



Biomedical Optics: Multichannel Spectroscopy

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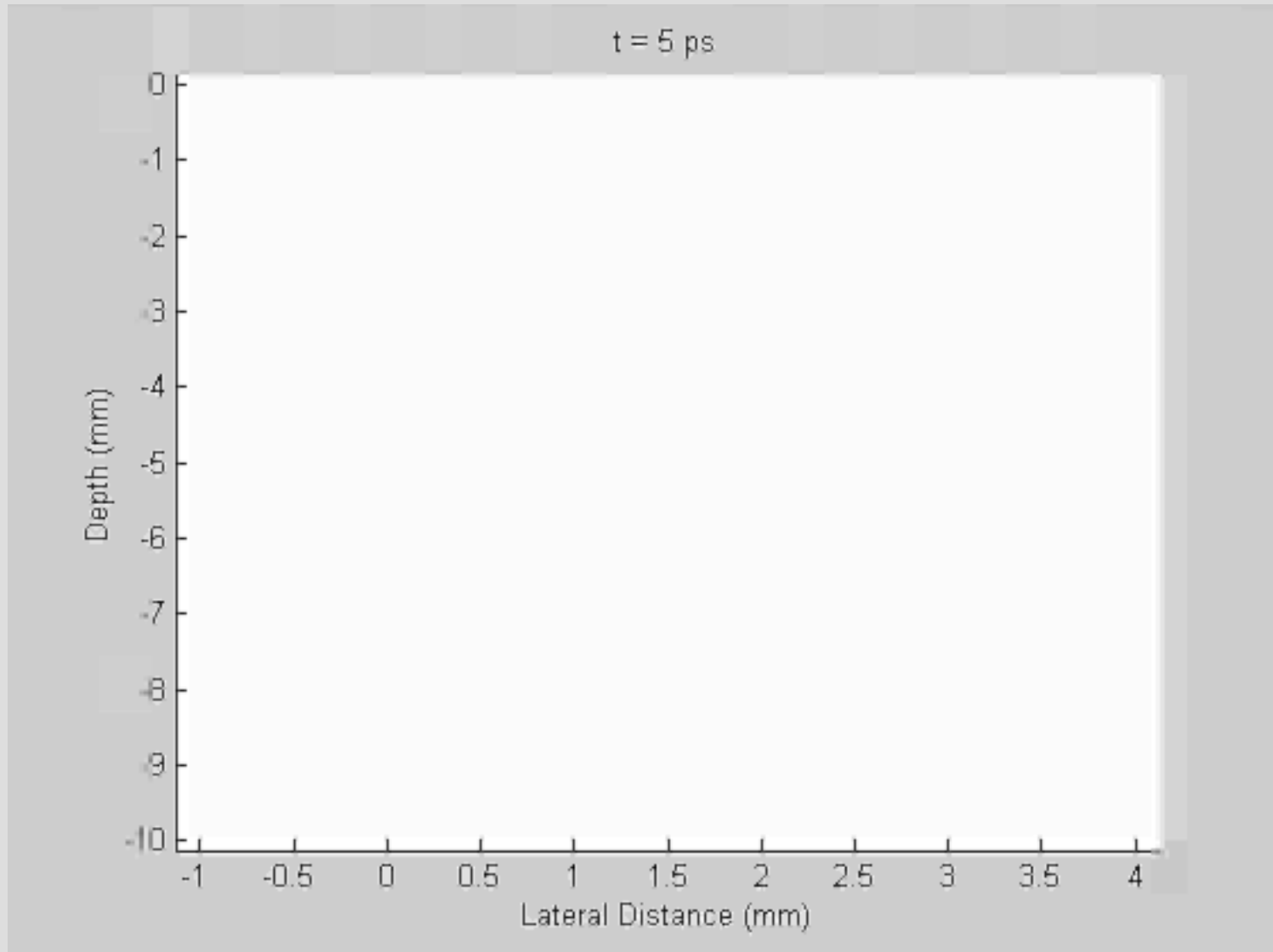
Quantum-Limited Imaging Detectors Symposium
Rochester Institute of Technology
March 2, 2009

