

Elucidating the Mechanism Governing the Cell Rotation Behavior under DEP

¹Yu Zhao, ¹Johnie Hodge, ⁴Jozef Brcka, ⁴Jacques Faguet, ⁴Eric Lee, ^{1,2,3}Guigen Zhang

¹Dept. of Bioengineering, ²Dept. of Electrical and Computer Engineering

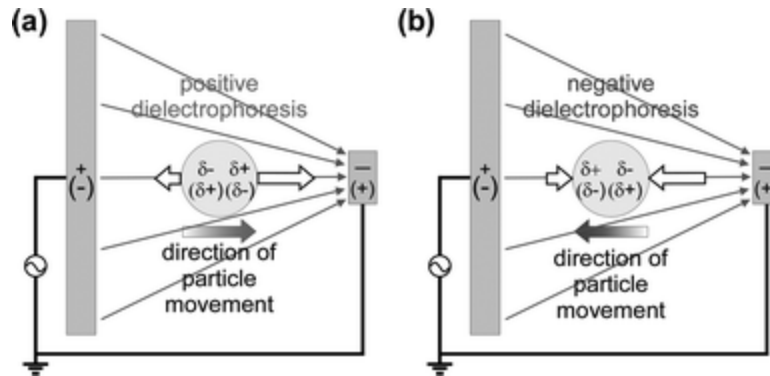
³Institute for Biological Interfaces of Engineering, Clemson, SC 29634

⁴Tokyo Electron U.S. Holdings, Inc., U.S. Technology Development Center, Austin,
TX 78741

guigen@clemson.edu



- What is dielectrophoresis (DEP)



<http://pubs.rsc.org/en/content/articlelanding/2007/lc/b712784g#!divAbstract>

- Application of DEP

- Particle Separation

- Cell sorting



Well explained by current theory

- Particle alignment

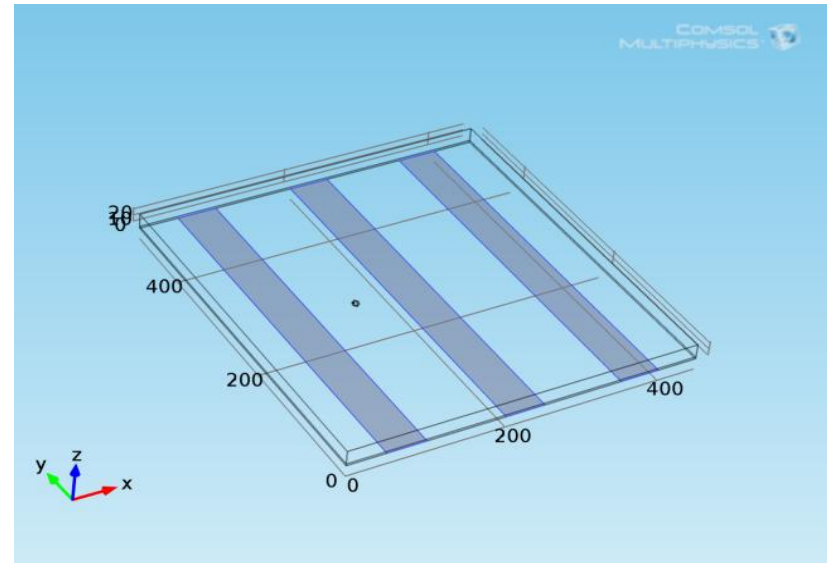
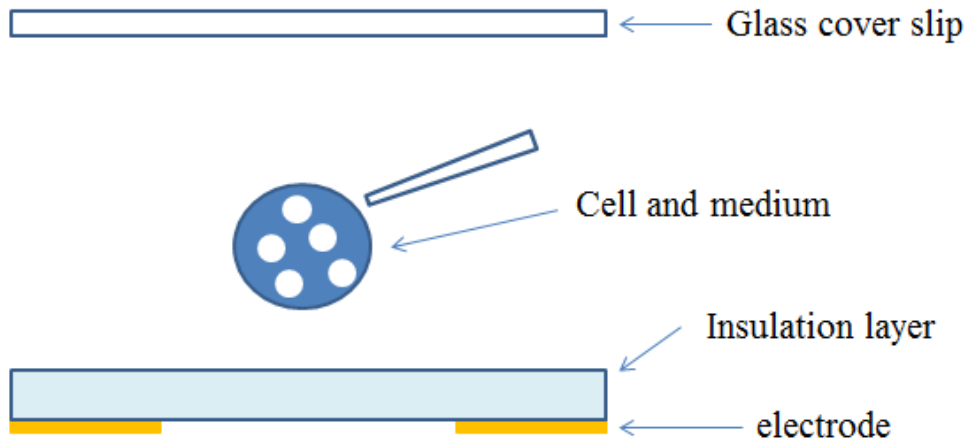
- Cell patterning

