

Proučavanje ravnoteže sila i momenata u diobenom vretenu u aproksimaciji srednjeg polja

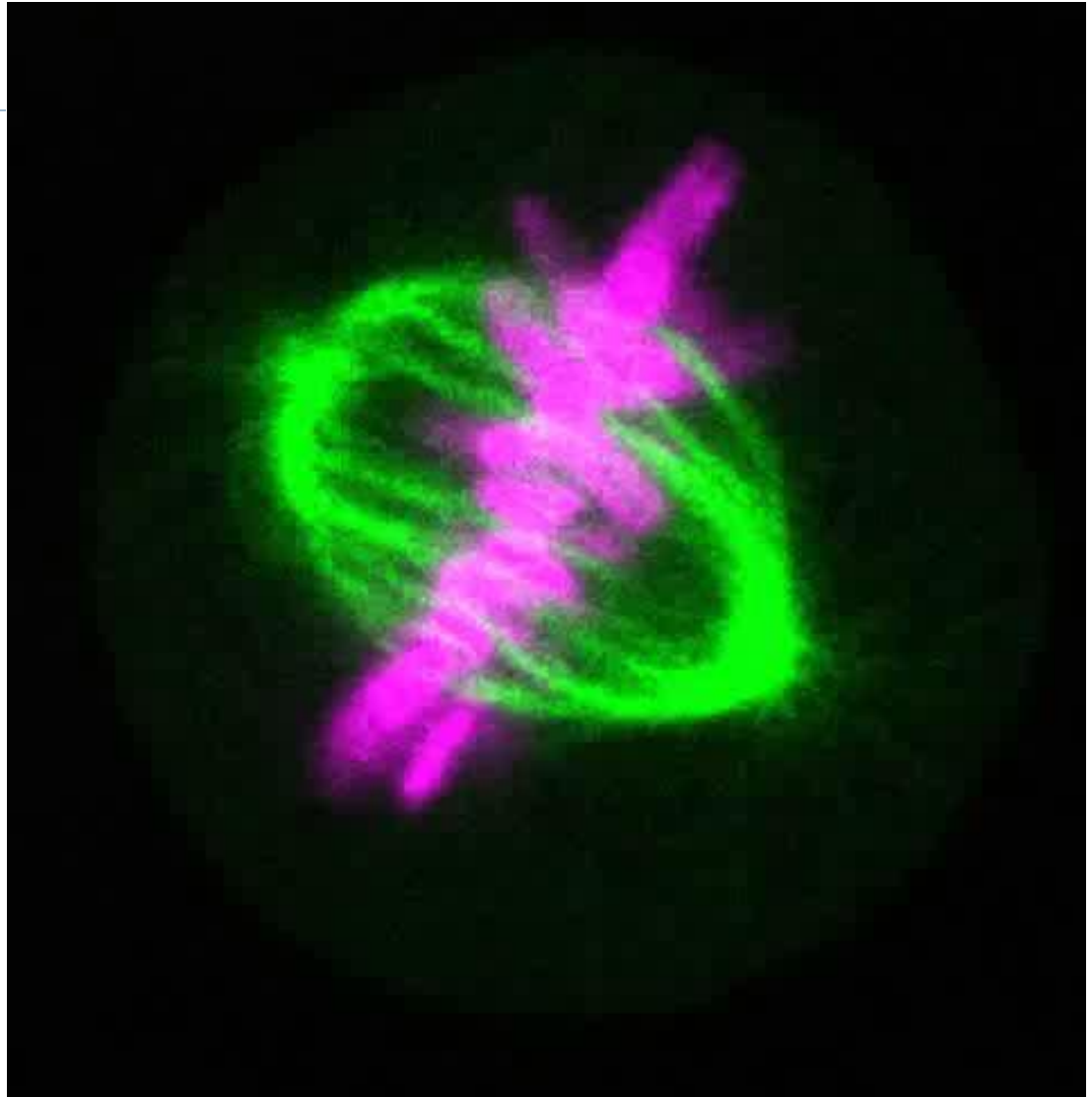
Arian Ivec

Mentor: izv. Prof. dr. sc. Nenad Pavin

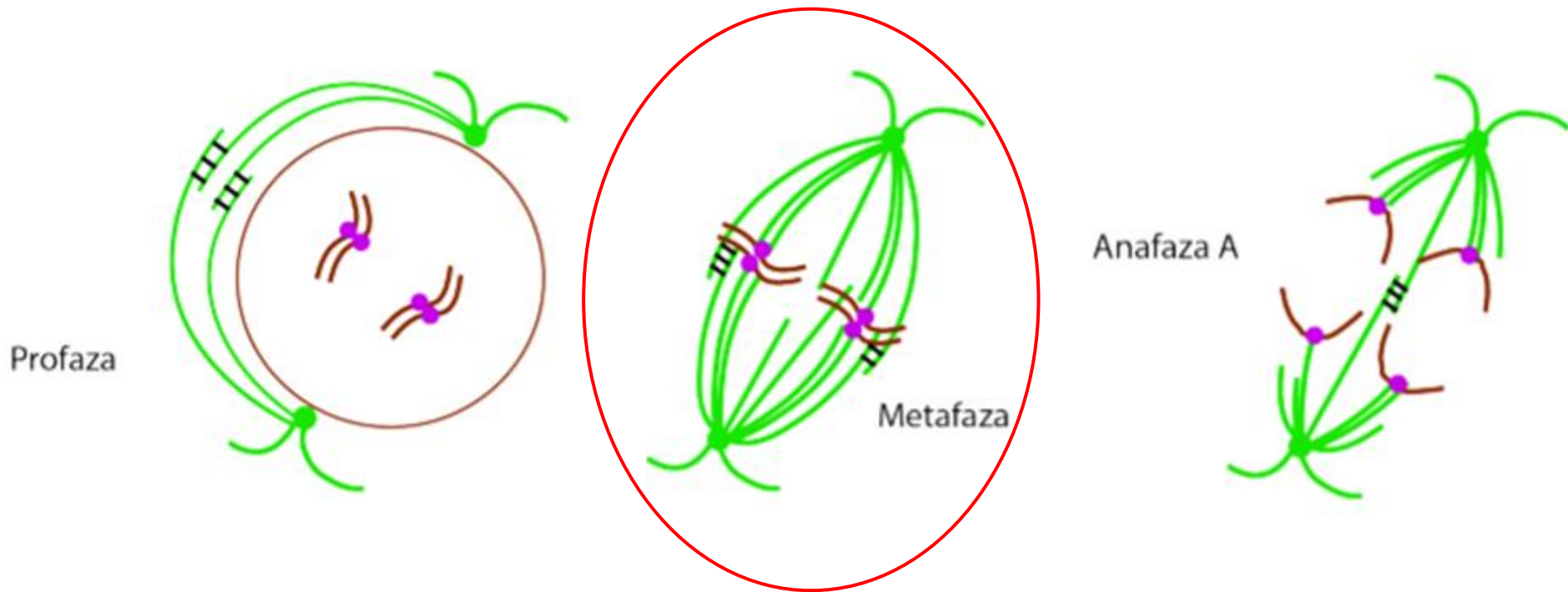
Fizički odsjek, PMF, Bijenička cesta

Stanična dioba

- ▶ Dioba tjelesnih stanica - mitozna
- ▶ Stanica majka se djeli na dvije stanice kćeri

