

# Intracellular microlasers

Matjaž Humar and Seok Hyun Yun

Journal club (2015. 12. 08)

Nature Photon. **9**, 527 (2015)

# About the group

## Faculty



Seok-Hyun (Andy) Yun, PhD

*Director*  
*Associate Professor*  
*Harvard Medical School*  
617-768-8704  
syun@hms.harvard.edu

## Research Fellows



Matjaz Humar, PhD

*Research Fellow*  
*Harvard Medical School*  
mhumar@mgh.harvard.edu



Seonghoon Kim, PhD

*Research Fellow*  
Expected to start on Jan 2016  
seonghoona@gmail.com



Ki Su Kim, PhD

*Research Fellow*  
*Harvard Medical School*  
kkim18@mgh.harvard.edu



Moonseok Kim, PhD

*Research Fellow*  
optiker01@gmail.com



# About the group



Seok Hyun (Andy) Yun

Professor, [Harvard University](#)

[Optics](#), [Imaging](#), [Biomedical Engineering](#), [Photomedicine](#)

Verified email at [hms.harvard.edu](mailto:hms.harvard.edu) - [Homepage](#)

Follow

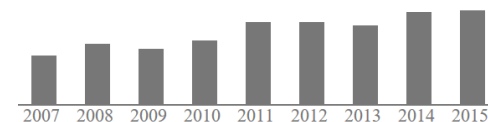
Title	1–20	Cited by	Year
<a href="#">High-speed optical frequency-domain imaging</a>	S Yun, G Tearney, J de Boer, N Iftimia, B Bouma Optics Express 11 (22), 2953-2963	863	2003
<a href="#">In vivo high-resolution video-rate spectral-domain optical coherence tomography of the human retina and optic nerve</a>	N Nassif, B Cense, B Park, M Pierce, S Yun, B Bouma, G Tearney, T Chen, ... Optics Express 12 (3), 367-376	557	2004
<a href="#">In vivo human retinal imaging by ultrahigh-speed spectral domain optical coherence tomography</a>	N Nassif, B Cense, B Hyle Park, SH Yun, TC Chen, BE Bouma, ... Optics letters 29 (5), 480-482	543	2004
<a href="#">Ultrahigh-resolution high-speed retinal imaging using spectral-domain optical coherence tomography</a>	B Cense, N Nassif, T Chen, M Pierce, SH Yun, B Park, B Bouma, ... Optics Express 12 (11), 2435-2447	484	2004
<a href="#">High-speed spectral-domain optical coherence tomography at 1.3 <math>\mu\text{m}</math> wavelength</a>	S Yun, G Tearney, B Bouma, B Park, J de Boer Optics Express 11 (26), 3598-3604	420	2003
<a href="#">High-speed wavelength-swept semiconductor laser with a polygon-scanner-based wavelength filter</a>	SH Yun, C Boudoux, GJ Tearney, BE Bouma	406	2003

Google Scholar



Get my own profile

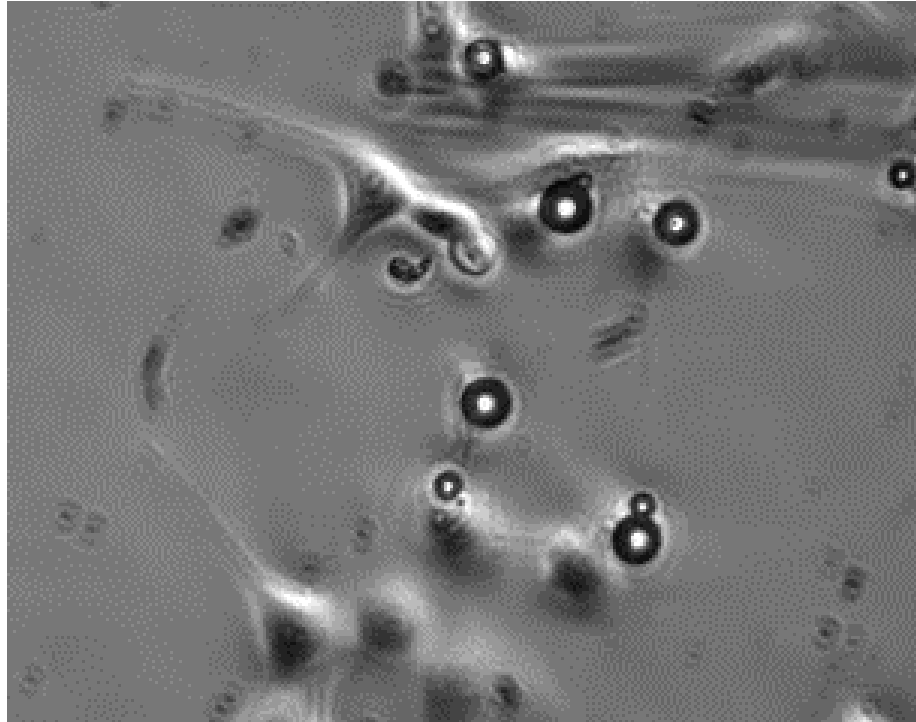
Citation indices	All	Since 2010
Citations	10395	6423
h-index	47	43
i10-index	108	91