- Unique single cell behavior

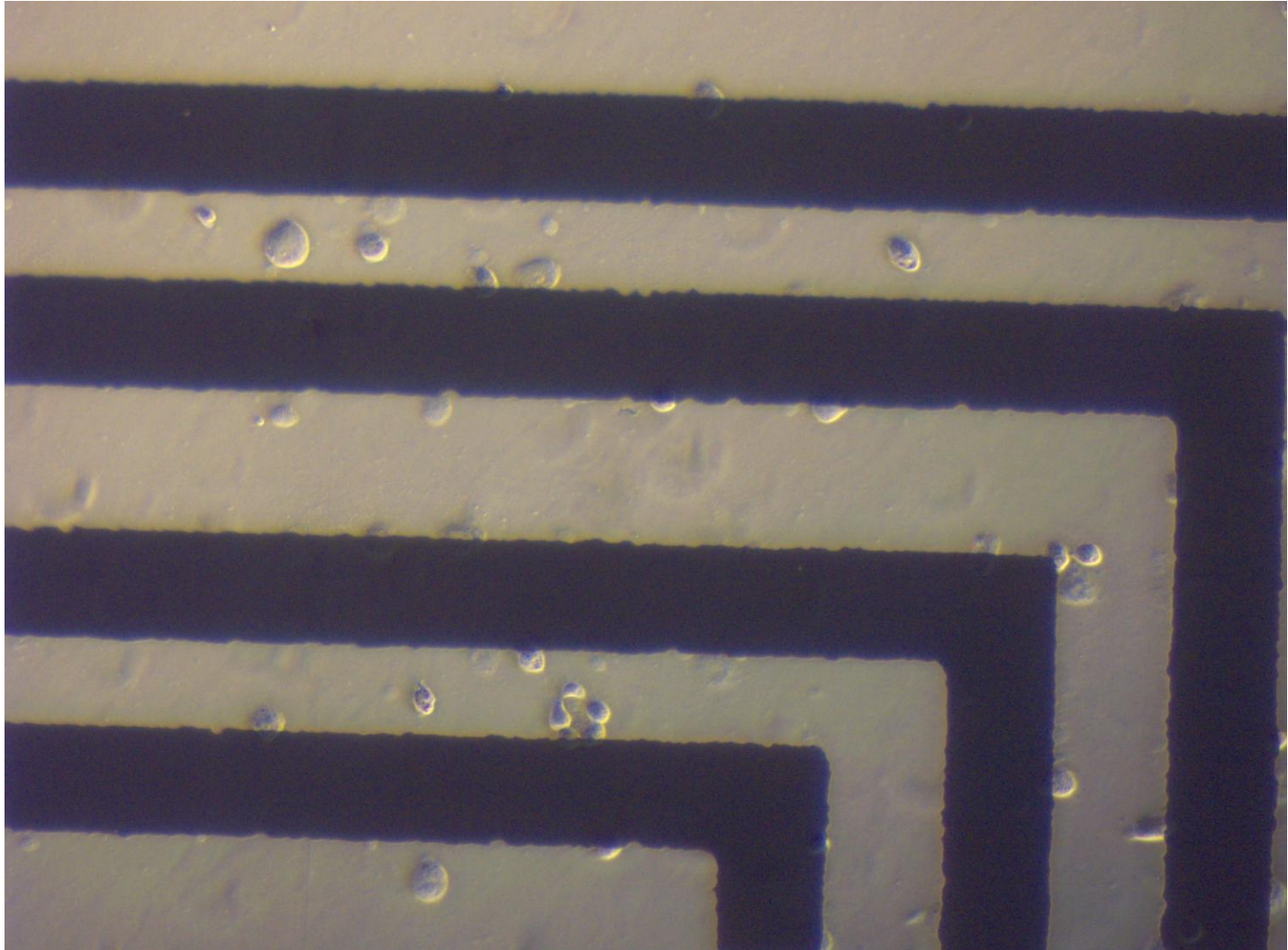


No complete theory

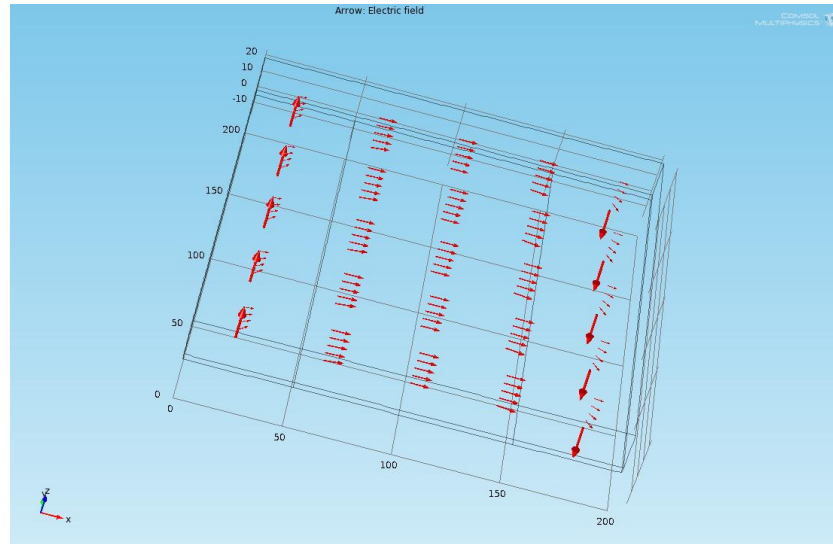
Experimental setup and modeling geometry



Experimental observation (rat adipose stem cell)



Conventional method



Particle is treated as a point dipole:

$$F = 2\pi a^3 \epsilon_0 \epsilon_m \text{Re}(f_{cm}) \nabla E_{rms}^2$$
$$\text{Re}(f_{cm}) = \frac{\omega^2 (\epsilon_p - \epsilon_m) (\epsilon_p + 2\epsilon_m) + (\sigma_p - \sigma_m) (\sigma_p + 