

# 1 overview

- What is bionanotechnology
- Bionanomachines, How do they work?
- Structural and functional principles of bionanotechnology
- Modern bionanotechnology

## 1.1 Suggested reading

E. Gazit, Plenty of room for biology at the bottom

Bionanotechnology lessons, Nature

Nanobiotechnology: Concepts, Applications and Perspectives.

Nanobiotechnology II: More Concepts and Applications.

Bionanotechnology: Proteins to Nanodevices

## 1.2 Grading scheme

- Final examination 50% (12 Dec. ?) - we need to pass it
- Final report (Due to Feb. 6 2017) 50%
- Physical chemistry knowledge is expected (free energy, Gibbs free energy, Helmholtz free energy etc.)

## 1.3 Workshop - second part of the semester

1. constructing synthetic nanopores 2. solvating sucrose butyrate nanopores  
3. applying ... ..

We will be supplied with tutorial and all necessary files. In the tutorial there will be 3 tasks - those are necessary. There is also one Challenge that is necessary. If you do any other challenge you might get extra points (5.5).

## 1.4 Grading seminar

- we start Nov. 28.
- List of all seminar topics is given on eportal.
- You have to declare time and topic of your speech on some blog.
- Oral presentation counts for 80% of grade.
- Participation is 20%
- Presence is obligatory.
- Presentation should be based on recent papers. (up to 5 yr. old)

- Speech should last 30 min +- 5 min. Discussion should last about 5-10 min.
- try to limit your presentation to 20-25 slides.
- presentation goal should be to enchant us.

Presentation goals:

1. deliver clear introduction and the purpose of the study
2. explain the necessary methodology
3. present the result/discussion
4. conclude the topic
5. provide the comprehensive analysis of the strong and weak points of the study
6. **very important** make a personal constructive criticism
7. Once you have a draft of your speech go ask profesor if it is enough.

## 2 2 class

Nanoscale is everything up to 100nm

Top-down approach ( Michael Angel craved his pieta in a big block. its about making something, starting with something bigger.

Nanotechnology (molecular nanotechnology) - a technology that manipulates matter at dimensions up to 100nm.

The study of nanotechnology includes molecular systems, molecular assemblies (such as quantum dots), and organized self-assembled devices and machines.

Richard Feynman - nice physics lectures

Macro- vs nano-scale Upon reducing the size and no change in substance, fundamental properties of material such as electrical conductivity, melting point, color etc. will can change.

For example something that is soft and malleable on the macro-scale may be 100 times stronger and 6 times lighter than steel at the nano-scale (nanotubes).

Biotechnology is technology that involves the use and manipulation of living organisms - genetic engineering.

Nanobiotechnology - the use of nanoscience for specific biological applications.

Bionanotechnology - subset of nanotechnology where biology provides the inspiration and or the ultimate goal.

Technology that uses biological assemblies for various applications that may not be directionally associated with biology.

## 3 3 class - Natural bionanomachines

### 3.1 Examples of bionanomachines

These are examples of bionanomachines that work outside the cells: \*pepsin \*lysozyme \*amylase They find their use in industry like laundry specific, corn sugar thing (syrup kukurydziany)

When you start to look at the nanoscale the fundamental properties are changing, for example consider gravity. If you have huge building you need some kind of support for material to last. On the other hand when you go to microscale droplets can stay on the leaf - that is because adhesion and random forces are greater than gravity meaning those small things do not obey gravity. Other example is moving bacteria once it will stop moving its flagellum? it will stop immediately - on opposite side consider a boat.

### 3.2 More examples

Nanocar build of fullerene wheels and alkynes axis. This car is placed on gold surface, once it is heated fullerene wheels start to rotate cause there is very low bond-bond rotational boundary. Few years later there were proposed another nanocar with engine this time, build out of photoactive compound that rotates under light. Structure of this motor is a bit harsh opposite to real car, that is because you can't play with atoms as you like.

### 3.3 Examples summary

In nanoscale there is no smooth atomic-scale motion (transition from one rotatory state to the next as utilized when the appropriate chemical energy is applied. Individual atomic properties (covalent bonds, steric interaction, electrostatic interactions, hydrogen bonds) are defined rather than bulk properties (e.g. viscosity, friction).

Natural biomachines operate in cellular space - that is very special conditions, that are very important to specific nanomachine

Individual parts interact through random motion and diffusion

Q: Is diffusive motion sufficient to allow interaction between the two bionanmachines in cell containing millions of other biomolecules. This may happen on the exam and should be answered with some equations about diffusion.

### 3.4 Bionanomachines in water environment

\*form and function of biological system are linked to chemical properties of individual components and water environment : hydrophobic effect is largely responsible for it \* in water biomolecules are able to form a single form?? .. missed it

Hydrophobic effect narrows number of possible protein conformations \* Carbon rich parts of the protein are hydrophobic Q: Think about hydrophobic effect in terms of thermodynamic what you gain and lose in terms of enthalpy, what about entropy? Final goal will be to tell what happens with Gibbs energy. This process is spontaneous  $\Delta G < 0$ .

