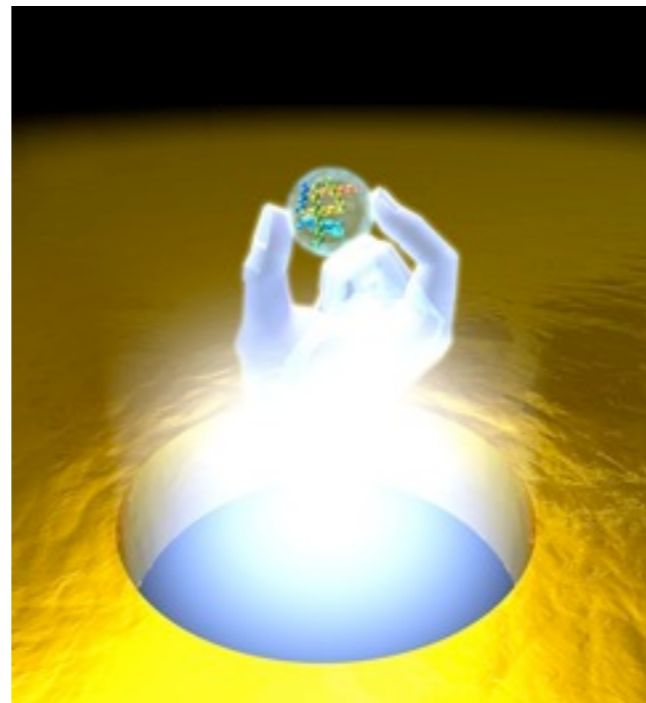


# BioPlasmonics:

Developing novel nanotools for biosciences & medicine



Romain Quidant

ICFO-Institut de Ciències Fotòniques &  
ICREA-Institució Catalana de Recerca i Estudis Avançats

# ICFO - The Institute of Photonic Sciences

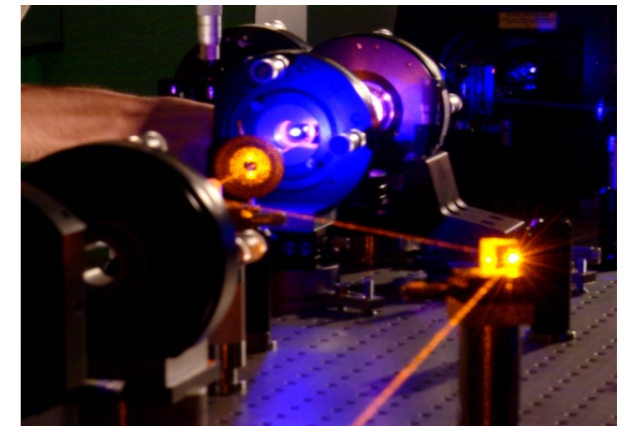


Advancing the limits of scientific and technological knowledge in optical sciences



A few figures...

- 250 people (200 researchers)
- 20 groups (15 senior and 5 junior)
- 14000 square meters (250 clean room)





Advancing the limits of scientific and technological knowledge in optical sciences

For more details please visit: [www.icfo.es](http://www.icfo.es)

## BIO PHOTONICS

Optical Tweezers; Plasmonics Oncology; Nano-Surgery & Neuro-Photonics; Chemically Selective Imaging; Sensors; Lab-on-a-Chip; Single-Molecule Photonics; Ultrafast Microscopy; Cell Spectroscopy.

## QUANTUM OPTICS

Quantum Information Processing & Communication; Quantum Cryptography; Quantum Imaging & Metrology; Coherent Control & Quantum Biology; Single-Atom Photonics; Quantum Dynamics.

## NONLINEAR OPTICS

All-Optical Photonic Devices; Frequency Conversion and OPOs; Photonic Crystals; Ultrafast & Atto-Optics; Tunable Lasers; Electro & Acousto-Optic Devices; Fiber & Integrated Sensors; Photonics Clean-Tech.

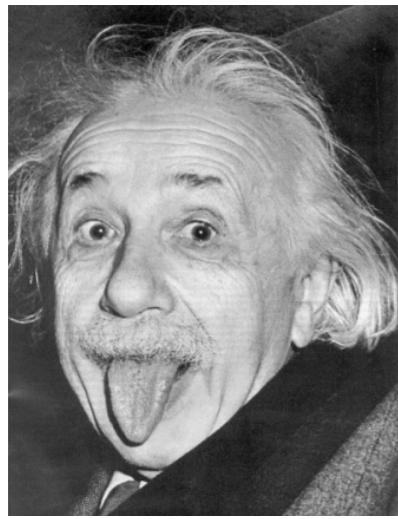
## NANO PHOTONICS

Nano-Photonic Devices; Organic LEDs; Light Harvesting; Solar Cells; Plasmonics; Molecular NanoPhotonics; Nano-Optical Tweezers; Nano-Optical Manipulation on a Chip; Nano-Antennas; Nano-Cavities.



# PROVIDING NOVEL NANOTOOLS TO MEDICAL DOCTORS

**Physicist**



10 – 100 nm



**Medical doctor**



- Earlier and more reliable diagnosis
- More efficient & less-invasive therapies

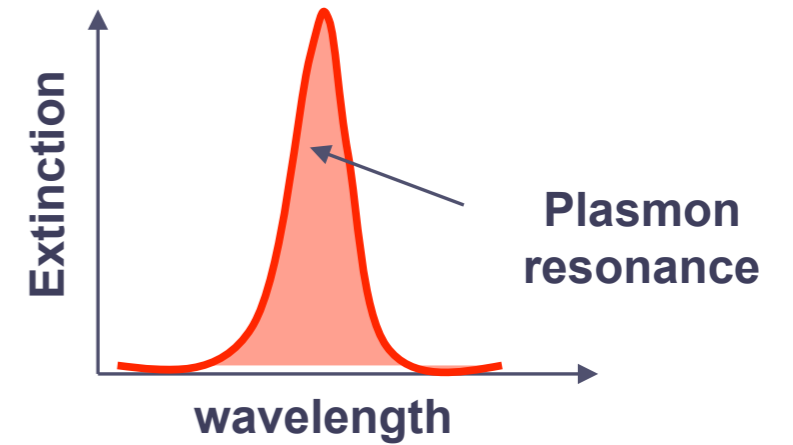
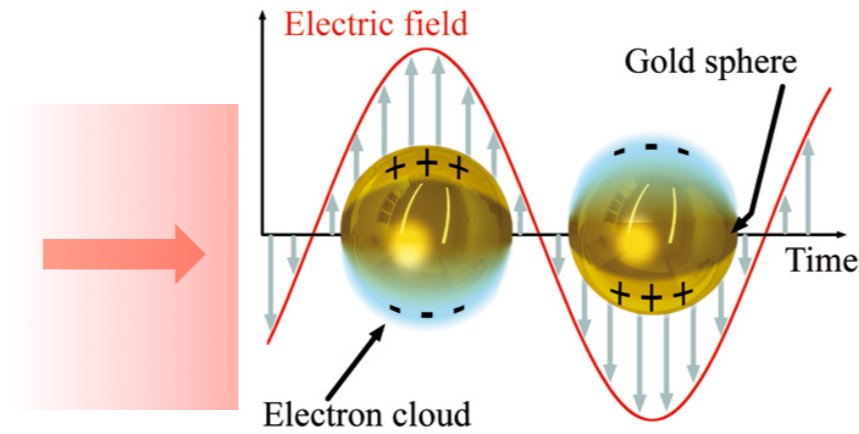


# The protagonist? A gold nanoparticle

## GOLD NANOPARTICLE



## LOCALIZED SURFACE PLASMONS



Enhanced Scattering

Enhanced Absorption

Nano-source  
of light

Nano-source  
of heat

# Outline

Part 1- Plasmon nano-optics for sensing and trapping  
- wed 9.45 am

Part 2 - ThermoPlasmonics: using metallic NP as heat  
nanosources - Thur 9.00 am

- Part 1 -

Plasmon nano-optics for sensing and trapping:

Towards an integrated plasmon-based analytical platform  
for early cancer diagnosis



# FROM THE MACRO- TO THE NANO-SCALE

- ✓ Cancer is usually diagnosed based on the detection of a **macroscopic** tumor

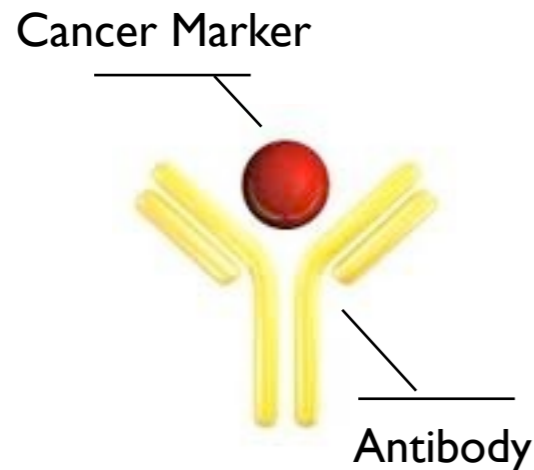


## An example: Breast cancer



- Screening through Mammograms and MRI
- Diagnostic through biopsy (taking a piece of tissue and inspect it below the microscope)

- ✓ **Earlier detection** could be achieved by detecting low concentration of **cancer markers** over-expressed at the membrane of cancer cells or in blood.

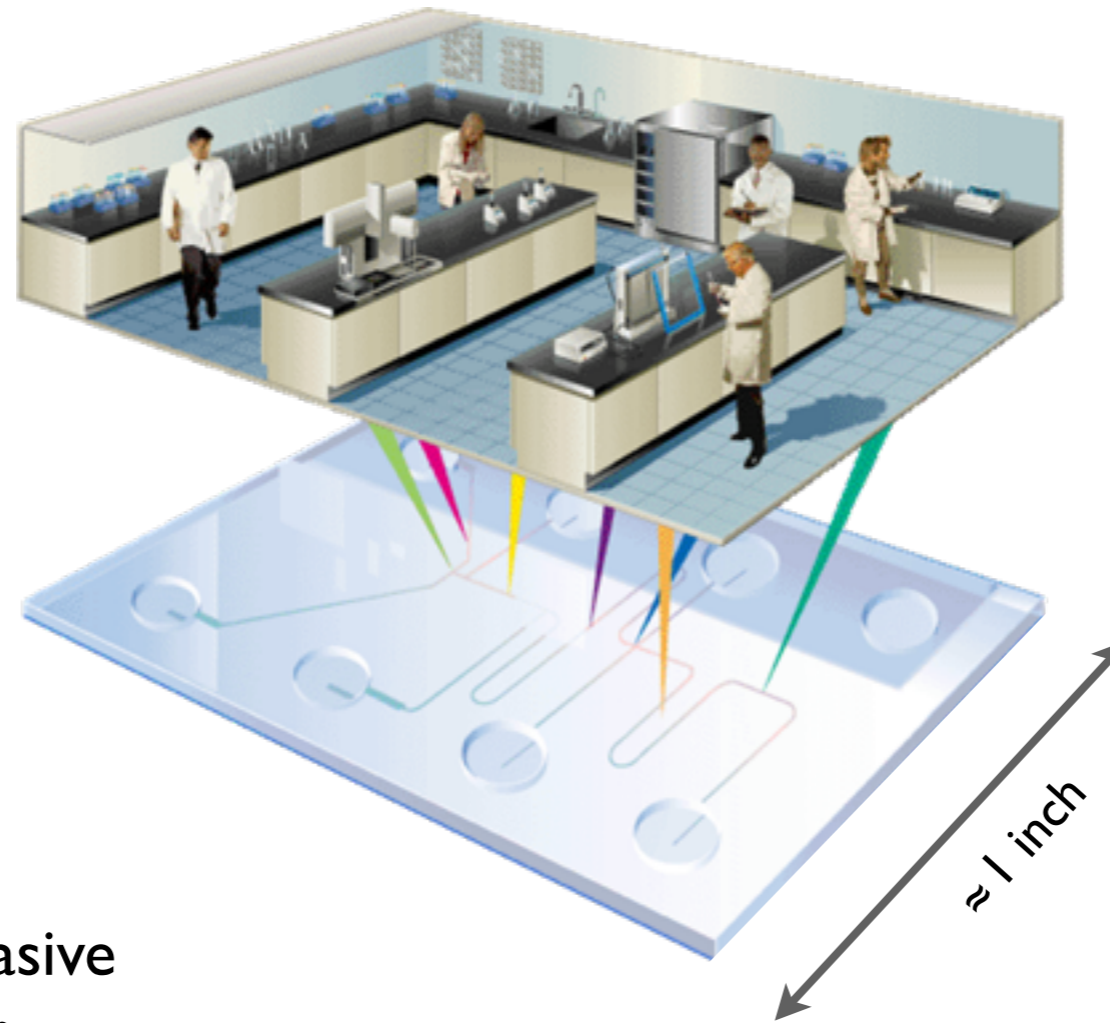


### Cancer Marker detection must combine:

- High sensitivity and specificity
- Label-less
- Parallel assays
- Cost effective

# LAB-ON-A-CHIP

Honey, I shrunk the lab....

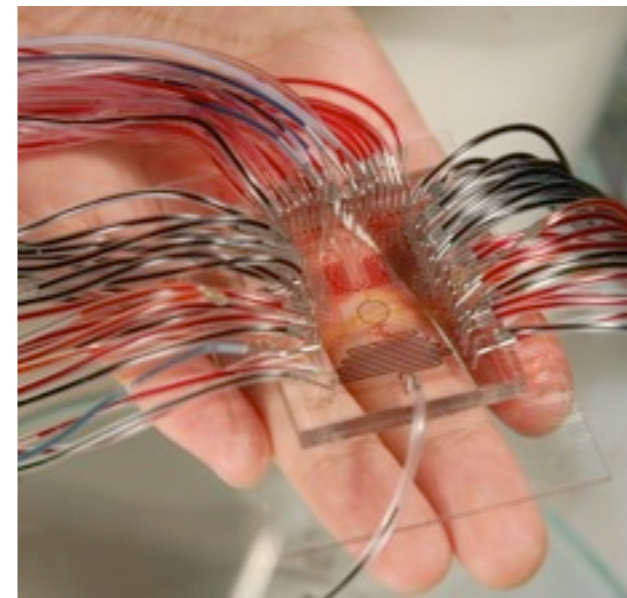
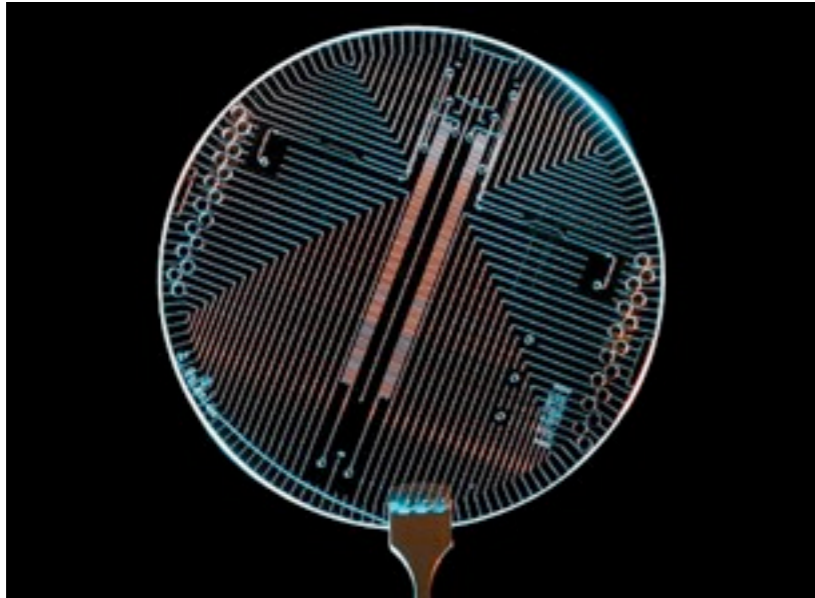


But it is now...

- Less invasive
- Cheaper
- Transportable
- More sensitive

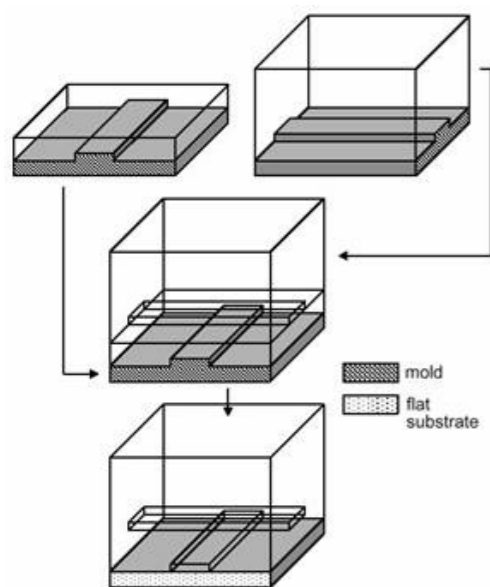
# MICROFLUIDICS: THE WALL OF THE LABORATORY

## The corridors and the lab space...

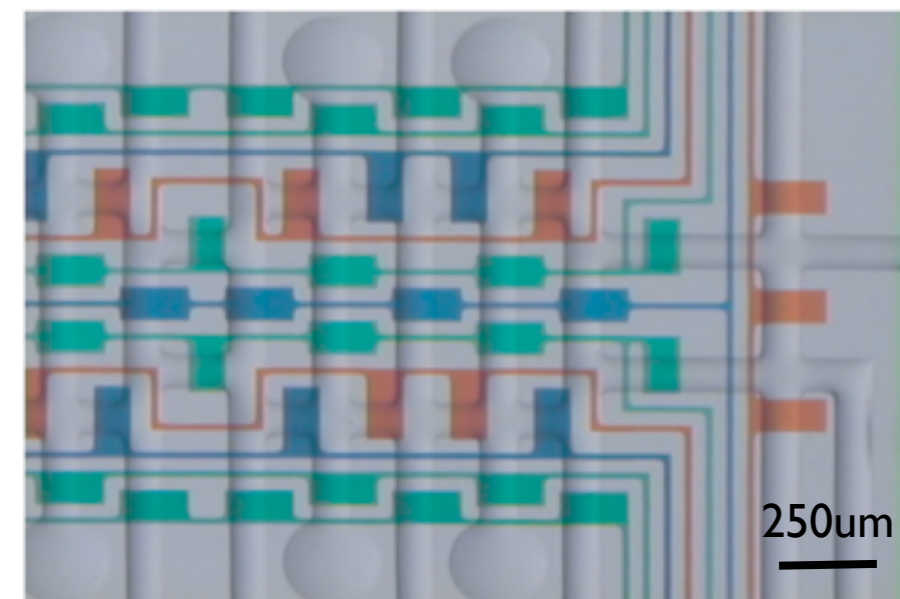
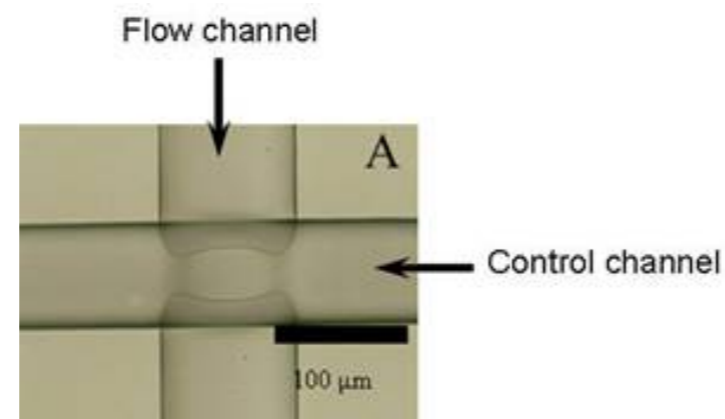