*3 biomedical spectroscopy arenas
detectors used
daring to dream*

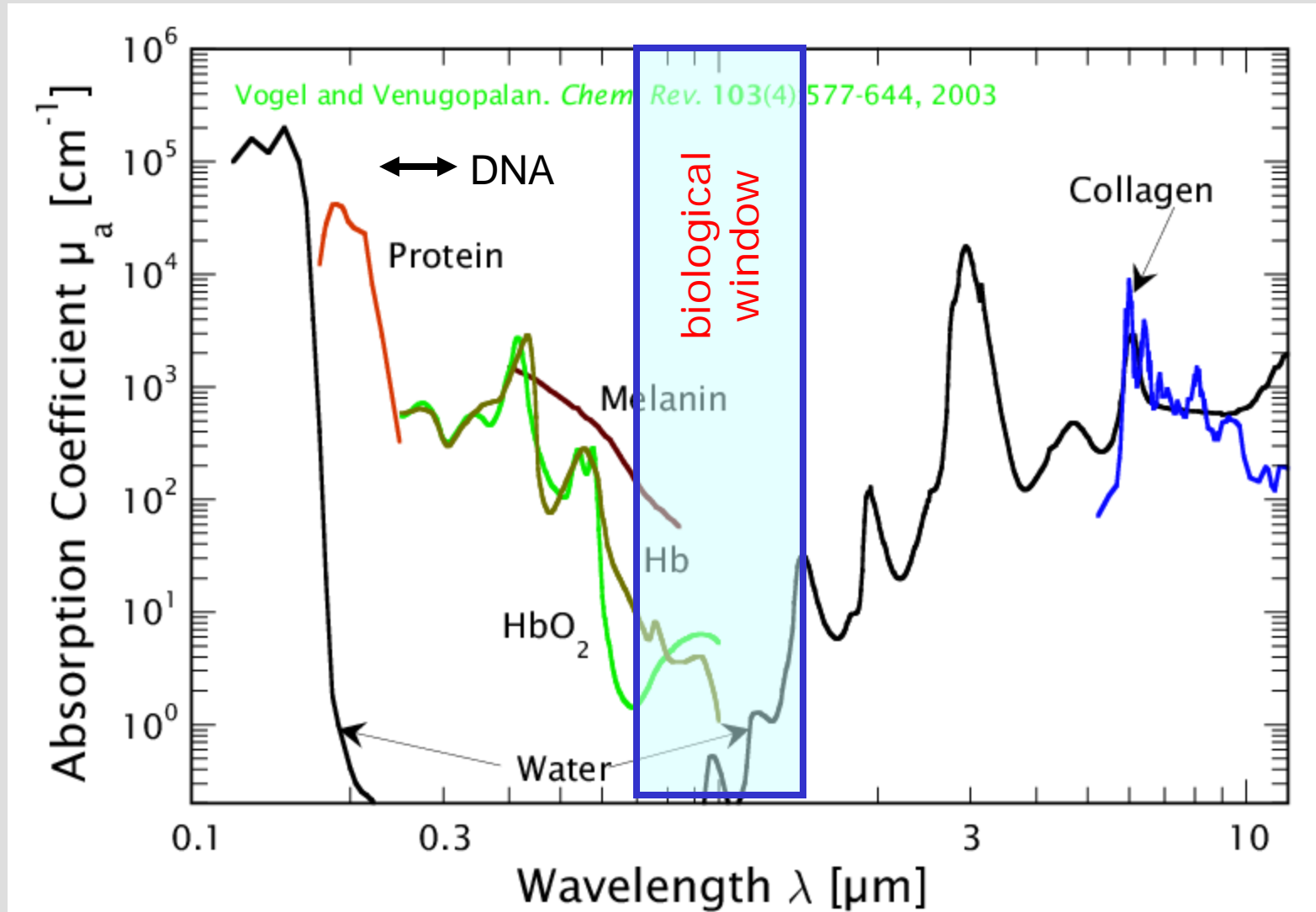
Biomedical Optics: Application Areas

- diffuse photon propagation
- fluorescence lifetime spectroscopy
- Raman spectroscopy
- ***barely imaging!!!***