2\sigma_m)}{\omega^2 (\epsilon_p + 2\epsilon_m)^2 + (\sigma_p + 2\sigma_m)^2}$$

Cannot explain why cell rotates (No phase difference under experimental condition)

New method to overcome limitations

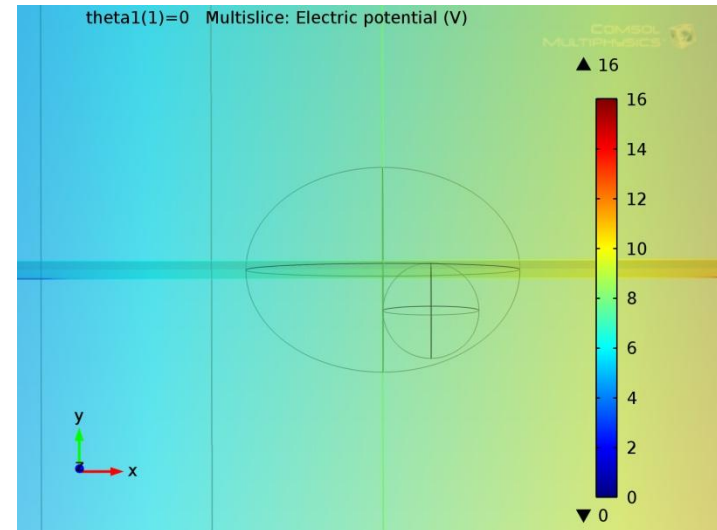
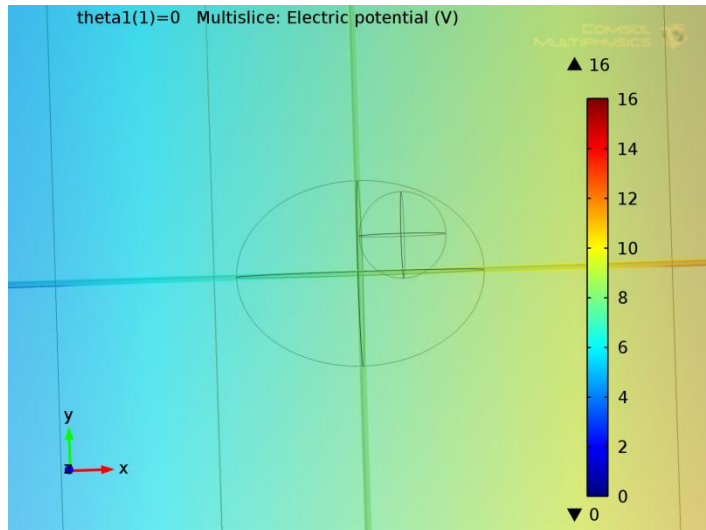
$$F = \frac{\epsilon_p - \epsilon_m}{\epsilon_p + 2\epsilon_m} \epsilon_0 \int (P \cdot \nabla) E dV$$
$$T = \frac{\epsilon_p - \epsilon_m}{\epsilon_p + 2\epsilon_m} \epsilon_0 \int r \times [(P \cdot \nabla) E] dV$$