- Small fluid volumes (nl, pl)
- Cheap
- Transparent

## Please close the door when leaving...



The Quake's valve

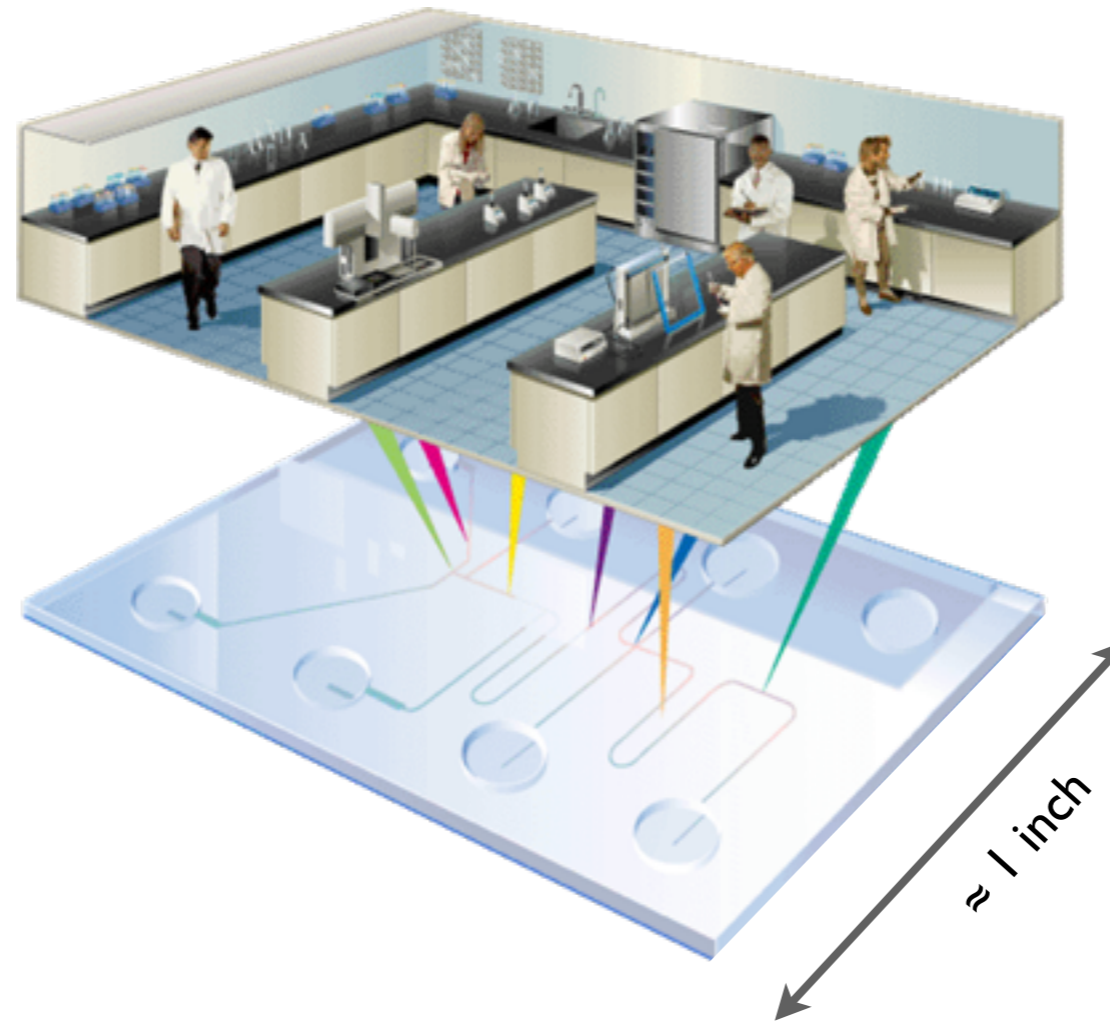


Courtesy of S. Maerkl



# LAB-ON-A-CHIP

Honey, I shrunk the lab....



But it is now...

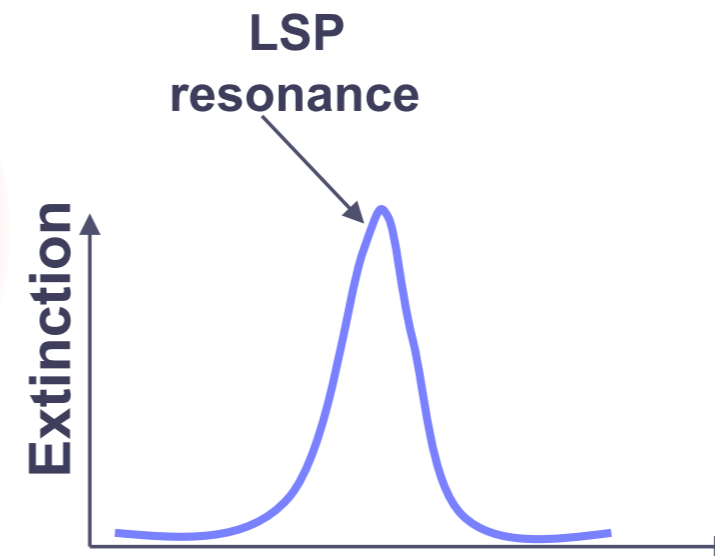
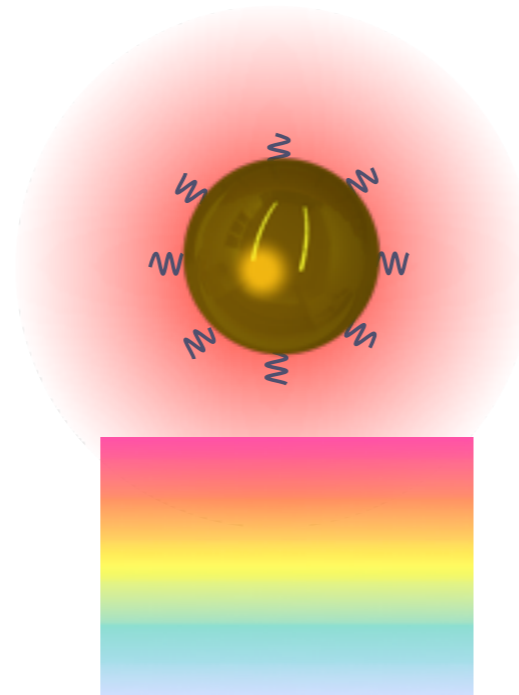
- Less invasive
- Cheaper
- Transportable
- More sensitive

# LSPR-sensing (Resonance shift sensing)

LSP resonance are highly sensitive to a small change of the dielectric surrounding

$$\alpha(\omega) = 4\pi\epsilon_{ref}a^3 \frac{\epsilon_{metal}(\omega) - \epsilon_{ref}}{\epsilon_{metal}(\omega) + 2\epsilon_{ref}}$$

- ≡ receptor
- Target molecule



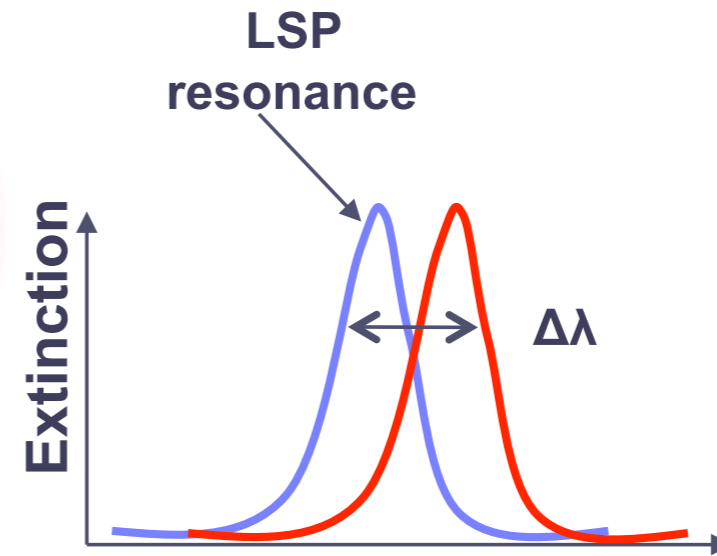
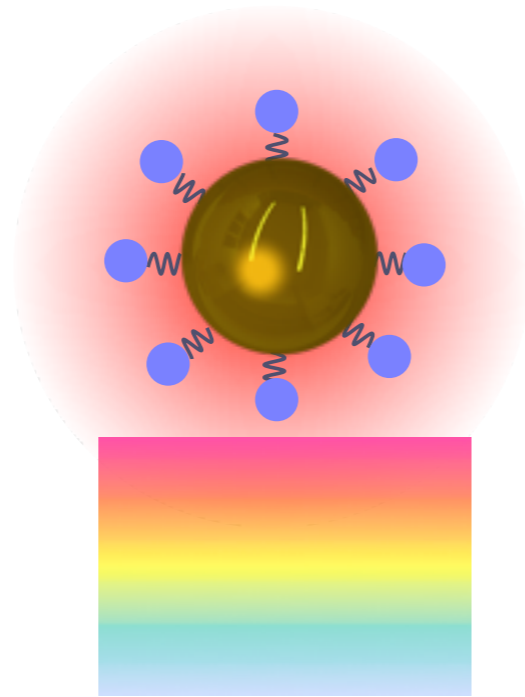
By monitoring  $\Delta\lambda$  one gets a direct information about the number of binding events occurring at the particle surface

# LSPR-sensing (Resonance shift sensing)

LSP resonance are highly sensitive to a small change of the dielectric surrounding

$$\alpha(\omega) = 4\pi\epsilon_{ref}a^3 \frac{\epsilon_{metal}(\omega) - \epsilon_{ref}}{\epsilon_{metal}(\omega) + 2\epsilon_{ref}}$$

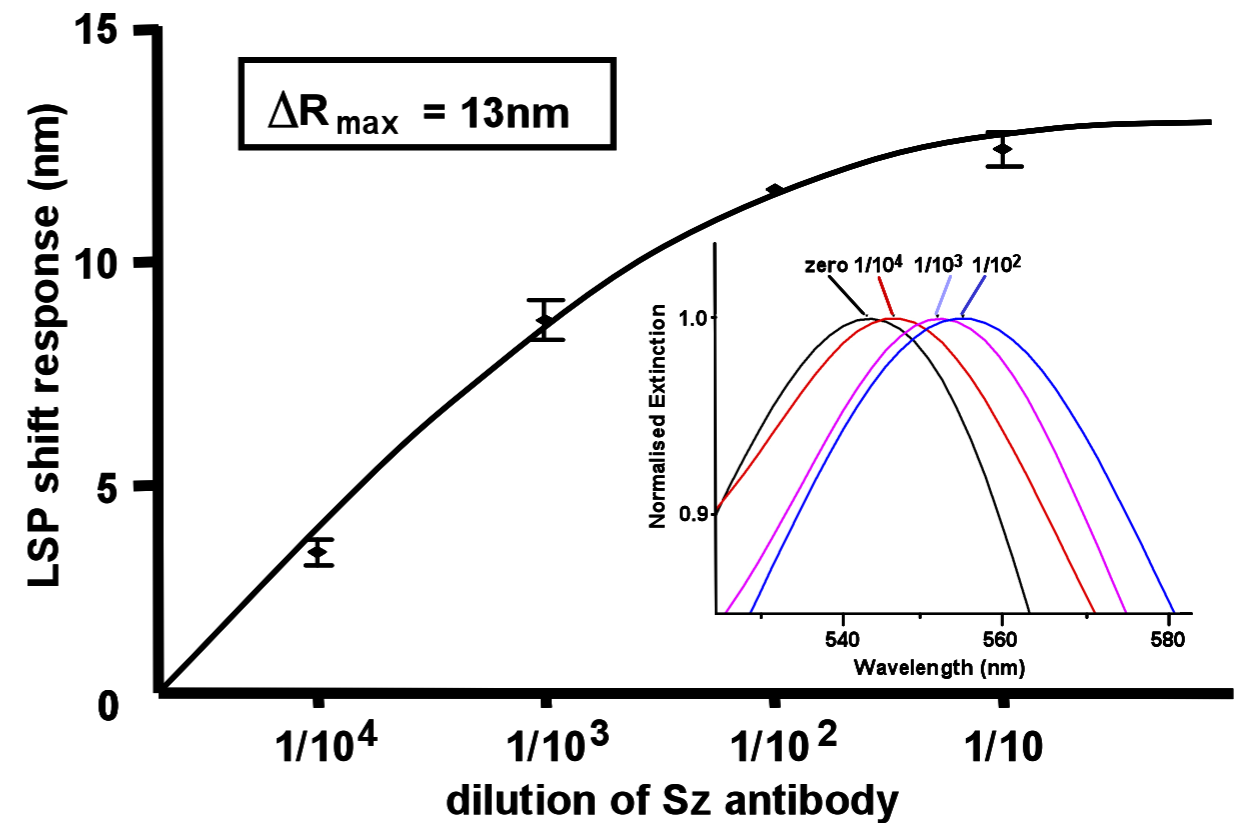
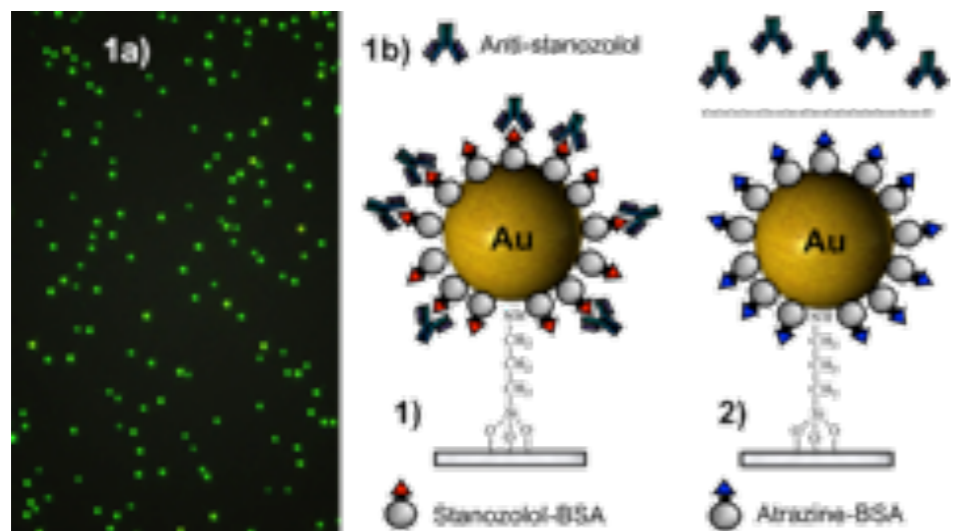
- ≡ receptor
- Target molecule



By monitoring  $\Delta\lambda$  one gets a direct information about the number of binding events occurring at the particle surface



# LSPR sensing with randomly arranged gold colloids



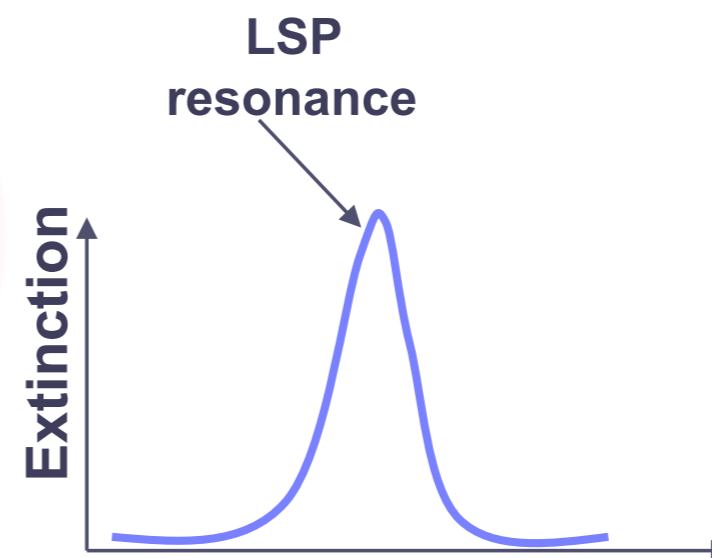
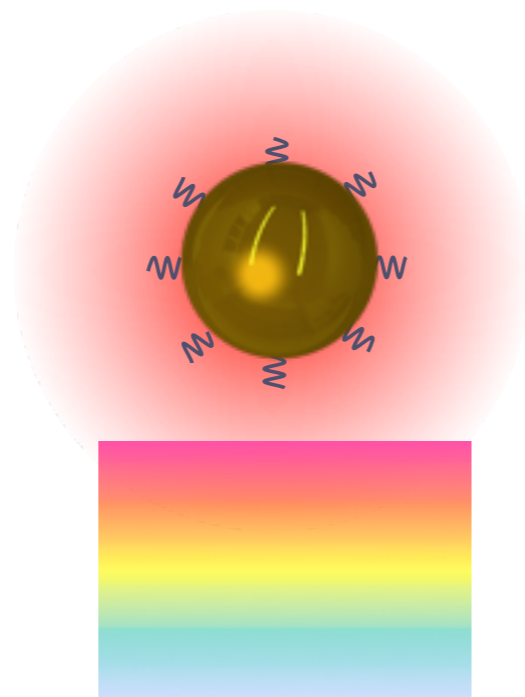
- ✓ Only 0.6 nm (4% of max shift) of non-specific binding
- ✓ Sensitivity in the nM range (higher sensitivity is expected with patterned samples)

# What dictates the Sensitivity of an optical sensor?

LSP resonance are highly sensitive to a small change of the dielectric surrounding

$$\alpha(\omega) = 4\pi\epsilon_{ref}a^3 \frac{\epsilon_{metal}(\omega) - \epsilon_{ref}}{\epsilon_{metal}(\omega) + 2\epsilon_{ref}}$$