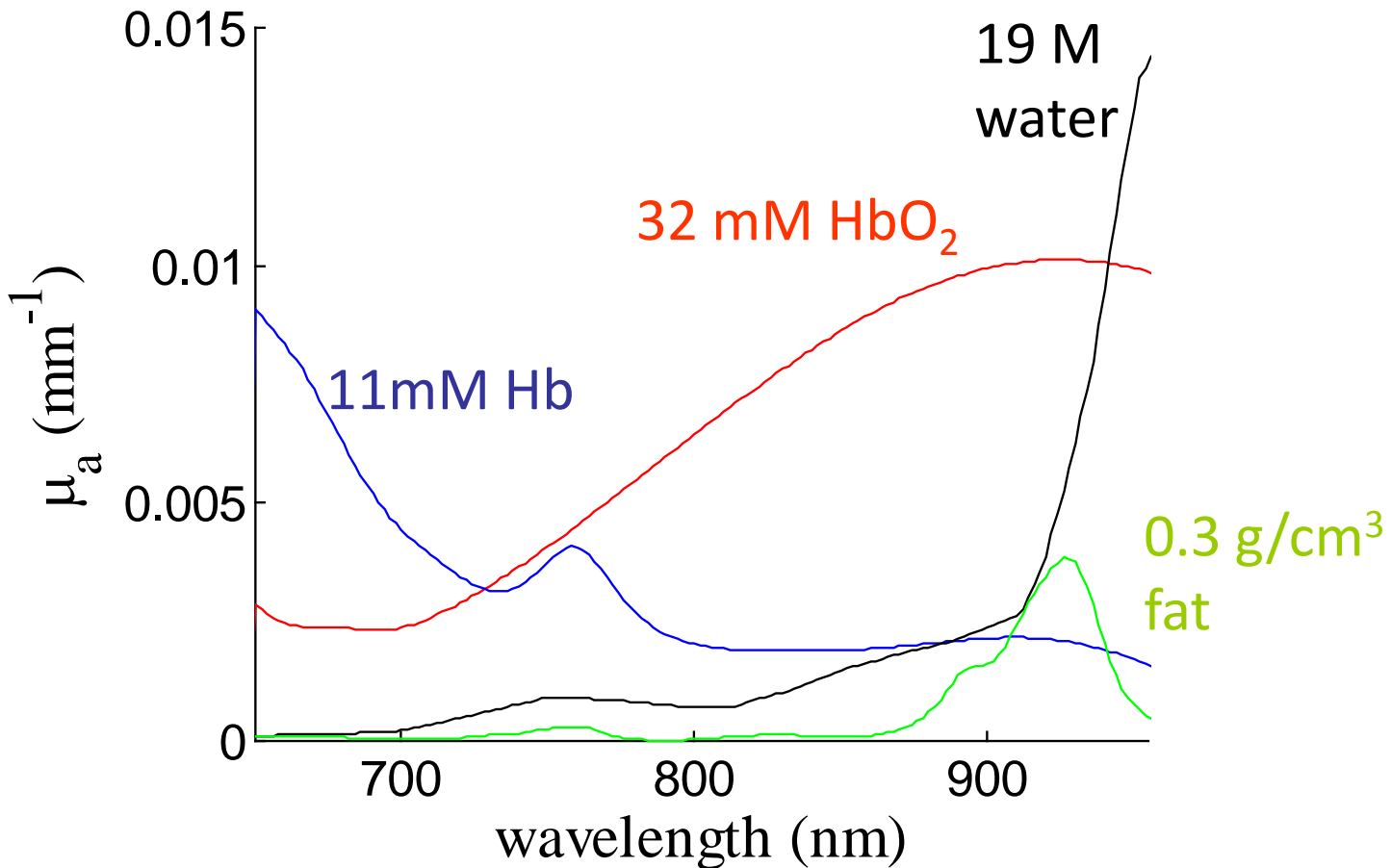
Area #1: Diffuse photon propagation



Where biomedical optics lives....