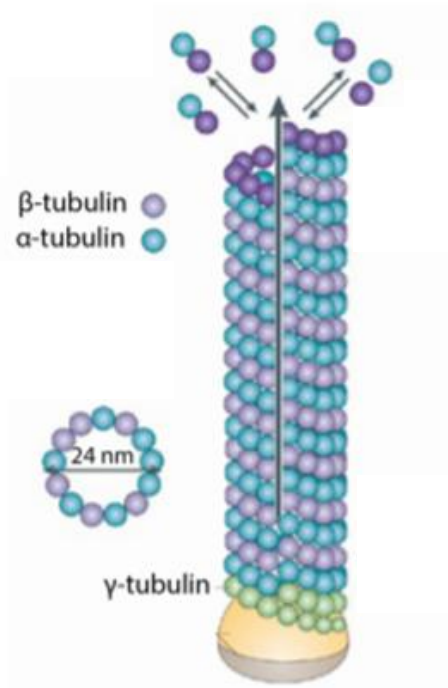


Faze diobe



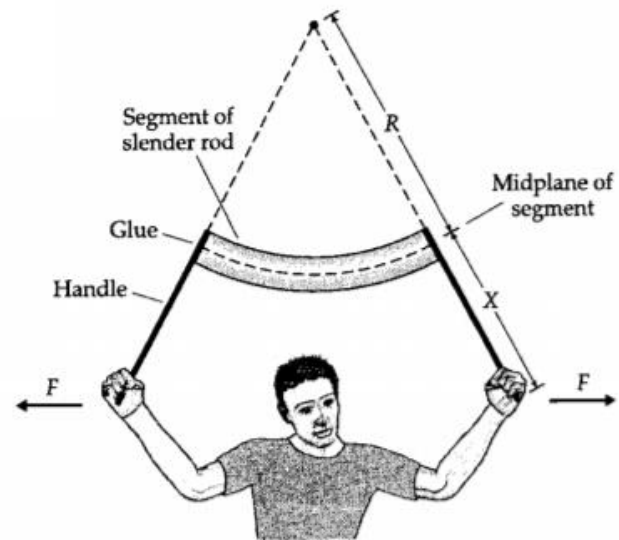
Alberts, B., Johnson, A.
Molecular Biology of the Cell

Svojstva mikrotubula



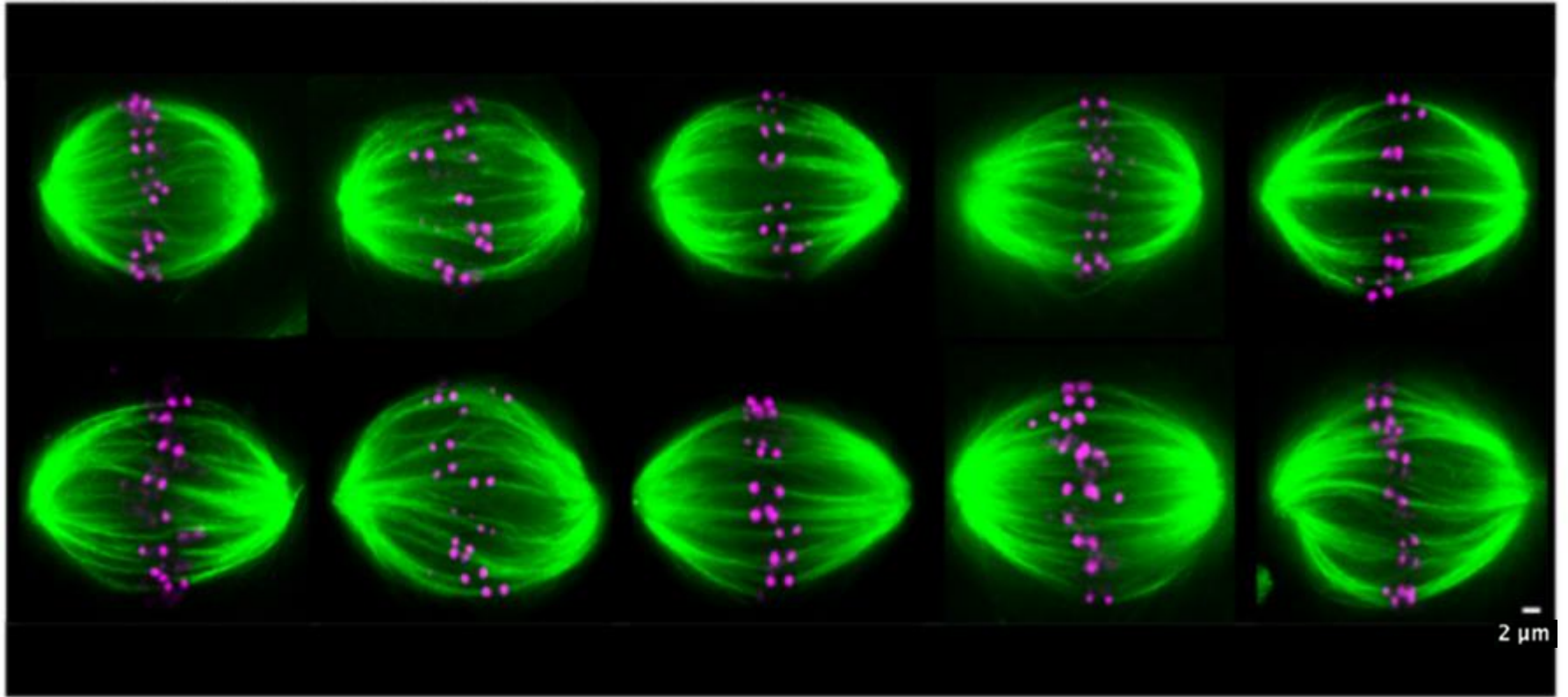
Rochlin, M. W., Dailey, M. E. & Bridgman, P. C. Polymerizing microtubules activate site-directed F-actin assembly in nerve growth cones. *Mol. Biol. Cell* **10**, 2309-2327 (1999)