# Introduction

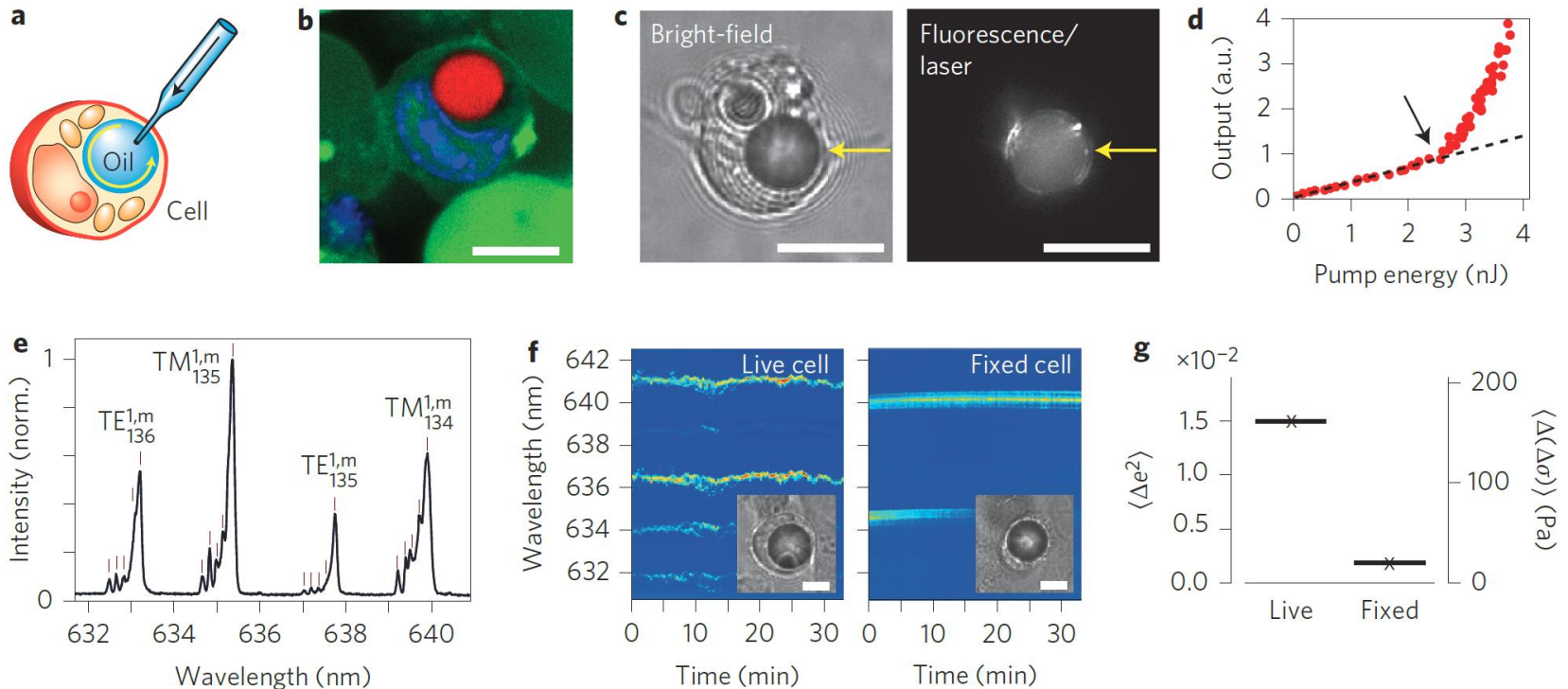
- Luminescent probes (dyes, QDs, bioluminescent molecules...) are indispensable tools in cell biology and medical sciences.
- Broad emission spectra limit the number of simultaneously usable probes.
- Optical resonances and stimulated emission allow spectral narrowing.
- Cellular lasers with external cavities demonstrated by the authors and others.
- In this paper, effective approaches to generate stand-alone cell lasers based on intracellular whispering-gallery mode (WGM) resonator.

