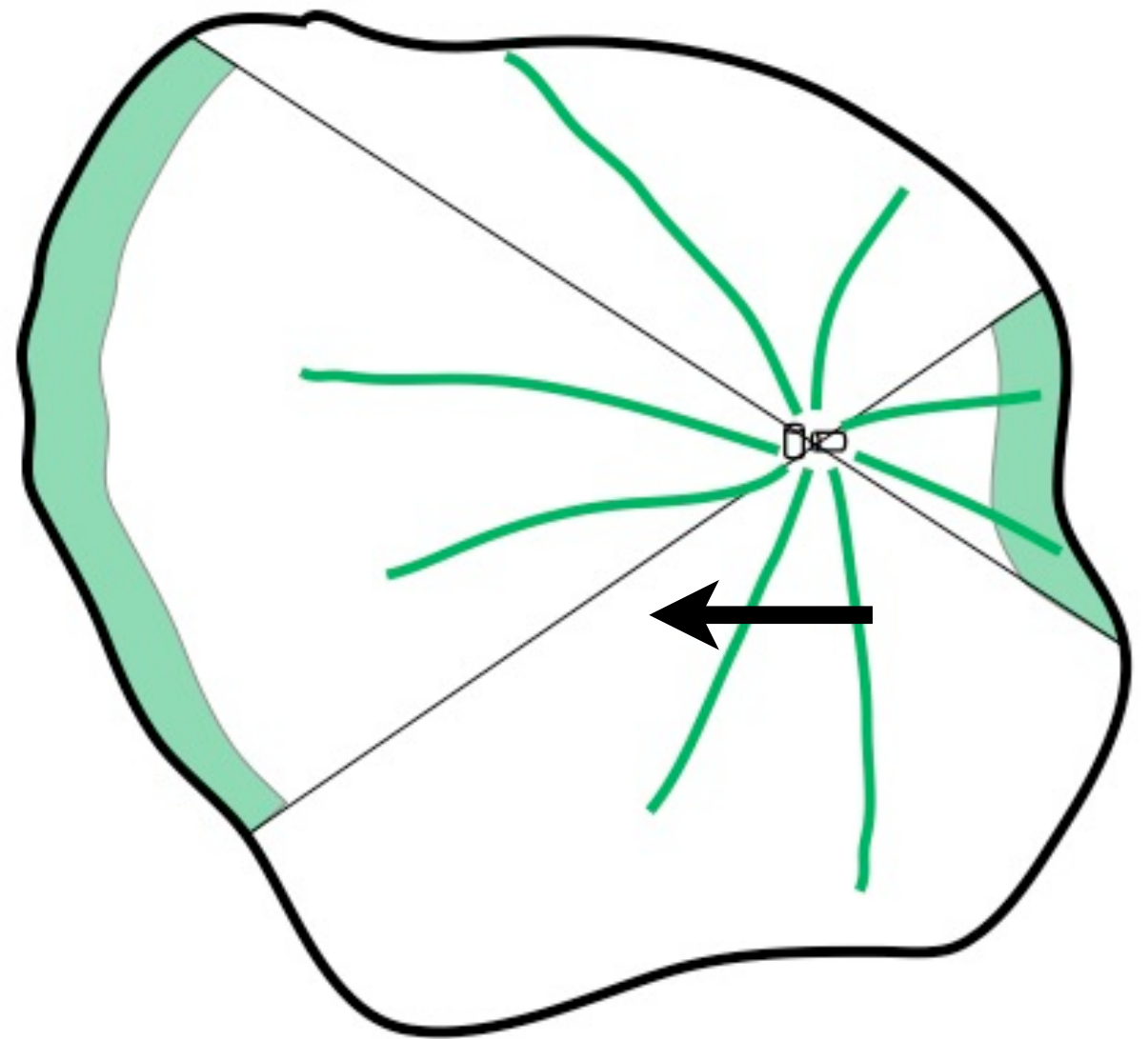
- Cortical pulling - destabilizing.