- ≍ receptor
- Target molecule



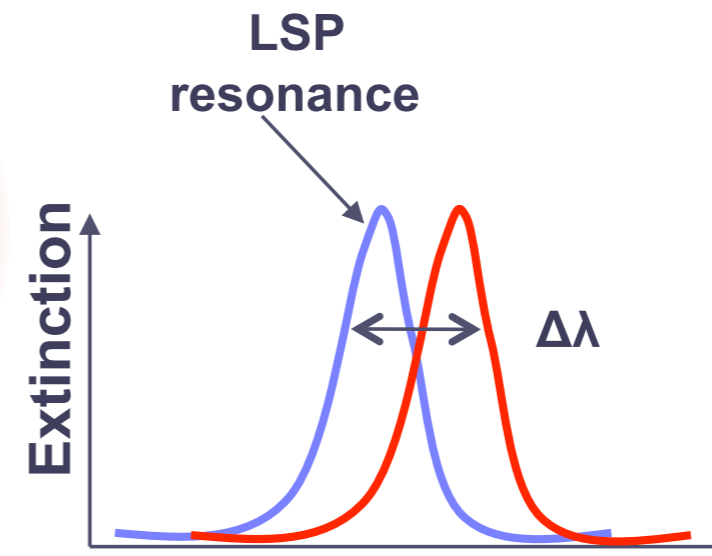
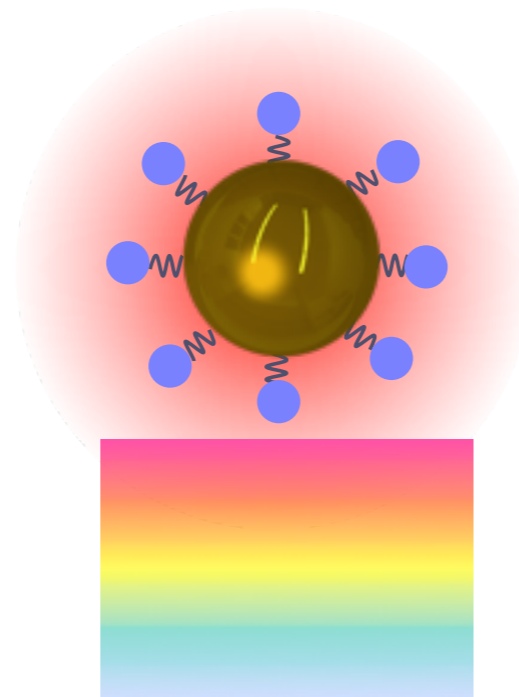
By monitoring  $\Delta\lambda$  one gets a direct information about the number of binding events occurring at the particle surface

# What dictates the Sensitivity of an optical sensor?

LSP resonance are highly sensitive to a small change of the dielectric surrounding

$$\alpha(\omega) = 4\pi\epsilon_{ref}a^3 \frac{\epsilon_{metal}(\omega) - \epsilon_{ref}}{\epsilon_{metal}(\omega) + 2\epsilon_{ref}}$$

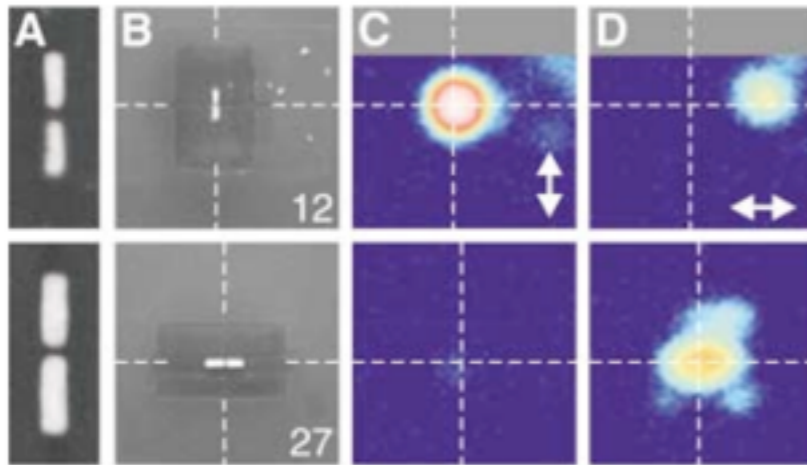
- ≡ receptor
- Target molecule



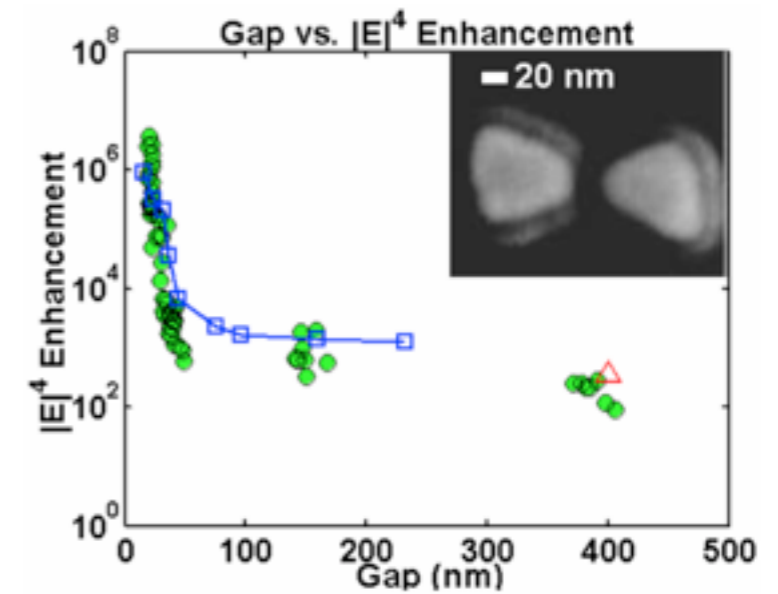
By monitoring  $\Delta\lambda$  one gets a direct information about the number of binding events occurring at the particle surface

# From Radiowave to Optical antennas...

RW Half-wave dipole antenna



P. Muhlschlegel et al, Science **308**, 1607 (2005)



P. J. Schuck et al, Phys. Rev. Lett. 94, 017402 (2005)

# Optical Gap Antenna

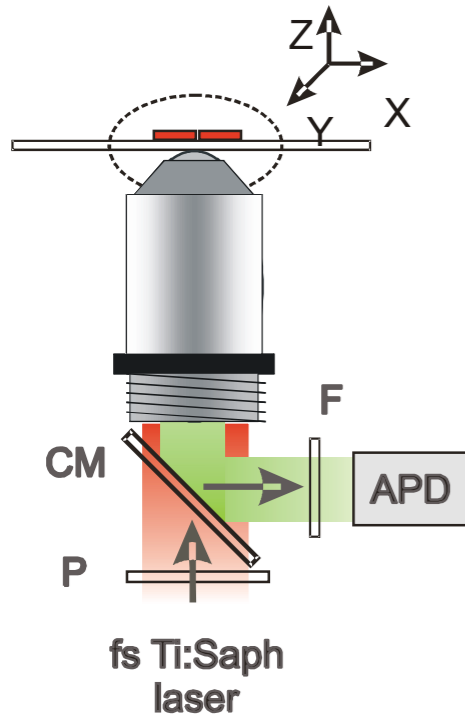


Petru  
Ghenuche

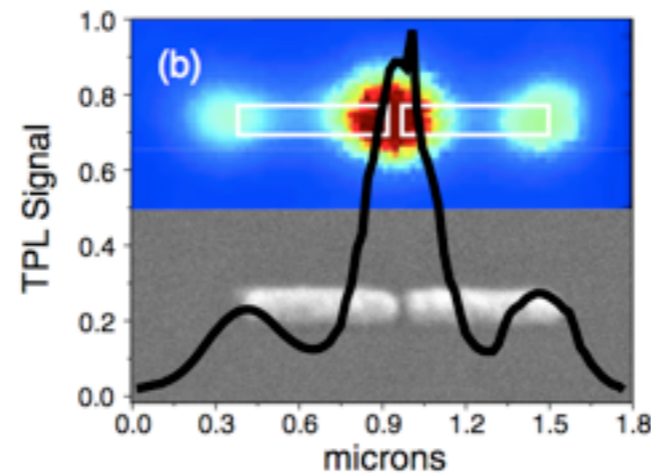


Sudhir  
Cherruku-  
lappurath

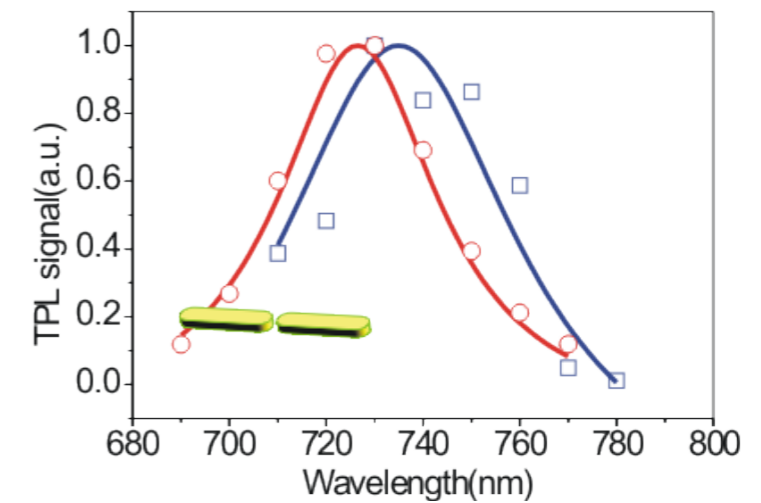
Two photon Luminescence in gold (TPL)



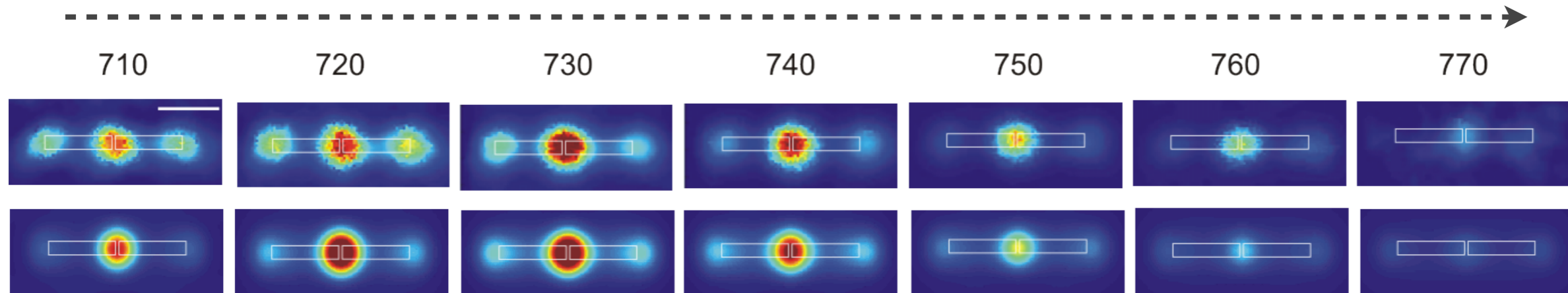
TPL map at resonance



Near field Spectroscopy in the gap



Wavelength (nm)

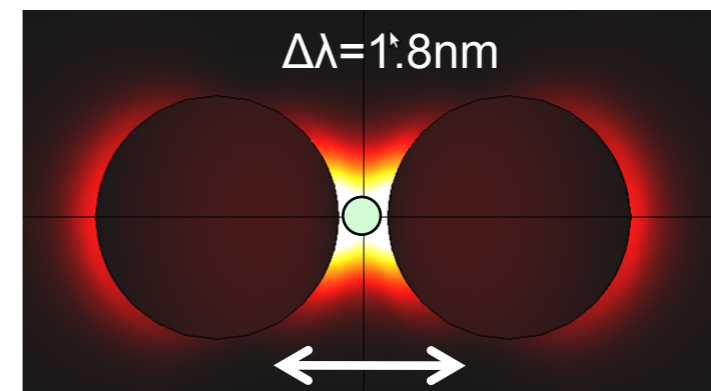
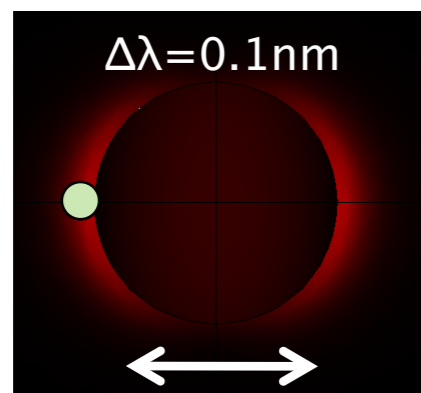


P. Ghenuche et al, Phys. Rev. Lett. **101**, 116805 (2008)



# Shaping plasmonic modes for higher sensitivity

The sensitivity of a plasmonic sensor is mostly governed by spatial overlap between the plasmonic mode and the analyte



● 8nm dielectric sphere (n=1.5)

Low spatial overlap



small resonance shift

high spatial overlap



increased resonance shift

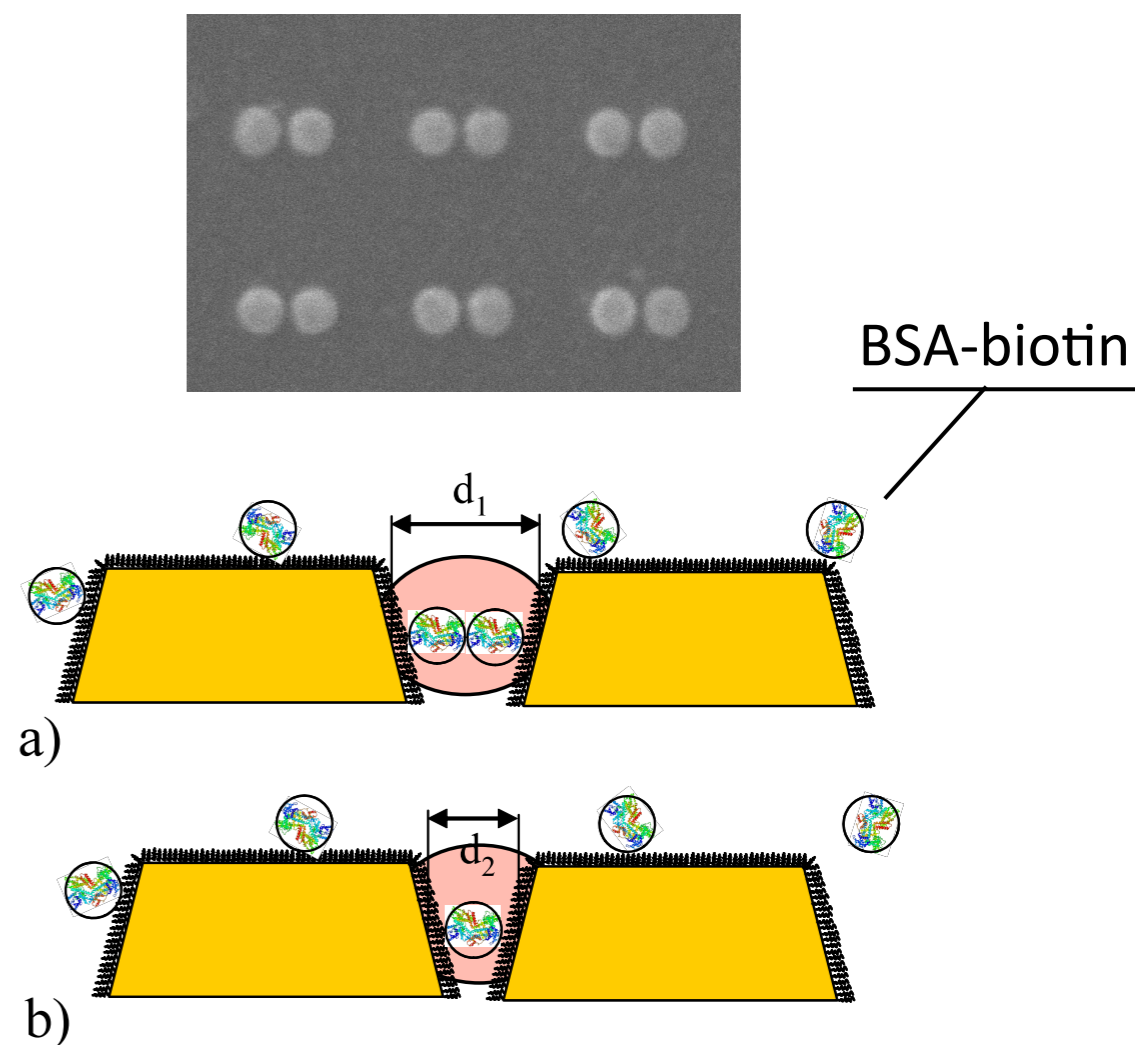
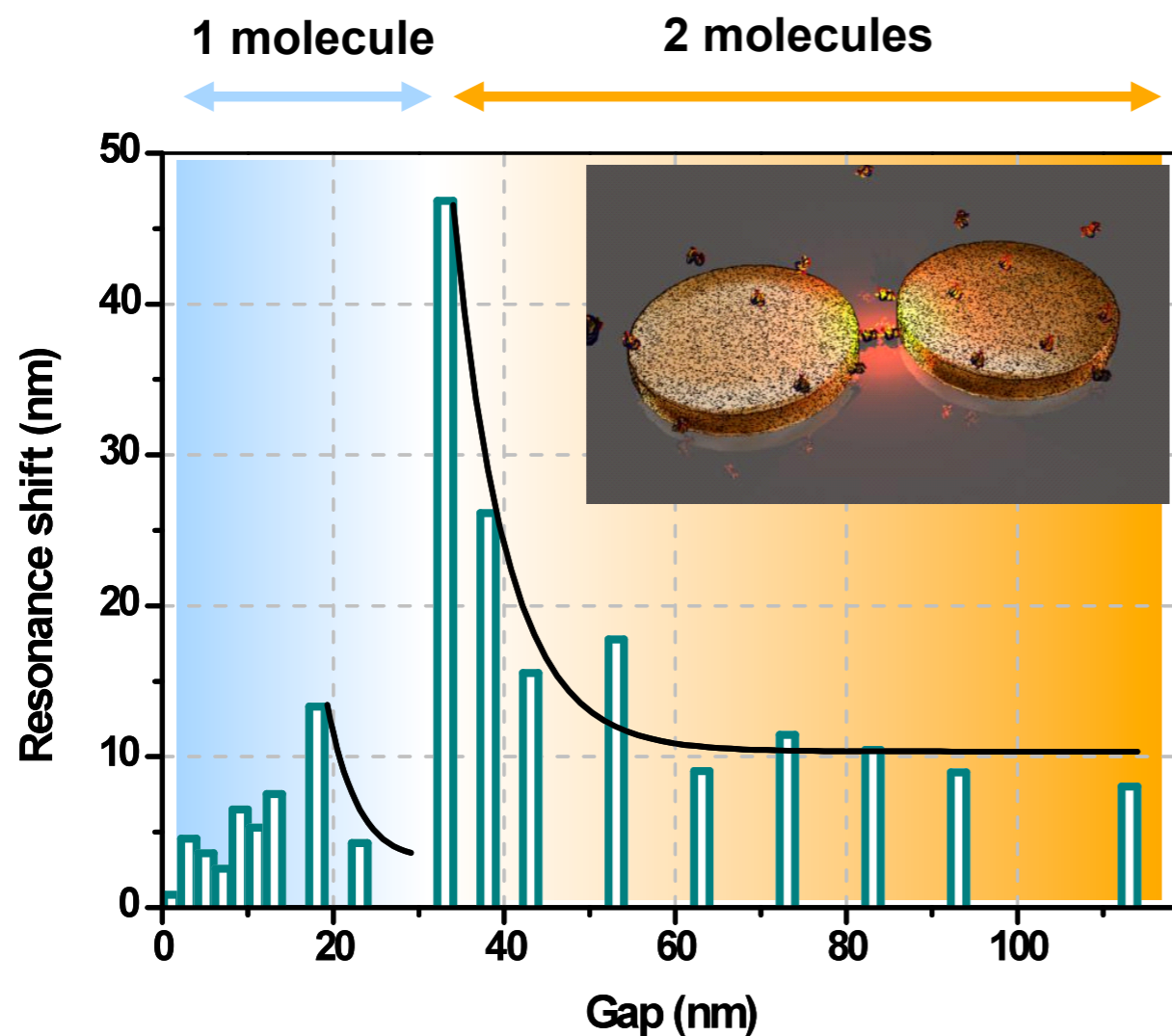
# Enhanced molecular detection in plasmonic dimers



