### Physical aspects in the self-assembly of biological complexes



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# Self-assembling supra-molecular systems

 Example of supramolecular hierarchy: the skeleton of a sea sponge (Eupectella, J. Aizenberg, 2003).



### What questions can physicists answer?

- What is the mechanism of self-assembly?
  - What intermediates, kinetic bottlenecks?
  - How do the assembly pathways depend on environment?
- What are the dynamics of the final complex?
- What physical properties emerge from a given hybrid structure?
  - How to design a biological metamaterial.

#### Viruses: best size/complexity ratio?

- Most viruses are symmetric: icosahedral or helical capsid structures.
- Most capsids are composed of a small number of types of proteins.

CAPSID \*



Session L34: Virus-Inspired Supramolecular Structures Tuesday afternoon, March 6, 2007, APS Meeting.

### Mechanism of self-assembly: theory



### Mechanism of self-assembly: experiment

#### Thermodynamics



Gelbart et coll., PNAS, 2003  $\lambda$ -phage packages DNA at a pressure of 20 atm.

Kinetics (DLS)

- Cryo-TEM
  - Zlotnick et coll.: Capsid formation is a nucleated process and the nucleus is a pentamer of dimers.

NOTE: ensemble averaging methods do not provide direct insights into the pathways of assembly.

#### Assembly and dynamics – single virus

Optical tweezers



 $\Phi$ 29: the strongest motor known Characterization of the mechanical work cycle Bustamante et coll., Cell, 2005 Atomic force microscopy



G. Wuite et coll., PRL, 2006 Continuum theory holds – important for interpretation of dynamics

# Detection and manipulation of single viruses – a need for new methods



Klenerman et coll., JACS, 2006

 $\mathbf{F} = -\alpha \nabla \mathbf{E}^2$ 

Two ways to reach subwavelength particle manipulation

- Increase polarizability
  - Protein coat (capsid)



VLP's C. Chen et al., Nano Lett., 2006 Use near-field: sharper gradients and field enhancements



Okamoto and Kawata, PRL, 1999

### The first questions

- Will VLP models be relevant for studying viral processes?
- What forces and what optical signals can be expected from near-field?



### Aperture-trap: experimental setup



### First results



### 3D field intensity mapping



Amarie D, Rawlinson ND, Schaich WL, et al. <u>Three-dimensional mapping of the light intensity transmitted through nanoapertures</u> NANO LETTERS 5: 1227-1230 JUL 2005

## Tailoring field distributions and enhancements

SP Modes

Max. Enhancement: 3000



### On-going projects

#### Optothermal ratchet





### Virus-like particles -from probes to metamaterials-



Chen, C. *et al.* Nanoparticle-Templated Assembly of Viral Protein Cages. *Nano Letters* **6**, 611-616 (2006).

### Particle acts as a template promoting self-assembly



~ 90% encapsulation efficiency. (As good as viral RNA)

### Core-induced polymorphism



Sun et al., PNAS, 2007

# Photothermal in-vivo tracking of metal nanoparticles



### Towards bio-metamaterials

- Metamaterials: optical properties determined by their organized structure rather than inherited directly from the material properties of individual subunits.
- Metallodielectric metamaterials are composed of resonant metal inclusions in a dielectric matrix and have subwavelength lattice periods.
  - Smith et al., Metamaterials and negative refractive index. Science 305, 788-792 (2004).
  - Brown et al., Large Electromagnetic Stop Bands in Metallodielectric Photonic Crystals. *Applied Physics Letters* 67, 2138-2140 (1995).
  - Fan et al., Large omnidirectional band gaps in metallodielectric photonic crystals. *Physical Review B* 54, 11245-11251 (1996).

#### Symmetric protein shells: 3D structures with sub-100 nm lattice parameter



# Plasmon hybridization (Nordlander, 2003)



### 3D metallodielectric plasmonic band structure





### Why VLPs?Colloidal crystal:To preserve long range order $\sigma_{position} = N^{1/2} \cdot \sigma_{Au}$



Characteristic lengths:  $\lambda_0 \approx 0.6 \ \mu m$   $\lambda_{SP} \approx ?$  $\lambda_{decay} \approx 20 \ \mu m$ 

VLP crystal:



# VLPs crystallize in 3D metallodielectric arrays



### Band splitting observed



### Conclusions

- There is a vast unexplored territory in the area of mechanisms of self-assembly.
- New Tools (both theoretical and experimental) are needed to access:
  - Dynamics,
  - Intermediates,
  - Relationship between structure and properties.
- Optics provides convenient ways to manipulate, probe and build.
- Examples: near-field forces for trap integration, virusmimic probes, biological metamaterials.

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