Important near-IR absorbers

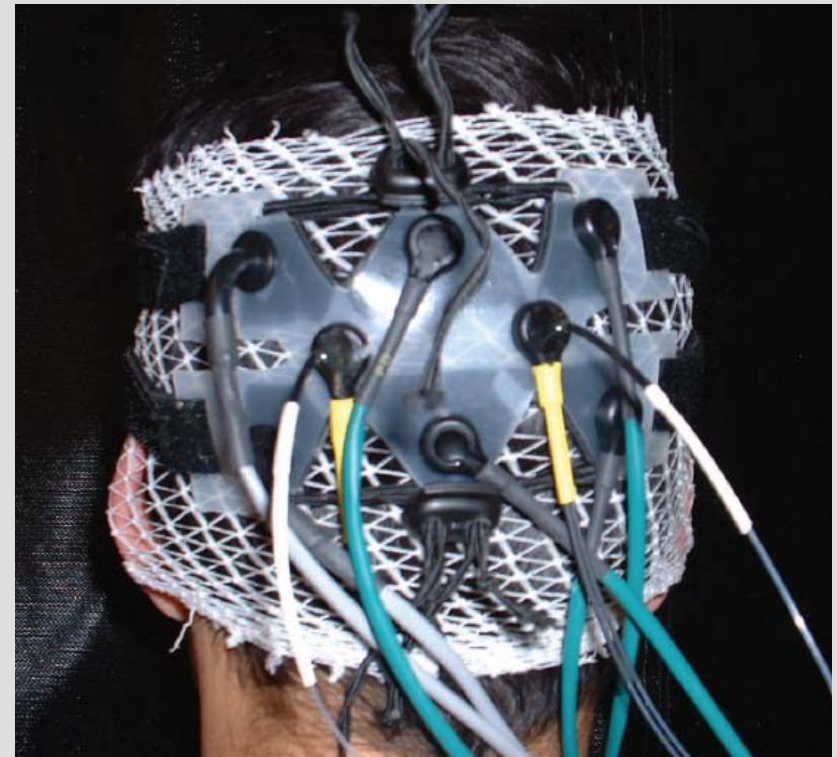


Near-infrared cerebral blood monitoring



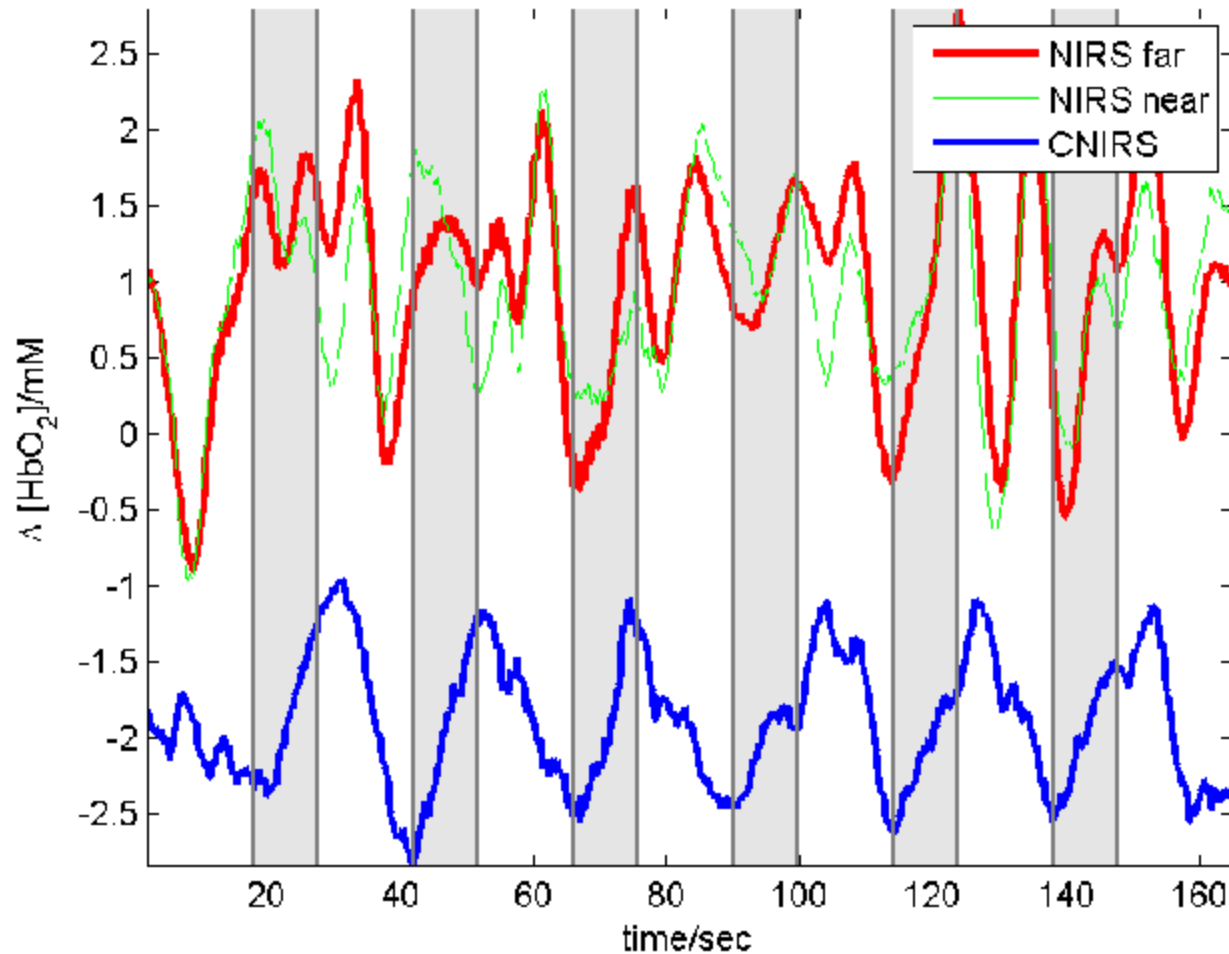
light in (690, 830 nm)