Srdjan  
Acimovic



Mark  
Kreuzer



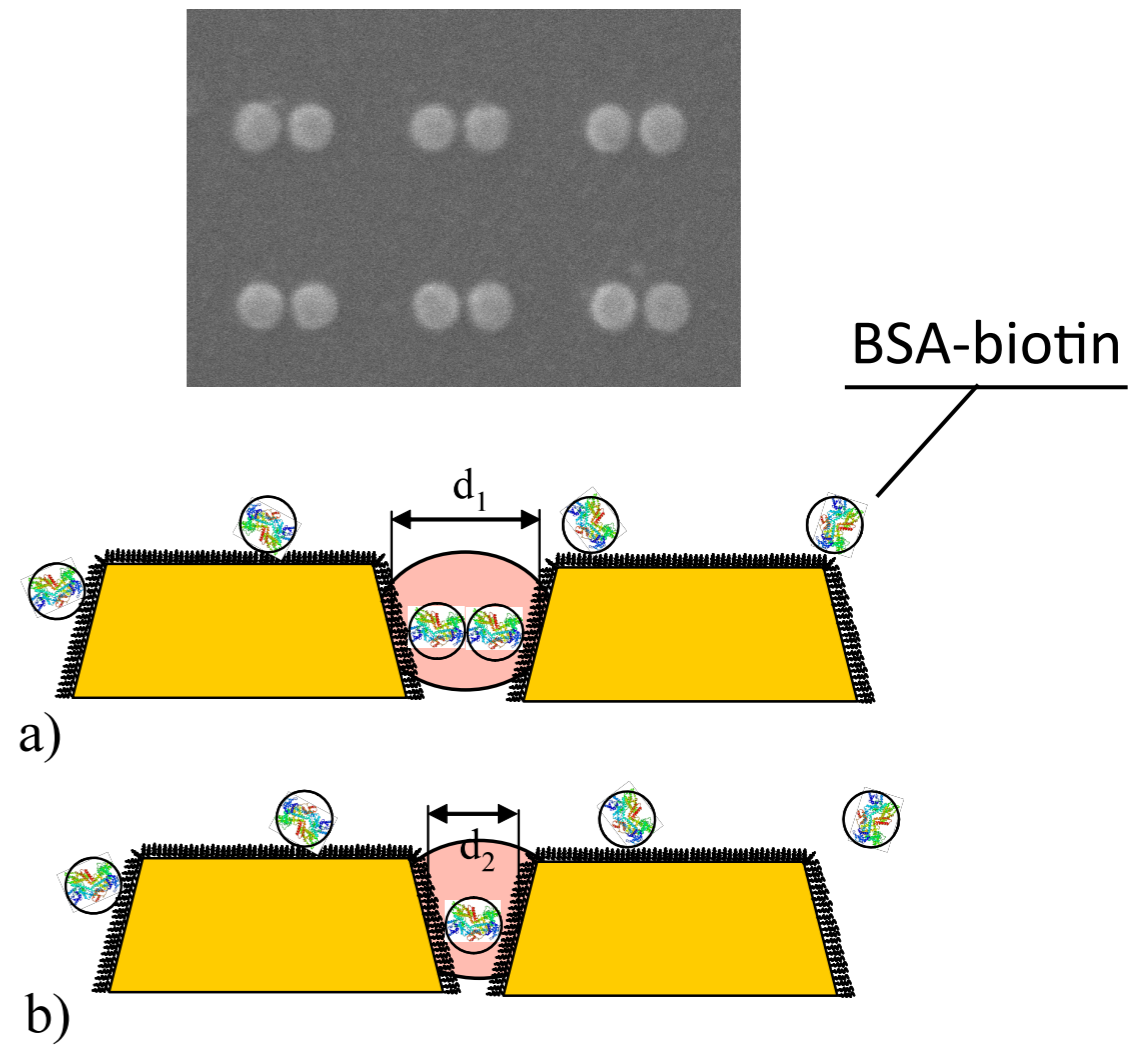
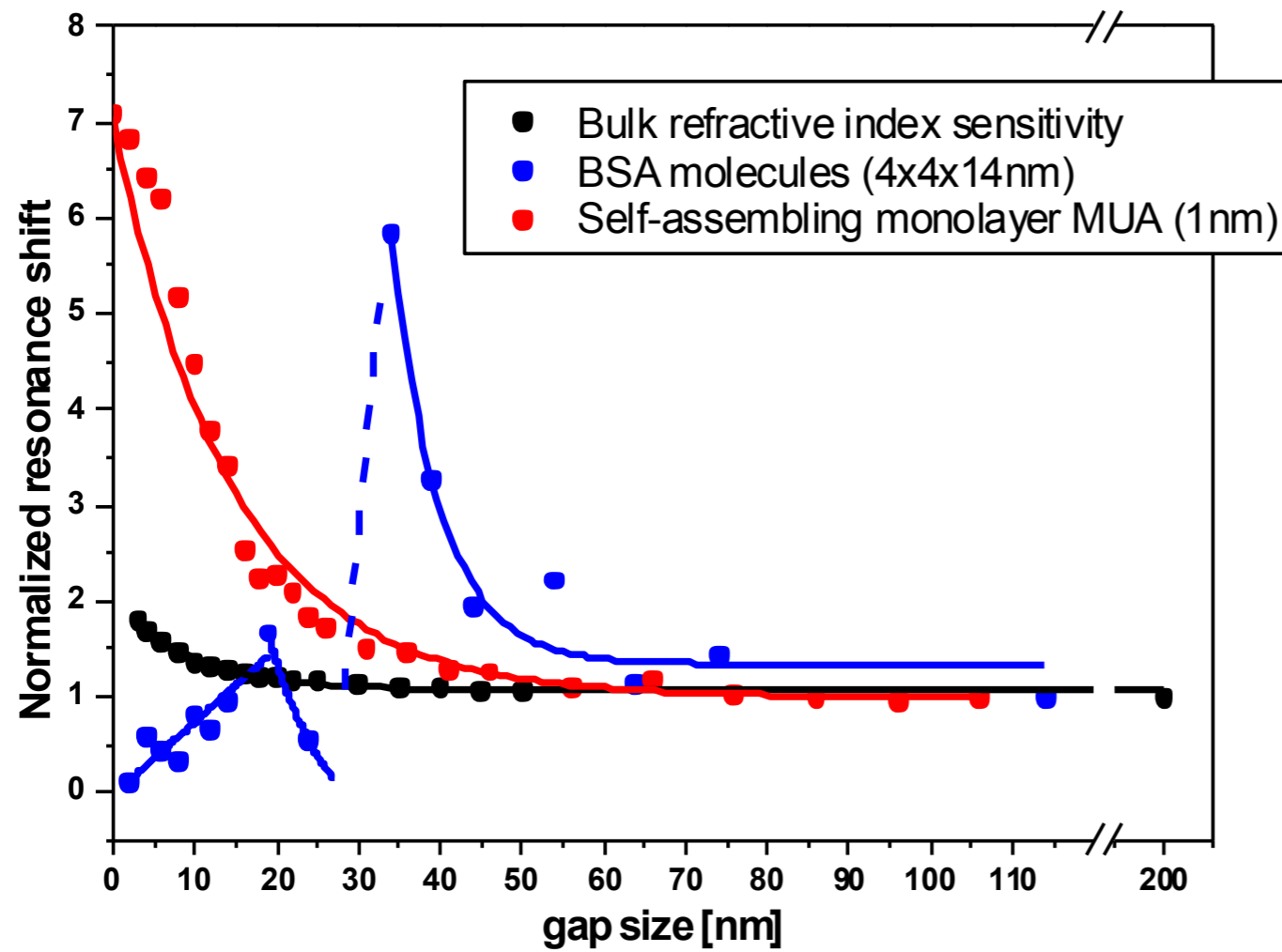
# Influence of the size of the target molecule



Srdjan  
Acimovic



Mark  
Kreuzer





# A short Lab tour



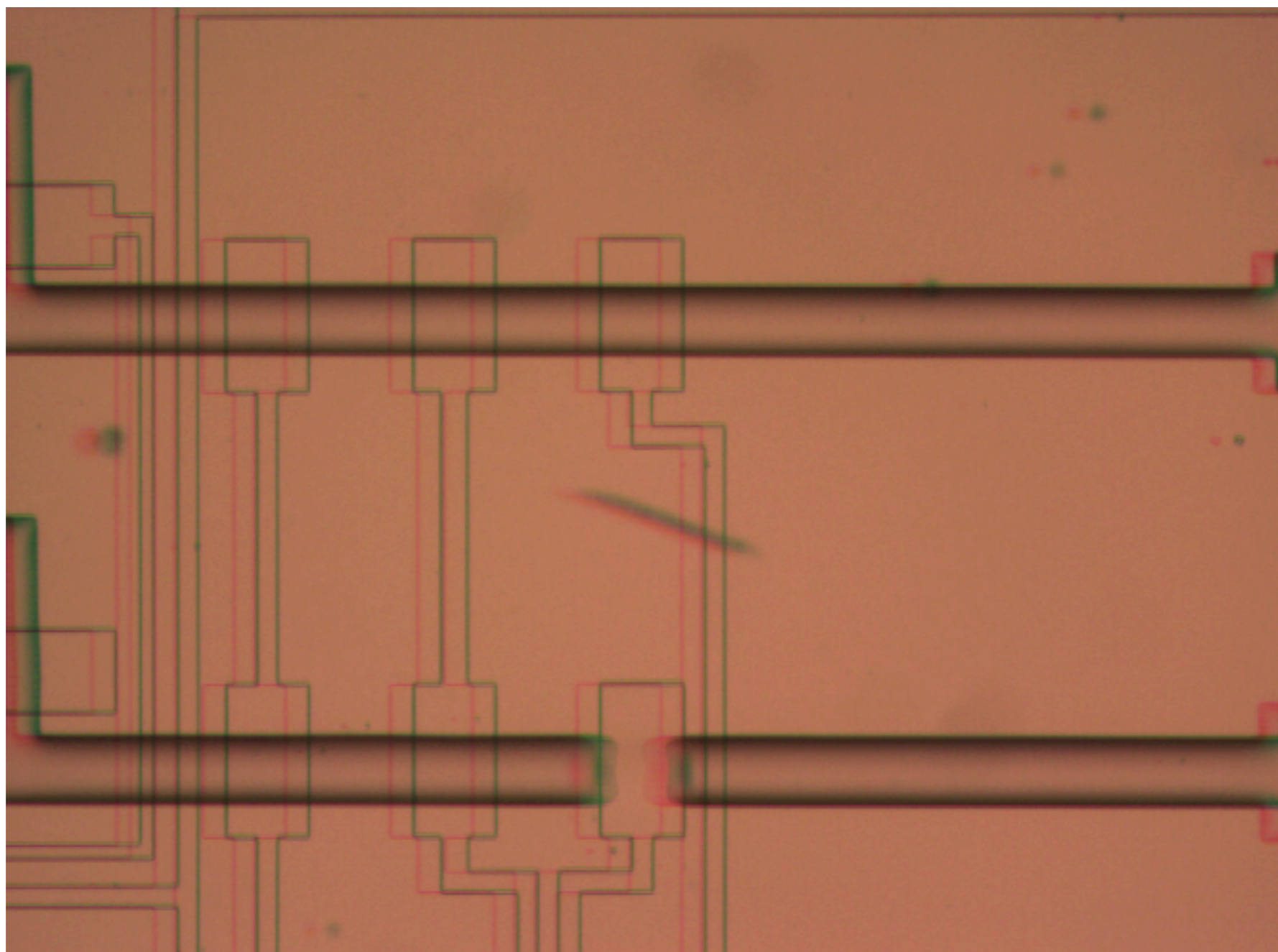
Srdjan  
Acimovic



Mark  
Kreuzer



Maria Ale  
Ortega





# Real time monitoring of LSP shift



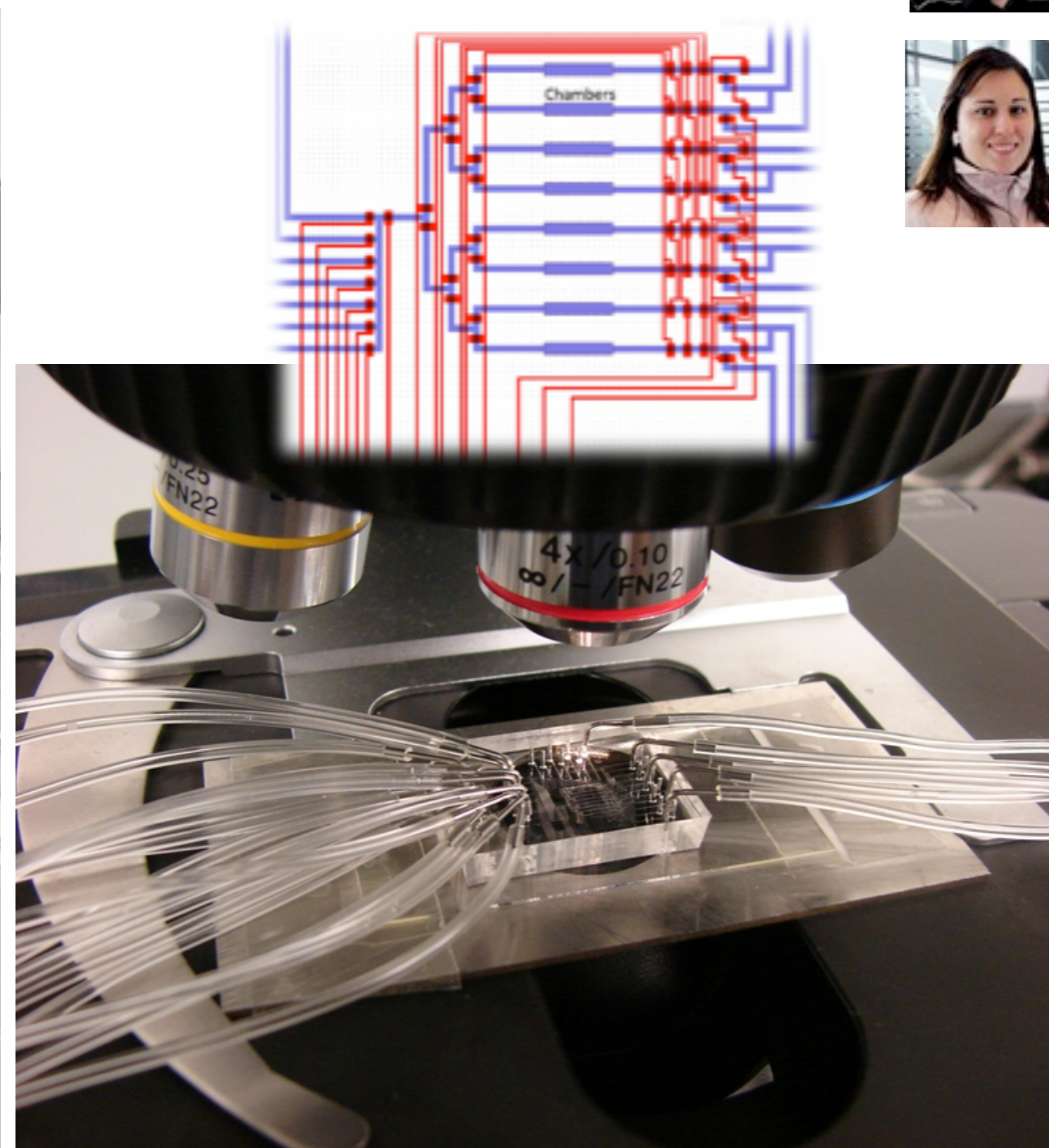
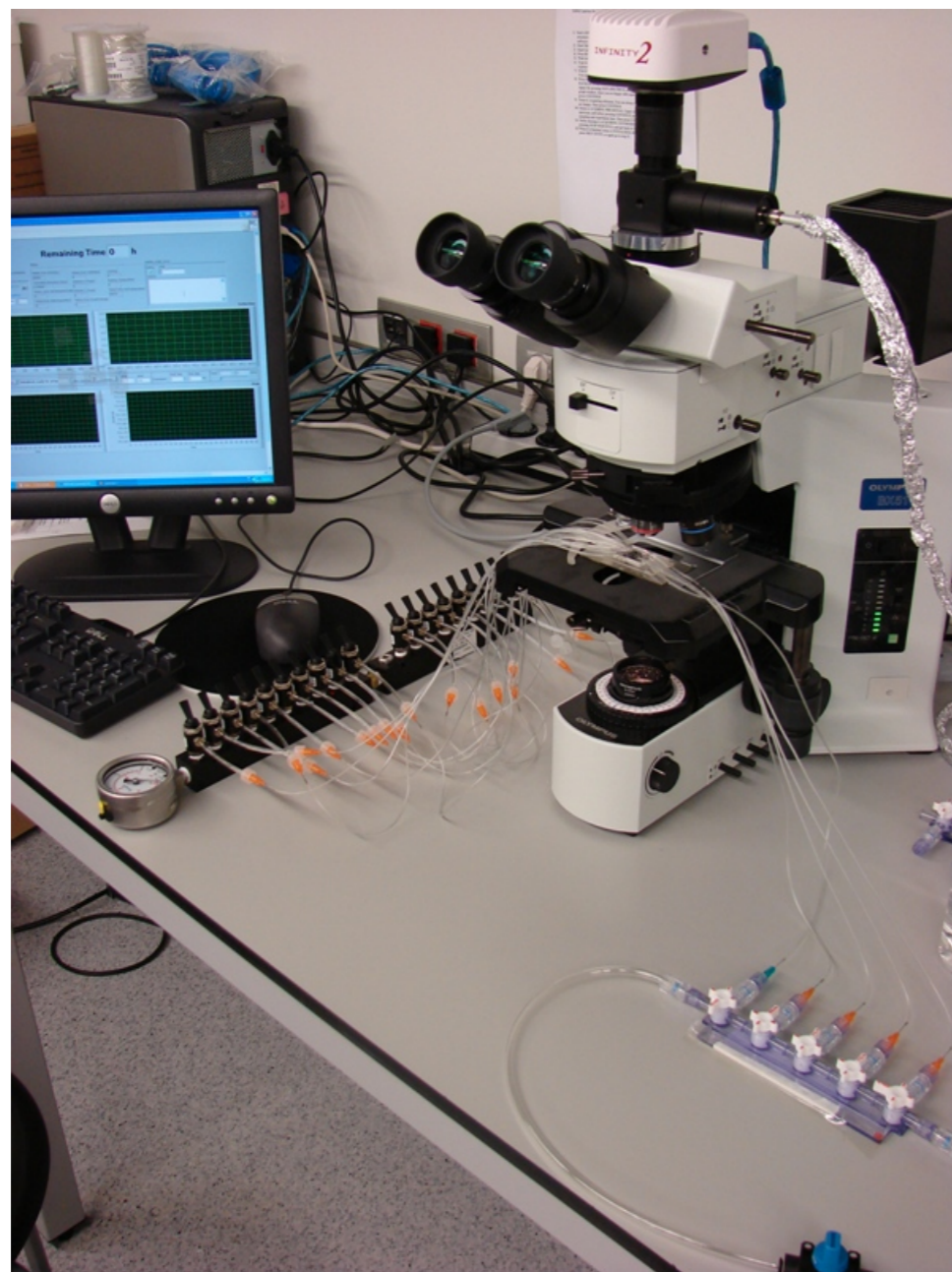
Srdjan Acimovic