# Forces, steady states and their stability

---

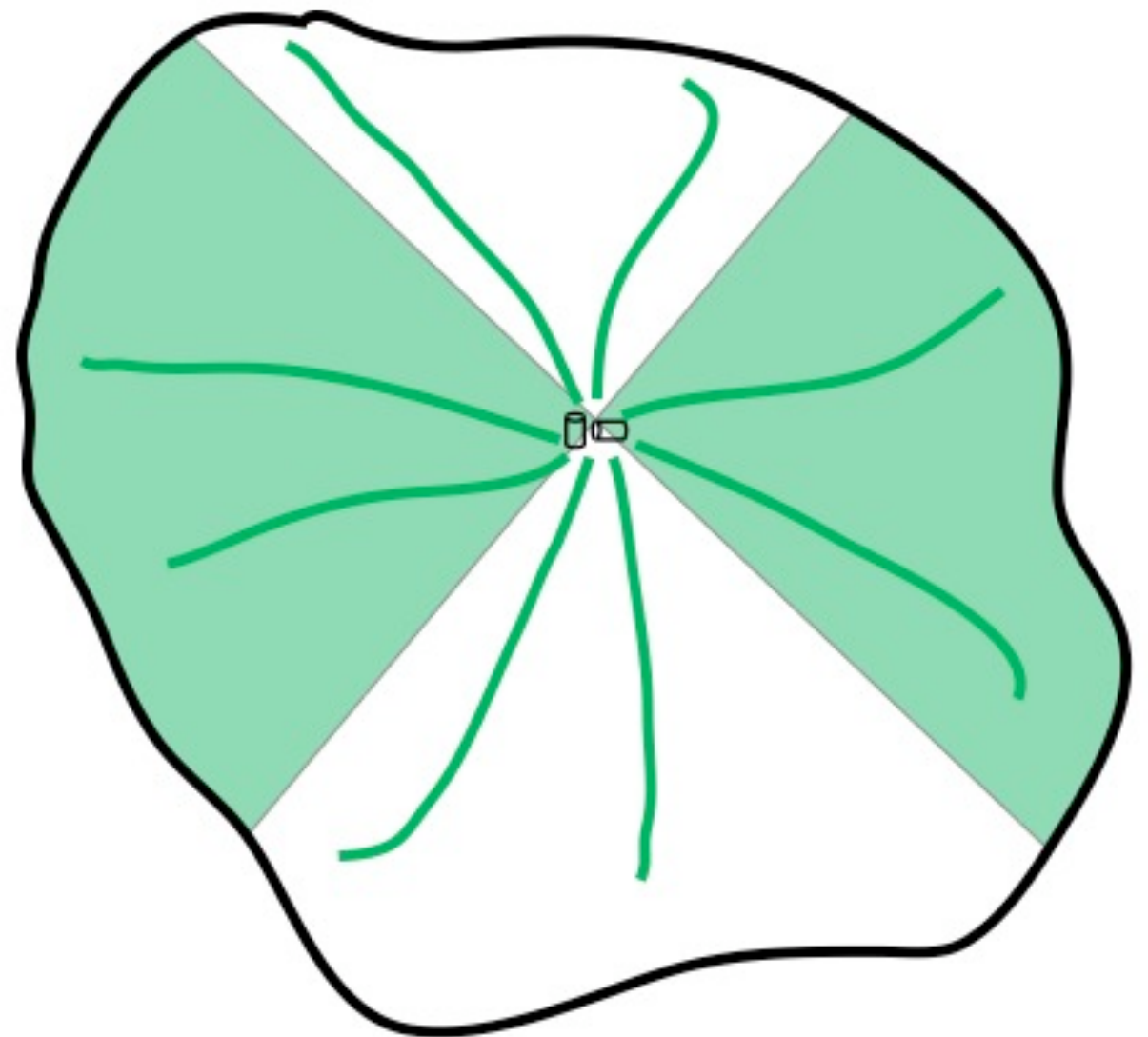
- Cortical pulling - destabilizing.
- Cortical pushing - stabilizing.



# Forces, steady states and their stability

---

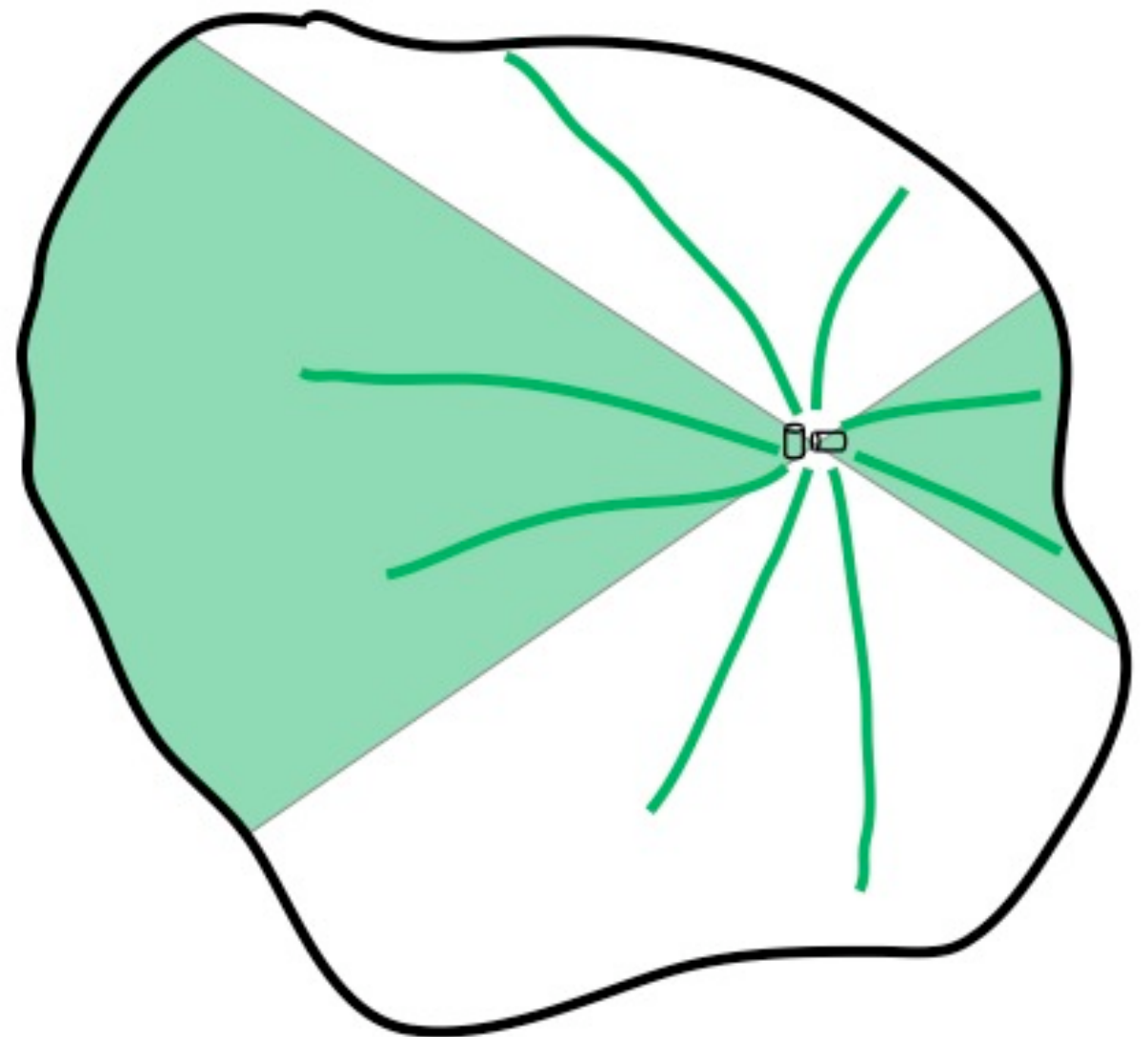
- Cortical pulling - destabilizing.
- Cortical pushing - stabilizing.
- Cytoplasmic pulling - stabilizing.