- Size effect is correctly reflected
- The dipole-dipole interaction is included
- Works for particles with non-uniform electric property (e.g. cell)

Cell property

- Non-sphere shape
- Nucleus with high permittivity due to greater hydrated free ion content
- Nucleus usually at fixed position

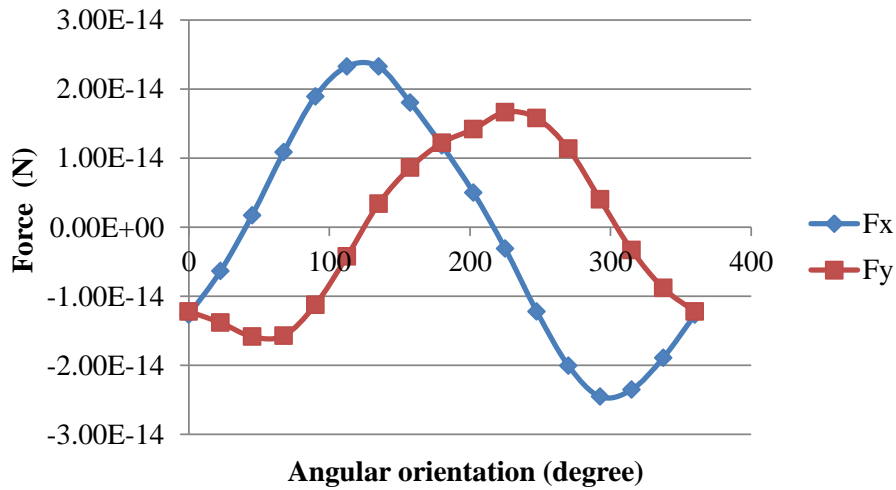
Modeling of cell structure



COMSOL model of an ellipsoid shaped cell with an off-centered nucleus; upper(left), lower(right)

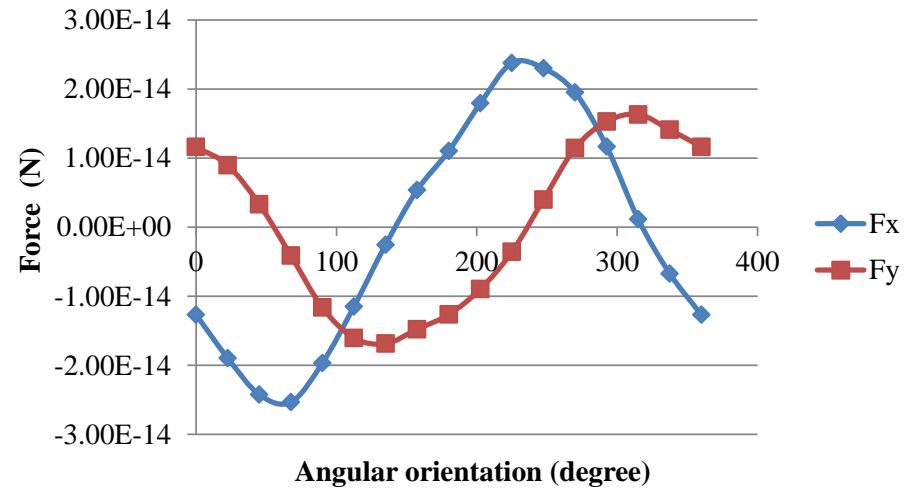
Net DEP force on cell at different orientations