Mark Kreuzer



Maria Ale Ortega





# Real time monitoring of LSP shift



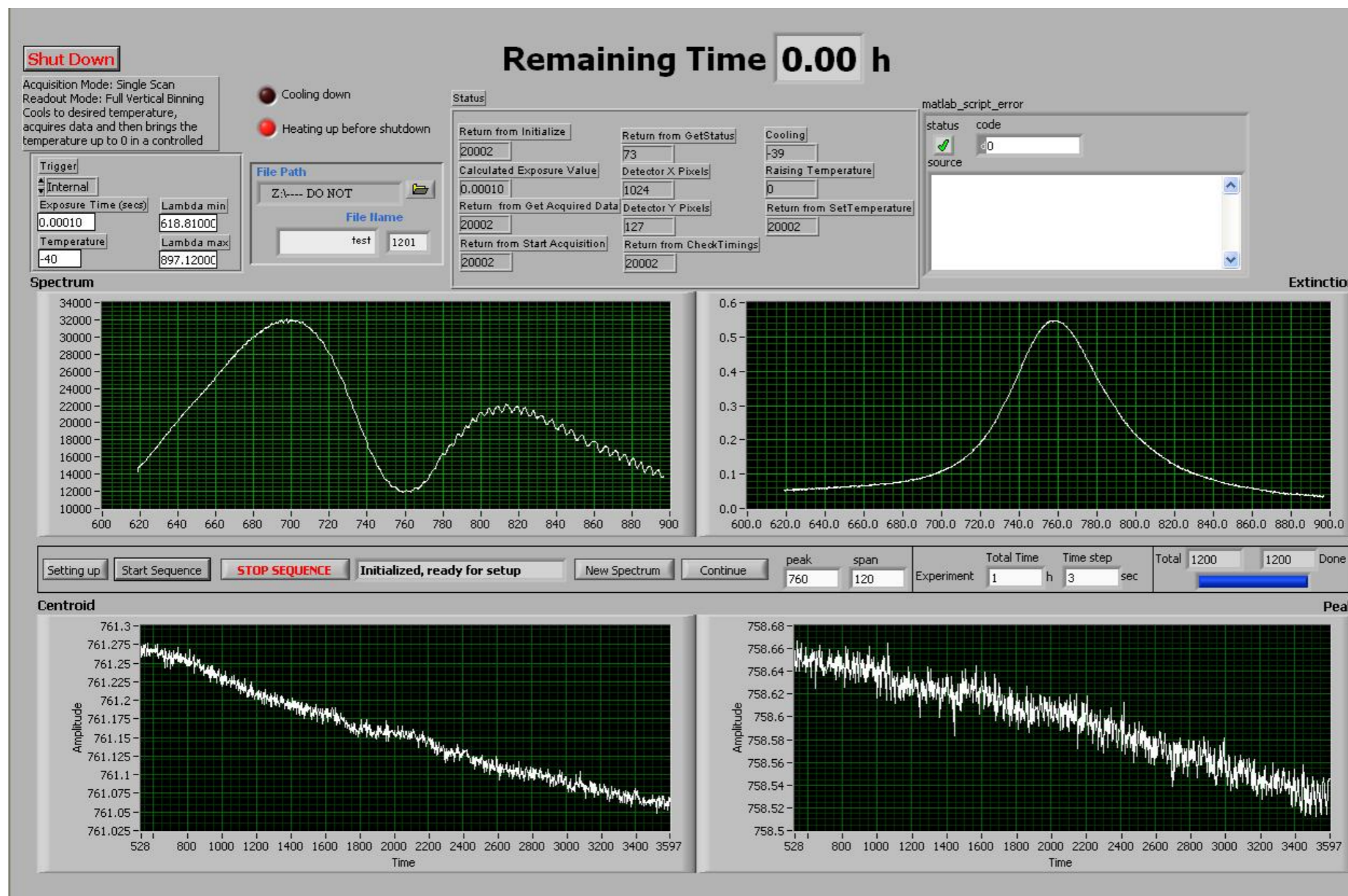
Srdjan Acimovic



Mark Kreuzer



Maria Ale Ortega



# Bulk refractive index sensing



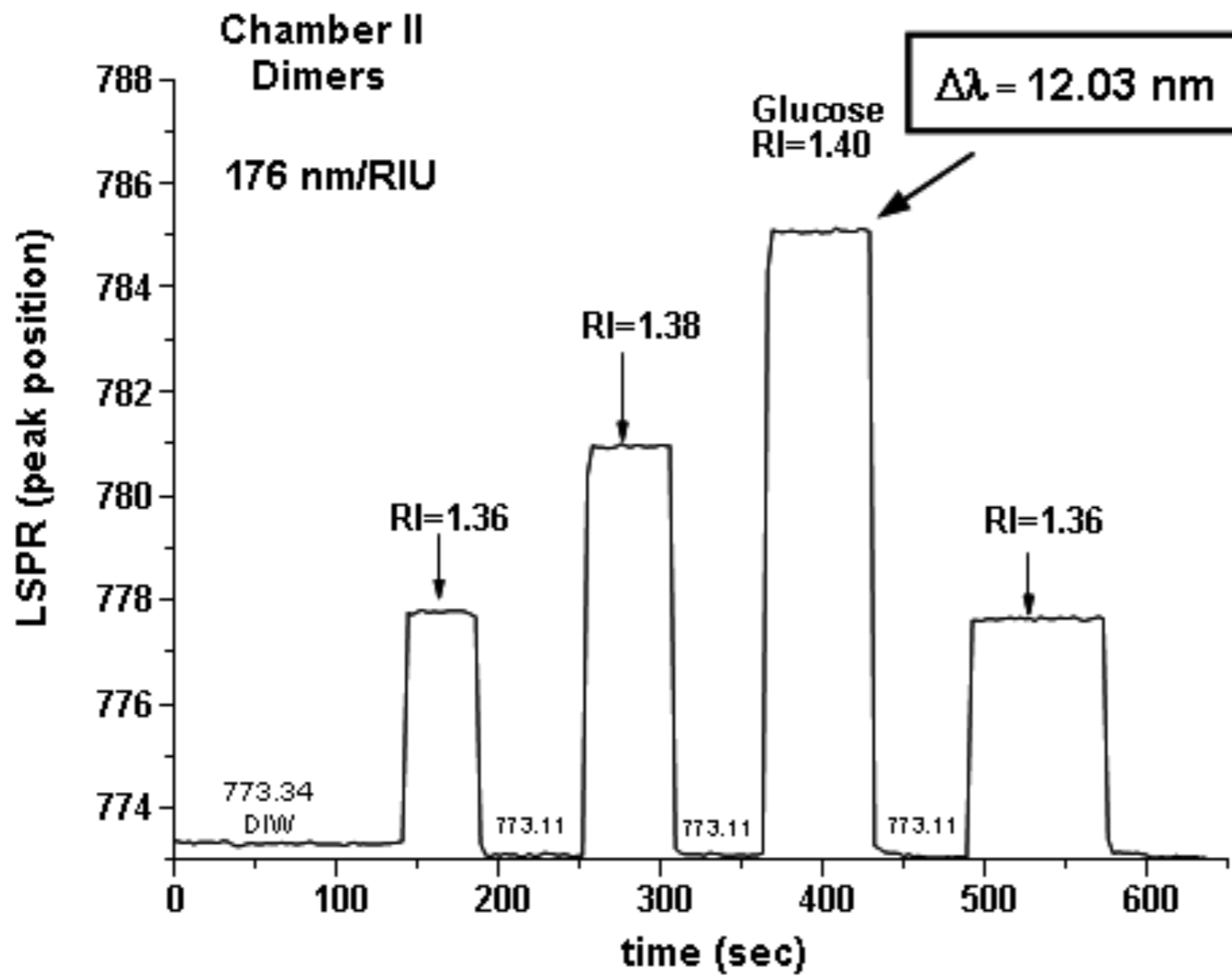
Srdjan Acimovic



Mark Kreuzer



Maria Ale Ortega



# IgG detection in a microfluidic chip



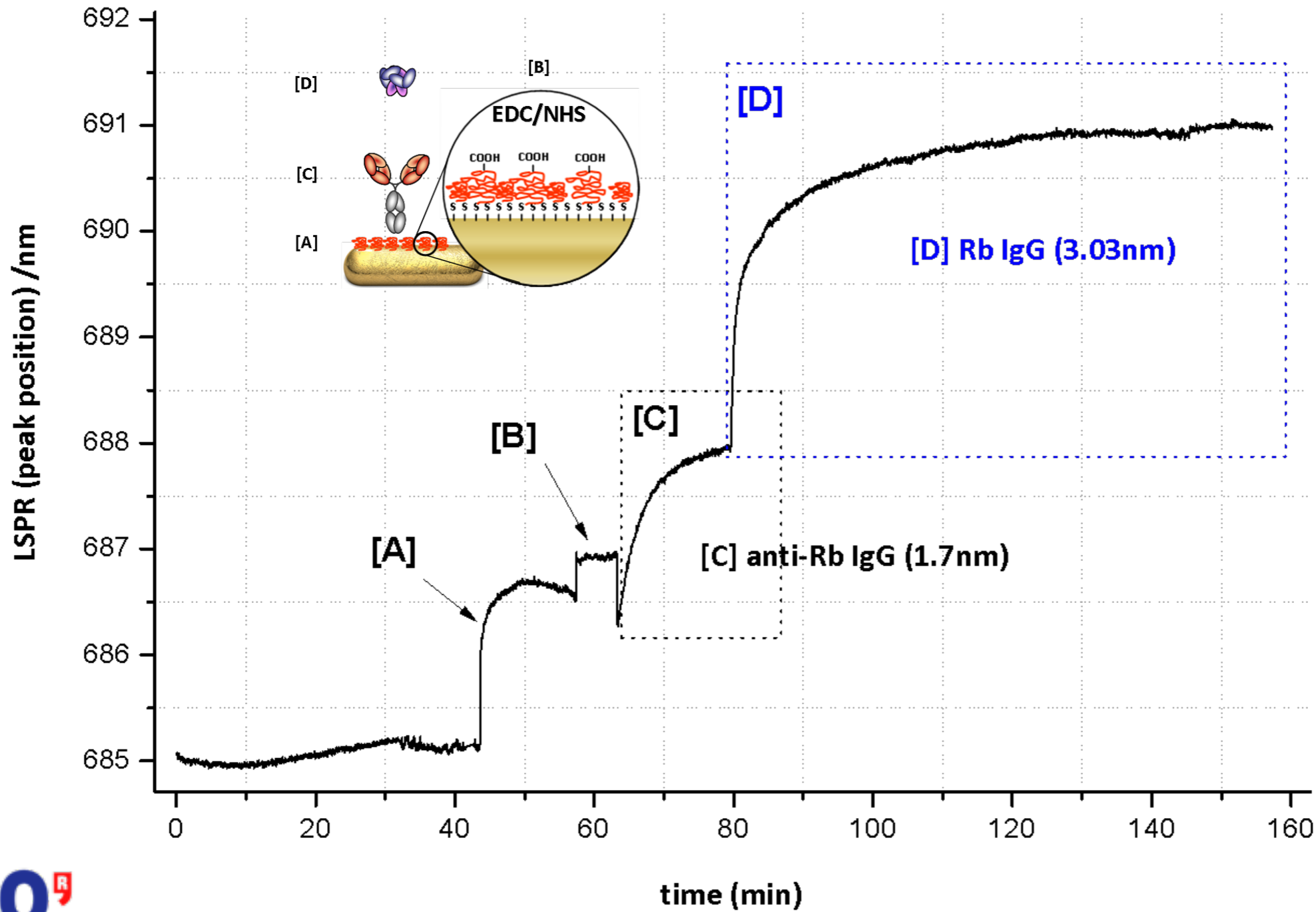
Srdjan Acimovic



Mark Kreuzer



Maria Ale Ortega



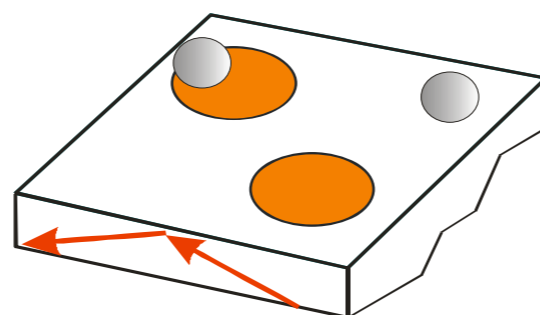
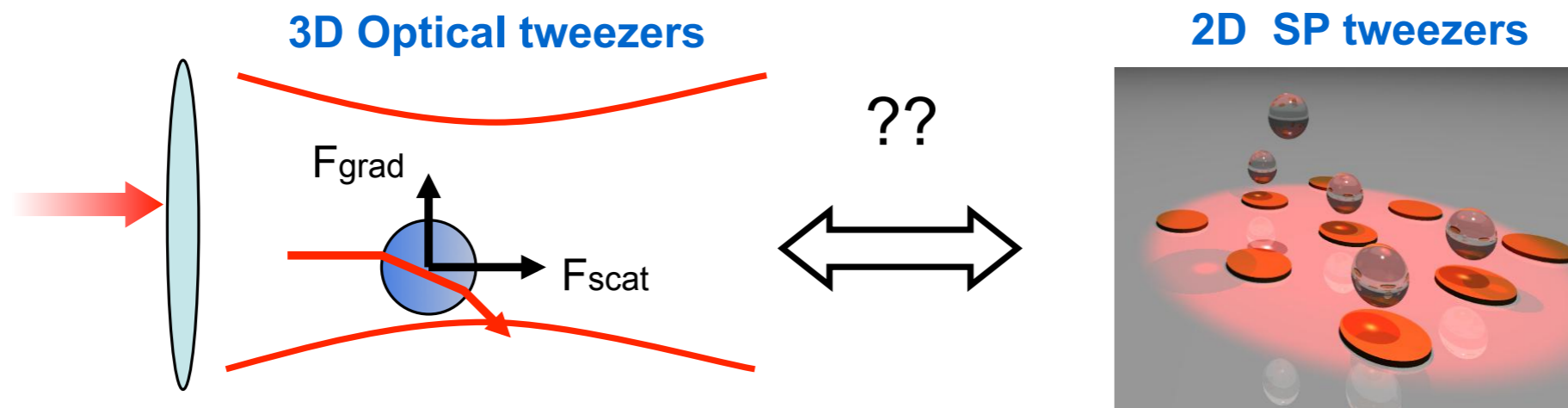


# Surface Plasmon-based trapping

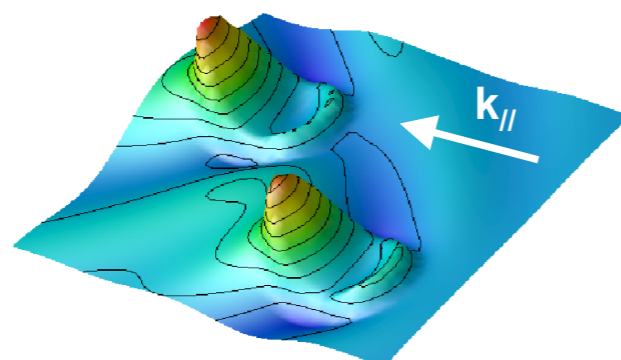


Maurizio Righini

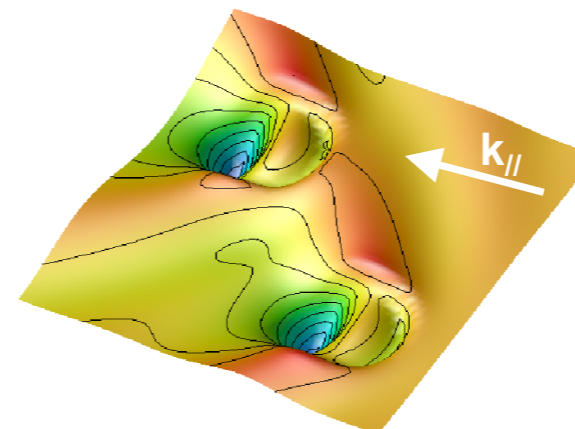
Now@Berkeley



ELECTRIC NEAR-FIELD INTENSITY



NEAR-FIELD OPTICAL POTENTIAL



Quidant et al, Optics Letters 30, 1009-1011 (2005)

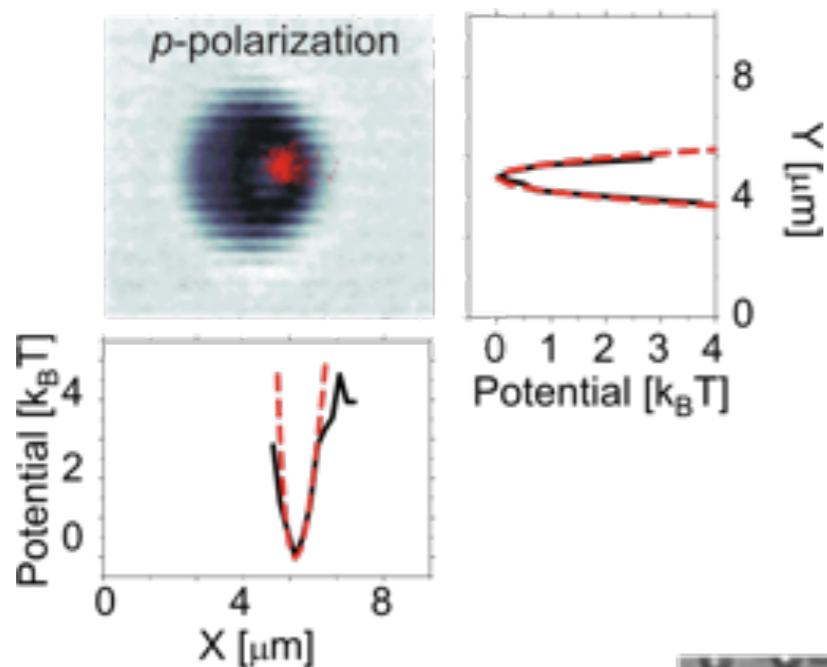
# Parallel trapping of micro-object with SP traps