# Forces, steady states and their stability

---

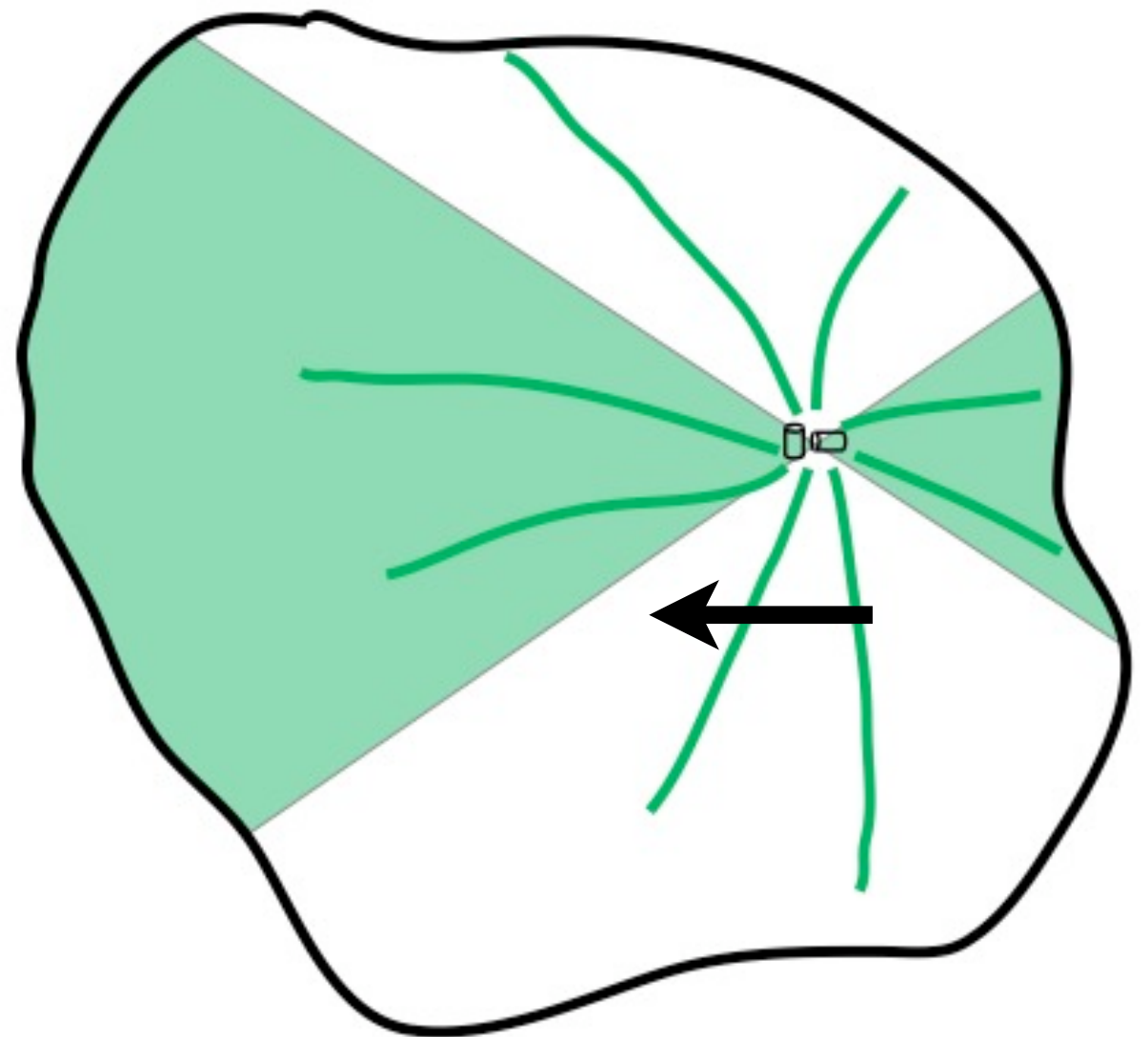
- Cortical pulling - destabilizing.
- Cortical pushing - stabilizing.
- Cytoplasmic pulling - stabilizing.



# Forces, steady states and their stability

---

- Cortical pulling - destabilizing.
- Cortical pushing - stabilizing.
- Cytoplasmic pulling - stabilizing.

