



Parallel tutorial session # 1: BASIC MECHANICS

08 / 07 / 2006

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Welcome and introduction to GEM⁴

- international, interdisciplinary, inter-institutional initiative started early 2006
nucleus: MIT, NSU, Harvard, Institut Pasteur
engineering, biology, medicine & public health intersected
paradigm for global cooperation through research, training & translation
networking opportunities
infrastructure capabilities
- training program: summer school (2007 in Singapore, focus on cancer)*
also offered by GEM⁴: distinguished lecture series
junior scientist program
- * with the newly announced SMART: Singapore-MIT Alliance for Research and Technology Center in Singapore
- GEM⁴ co-sponsors the next world-wide meeting on biomechanics in 2010.
- Cell and molecular mechanics in biomedicine, with a focus on infectious diseases.
from a course developed at MIT by Profs. Patrick Doyle, Alan Grodzinsky & R. Kamm.
lab info: http://www.openwetware.org/wiki/GEM4_labs
breakdown of participants: 30% life scientists, 70% engineers
poster sessions to share research knowledge.
research proposals to be developed during the summer schools, presented in 3-4 pages
direct questions to Maggie Sullivan or Roger Kamm: sullmag@mit.edu rdkamm@mit.edu

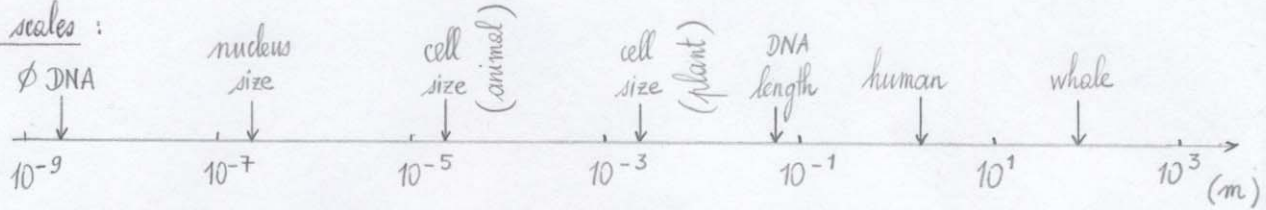
Simple statistical mechanics for biological systems

L. Mahadevan

- Questions: What is the goal?
starting with the central dogma in biology: DNA → RNA → proteins → organelles
(Crick) ecosystem ← organism ← tissue ← cells ←
- (and feedback!), essentially increasing length scale

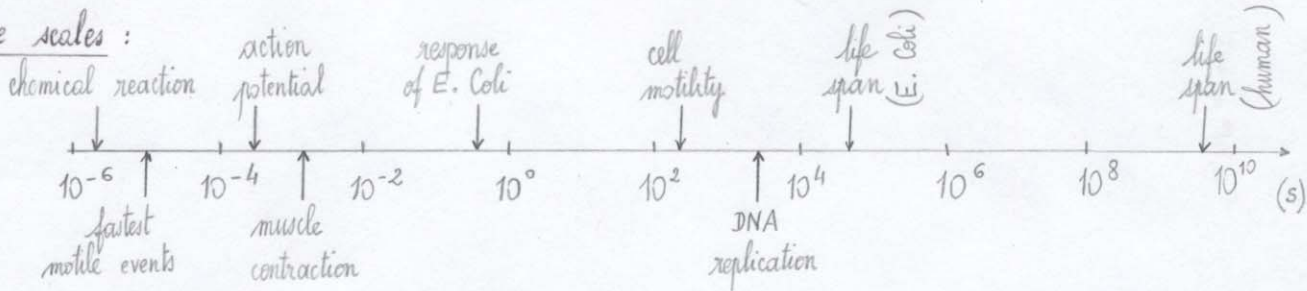
Hence, length and time scales and "out of equilibrium" principles matter.

Length scales:



cells are the fundamental units of life: smallest to function independently
 cell size \propto amount of DNA enclosed.