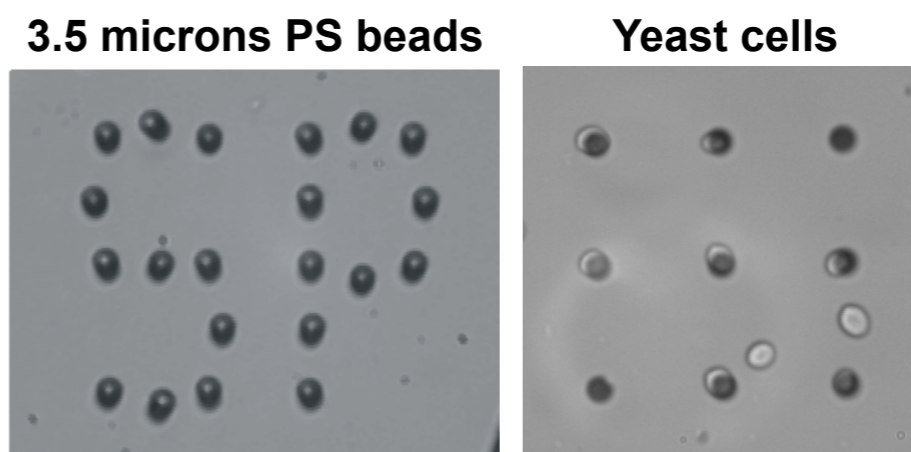
Maurizio Righini

Now@Berkeley

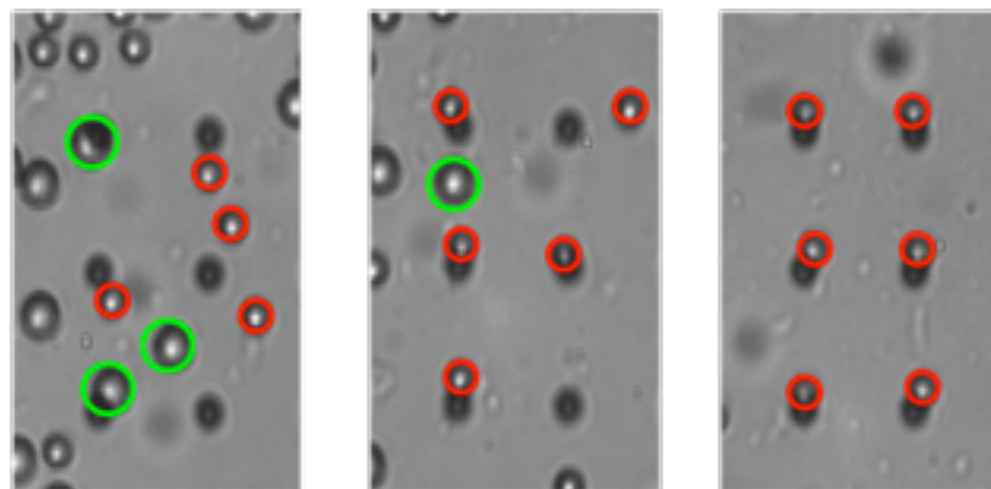
## SP TRAP FEATURES



## PARALLEL TRAPPING



## SELECTIVE SP TRAPPING



M. Righini et al, Nature Physics **3**, 477 (2007)

M. Righini et al, Phys. Rev. Lett **100**, 183604 (2008)

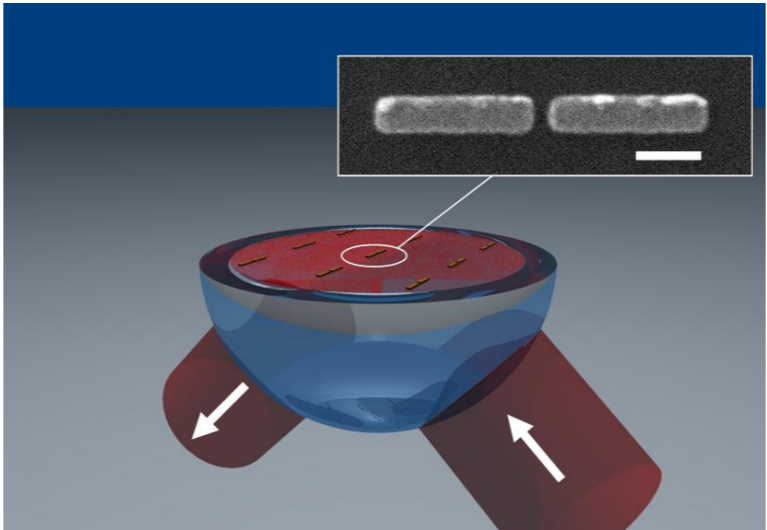


# Trapping 200 nm PS beads with optical gap antennas

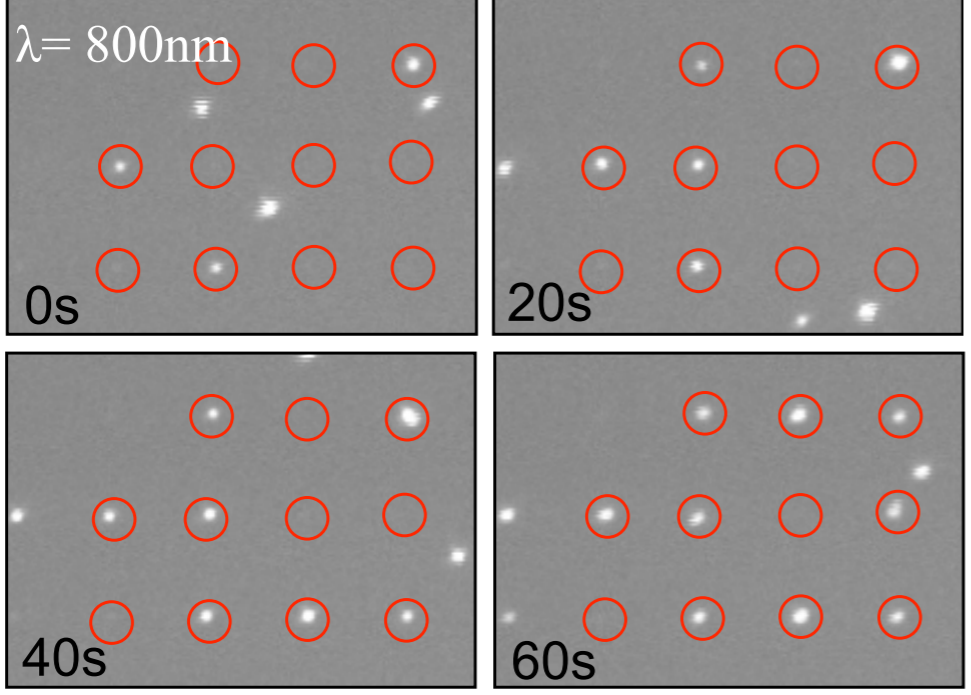
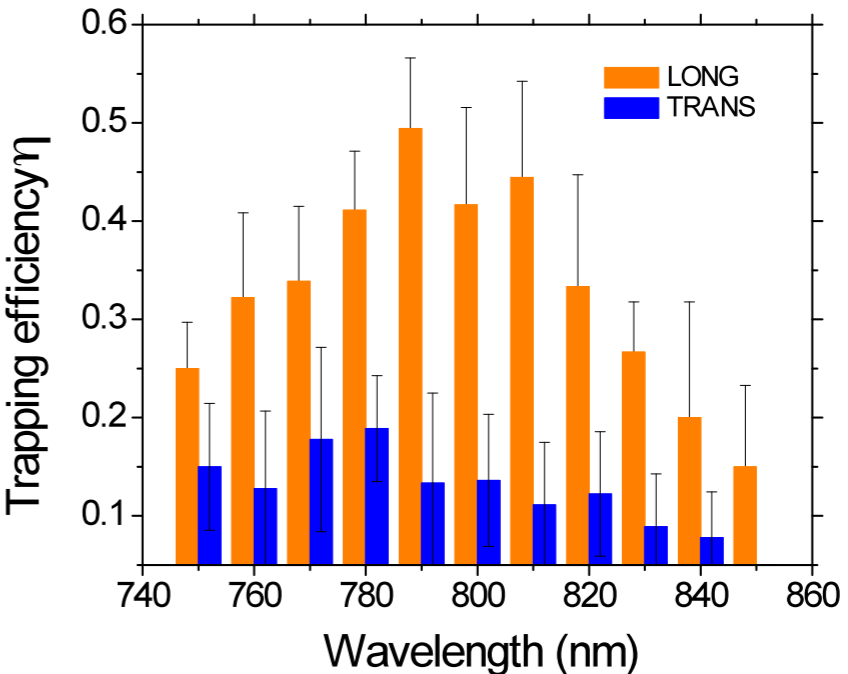


Maurizio Righini

Now@Berkeley



Trapping efficiency vs wavelength



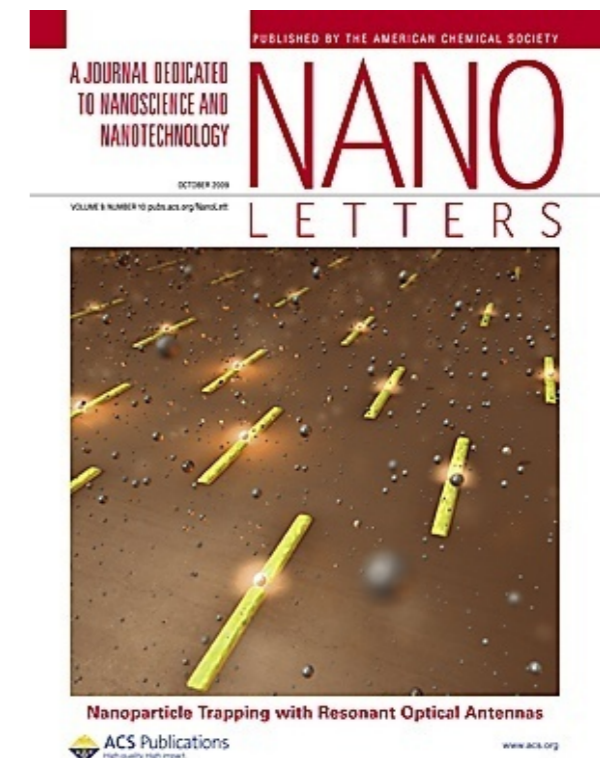
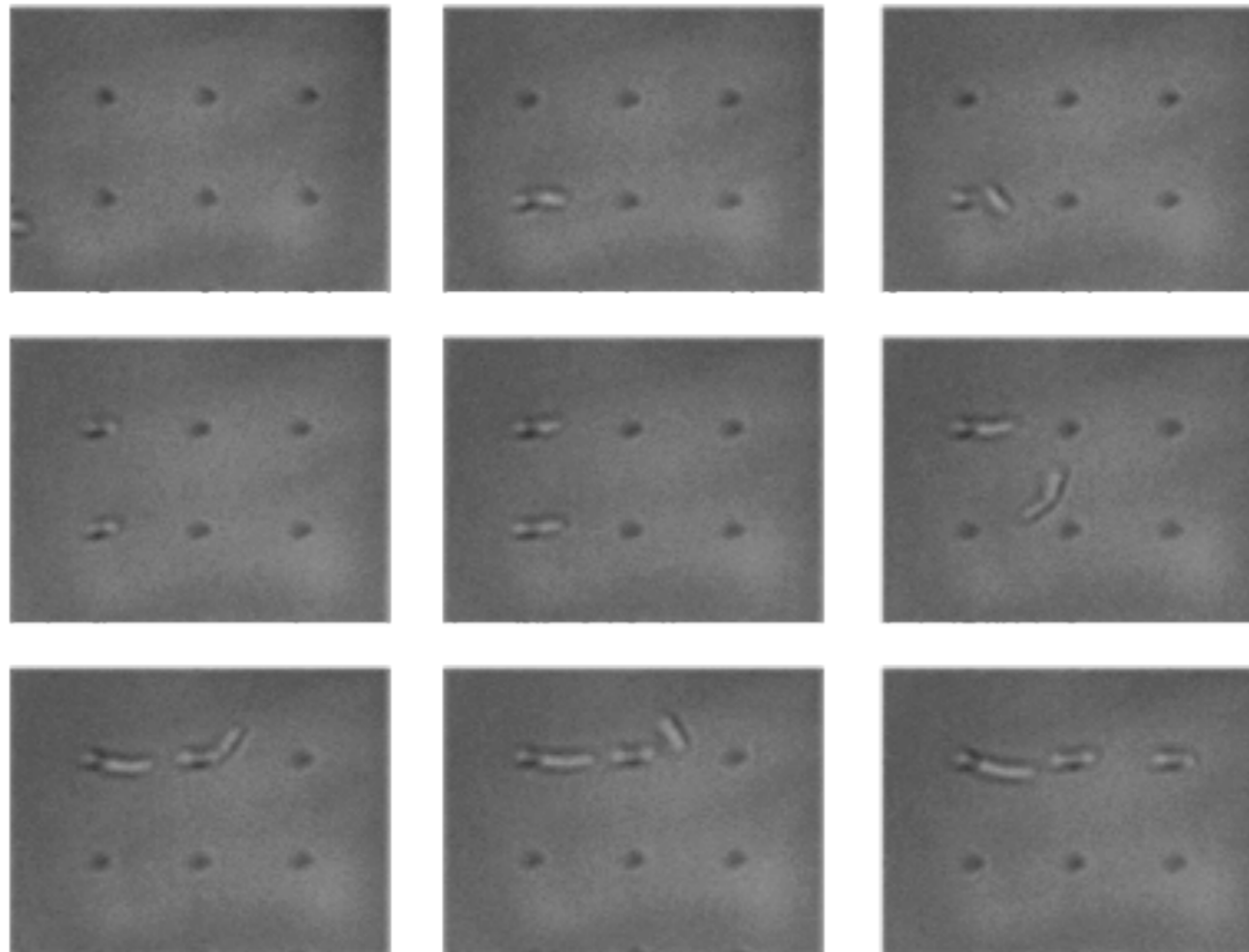
M. Righini et al, NanoLetters **9**, 3387–3391 (2009)

# Trapping and alignment of E-coli bacteria



Maurizio Righini

Now@Berkeley



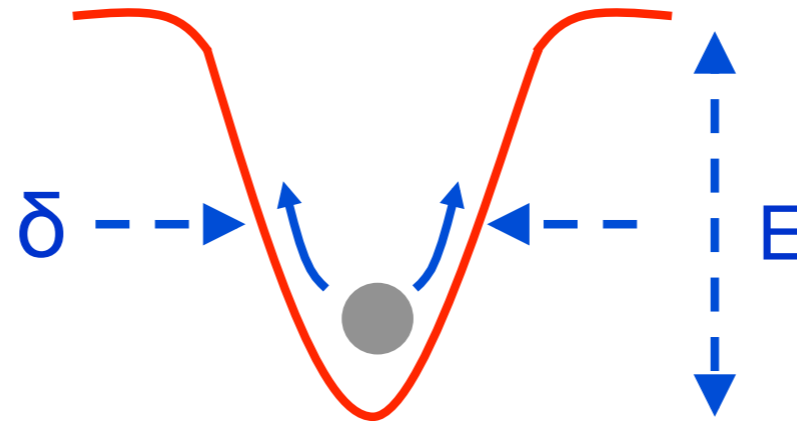
- ◆ Division time ~ 60 min both for the trapped and untrapped bacteria
- ◆ Estimated local intensity experienced by the trapped bacteria is 10 times below the damage threshold  $10^{10} \text{W.m}^{-2}$

# Where is the limit?

When downsizing the size of the trapped specimen ...

↘ Optical gradient force ( $R^3$ )

$$\mathbf{F}_{\text{gradient}} = \frac{2\pi n_m R^3}{c} \left( \frac{m^2 - 1}{m^2 + 2} \right) \nabla I$$



↗ Brownian motion ( $1/R$ )

$$\langle x^2(t) \rangle = \frac{2k_B T}{6\pi\eta R} |t|$$

Increasing the trapping efficiency requires ...