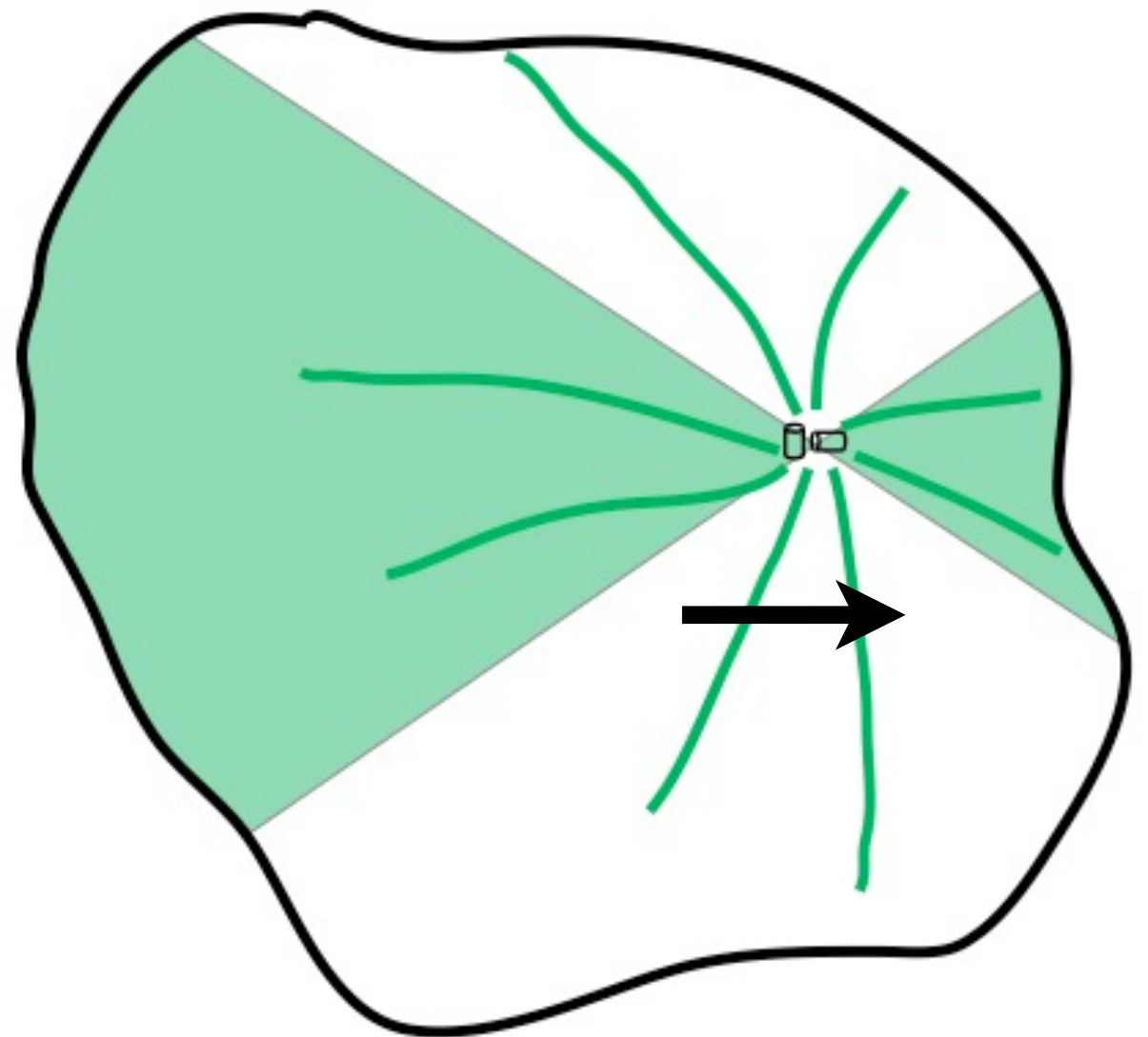




# Forces, steady states and their stability

---

- Cortical pulling - destabilizing.
- Cortical pushing - stabilizing.
- Cytoplasmic pulling - stabilizing.
- Cytoplasmic pushing - destabilizing.