## Soft microresonator (1/2) – Injecting oil into the cells



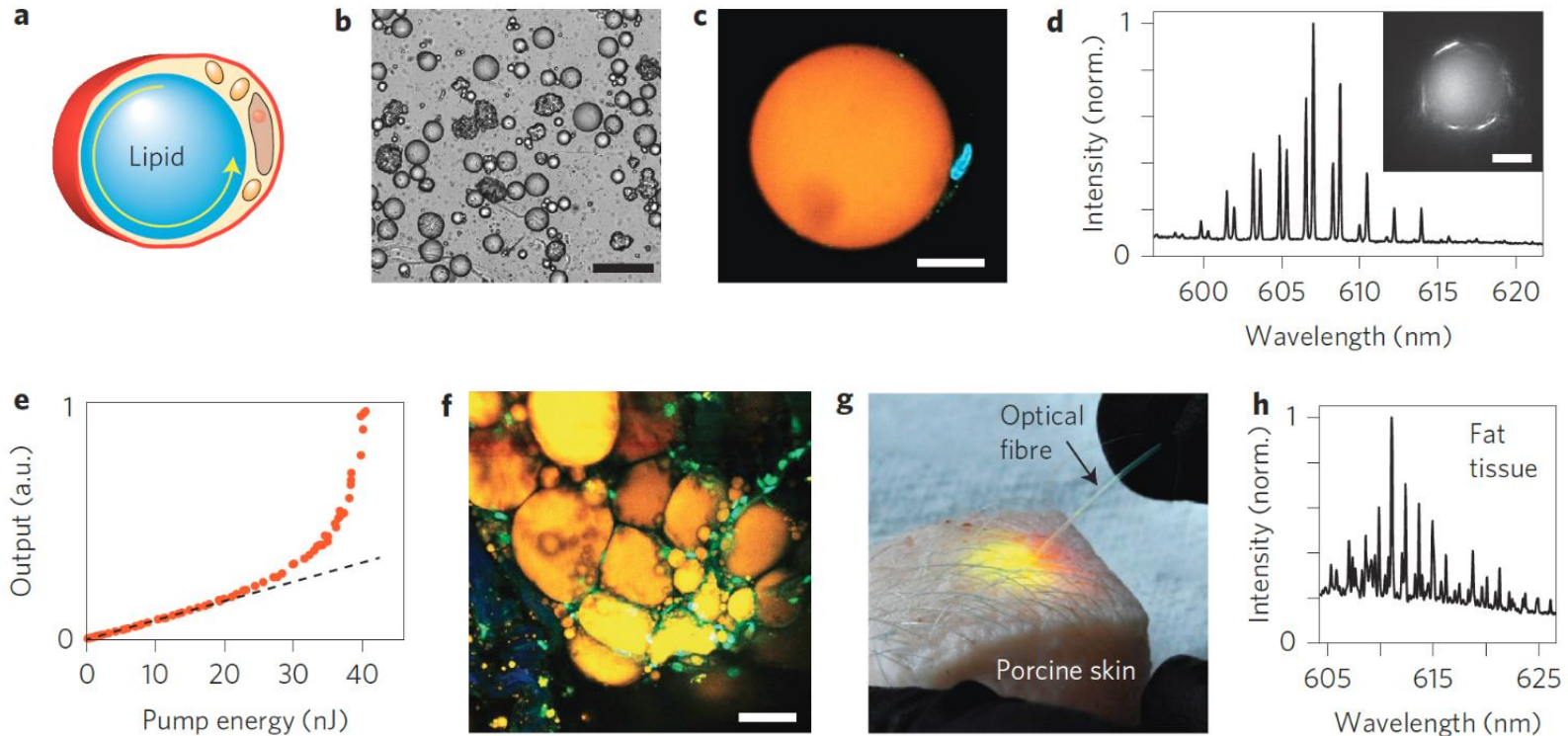
- Dye-mixed fluid injected into the cells.
- Cytoplasmic internal stress  $\Delta\sigma = 500\text{pN/m}^2$  with  $20\text{pN/m}^2$  sensitivity measured.
- Dynamic variations of the cellular stress in live and dead cells revealed by time-lapse traces of the output spectra.