- Elastičan štap



Howard J.,
Mechanics of Motor Proteins & the
Cytoskeleton

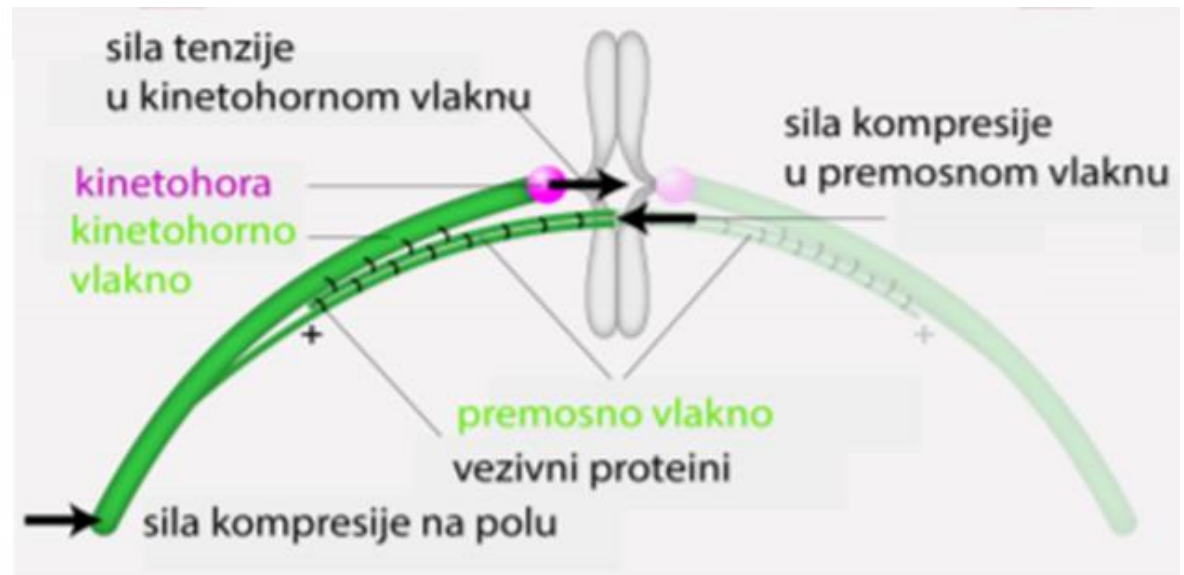
Izgled stanice u metafazi



Tolić Lab

Premosni mikrotubuli

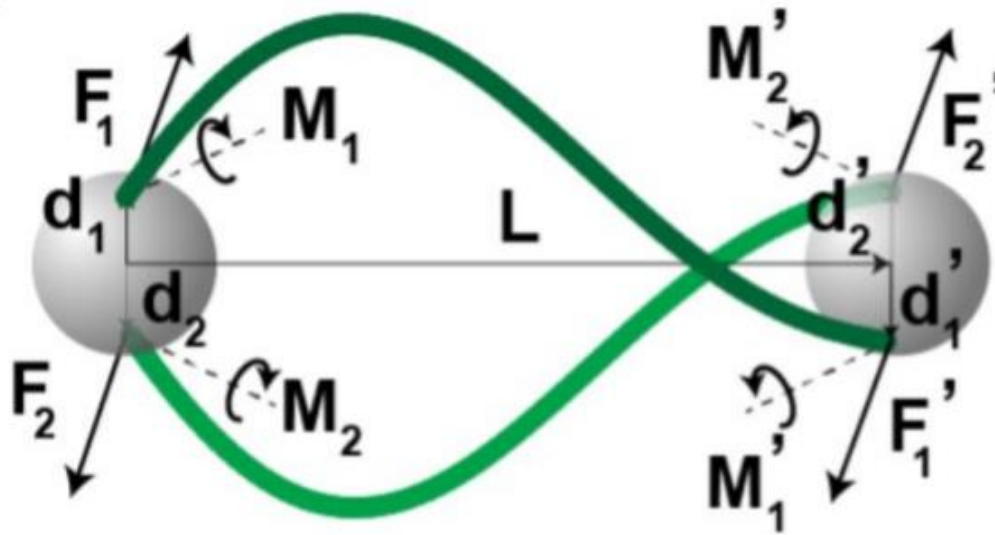
- ▶ Dio mikrotubula se spaja i formira premosno vlakno



