# biophysics

PHYS 320 / 420: Intro to Biological Physics



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- ▶ Problem sets: 100% of the course grade. Each problem set will be based on a recent paper in the literature, and consist of two parts: an intro explaining why the topic is important and interesting, and then a guided, step-by-step walk through some calculations in the paper (either analytical or numerical). Working in groups is fine (even encouraged), but please hand in your own write-up.

# The arrogance of physicists



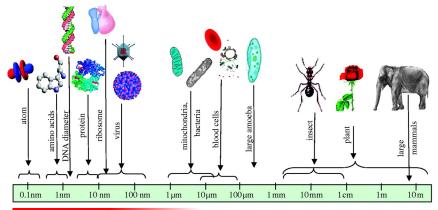
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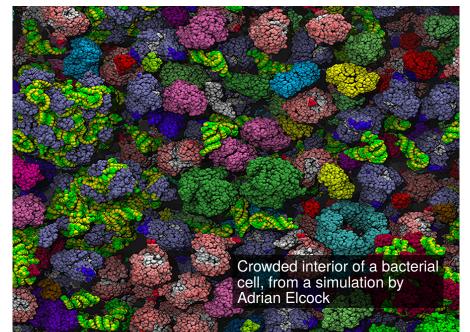
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# Fluctuations rule at scales below 1 $\mu \mathrm{m}$



thermal fluctuations matter

# The not-so-primordial soup



Shaken, not stirred...

See movie in video link on website.

Virus assembly [Perlmutter et al., eLife (2013)]

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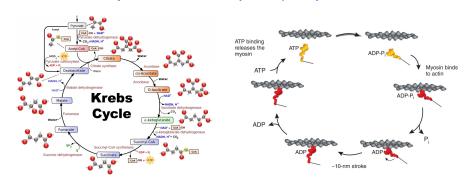
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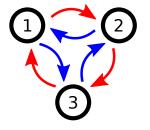
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Biology provides a spectacular guide to the **how** of life. Can physics help provide a non-trivial answer to the **why**?

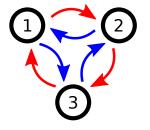
The bane of every intro bio course: cycles upon cycles...



For us, all these will be specific cases of a **universal mathematical framework**: Markov state models, describing stochastic transitions between states.

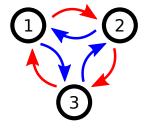


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Versatility of the approach: the states can be molecule numbers in a chemical reaction, different structures of a protein, the populations of a genetic variant...

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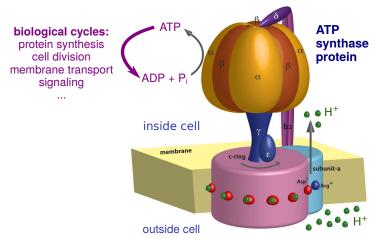


Two broad categories of transition networks:

- equilibrium: any random path on the network and its reverse are equally likely
- ► non-equilibrium: the symmetry between paths and their reverses is broken ⇒ requires external fuel source (we will prove this!)

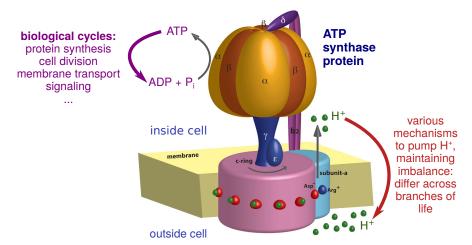
## Non-equilibrium driving in living systems: ATP

ATP as the prime biological fuel source is as universal as the genetic code.



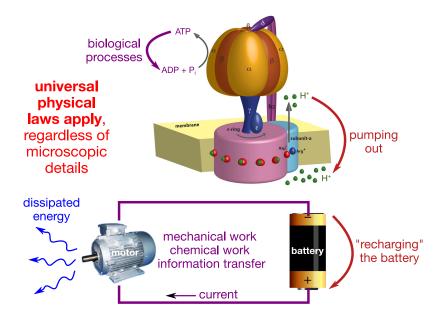
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Synthesizing ATP is a **nonequilibrium** process, driven by ion imbalances.

# The role of physics: nonequilibrium thermodynamics



#### The second law of thermodynamics



Evolutionary conundrum: Self-recharging batteries cannot spontaneously arise from an equilibrium primordial soup.

# DAMN YOU THERMODYNAMICS

Always ruining everything

### The second law of thermodynamics



DAMN YOU THERMODYNAMICS

Always ruining everything

Evolutionary conundrum:

Self-recharging batteries cannot spontaneously arise from an equilibrium primordial soup.

Every persistent imbalance in nature is driven by a more fundamental imbalance:

If you see a current, search for the power outlet.

### The fundamental imbalances

All nonequilibrium processes on earth:

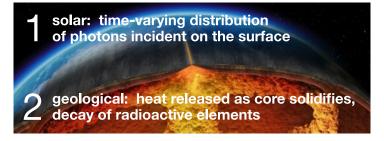


#### The fundamental imbalances

All nonequilibrium processes on earth:



are ultimately "plugged into" two major imbalances:



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True universality: The molecular details of potential life-forms on exoplanets will be different, but the constraints of nonequilibrium physics will be the same.