light out





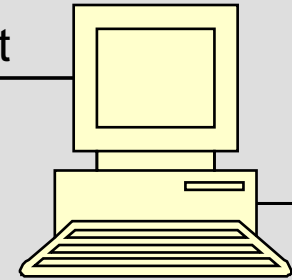
Seeing functional responses: visual stimulation



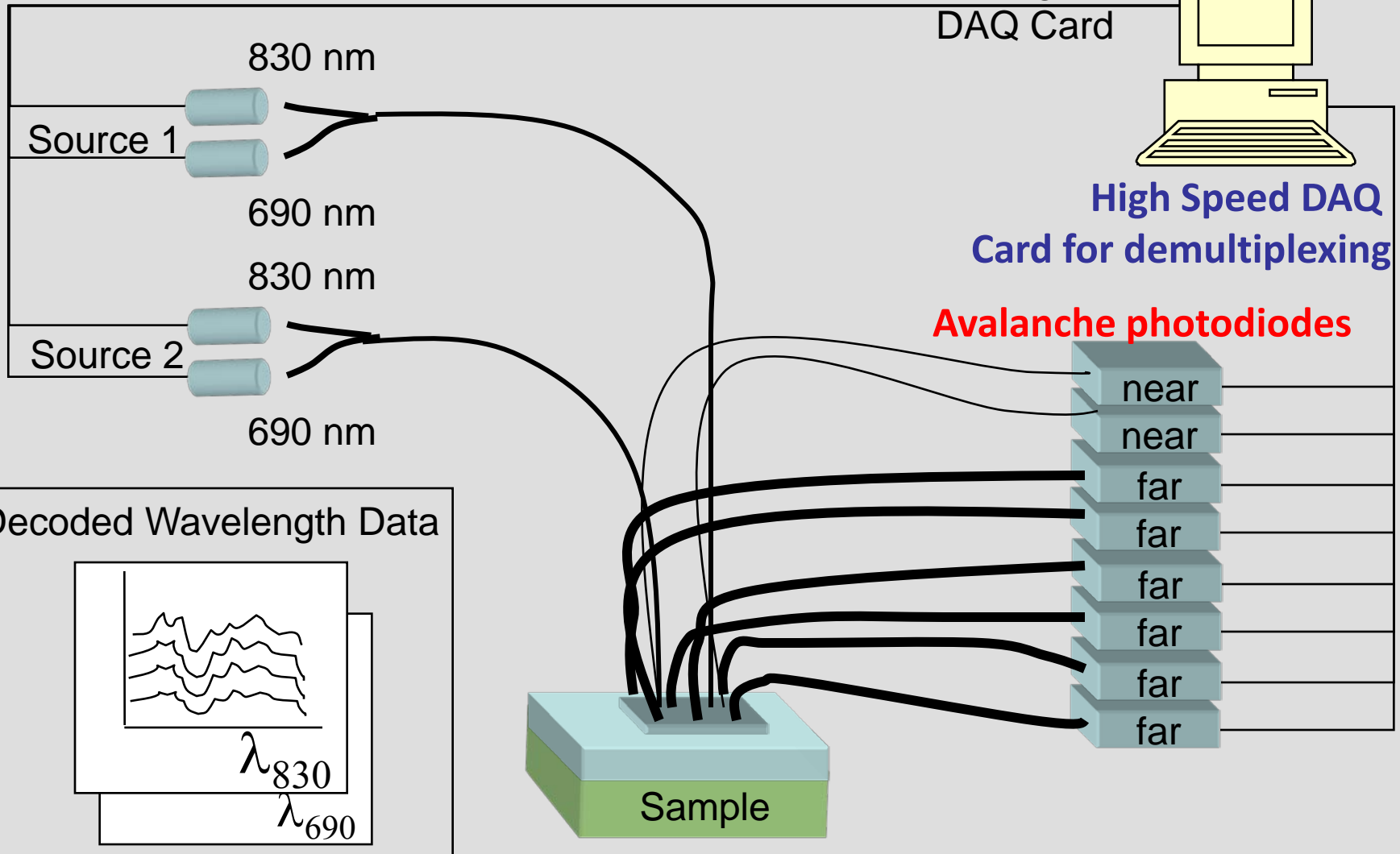
Brain monitoring system layout

1-10 kHz modulation
for **wavelength encoding**

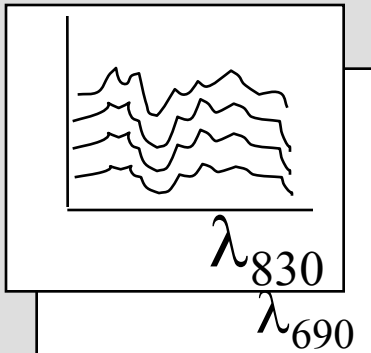
Analog Out
DAQ Card