- ▶ Tretiramo kao jedan jedinstveni objekt

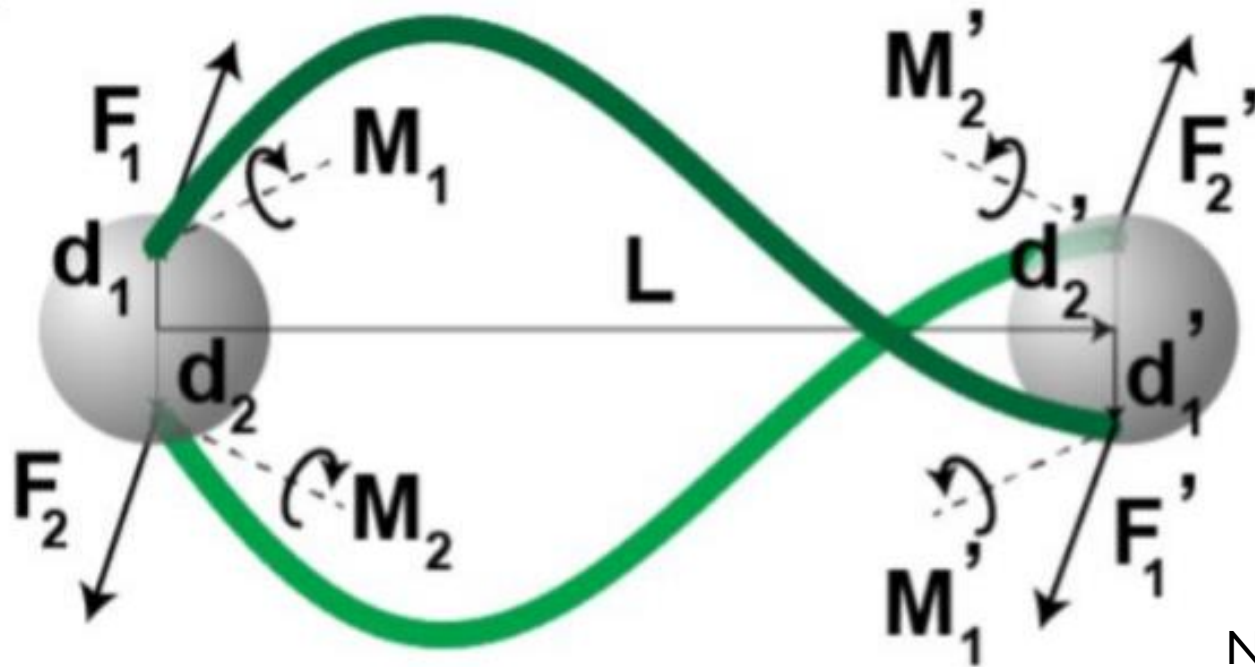
Tolić, I. M. and Pavin, N. Bridging the gap between sister kinetochores *Cell Cycle* 15 1169-1170 (2016)

Diskretni model ravnoteže



- ▶ Novak *et al.* : statička ravnoteža u metafazi
- ▶ Izjednačene sile i momenti na centrosomu i na mikrotubulu

Jednadžbe ravnoteže



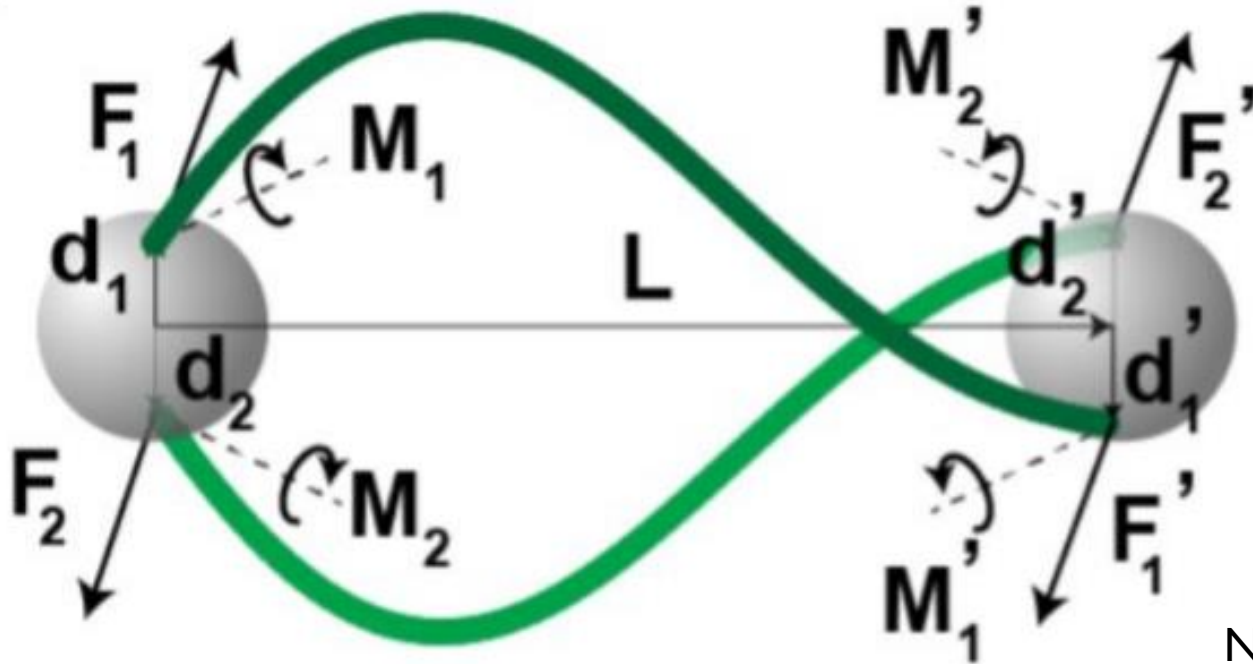
Novak et al.