x and y component of DEP force on cell with nucleus on upperside



an ellipsoid-shaped cell with upper side off-centered nucleus

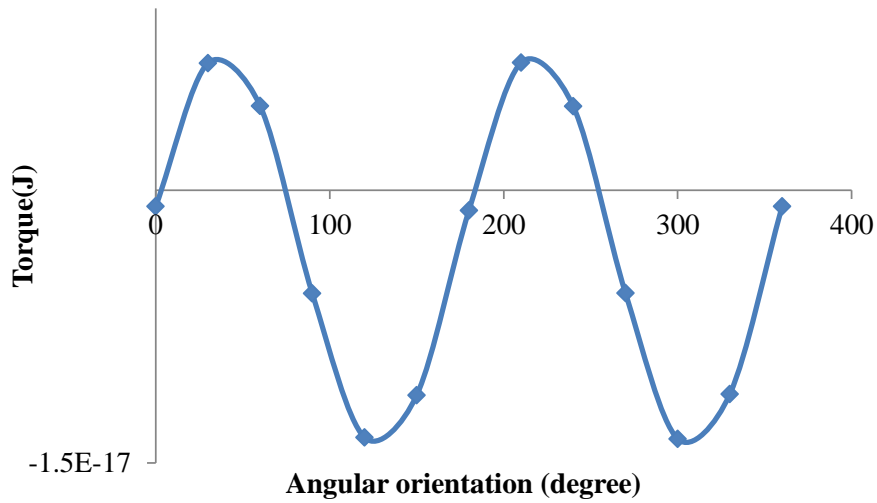
x and y component of DEP force on cell with nucleus on lowerside



an ellipsoid-shaped cell with lower side off-centered nucleus

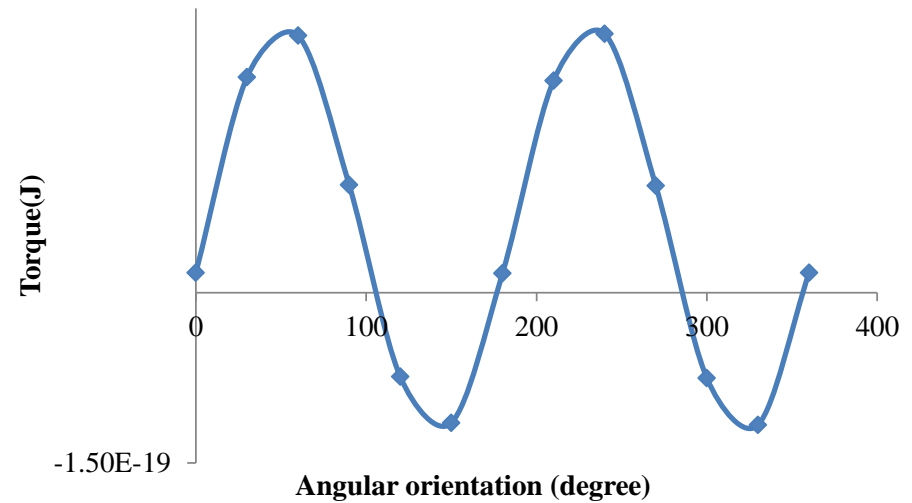
Net torque on cell at different orientation

Torque on cell with nucleus on upside



**an ellipsoid-shaped cell with upper side off-centered nucleus
(Average negative torque suggests clockwise rotation)**

Torque on cell with nucleus on downside



**an ellipsoid-shaped cell with lower side off-centered nucleus
(Average positive torque suggests counterclockwise rotation)**

Conclusion

- A new numerical method is developed which will overcome the limitations in the calculation of DEP force and torque.
- Our hypotheses that the cell rotation is caused by the non-circular shape of the cell body and the off-centered location of its nucleus and that the rotation direction depends on the relative location of nucleus with respect to the electrical field are confirmed by the simulation results.

Acknowledgement:

- Tokyo Electron U.S. Holdings, Inc., U.S. Technology Development Center
- Dr. guigen Zhang
- Johnie Hodge
- Institute for biological interfaces of engineering
- Clemson computing and information technology