Increasing the trap  
Confinement ( $\downarrow \delta$ )

=

Concentrating light  
Fields down to the sub- $\lambda$  scale

&

Increasing the  
Potential depth ( $\uparrow E$ )

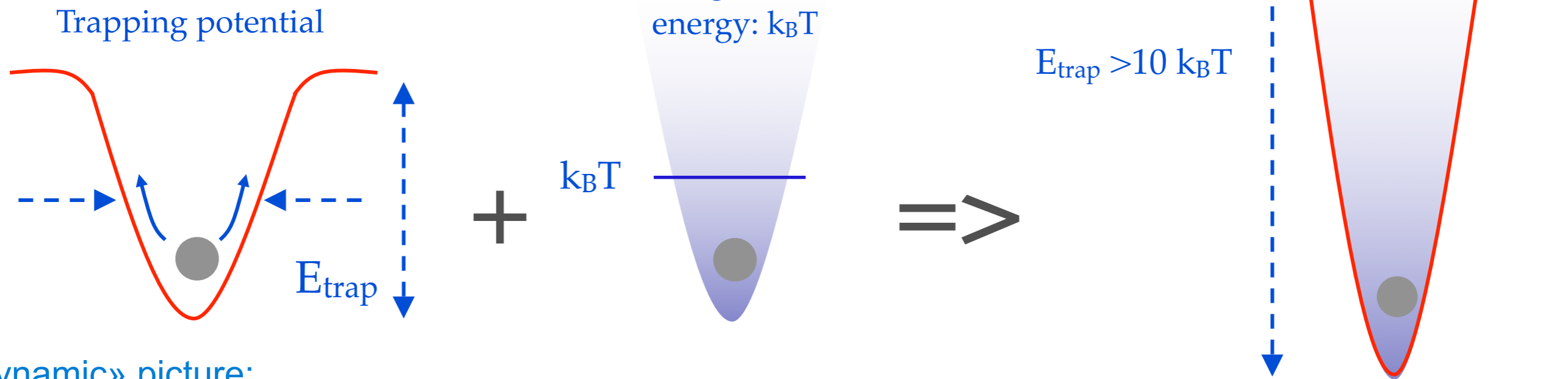
=

Increasing the local  
Field intensity within the trap

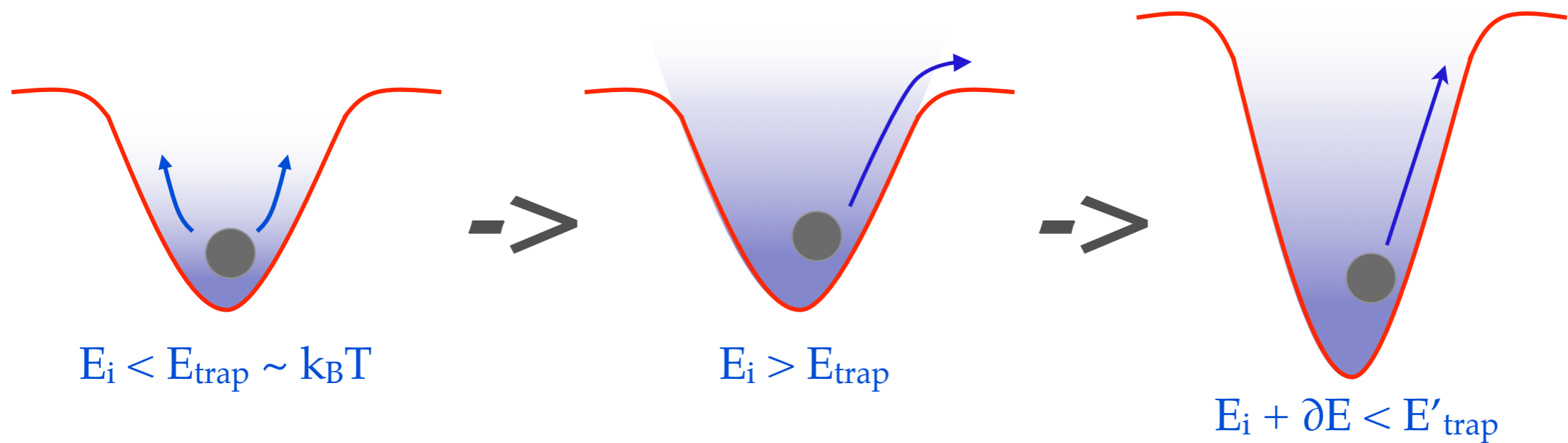
Risks of photo-damage

# Relaxing the trapping conditions using a dynamic trap

«Static» picture:



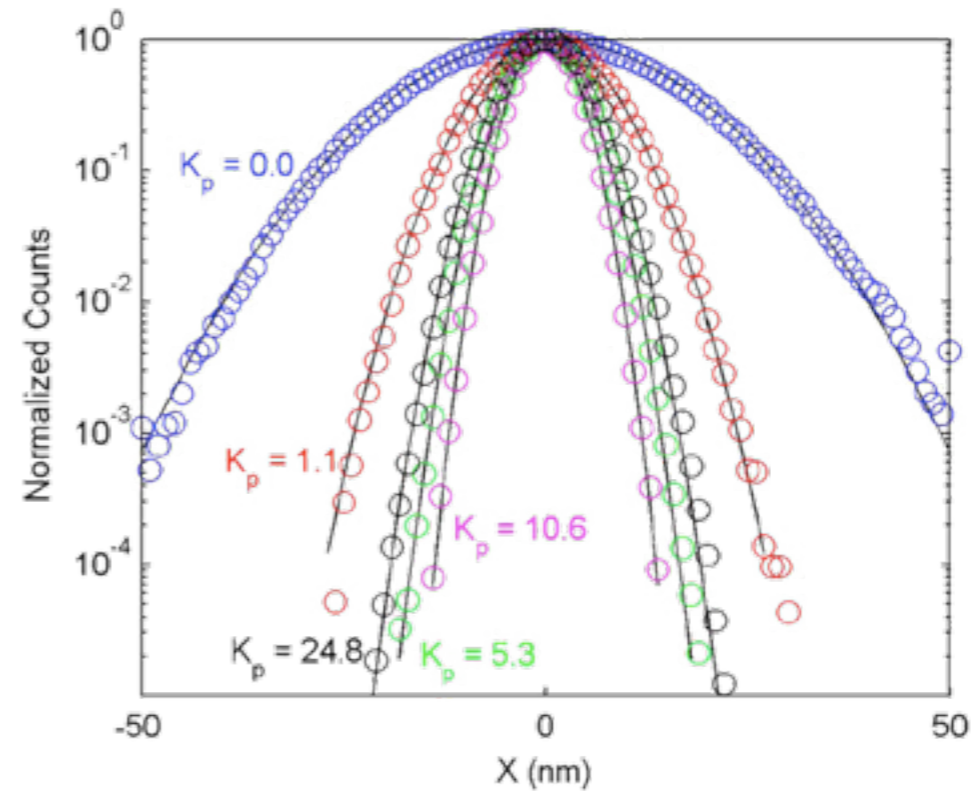
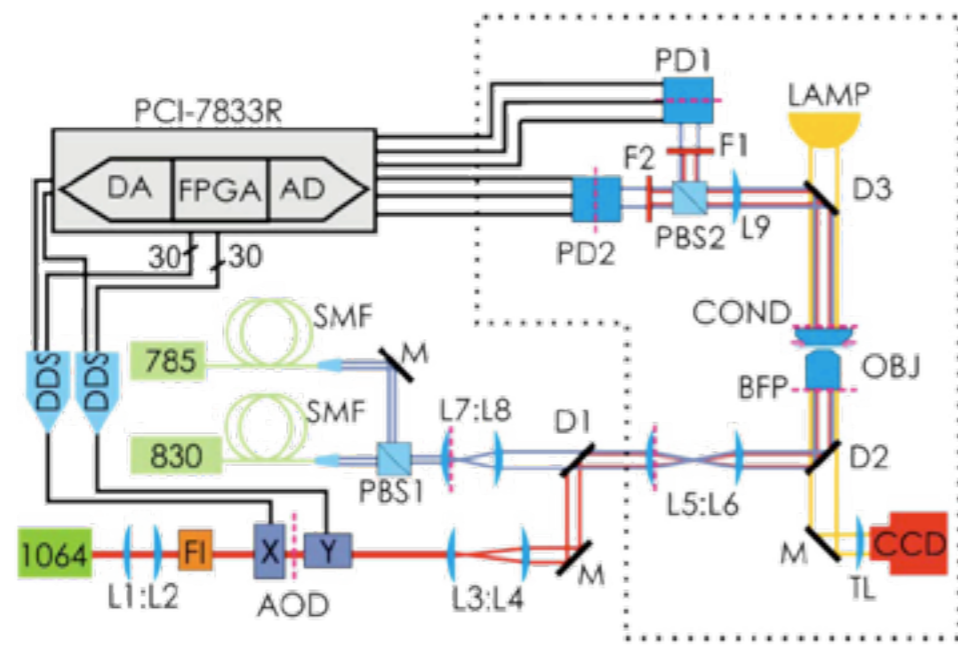
«Dynamic» picture:



# Relaxing the trapping conditions using a dynamic trap

«Static» picture:

Average kinetic



Wallin, A. E., et al, Stiffer optical tweezers through real-time feedback control. *Appl. Phys. Lett.* **92**, 224104 (2008).

$$E_i < E_{\text{trap}} \sim k_B T$$

$$E_i > E_{\text{trap}}$$

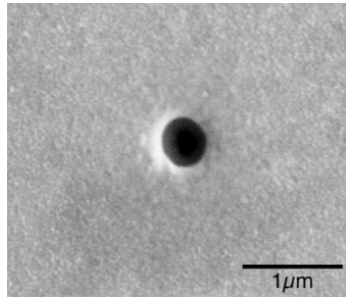
$$E_i + \partial E < E'_{\text{trap}}$$



# Self-Induced Back-Action (SIBA) Trapping in metallic nanoapertures

## Single nano-aperture:

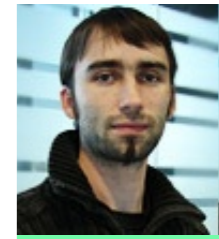
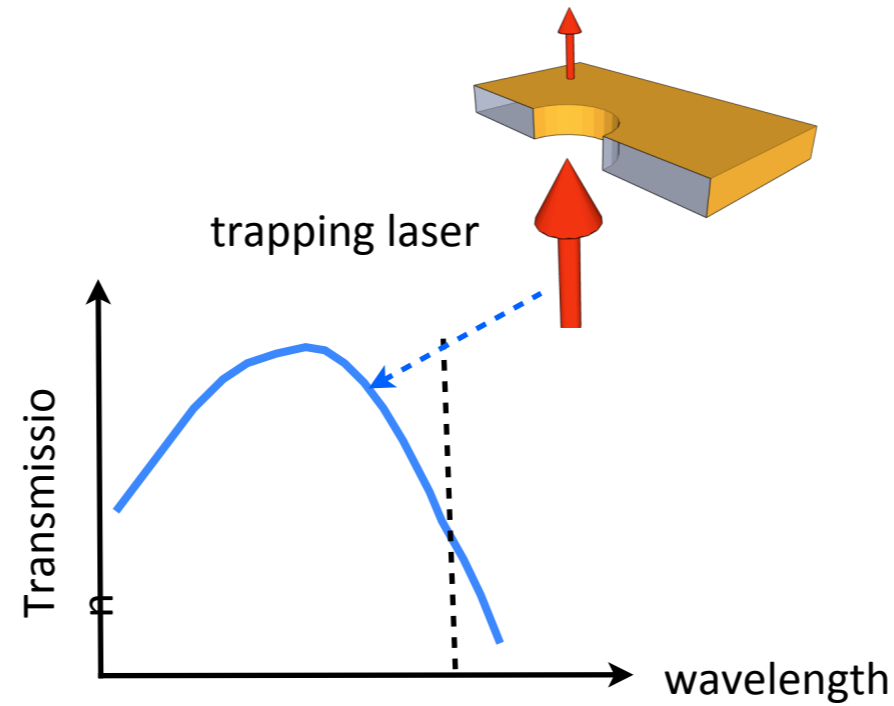
310 nm diameter, 100 nm thick gold layer



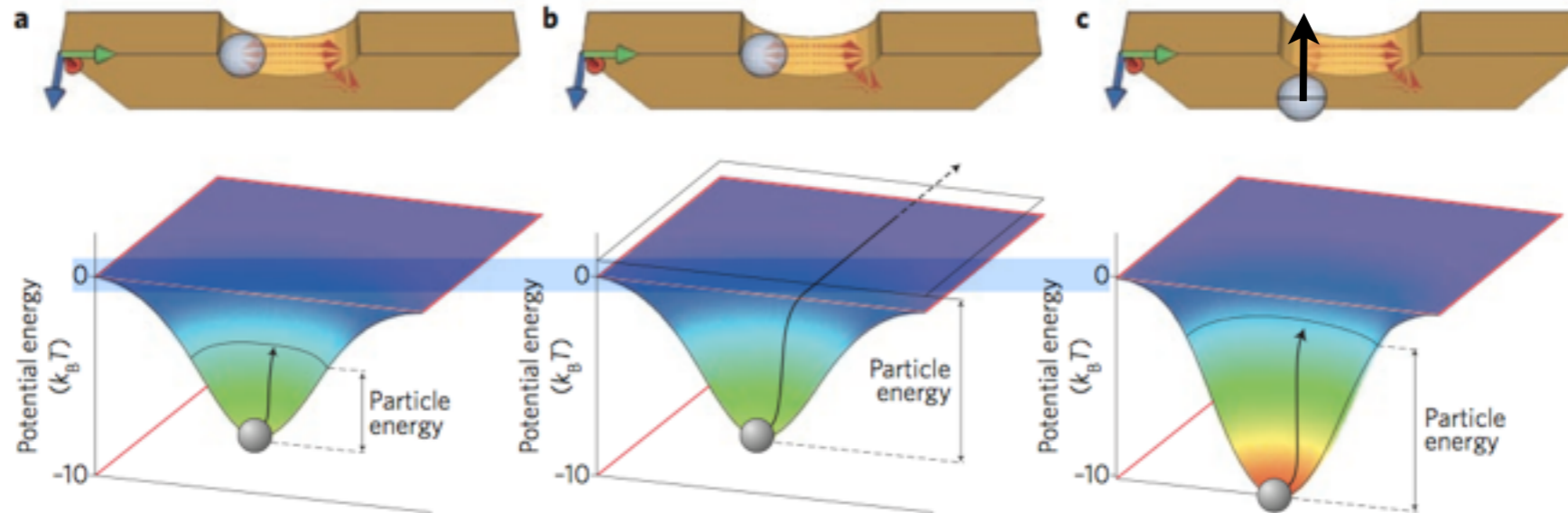
Trapping with a nano-hole:

Okamoto et al (1999)

Kwak et al (2004)



Mathieu  
Juan  
now @Sydney

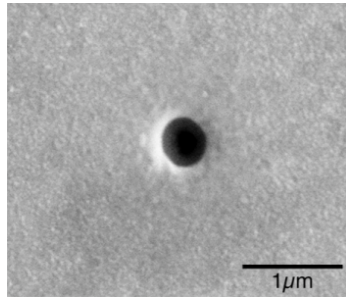


In the SIBA approach, trapping not only relies on the gradient force but is assisted by the change in the momentum of transmitted photons

# Self-Induced Back-Action (SIBA) Trapping in metallic nanoapertures

## Single nano-aperture:

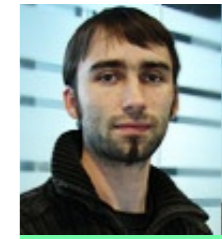
310 nm diameter, 100 nm thick gold layer



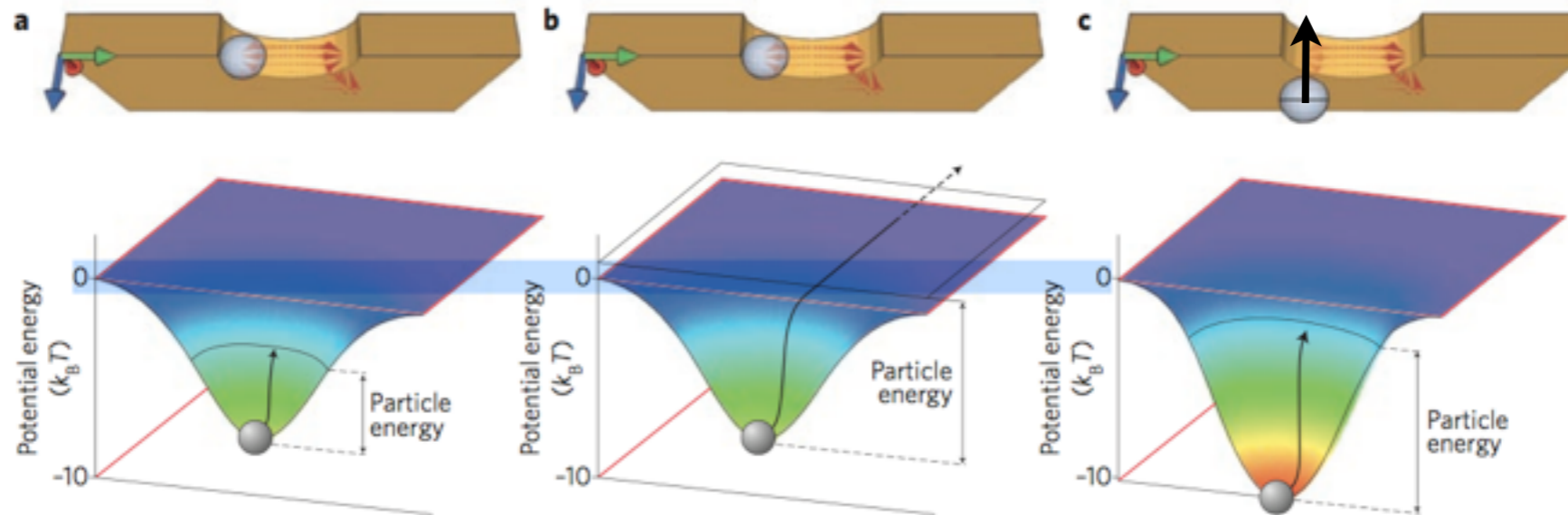
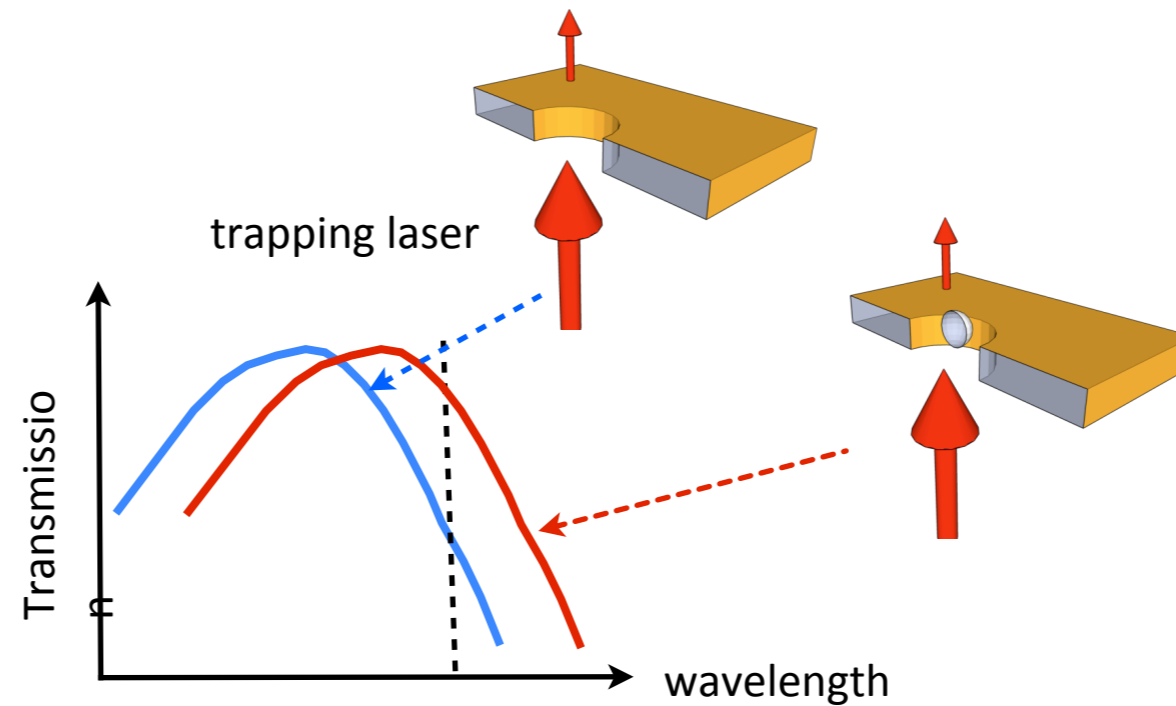
Trapping with a nano-hole:

Okamoto et al (1999)

Kwak et al (2004)