# Soft microresonator (1/2) – Injecting oil into the cells



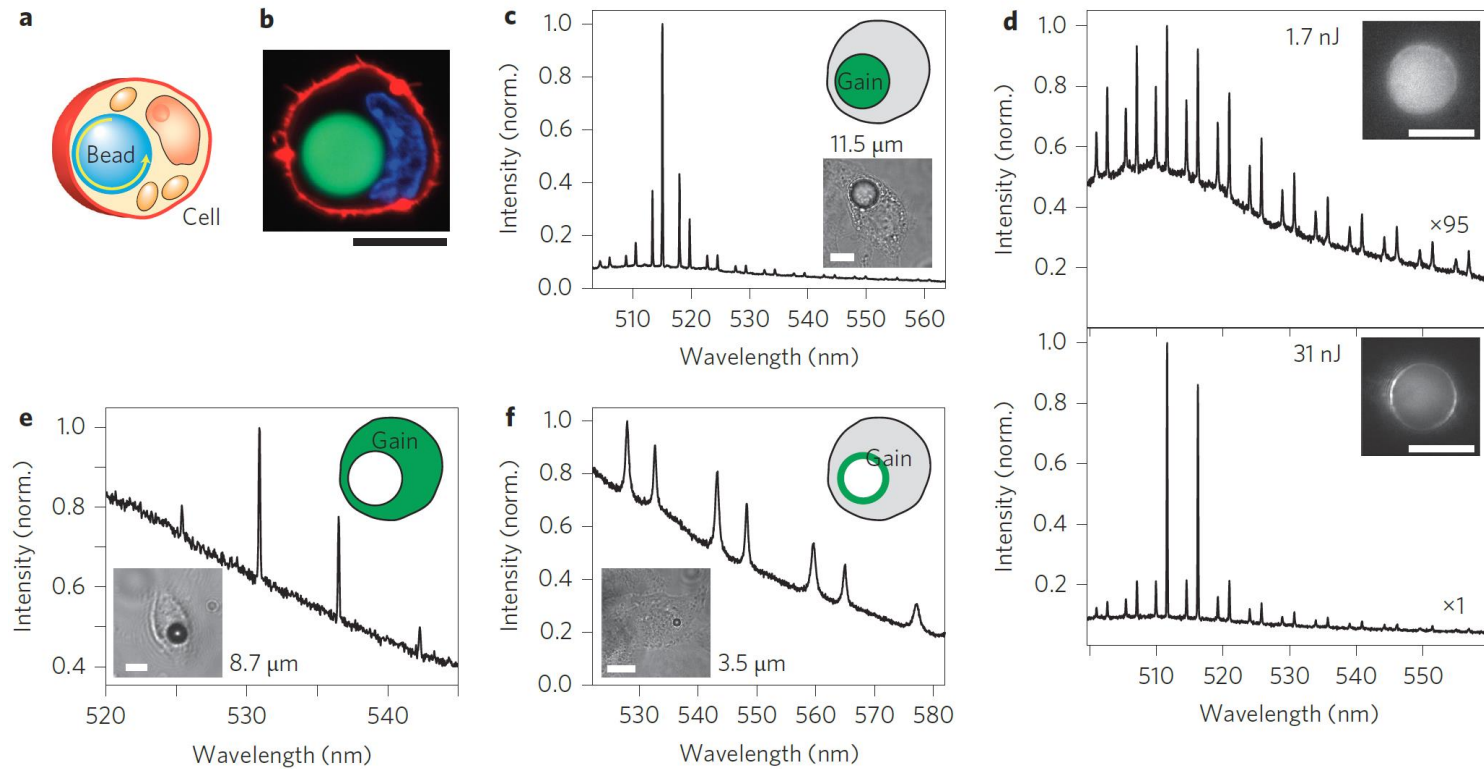
- Dye-mixed fluid injected into the cells.
- Cytoplasmic internal stress  $\Delta\sigma = 500\text{pN/m}^2$  with  $20\text{pN/m}^2$  sensitivity measured.
- Dynamic variations of the cellular stress in live and dead cells revealed by time-lapse traces of the output spectra.