Time scales:



Out of equilibrium biology:

soft, wet, dynamic, warm
 information (encoded in DNA, ...), energy, matter
 we wish to couple energy and matter: biology is warm, hence in motion, hence energy.
 (dynamic & directed)

Energy scales:

thermal energy used as a ruler
 energy / mole / $^{\circ}\text{K}$ \sim RT
 (of ideal gas) $= \frac{1}{2} RT$ per degree of freedom
 (3 d.o.f. for ideal gas)
 $= 8.3 \text{ J / mole / } ^{\circ}\text{K}$ for ideal gas
 energy / molecule $\sim RT / c_{pA} = k_B T$
 (and per degree of freedom)

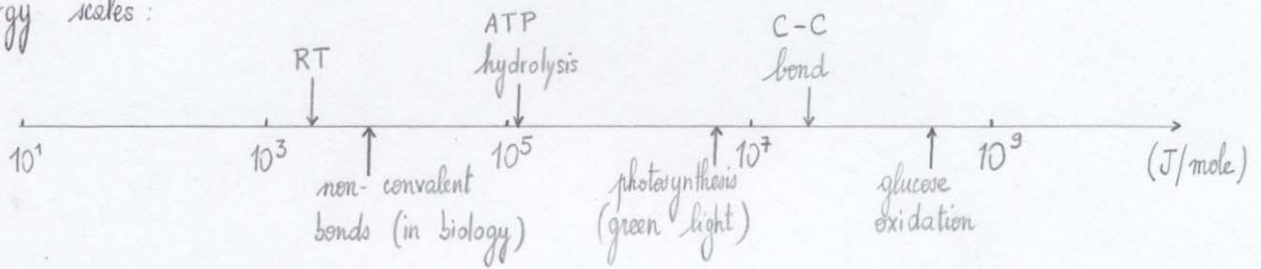
with Avogadro's number $c_{pA} \sim 6 \times 10^{23}$ and Boltzmann's constant $k_B = 1.38 \times 10^{-23} \text{ J / molecule / } ^{\circ}\text{K}$

$U_{\text{molecule}} \sim 4 \times 10^{-21} \text{ J}$
 $\sim 4 \times 10^{-12} \text{ N} \times 10^{-9} \text{ m}$

hence force scale of pN and length scale of nm. (pico Newtons, nanometers)

Coupled interactions: chemical (physical) } grand goal is to understand
 electrical } how these interact (are organized)
 mechanical } in space and time

Energy scales:



Over the past decade, much progress has been made experimentally and technically. Down on small scales, biology is geometrically dominated by filaments and membranes which is connected with chemical malleability which makes for physical complexity (nonlinearity)

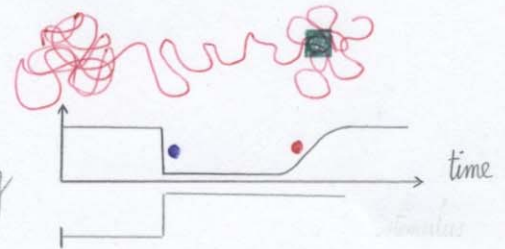


Outline:

- random walks & diffusion
- drag, mobility, Boltzmann's law, Stokes-Einstein
- biological forces & energies
- physics, mechanics and mechano-chemistry of polymers and membranes

Book reference: *E. Coli in motion* (2005) by H. C. Berg

- a bacteria tumbles randomly in a homogeneous environment, then more directed in the presence of a chemoattractant (response to a stimulus); when the chemoattractant diffuses away, randomness reappears (adaptation)



sensing and movement are coupled:

- in the absence of any active processes, what happens to a bolus of chemoattractant?

Carnot, Helmholtz, Boltzmann, Gibbs: maximize the disorder statistical probability

random walks (in 1-D) } and diffusion }
microscopic } macroscopic }



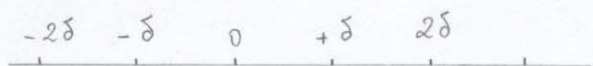
$t=0$

$t \rightarrow \infty$

- from kinetic energy: $\frac{1}{2} k_B T = \frac{1}{2} m \langle v^2 \rangle$
 $\langle v^2 \rangle = k_B T / m$ mean squared velocity

for a lysosyme $m \sim 14 \text{ kg / mole}$ and 6×10^{23} molecules / mole
 $\sqrt{\langle v^2 \rangle} \sim 14 \text{ m / s}$ very high!

but collisions and disipation \Rightarrow no net motion of the lysosyme
 velocity v , step size $\delta = \pm v \tau$ with τ : time between collisions.