**High Speed DAQ
Card for demultiplexing**

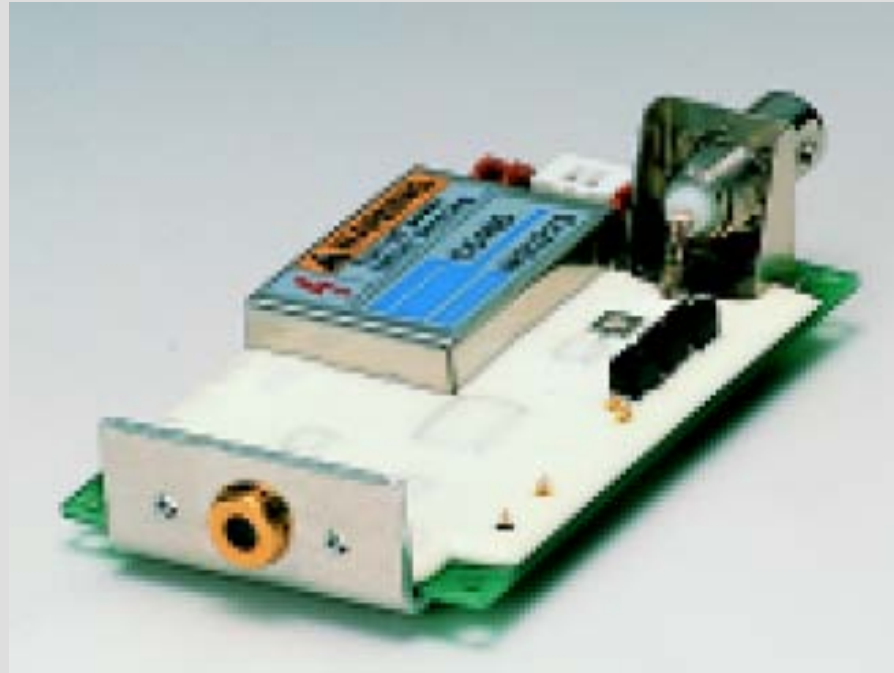


Decoded Wavelength Data



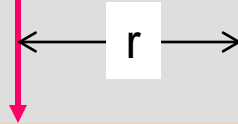
Typical detector for NIRS work

- Hamamatsu silicon avalanche photodiode modules
- Frequency rolloff in low MHz to GHz
- Spectral response out to 1000 nm