Lijevi centrosom

$$\sum_i^n \vec{F}_i = 0,$$

$$\sum_i^n (\vec{M}_i + \vec{d}_i \times \vec{F}_i) = 0.$$

Jednadžbe ravnoteže



Novak et al.

Lijevo centrosom

$$\sum_i^n \vec{F}_i = 0,$$

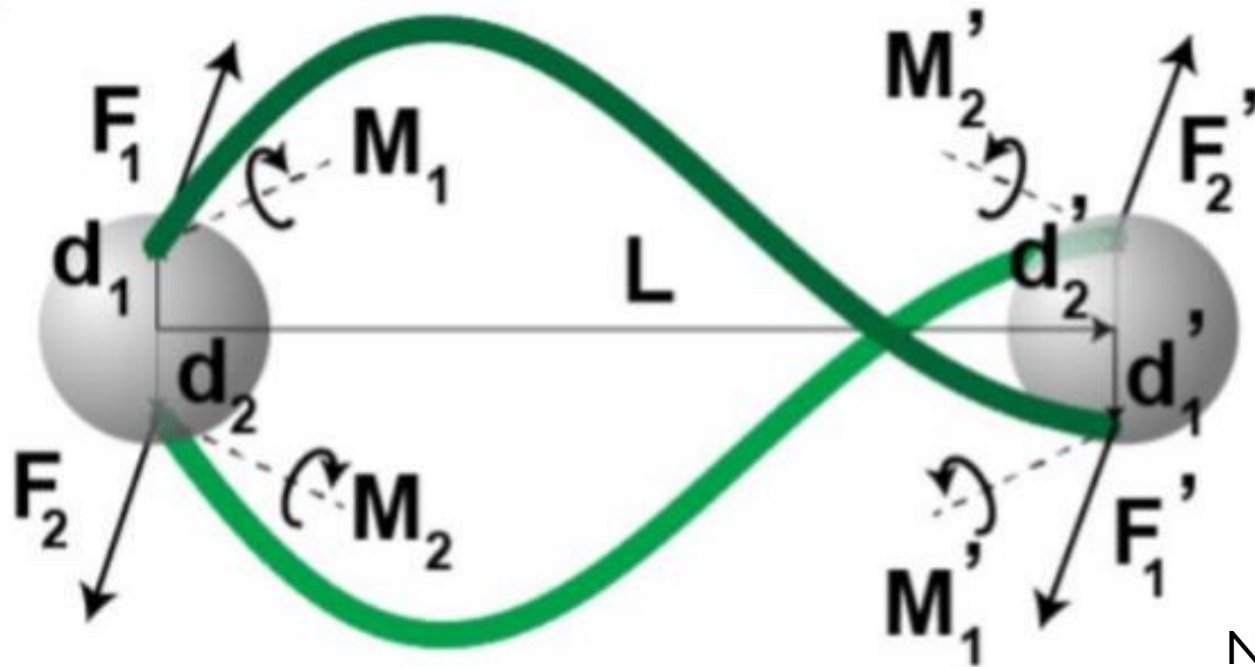
$$\sum_i^n (\vec{M}_i + \vec{d}_i \times \vec{F}_i) = 0.$$

Desno centrosom

$$\sum_i^n \vec{F}_i' = 0,$$

$$\sum_i^n (\vec{M}_i' + \vec{d}_i' \times \vec{F}_i') = 0.$$

Jednadžbe ravnoteže



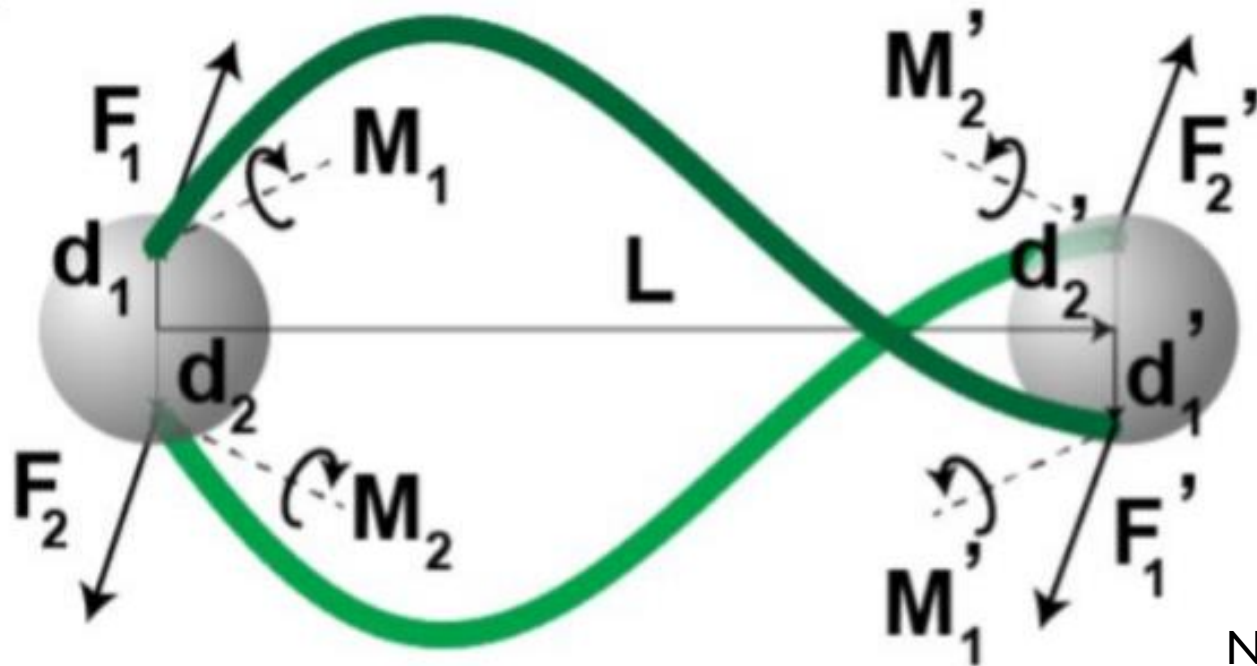
Novak et al.