① probability of going in either direction $p = 1/2$

② each step independent of others

- with N particles at origin at $t=0$

$$x_i(n) = x_i(n-1) \pm \delta \quad \begin{cases} i: \text{particle label} \\ n: \text{number of steps} = t/\tau \end{cases}$$

$$\langle x(n) \rangle = \frac{1}{N} \sum_{i=1}^N x_i(n) = \langle x_i(n-1) \rangle = \dots = \langle x_i(0) \rangle = 0$$

$$\langle x(n) \rangle = 0 \quad \text{mean location of particles}$$

- spread of distribution: variance

$$x_i^2(n) = x_i^2(n-1) + \delta^2 \pm 2\delta x_i(n-1)$$

$$\langle x_i^2(n) \rangle = \langle x_i^2(n-1) \rangle + \delta^2 = n \delta^2$$

$$\langle x^2(t/\tau) \rangle = \frac{1}{2} \cdot 2\delta^2/\tau \cdot t = 2 D t \quad \text{with } D = \frac{1}{2} \frac{\delta^2}{\tau}$$

mean square proportional to time

$$\sqrt{\langle x^2(t/\tau) \rangle} \propto t^{1/2} \quad \text{slow (diffusion)}$$

$$D \sim \sqrt{\langle v^2 \rangle} \delta \sim 10 \text{ m/s} \cdot 10^{-10} \text{ m} \sim 10^{-9} \text{ m}^2/\text{s} \quad \text{or } 10^{-5} \text{ cm}^2/\text{s}$$

$$t \mid_{10^{-4} \text{ cm} \sim 1 \mu\text{m}} \sim (10^{-6} \text{ m}) \div (10^{-9} \text{ m}^2/\text{s}) \sim 10^{-3} \text{ s} \quad \text{quick on small scale}$$

$$t \mid_{1 \mu\text{m}} \sim 10^4 \text{ s or } 10 \text{ hours} \quad \text{slow on large scale}$$

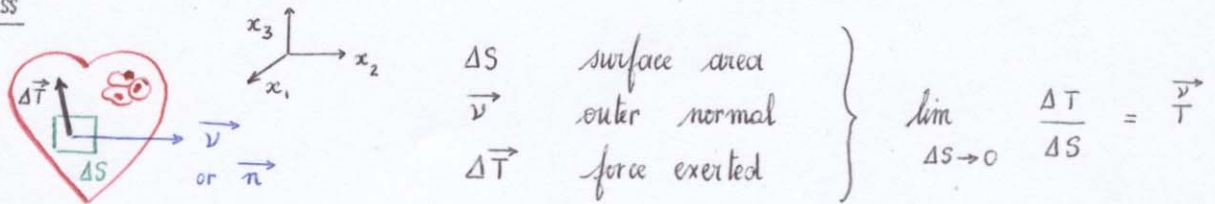
Foundations of continuum mechanics

Geert Schmid - Schönbein

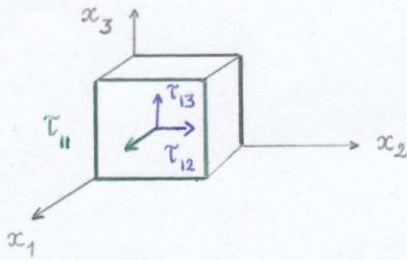
Newton's law $\vec{F} = m \frac{d\vec{v}}{dt}$ was voted to be the most important human discovery. is at the core of biology (force & velocity related).
 The major part of mechanics today is to understand and measure forces.

Forces what should the mass m be in biology? neighboring cells influence one cell
 Newton's law is not as useful in this discrete form as in engineering
force: interaction between discrete identifiable objects
 but another definition of force / interaction is needed in biology

Stress



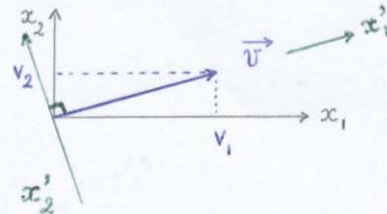
and with $\vec{v} \parallel x_1$ axis : $\frac{\vec{v}}{T}^{(1)} : \tau_{11} \quad \tau_{12} \quad \tau_{13}$
 $\vec{v} \parallel x_2$ axis : $\frac{\vec{v}}{T}^{(2)} : \tau_{21} \quad \tau_{22} \quad \tau_{23}$
 $\vec{v} \parallel x_3$ axis : $\frac{\vec{v}}{T}^{(3)} : \tau_{31} \quad \tau_{32} \quad \tau_{33}$
 stress tensor τ_{ij}
 force per unit area