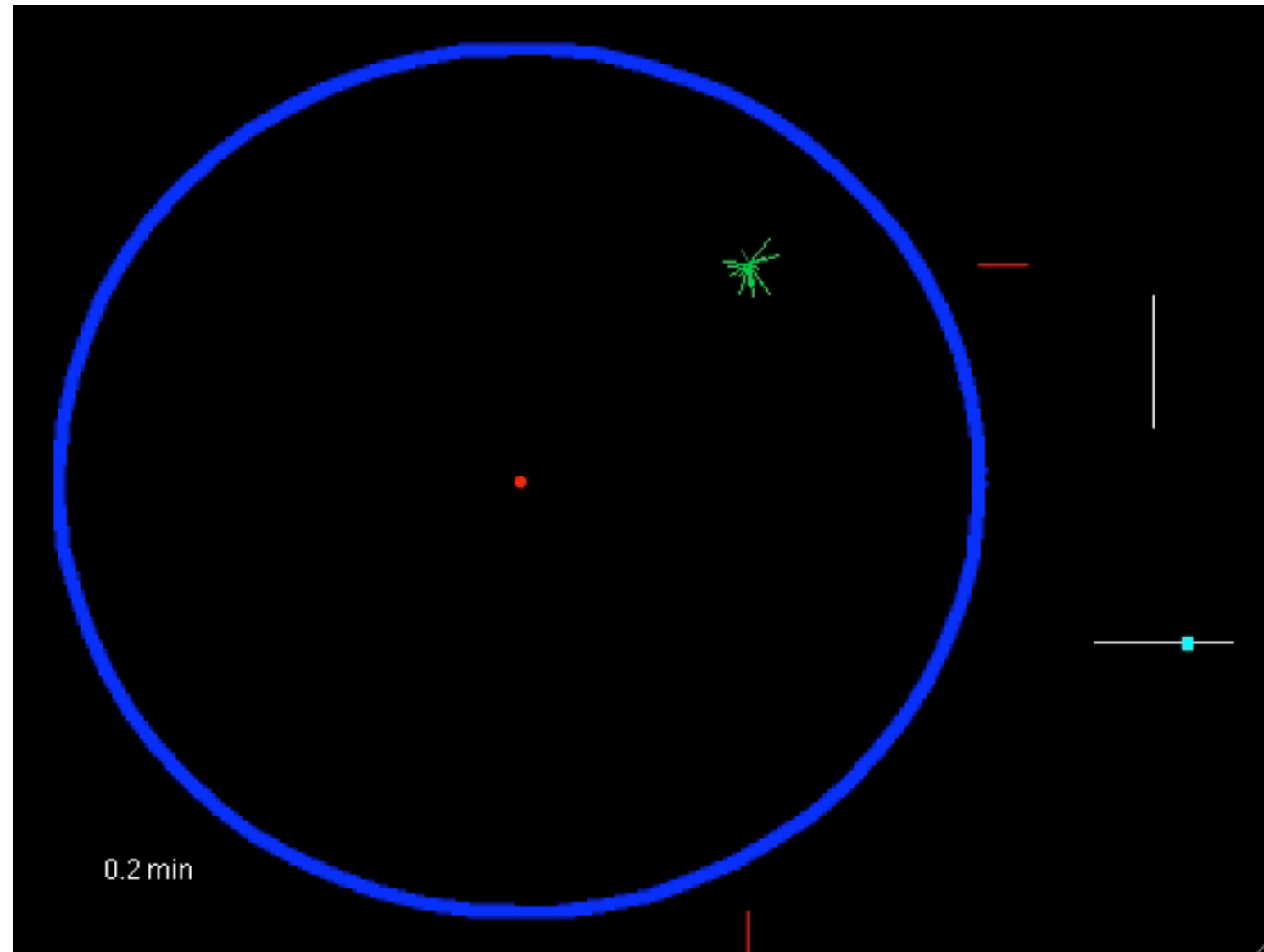


# Two-D simulation - cytoplasmic pulling

---

# Two-D simulation - cytoplasmic pulling

---



# One-D model of motor-based centering

---

Microtubule (MT) dynamics - stochastic transitions between growing and shrinking states

$$\frac{\partial g}{\partial t} = -v_g \frac{\partial g}{\partial l} - cg + rs \quad \text{(density of growing MTs)}$$

$$\frac{\partial s}{\partial t} = v_s \frac{\partial s}{\partial l} + cg - rs \quad \text{(density of shrinking MTs)}$$

# One-D model of motor-based centering

---

Microtubule (MT) dynamics - stochastic transitions between growing and shrinking states

$$\frac{\partial g}{\partial t} = -v_g \frac{\partial g}{\partial l} - cg + rs \quad (\text{density of growing MTs})$$

$$\frac{\partial s}{\partial t} = v_s \frac{\partial s}{\partial l} + cg - rs \quad (\text{density of shrinking MTs})$$

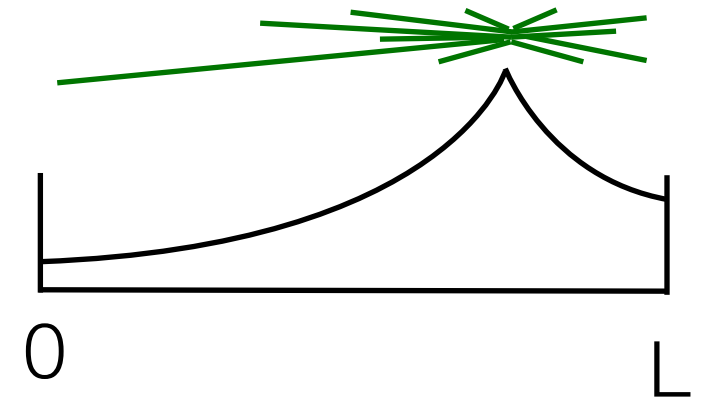
$$g(l) + s(l) = Ae^{-\lambda l} \quad \left( \text{where } \lambda = \frac{v_s c - v_g r}{v_g v_s} \right)$$

(density of MT of length  $l$ )