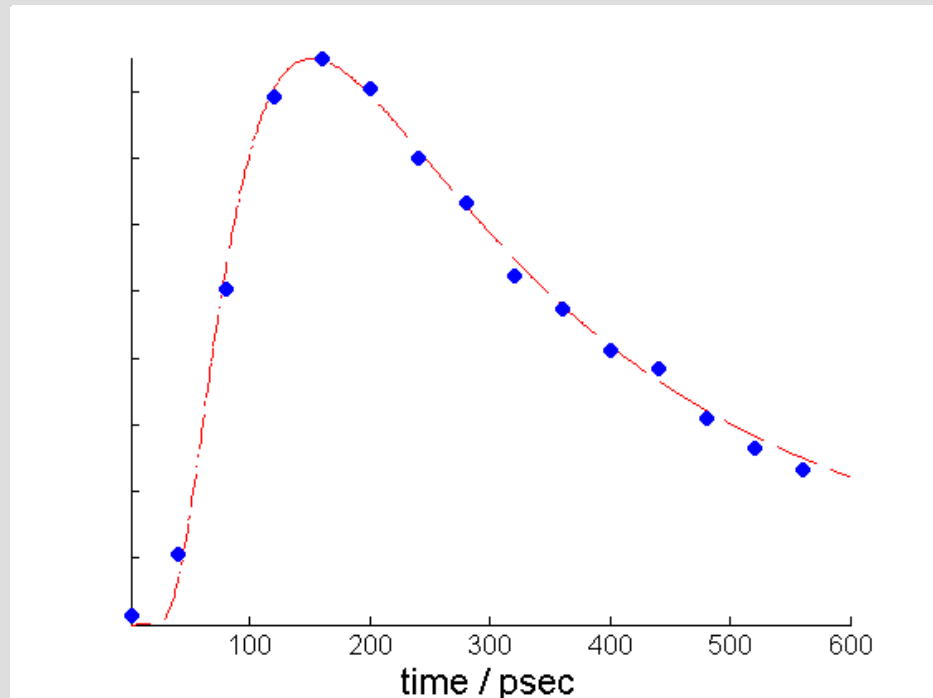


Time-resolved measurements

pulse at $t=0$ remitted light at $t > 0$



absorption and scattering



Hand-Held Optical Breast Scanner

Measurement of breast tissue optical properties B. J. Tromberg and others 663

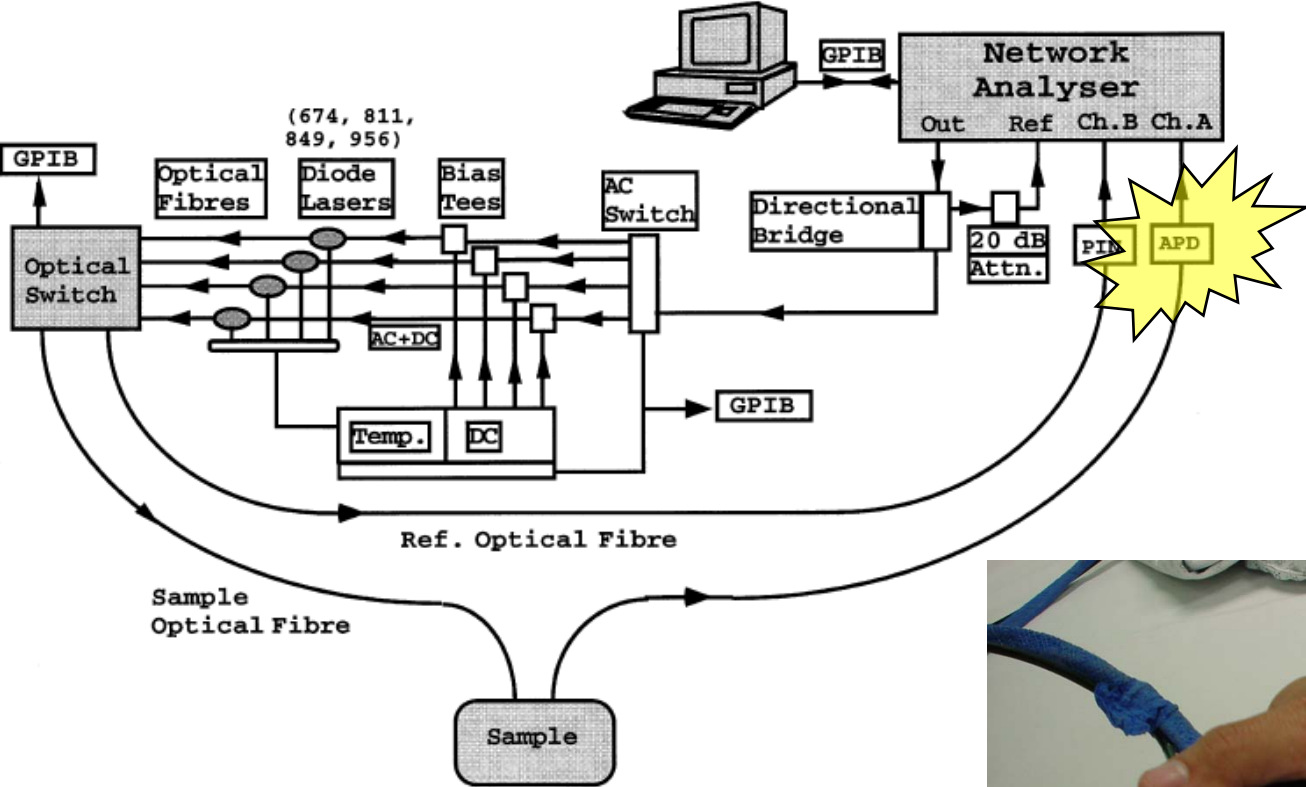


Figure 1. Multiwavelength, multifrequency FDPM instrument.