$\tau_{11} \quad \tau_{22} \quad \tau_{33}$ normal stress
 $\tau_{12} \quad \tau_{13} \quad \tau_{21} \dots$ shear stress
 pressure = $-\frac{1}{3} (\tau_{11} + \tau_{22} + \tau_{33})$
 from outer world onto cubic element

Principal stress

Starting with an analogy with vectors
 coordinates depend on set of axes
 in principal coordinates system (one and only one exists):



$$\begin{pmatrix} \tau_{11} & 0 & 0 \\ 0 & \tau_{22} & 0 \\ 0 & 0 & \tau_{33} \end{pmatrix}$$

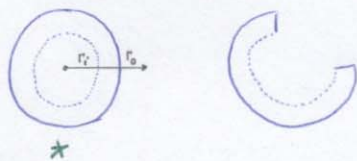
mean normal stress

Stress deviators

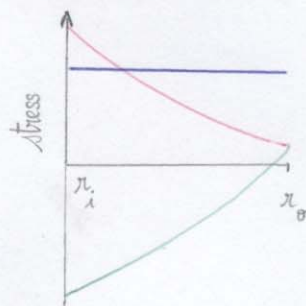
the solutions to certain problems are independent of normal stresses.

$$\begin{cases} \tau_{ij}^* = \tau_{ij} - \frac{1}{3} (\tau_{11} + \tau_{22} + \tau_{33}) \delta_{ij} \\ \tau_{ij}^* = \tau_{ij} + p \delta_{ij} \\ \delta_{ij} = 0 \text{ if } i \neq j \text{ and } 1 \text{ if } i=j \end{cases}$$

Residual stress

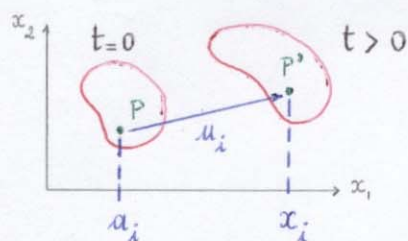


the unloaded* state is not necessarily stress-free



Deformation

strain
strain rate

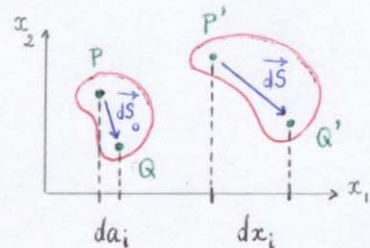


displacement $u_i = x_i - a_i$

final - initial

includes deformation AND translation

Hooke: change in length measures deformation alone

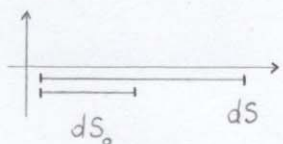


change of length $dS^2 - dS_0^2 = \text{strain} \times 2 \times \text{initial distance}$

strain $E_{ij} = \frac{1}{2} \left(\frac{\partial x_k}{\partial a_j} \cdot \frac{\partial x_k}{\partial a_i} - \delta_{ij} \right)$

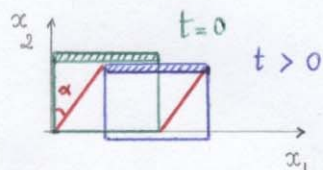
deformation gradients $\frac{\partial x_k}{\partial a_j}$

in 1-D motion



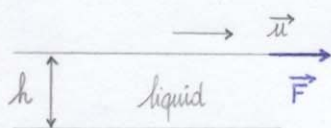
$E_{11} = \frac{1}{2} \left(\frac{dS^2}{dS_0^2} - 1 \right)$

stretch ratio $\lambda = \frac{dS}{dS_0}$

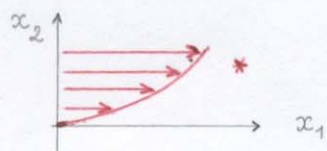


$E_{12} \approx \tan \alpha$

Reference states in biology are debatable, difficult to define (because deformable materials).
In a fluid, what quantity is proportional to the applied stress? strain rate



$F \sim \frac{u}{h}$ velocity gradient



spatial velocity gradients
or strain rate

$$\begin{matrix} \frac{\partial v_1}{\partial x_1} & \frac{\partial v_1}{\partial x_2} & \frac{\partial v_1}{\partial x_3} \\ \vdots & \vdots & \vdots \\ \frac{\partial v_3}{\partial x_3} \end{matrix}$$