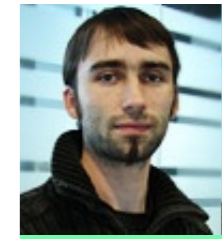
Mathieu  
Juan  
now @Sydney



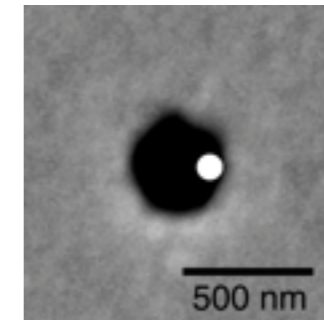
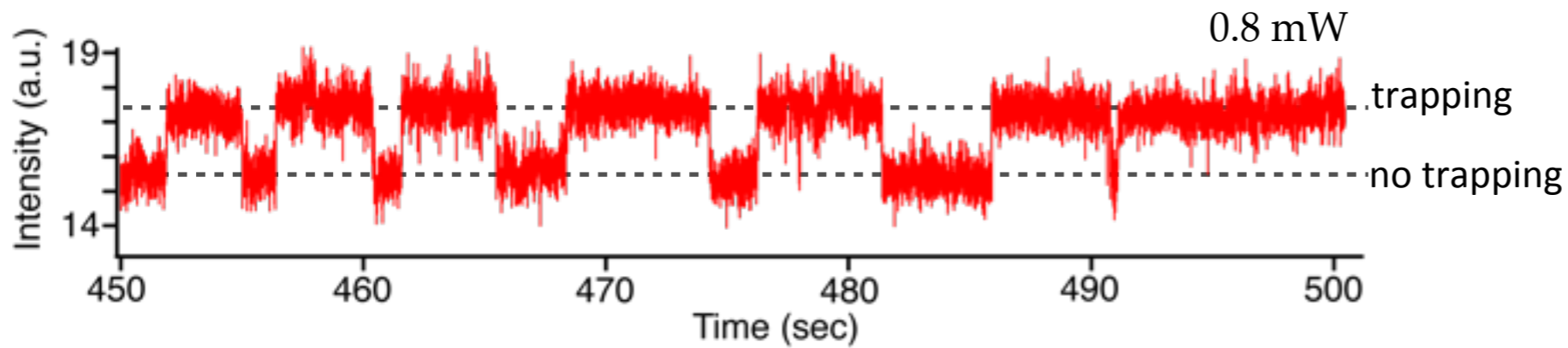
In the SIBA approach, trapping not only relies on the gradient force but is assisted by the change in the momentum of transmitted photons

# SIBA Trapping of 100nm and 50 nm PS beads in spherical nanoapertures

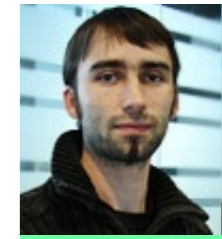
Trapping of 100 nm PS beads:  
0.8 mW for ~ 4 sec average trapping time



Mathieu  
Juan  
now @Sydney

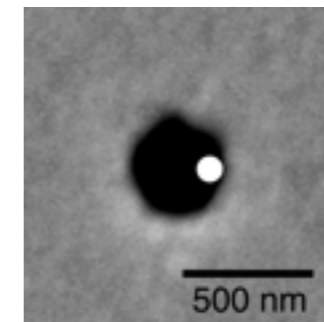
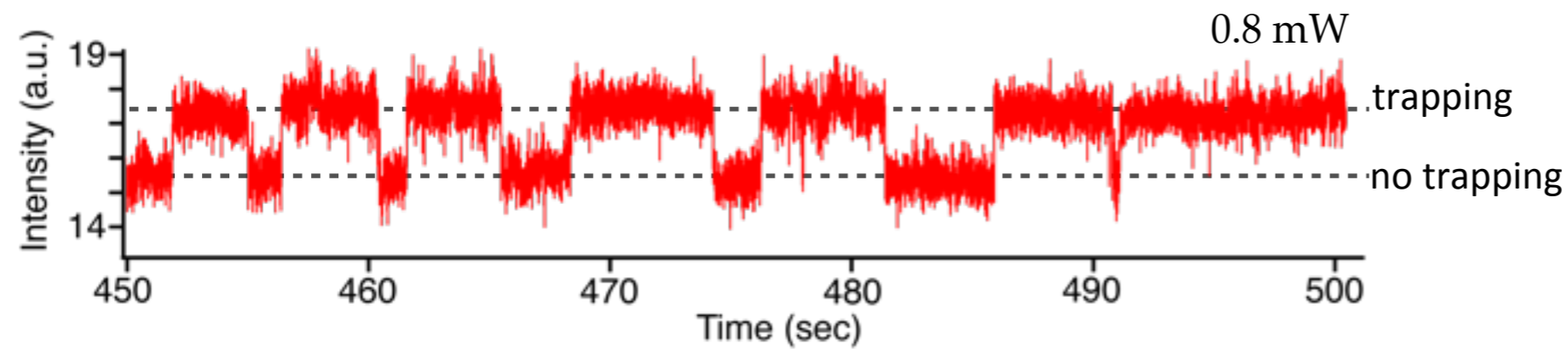


# SIBA Trapping of 100nm and 50 nm PS beads in spherical nanoapertures

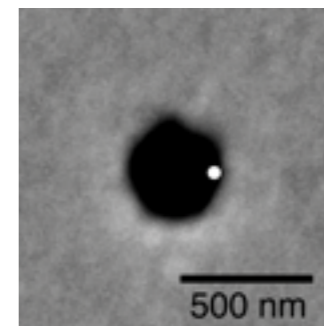
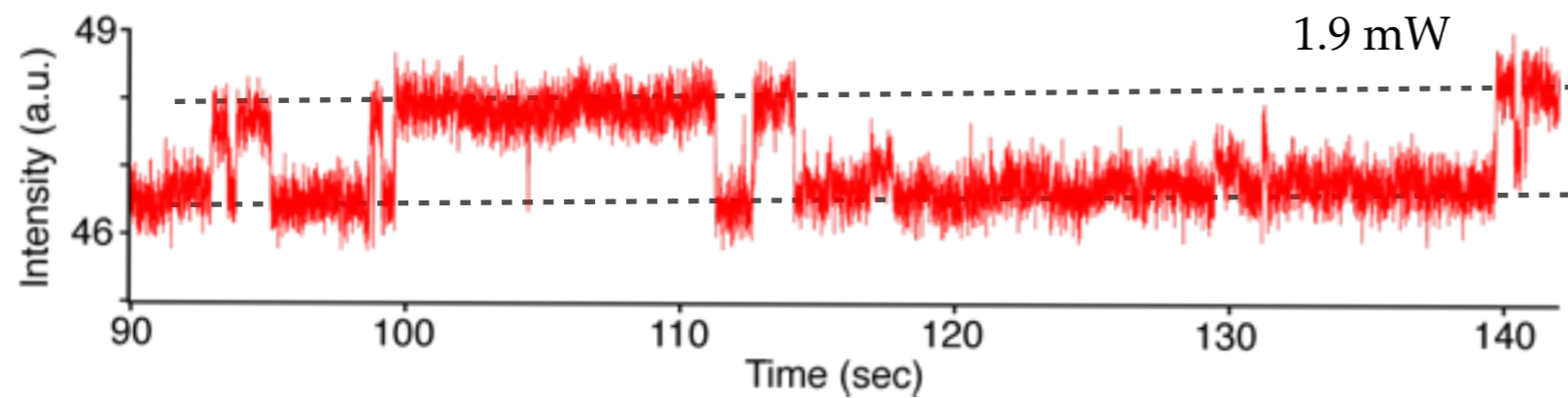


Mathieu Juan  
now @Sydney

Trapping of 100 nm PS beads:  
0.8 mW for ~ 4 sec average trapping time



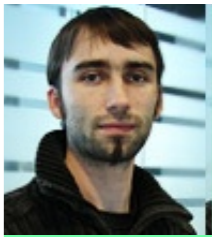
Trapping of 50 nm PS beads:  
1.9 mW for ~ 5 sec average trapping time



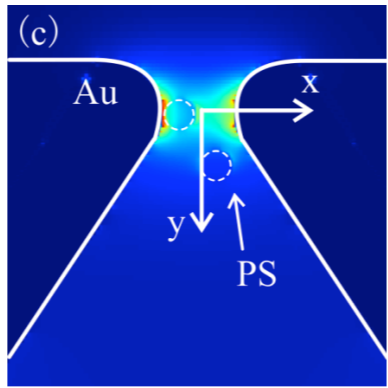
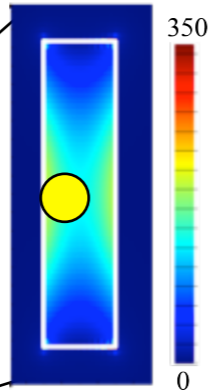
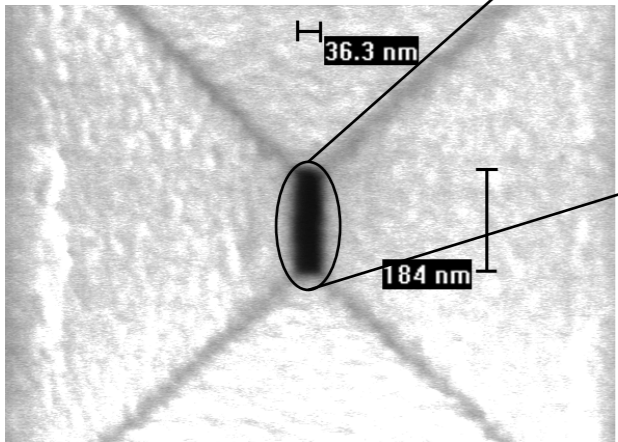


# Improving SIWA trapping through mode engineering

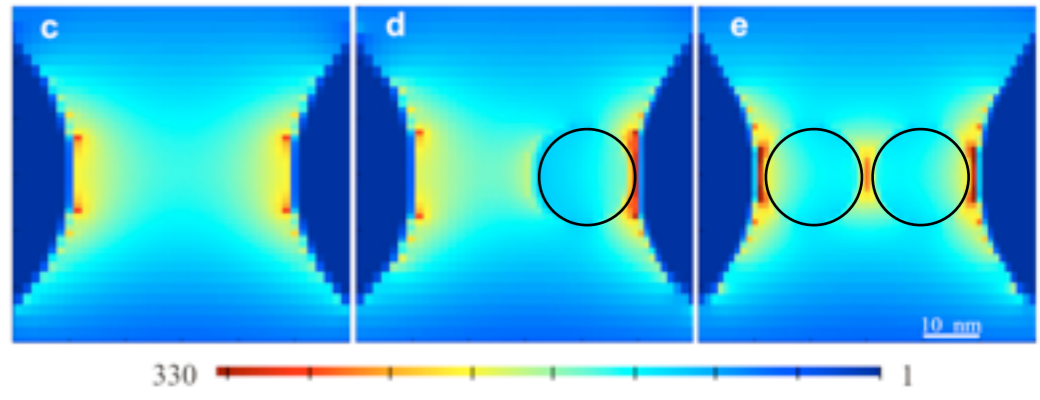
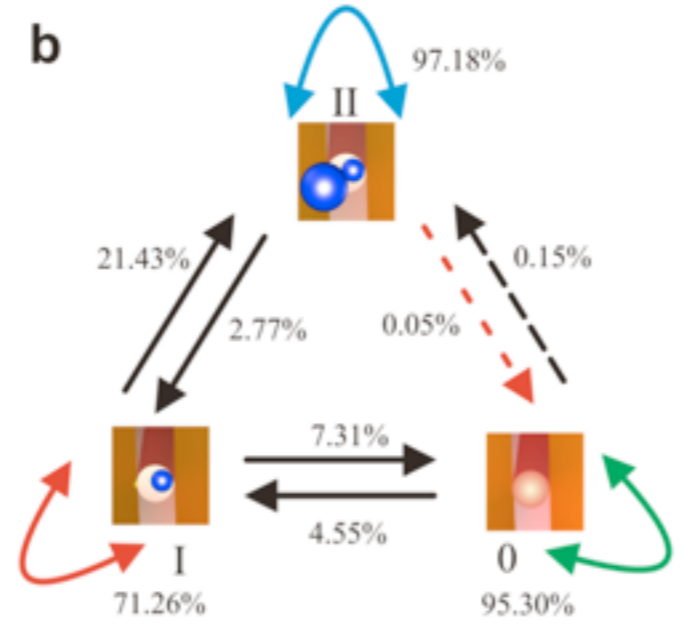
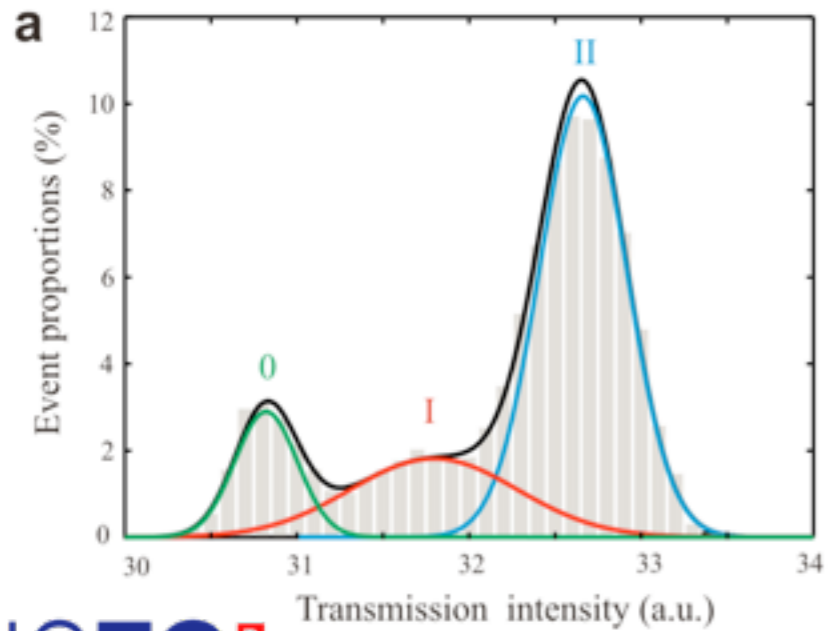
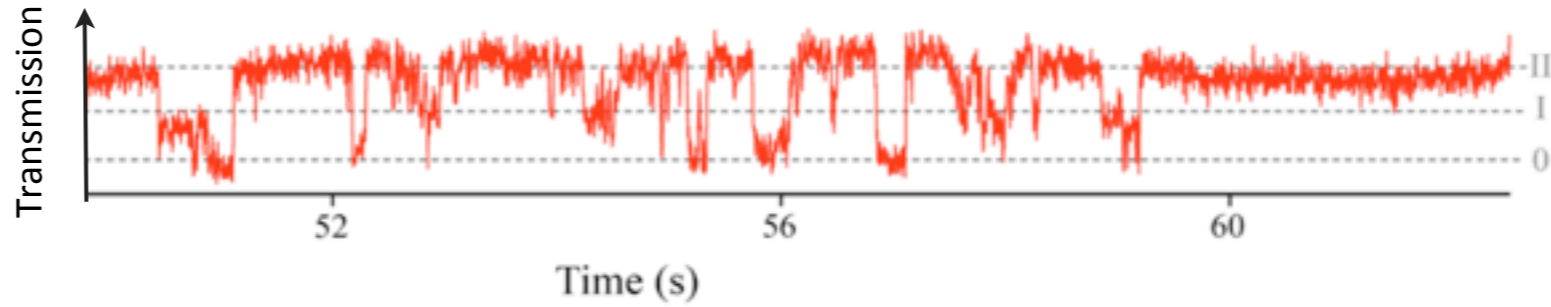
In collaboration with IMEC  
(C. Chen, P. van Dorpe)



Mathieu Juan  
now @Sydney



Trapping of 20nm PS beads



C. Chen, M. L. Juan et al, Nano Lett. **12**, 125-132 (2012)

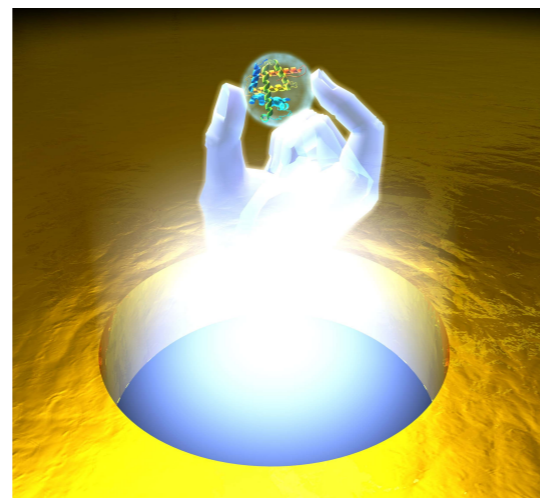


## Plasmon nano-optical tweezers

Mathieu L. Juan<sup>1</sup>, Maurizio Righini<sup>1</sup> and Romain Quidant<sup>1,2</sup>

Conventional optical tweezers, formed at the diffraction-limited focus of a laser beam, have become a powerful and flexible tool for manipulating micrometre-sized objects. Extending optical trapping down to the nanometre scale would open unprecedented opportunities in many fields of science, where such nano-optical tweezers would allow the ultra-accurate positioning of single nano-objects. Among the possible strategies, the ability of metallic nanostructures to control light at the subwavelength scale can be exploited to engineer such nano-optical traps. This Review summarizes the recent advances in the emerging field of plasmon-based optical trapping and discusses the details of plasmon tweezers along with their potential applications to bioscience and quantum optics.

Nature Photonics 5, 349–356 (2011)



Towards single protein trapping??



## Surface Plasmon Early Detection of Circulating Heat Shock Proteins & Tumor Cells (SPEDOC)



### OBJECTIVE

Combining the last advances of plasmon nano-optics (sensing, tweezers, SERS) microfluidics and oncology to develop a new integrated platform for early stage cancer diagnosis and treatment monitoring

### PARTNERS

**UB** - Universite de Bourgogne  
**INSERM** - Institut National de la Sante et de la Recherche Medicale  
**EPFL** - Ecole Polytechnique Federale de Lausanne  
**COSINGO** - Image Optic Spain S.L.  
**ICFO** - Institute of Photonic Sciences (Coordinator)

[spedoc@icfo.es](mailto:spedoc@icfo.es)

**CONTACT**

**WEB**

[www.spedoc.eu](http://www.spedoc.eu)

Initiated the 1<sup>st</sup> of January 2010



# THE PLASMON NANOOPTICS GROUP@ICFO



[romain.quidant@icfo.es](mailto:romain.quidant@icfo.es)





# The Plasmon Nano-Optics Group @ ICFO.es



Openings for both  
postdocs and PhDs !

## PhD students

Giorgio  
Volpe



Srdjan  
Acimovic



Michael  
Geiselmann



Jan  
Gieseler



Jon  
Donner



María Ale  
Ortega



Valeria  
Rodríguez



Soon@ESPCI

## Postdocs

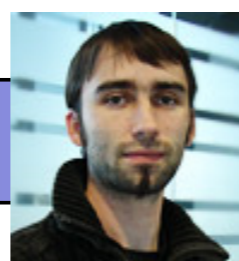
Jan Renger



Mark Kreuzer



Mathieu Juan



Christopher  
Galloway



Elisabet Xifre



Sebastian  
Thompson



Now @Sydney