# Soft microresonator (2/2) – Natural lipid droplets



- Incubated adipocytes with a lipophilic fluorescent dye
- Extracted adipocyte contain a single lipid droplet with nearly perfect spherical shape.
- Excited adipocytes in fat showed lasing.

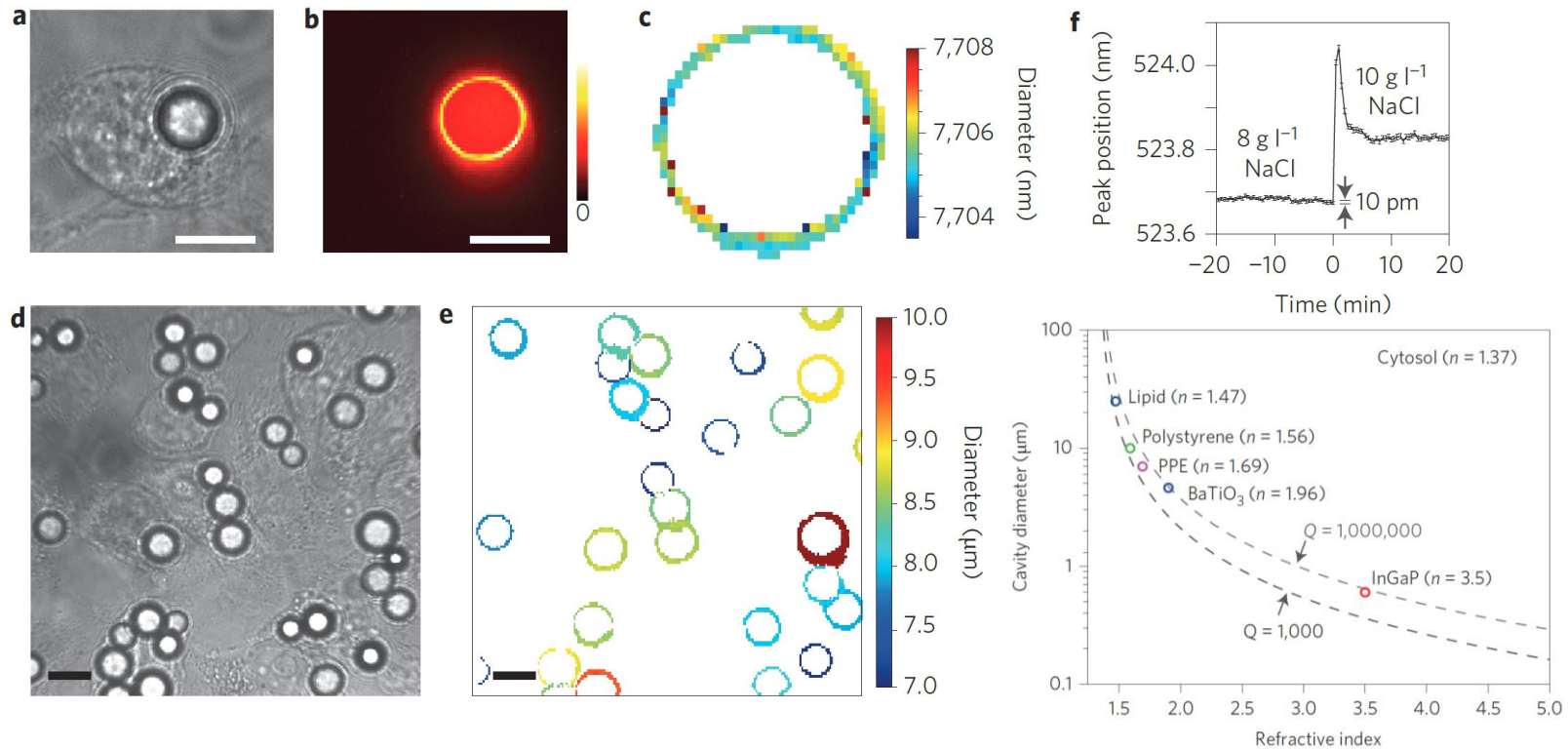
# Hard microresonator – Phagocytized polystyrene beads