Ravnoteža jednog mikrotubula

$$\vec{F}_i + \vec{F}'_i = 0,$$

$$\vec{M}_i + \vec{M}'_i + \vec{d}_i \times \vec{F}_i + (\vec{d}'_i + \vec{L}) \times \vec{F}'_i = 0.$$

Jednadžbe ravnoteže



Novak et al.

Nije pogodno za opis čitavog vretena \rightarrow kontinuirani model

Kontinuirani model

- Prelazak na kontinuirane varijable

$$i = 1, \dots, n, \quad \longrightarrow \quad \rho(\theta, \phi)$$

$$\vec{F}_i \quad \longrightarrow \quad \vec{F}(\rho(\theta, \phi))$$

$$\vec{M}_i \quad \longrightarrow \quad \vec{M}(\rho(\theta, \phi))$$

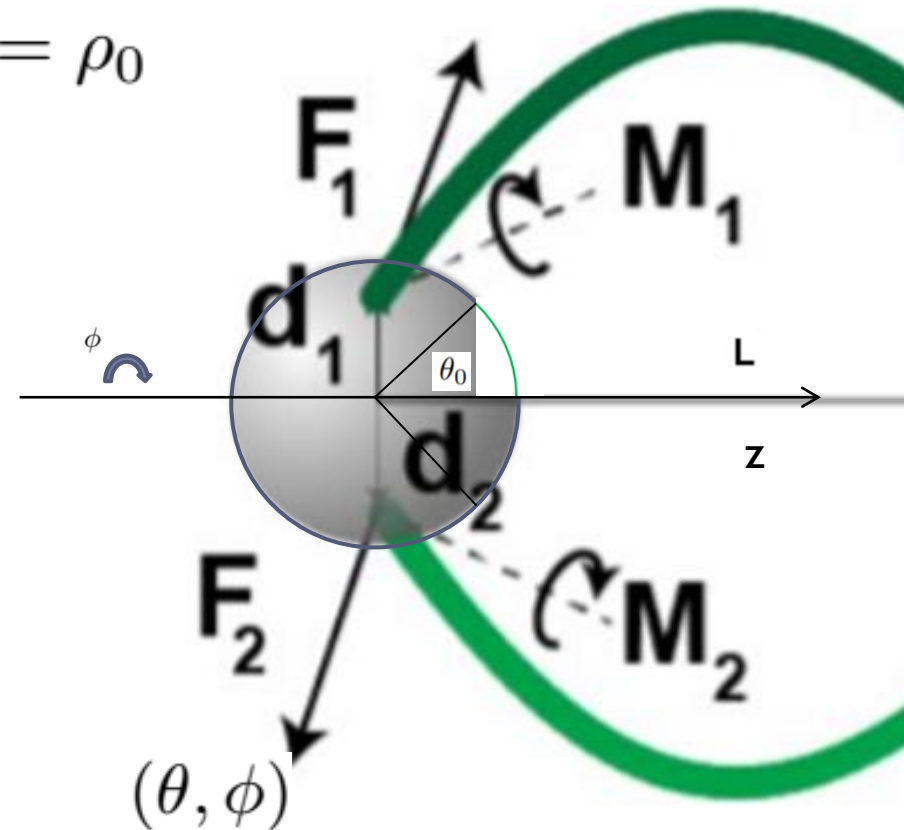
Gustoća mikrotubula

- ▶ Homogeno raspodjeljeni po površeni sfere
- ▶ Granični kut populiranosti

$$\rho(\theta, \phi) = \rho_0$$

θ_0

$\rho(\theta, \phi)$



Simetrije

- Rotaciona simetrija oko osi spojnice

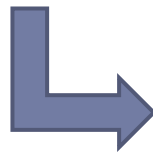


$$\vec{F}^{\perp}(\theta, \phi) = -\vec{F}^{\perp}(\theta, \phi + \pi),$$

$$\vec{F}^{\parallel}(\theta, \phi) = \vec{F}^{\parallel}(\theta, \phi + \pi),$$

$$|\vec{F}(\theta, \phi)| = |\vec{F}(\theta)|,$$

- Invarijantnost na zamjenu lijevog i desnog centrosoma

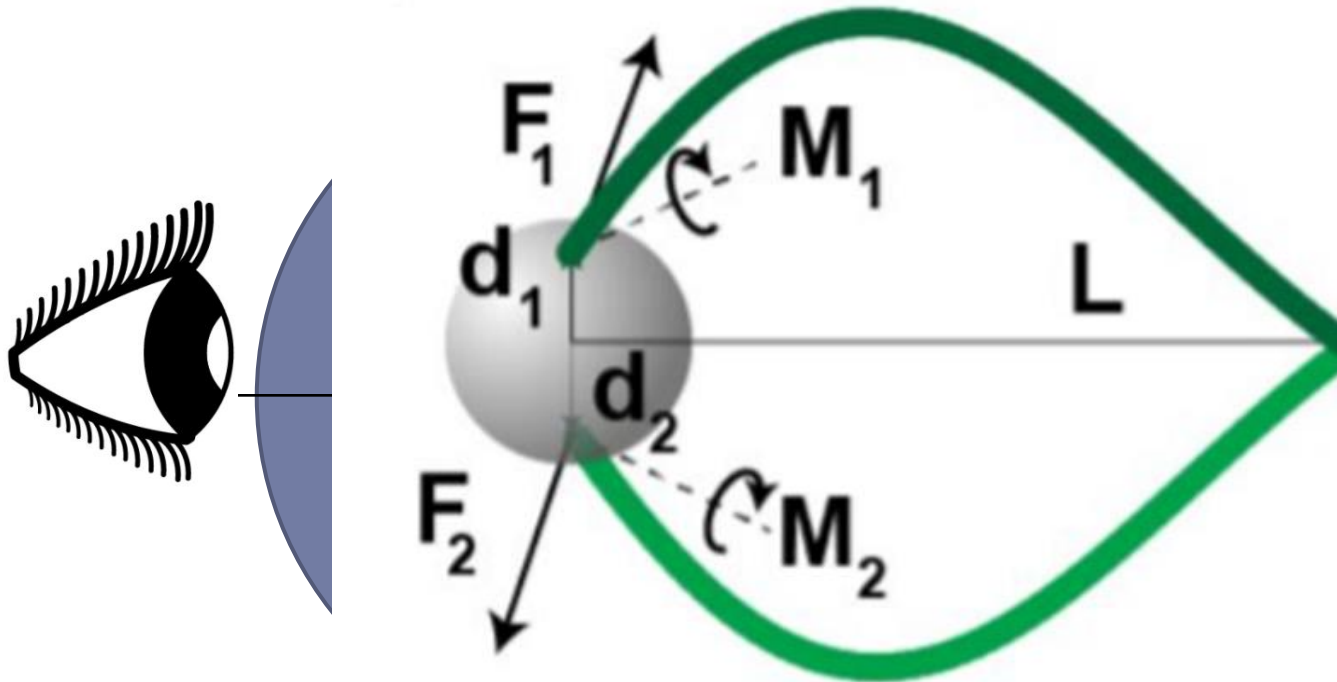


$$\theta' = +\theta$$

Implementiranje simetrija

$$\vec{F}(\theta, \phi) = f^{\parallel}(\theta)\hat{z} + f^{\perp}(\theta)(\cos(\phi + \delta)\hat{x} + \sin(\phi + \delta)\hat{y}),$$

$$\vec{M}(\theta, \phi) = m^{\parallel}(\theta)\hat{z} + m^{\perp}(\theta)(\cos(\phi + \delta)\hat{x} + \sin(\phi + \delta)\hat{y}),$$



Jednadvžbe ravnoteže - centrosom

$$\sum_i^n \vec{F}_i = 0,$$



$$\int_0^{2\pi} \int_0^{\theta_0} \vec{F}(\theta, \phi) d\theta d\phi,$$



$$\hat{z} : \int_0^{2\pi} \int_0^{\theta_0} f^{\parallel}(\theta) d\theta d\phi = \int_0^{\theta_0} f^{\parallel}(\theta) d\theta = 0.$$

