

**Лекция профессора Кирилла Ларина (Университет
Хьюстона)**

В рамках программы Фонда Дмитрия Зимина «Династия»
«Краткосрочные визиты иностранных ученых в российские
научные центры»

19 сентября 2015г. – 28 сентября 2015 г.

**Мониторинг оптического просветления биологических
тканей и обзор последних достижений**

Introduction

- Light can be **absorbed, reflected, scattered** and **remitted** by a tissue and each of these processes carries information about the **micro- and macrostructure, the movement and shape** of its components
- The **functional imaging** and monitoring of the state of human organs and tissues is a very attractive feature of the optical method
- The **microscopic, spectroscopic, polarimetric, interferometric, and speckle** methods are key techniques for tissue optical **imaging and medical diagnosis**

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Optics in Medicine

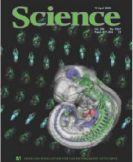
Radiology		Pathology	
1 cm	1 mm	100 μm	10 μm
Radio nucleotide DOT PET	MRI CT US	HFUS MRI X-Ray, CT	LIGHT, e.g., Microscopy OCT
Low resolution		Organ Level Tumor Staging	Architectural, Cellular, Optical Biopsy
<p>..... Diagnostic capability</p> <p>Histopathology is the golden standard especially for cancer diagnosis</p> <p>Only optical technique approach cellular resolution</p>			

DOT: Diffuse Optical Tomography; PET: Positron Emission Tomography; MRI: Magnetic Resonance Imaging; CT: Computed Tomography; US: Ultra Sound; HFUS: High Frequency Ultra Sound; OCT: Optical Coherence Tomography.

Introduction

➤ **Light in Medicine and Biology** (broader than "just imaging"....)

- Therapeutics: photo-thermal, -mechanical, -chemical modes
- Diagnostics: imaging, spectroscopy, "remote sensing"
- Analytics: advanced optical microscopies



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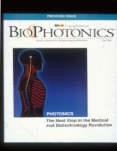
Basic definitions

- **Photonics:**
science & technology of light generation, manipulation, transmission, and measurement
ultraviolet-visible-infrared
- **Biophotonics:**
applications of photonics to life sciences and medicine

Why light?

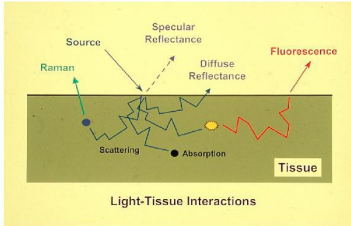
- photon energies = molecular energy levels
- wavelengths = cell, tissue micro-structures

→ Probe for diagnostics+analytics
 → Modify for therapeutics



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What happens to light when you shine it onto tissue? (where does it go? what does it do when it gets there?)

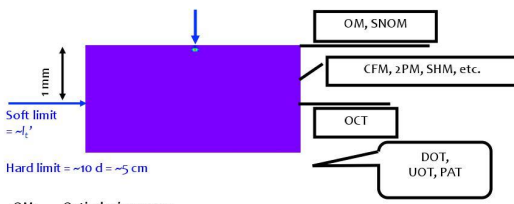


Light-Tissue Interactions

- relative probability/importance λ -dependent
- therapy (PDT, thermal, ablative) follows absorption
- (multiple) scattering dominant 600-1300 nm

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Challenges in Optical Imaging




Soft limit = $\sim \lambda^2$

Hard limit = $\sim 10d = \sim 5\text{ cm}$

- OM: Optical microscopy
- SNOM: Scanning near-field optical microscopy
- CFM: Confocal microscopy
- 2PM: Two-photon microscopy
- SHM: Second harmonic microscopy
- OCT: Optical coherence tomography
- DOT: Diffuse optical tomography
- UOT: Ultrasound-modulated optical tomography
- PAT: Photoacoustic tomography

Courtesy of Dr. L. Wang

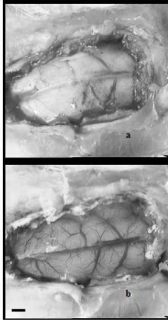
Optical Clearing



Transparent Not Transparent!

- being transparent would improve optical imaging and diagnostics!

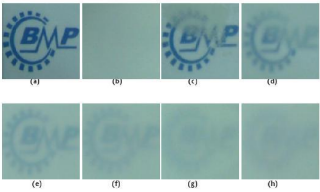
Optical Clearing



Visual changes and the measured optical changes for *in vivo* rabbit *dura mater* before and after treatment with glycerol epidurally.

Bashkatov et al, Biophysics J. 85, (2003).

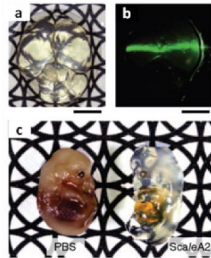
Optical Clearing



The photographic images of OCA-induced optical clearing of Intralipid-based tissue phantom in the sample cell: background (a), 5%-Intralipid (b), and 5%-Intralipid with 50%-PEG200 (c), -PEG400 (d), -DMSO (e), -glycerol (f), -1,4-butanediol (g), and -1,2-propanediol (h).

A.T. Yeh, et. al, J. Invest. Dermatol. 121 (2003).

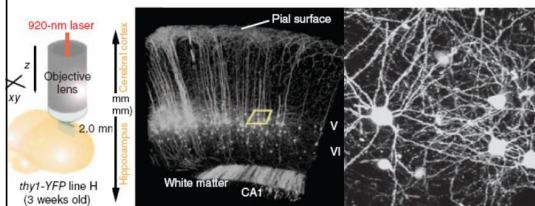
Optical Clearing



(a) Cleared brain was taken with patterned background. (b) A 1-mW, 532-nm laser beam irradiated at the cleared brain. (c) Two embryos placed in PBS or incubated in ScaleA2 solution for 2 weeks after fixation.

H. Hama, et al, Nat. Neurosci. 14 (2011).

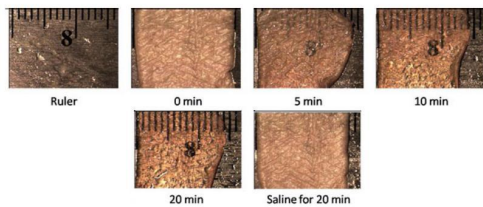
Optical Clearing



Reconstruction of ScaleA2-treated brain in the cerebral cortex and hippocampus of YFP-H mice. Right- An expanded view at a depth of 0.9 mm (a yellow box in b)

H. Hama, et al, Nat. Neurosci. 14 (2011).

Optical Clearing of Skin



Microvascular Therapeutics & Imaging Laboratory, UCI