# One-D model - calculating forces from MT densities

---

$$Tips(x) = Ae^{-\lambda|x-x_c|}$$

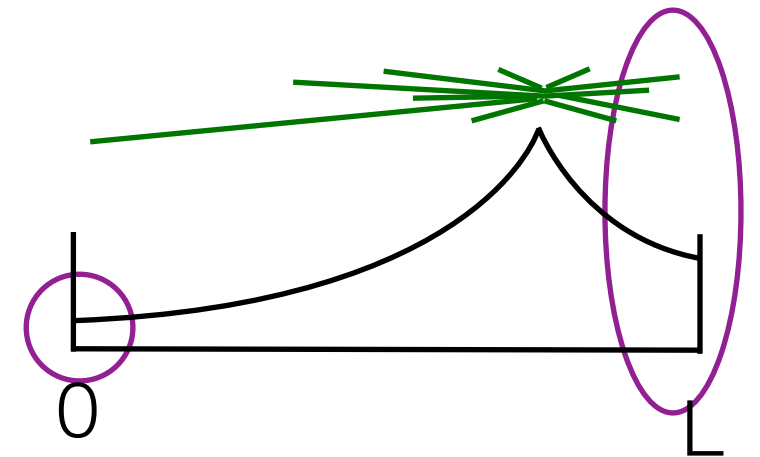




# One-D model - calculating forces from MT densities

---

$$Tips(x) = Ae^{-\lambda|x-x_c|}$$



If motors pull only at the cell periphery,

$$F_{pull}(x_c) = B(-e^{-\lambda x_c} + e^{-\lambda(L-x_c)})$$

# One-D model - calculating forces from MT densities

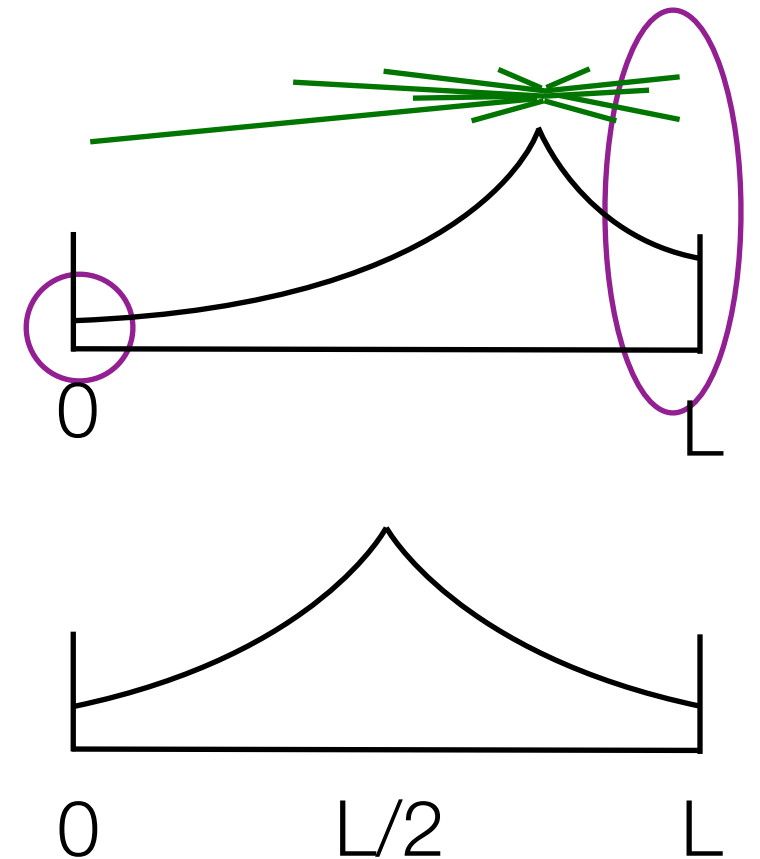
---

$$Tips(x) = Ae^{-\lambda|x-x_c|}$$

If motors pull only at the cell periphery,

$$F_{pull}(x_c) = B(-e^{-\lambda x_c} + e^{-\lambda(L-x_c)})$$

and the steady state position is at  $x_c = L/2$ .



# One-D model - calculating forces from MT densities

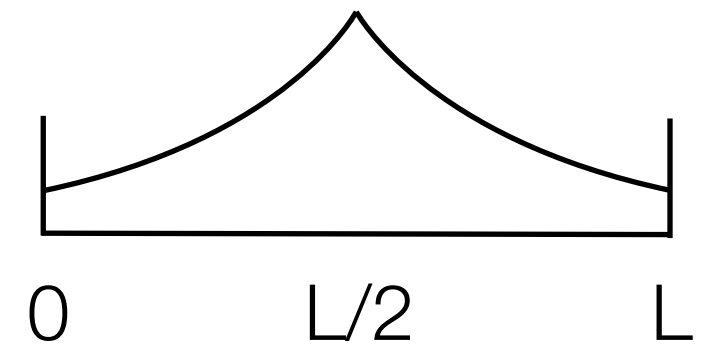
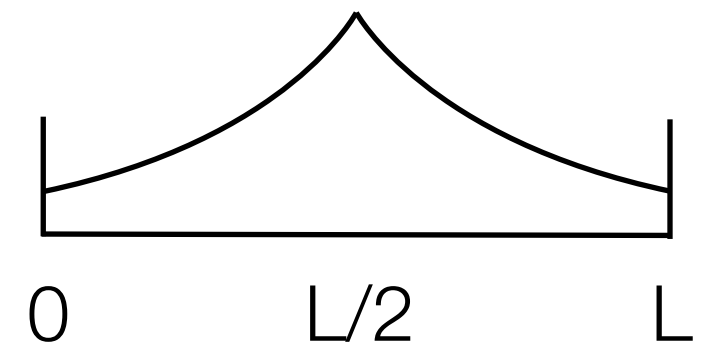
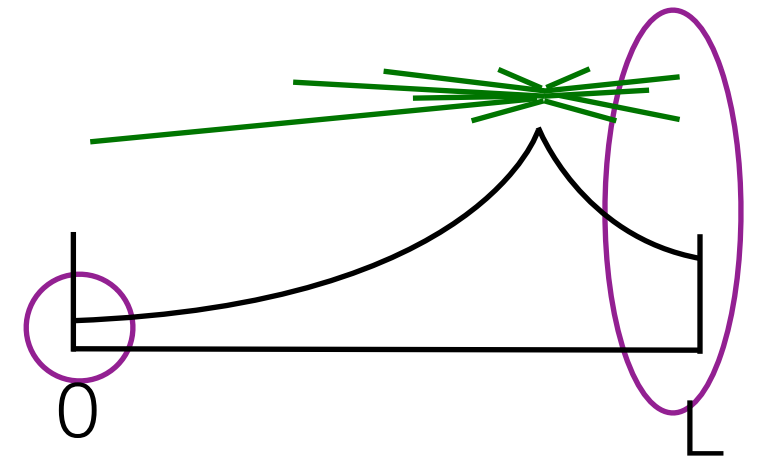
$$Tips(x) = Ae^{-\lambda|x-x_c|}$$

If motors pull only at the cell periphery,

$$F_{pull}(x_c) = B(-e^{-\lambda x_c} + e^{-\lambda(L-x_c)})$$

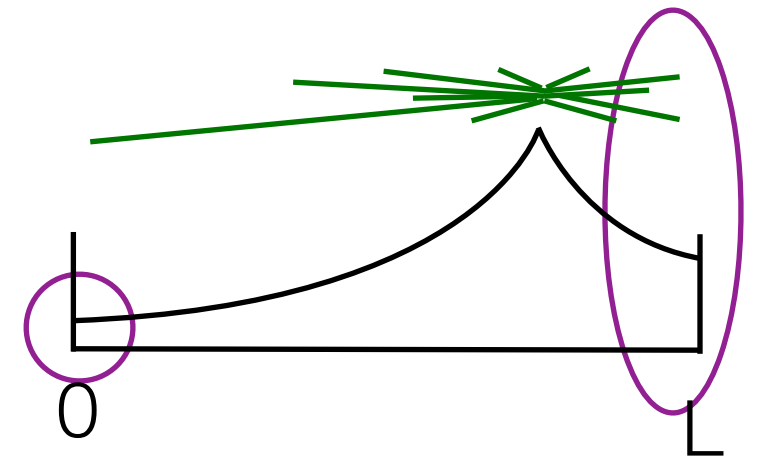
and the steady state position is at  $x_c = L/2$ .