$$\sum_i^n (\vec{M}_i + \vec{d}_i \times \vec{F}_i) = 0.$$



$$\int_0^{2\pi} \int_0^{\theta_0} (\vec{M}(\theta, \phi) + \vec{d}(\theta, \phi) \times \vec{F}(\theta, \phi)) d\theta d\phi = 0.$$



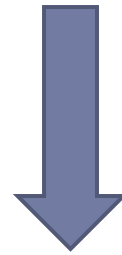
$$\int_0^{\theta_0} \vec{m}^{\parallel}(\theta) d\theta = 0$$

Jednadžbe ravnoteže – sile na mikrotubul

$$\vec{F}_i + \vec{F}'_i = 0,$$



$$\vec{F}(\theta, \phi) = -\vec{F}'(\theta', \phi'),$$



$$f^\perp(\theta) = \pm f^{\perp'}(\theta'),$$

$$\phi'_1 = \phi + (\delta - \delta'),$$

$$\phi'_2 = \phi + \pi + (\delta - \delta'),$$

- Veza položaja
lijevog i desnog
kraja MT-a

Jednadžbe ravnoteže – momenti na mikrotubul

$$\vec{M}_i + \vec{M}'_i + \vec{d}_i \times \vec{F}_i + (\vec{d}'_i + \vec{L}) \times \vec{F}'_i = 0.$$



$$\vec{M}(\theta, \phi) + \vec{M}'(\theta', \phi') + \vec{d} \times \vec{F}(\theta, \phi) + (\vec{d}' + \vec{L}) \times \vec{F}'(\theta', \phi') = 0.$$

Jednadžbe ravnoteže – momenti na mikrotubul

- Raspis po komponentama:

$$\begin{aligned}\hat{x} : m^\perp(\theta)\cos(\phi + \delta) + m^{\perp'}(\theta')\cos(\phi' + \delta') + d(\sin(\theta)\sin(\phi + \delta)f^\parallel \\ - \cos(\theta)\sin(\phi + \delta)f^\perp) + d'(\sin(\theta')\sin(\phi' + \delta')f^\parallel \\ - \cos(\theta')\sin(\phi' + \delta')f^\perp) - f^\perp L\sin(\phi' + \delta') = 0,\end{aligned}$$

$$\begin{aligned}\hat{y} : m^\perp(\theta)\sin(\phi + \delta) + m^{\perp'}(\theta')\sin(\phi' + \delta') + d(\cos(\theta)\sin(\phi + \delta)f^\perp \\ - \sin(\theta)\sin(\phi + \delta)f^\parallel) + d'(\cos(\theta')\sin(\phi' + \delta')f^\perp \\ - \sin(\theta')\sin(\phi' + \delta')f^\parallel) + f^\perp L\cos(\phi' + \delta') = 0,\end{aligned}$$