- Polystyrene beads are internalized into HeLa cells by endocytosis.
- Dye-embedded beads, gain-doped cytoplasm, dye-coated beads tested.

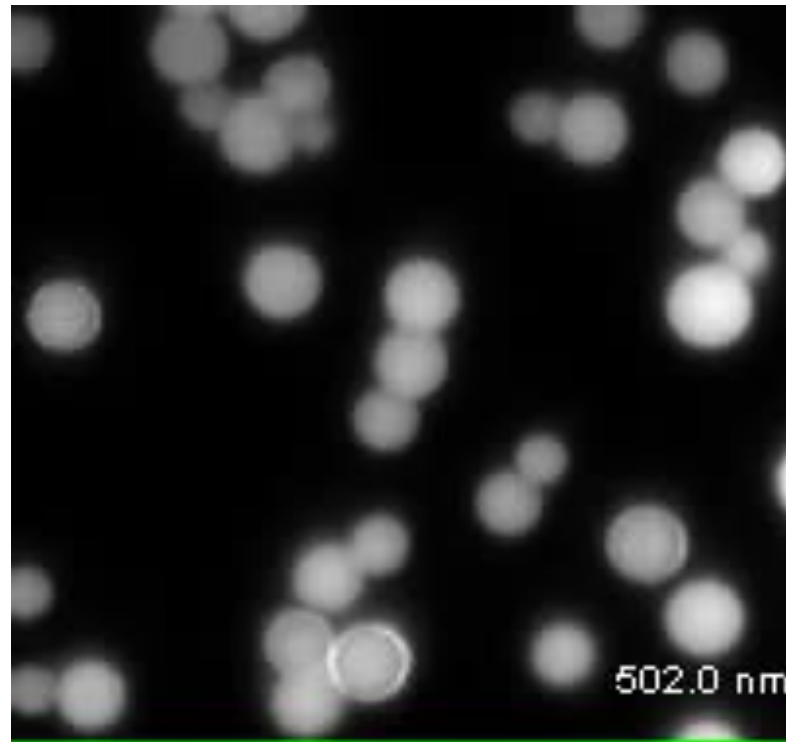


# Applications



- Effective bead diameter is calculated by fitting the spectra with WGM theory, exhibiting 50pm precision.
- Applied to tag individual cells and 2,000 individual cells distinguished.
- Using three beads per cell and five different dyes with non-overlapping spectra, tagging  $2 \times 10^{11}$  individual cells possible.

# Applications



- Effective bead diameter calculated by fitting the spectra with WGM theory, exhibiting 50pm precision.
- Applied to tag individual cells and 2,000 individual cells distinguished.
- Using three beads per cell and five different dyes with non-overlapping spectra, tagging  $2 \times 10^{11}$  individual cells possible.

# Summary

- Standalone cell lasers demonstrated by using intracellular WGM microresonators.
- The laser spectra from oil droplets charted cytoplasmic internal stress and its dynamic fluctuations.
- WGM within phagocytized beads of different sizes enabled individual tagging of thousands of cells, in principle, even much larger number possible.