### 3.5 Four 'molecules of life'

Q: Structures of proteins, lipids, polysaccharides and nucleic acids and impact of their structure on their properties. Q: 1st and 2nd law of thermodynamics. 3rd and 0th (thermometer)

### 3.6 Errors are natural part of protein synthesis

\*In bacterial cells, the genetic sequence is misread in about 1 in 2000 aa. The process; errors have often little effect on the function of the protein \* synthesis of the protein may terminate early and produce a truncated chain due to processivity errors. More common \* collagen as an example of a protein

### 3.7 Characteristics of nucleic acids

\*applied in nanoscale data storage and retrieval \*every ....

### 3.8 Lipids

\*Lipids are used for cellular infrastructure

### 3.9 Polysaccharides

Sacharose what is the conformation of each unit D or L? Carbon-hydrogen bonds are reservoirs of energy. Because glucose or sacharose is soluble in water it cannot be used for storing energy. Glycogen is much more appropriate for this purpose because it is insoluble, and more over there is simple access for enzymes to cut it down. \* different linear and branched polymers are created for different needs \* individual chains may associate with the large quantity of water forming gluttery gel \* carbohydrate chains may associate tightly side by side creating strong fibers with almost ....

### 3.10 Limits imposed by evolution

\* cells use a few synthetic techniques and rely on a few simple molecular plans to build their different bionanomachines \* Small steps are better - big evolutionary step might have disastrous results — evolution favors modification over innovation \* biomolecules require water environment, as well as proper temperature, pH and salinity. \* Biomolecules are constrained by: \* 20 amino acids alphabet \* ... \* bionanomachines have short life span and are build to perform only one task.

### 3.11 Aims of bionanotechnology

\* designing bionanomachinery from scratch. There is much problems with this task: \* no reliable folded structure of a protein from its chemical sequence \*

tools that are available today are don't give us an insight into chemical activity of folded protein structure.

## 4 Methods in bionanotechnology

### 4.1 Recombinant DNA technology

- \* using recombinant DNA technology one can construct any required protein.
- \* two natural enzymes: restriction enzymes and DNA ligase are utilized in technology.
- \* Restriction enzymes types: 7 classes
- \* Type I cleaves DNA at random sites far from its recognition sequence
- \* Then it gets better and more specific
- \* Restriction enzymes might produce blunt or sticky ends.
- \* Chemical synthesis of DNA, there are troubles with lengths of those.

### 4.2 Pros and cons of using bacteria

- \* easy to grow and cheap
- \* many animal and plant proteins have carbohydrate groups attached to their surfaces to be active and bacteria do not add these groups to engineered proteins.
- \* proteins tend to aggregate when they reach high concentration forming inclusion bodies. They are formed when new proteins associate randomly..

### 4.3 Cell-free methods of producing proteins

Why? It provides a controlled method for synthesizing proteins are difficult in engineered bacteria such as:

- \* membrane binding proteins
- \* proteins with

This is done *in vitro*

### 4.4 Site-directed mutagenesis

HOME WORK!

- \* may be used in determining the function of specific aa or regions within a protein
- \* may improve the stability of proteins, by engineering in cross-linking residues or improving the fitting of residues within the protein interior.

### 4.5 Fusion/chimeric proteins

Two proteins with different functions are combined creating a hybrid protein with both functions. For instance anticancer immunotoxins.

### 4.6 Antibodies

are interesting because there is a need for an effective method for recognizing individual molecules.

- \* Immune system produces high amount of different antibodies with hope that one will fit  $10^{15}$
- \* If we combine this natural library of molecules with modern method of synthesizing antibodies.
- \* It is now routinely

possible to obtain antibodies capable of high-affinity recognition of virtually any molecule. \* Monoclonal and heteroclonal antibodies production. \* antibodies can be found in: pregnancy test, heart protein in blood - heart attack, test for HIV viruses, test for patient with Lupus (autoimmune disease). \* Antibodies can be also used to treat diseases: neutralize toxins (snake toxin), effective in treating some types of cancer (Hodgkin lymphoma).

## 5 X-ray crystallography

\* X-ray technique provides the most detailed atomic structures \* resolution of the structure depends on the quality of the crystals \* typical X-ray studies of proteins are resolved within 1.5-3.0 Å \* at 3.0 Å final coordinates must be taken with care \* mobile regions of the structure may not be well resolved \* X-ray structure is average structure \* temperature factors (B-values) of the atomic positions are good indicators of the quality of coordinates. Large values > 40 might mean 2 things: this amino acid is mobile or resolution of this part of the protein is low. \* protein is bounded within oriented crystal lattice - single conformation \* to study functional aspects of proteins number of crystals obtained under varied conditions must be studied.