$$\hat{z} : m^\parallel(\theta) + m^{\parallel'}(\theta') = 0.$$

A
G



Taylorov razvoj

- ▶ Rješavamo razvojem u Taylorov red

$$f^{\perp/\parallel} = f_0^{\perp} + f_1^{\perp}\theta + o(\theta^2)$$

- ▶ Zadržavamo se na prvom redu

$$m^{\perp/\parallel} = m_0^{\perp} + m_1^{\perp}\theta + o(\theta^2)$$

Rješavanje modela - sile

$$\int_0^{\theta_0} \vec{f}^{\parallel}(\theta) d\theta = 0.$$



$$\frac{f_0^{\parallel}}{f_1^{\parallel}} = \frac{1 - \cos(\theta_0)}{\sin(\theta_0) - \cos(\theta_0)\theta_0}$$

$$\vec{F}(\theta, \phi) = -\vec{F}'(\theta', \phi'),$$



$$f^{\parallel} = f_0^{\parallel} + f_1^{\parallel} = -f^{\parallel'} = -(f_0^{\parallel'} + f_1^{\parallel'}),$$

Rješavanje modela - momenti

$$\int_0^{\theta_0} m^{\parallel}(\theta) d\theta = 0$$



$$\frac{m_0^{\parallel}}{m_1^{\parallel}} = \frac{1 - \cos(\theta_0)}{\sin(\theta_0) - \cos(\theta_0)\theta_0}$$

$$\vec{M}(\theta, \phi) + \vec{M}'(\theta', \phi') + \vec{d} \times \vec{F}(\theta, \phi) + (\vec{d}' + \vec{L}) \times \vec{F}'(\theta', \phi') = 0.$$



$$m^{\perp}(\theta) - m^{\perp}'(\theta) = -\frac{f^{\perp}L}{2} + \frac{1}{2}\sqrt{f^{\perp 2}L^2 - 4a^2 - 4af^{\perp}L},$$

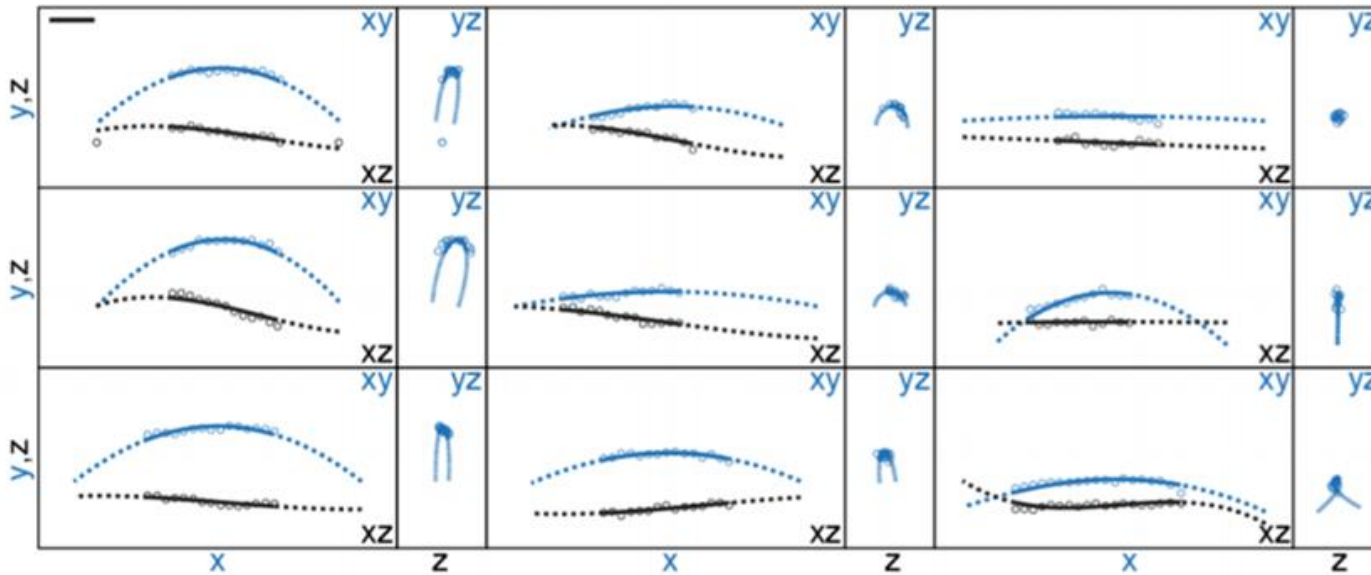
$$a = a(\theta) = 2\left| \vec{d}' \right| f^{\parallel}(\theta)\theta.$$

Oblik mikrotubula

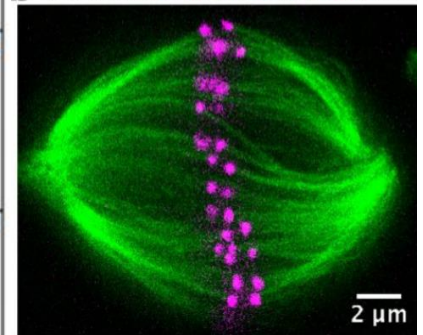
- ▶ Statička Kirchofova jednađba (SKJ)



$$\kappa \vec{t} \times \frac{d\vec{t}}{ds} + \tau \frac{d\phi}{ds} \vec{t} = \vec{r} \times \vec{F}_i - \vec{M}_i.$$



b U2OS cell, SiR-Tubulin, CENPA-GFP, horizontal



Tolić Lab

Novak et al.

Zaključak

- ▶ Od 16 koeficijenata ostaje njih 6 neovisno
- ▶ U nastavku rada treba dobiti oblike preko SKJ u usporedbi s mikroskopijom
- ▶ Steći uvid u statiku cjelokupnog vretena i kako do nje uopće dolazi

Zahvale

Zahvaljujem se Pavin-Tolić grupi na mogućnosti izradi seminara kao i svojim mentorima :



izv. prof. dr. sc.
Nenad Pavin



mag. phy. Ivana
Ban