For stability, check derivative of  $F_{pull}$ :



# One-D model - calculating forces from MT densities

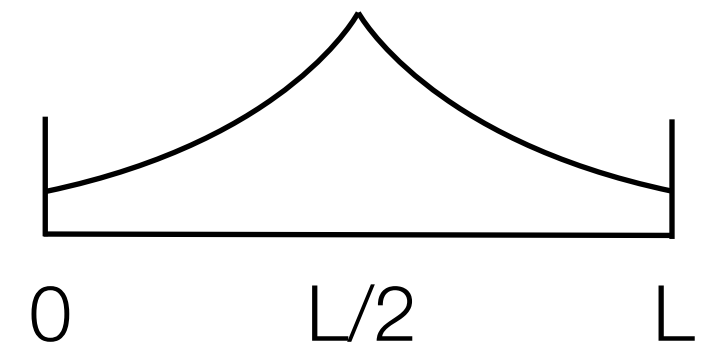
$$Tips(x) = Ae^{-\lambda|x-x_c|}$$



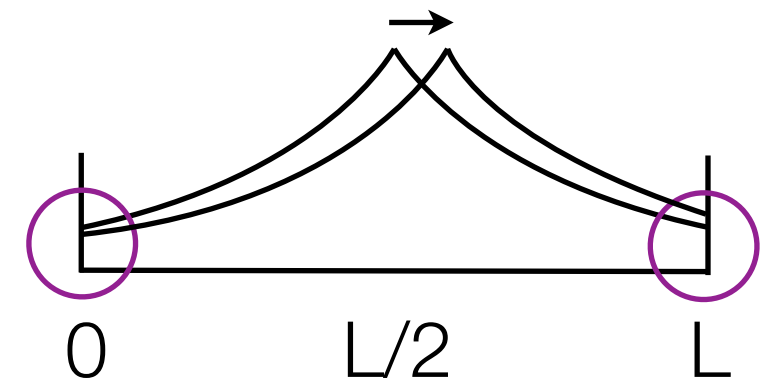
If motors pull only at the cell periphery,

$$F_{pull}(x_c) = B(-e^{-\lambda x_c} + e^{-\lambda(L-x_c)})$$

and the steady state position is at  $x_c = L/2$ .



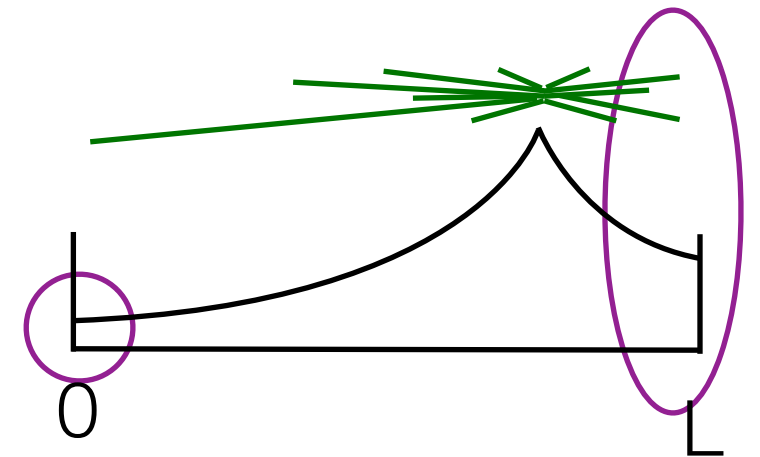
For stability, check derivative of  $F_{pull}$ :



$$F'_{pull}(L/2) = B\lambda(e^{-\lambda L/2} + e^{-\lambda L/2}) > 0$$

# One-D model - calculating forces from MT densities

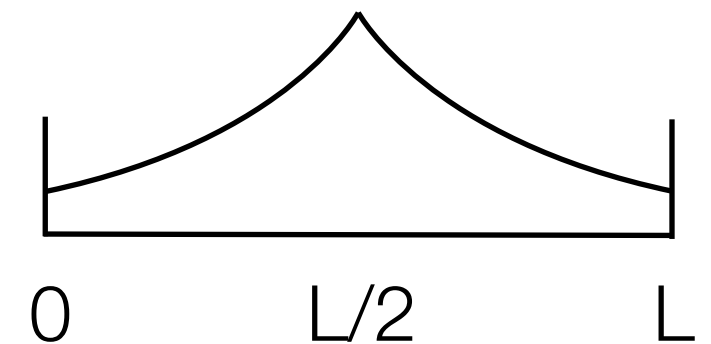
$$Tips(x) = Ae^{-\lambda|x-x_c|}$$



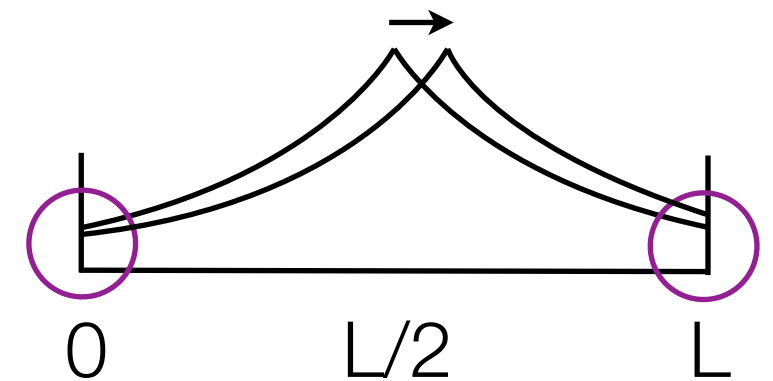
If motors pull only at the cell periphery,

$$F_{pull}(x_c) = B(-e^{-\lambda x_c} + e^{-\lambda(L-x_c)})$$

and the steady state position is at  $x_c = L/2$ .



For stability, check derivative of  $F_{pull}$ :



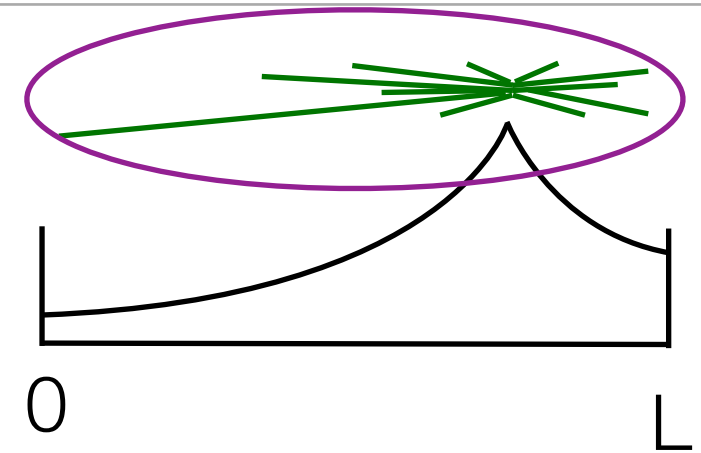
$$F'_{pull}(L/2) = B\lambda(e^{-\lambda L/2} + e^{-\lambda L/2}) > 0$$

Center is unstable!

# One-D model - calculating forces from MT densities

---

If motors pull all MT tips,



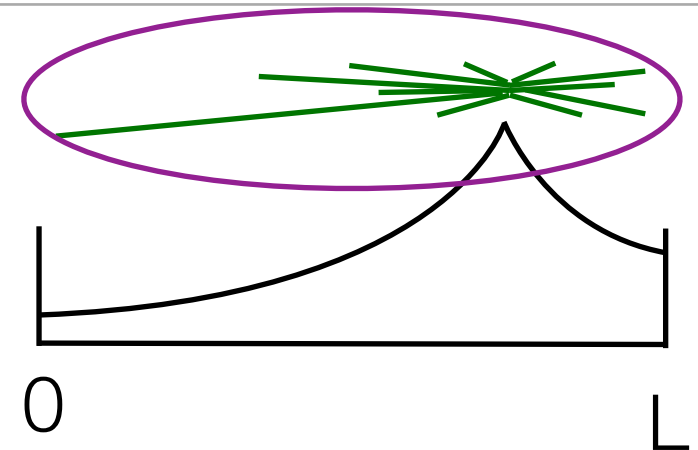
$$F_{pull}(x_c) = -C \int_0^{x_c} Tips(x) dx + C \int_{x_c}^L Tips(x) dx$$



# One-D model - calculating forces from MT densities

---

If motors pull all MT tips,

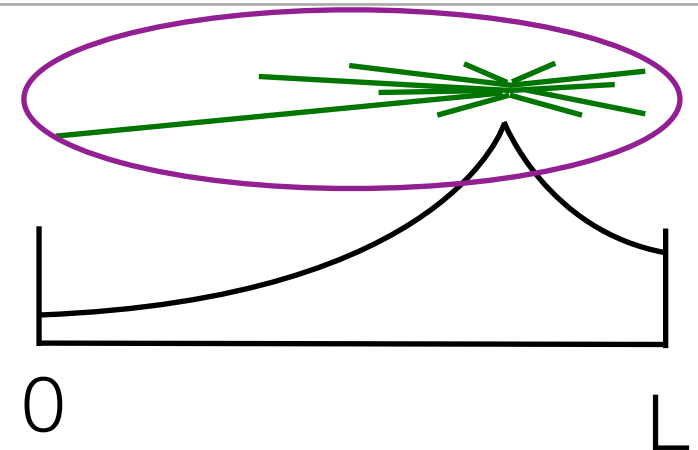


$$\begin{aligned} F_{pull}(x_c) &= -C \int_0^{x_c} Tips(x) \, dx + C \int_{x_c}^L Tips(x) \, dx \\ &= \frac{C}{\lambda} (e^{-\lambda x_c} - e^{-\lambda(L-x_c)}) \end{aligned}$$

# One-D model - calculating forces from MT densities

---

If motors pull all MT tips,



$$F_{pull}(x_c) = -C \int_0^{x_c} Tips(x) dx + C \int_{x_c}^L Tips(x) dx$$

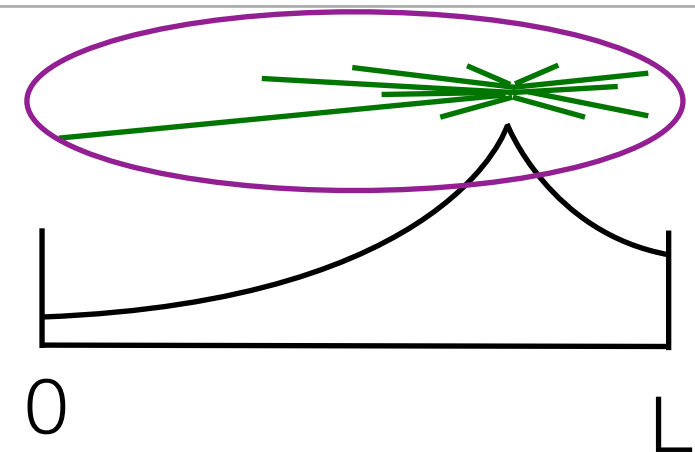
$$= \frac{C}{\lambda} (e^{-\lambda x_c} - e^{-\lambda(L-x_c)})$$

$$F'_{pull}(L/2) = C(-e^{-\lambda L/2} - e^{-\lambda L/2}) < 0$$

# One-D model - calculating forces from MT densities

---

If motors pull all MT tips,



$$F_{pull}(x_c) = -C \int_0^{x_c} Tips(x) dx + C \int_{x_c}^L Tips(x) dx$$

$$= \frac{C}{\lambda} (e^{-\lambda x_c} - e^{-\lambda(L-x_c)})$$

$$F'_{pull}(L/2) = C(-e^{-\lambda L/2} - e^{-\lambda L/2}) < 0$$

Center is stable!