Hand-Held Optical Breast Scanner



Pham, TH., et al. Review of Scientific Instruments, 71, 1 – 14, (2000).

Bevilacqua, F., et al. Applied Optics, 39, 6498-6507, (2000).

Jakobowski et al., J. Biomed. Opt., 9(1), 230-238 (2004).

(courtesy F. Bevilacqua)

Heavily multiplexed systems!

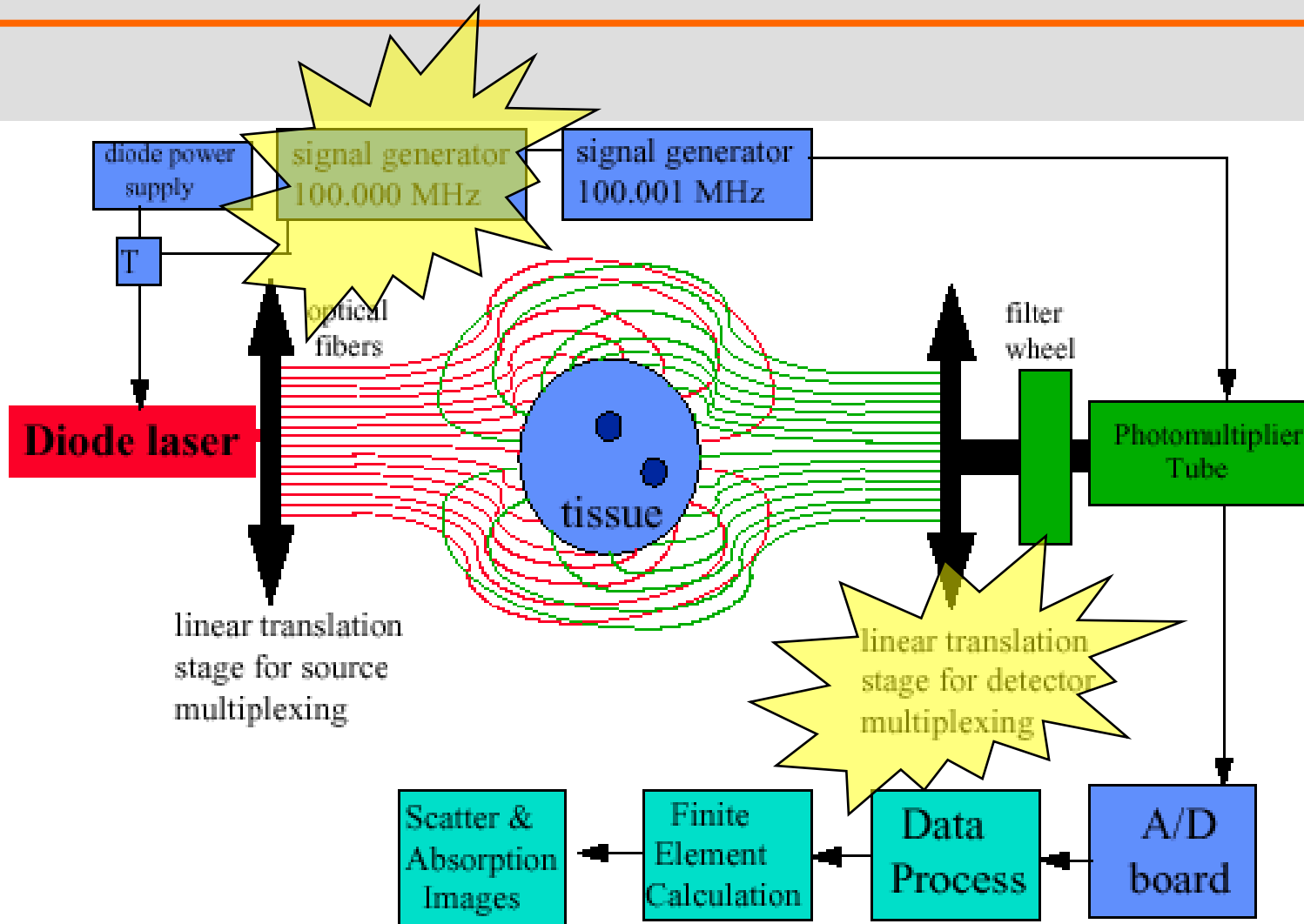


Fig. 1. Schematic of the automated imaging instrument including hardware and software processing. Source optical fibers are indicated in red and detector optical fibers in green.

Diffuse propagation: goals, requirements

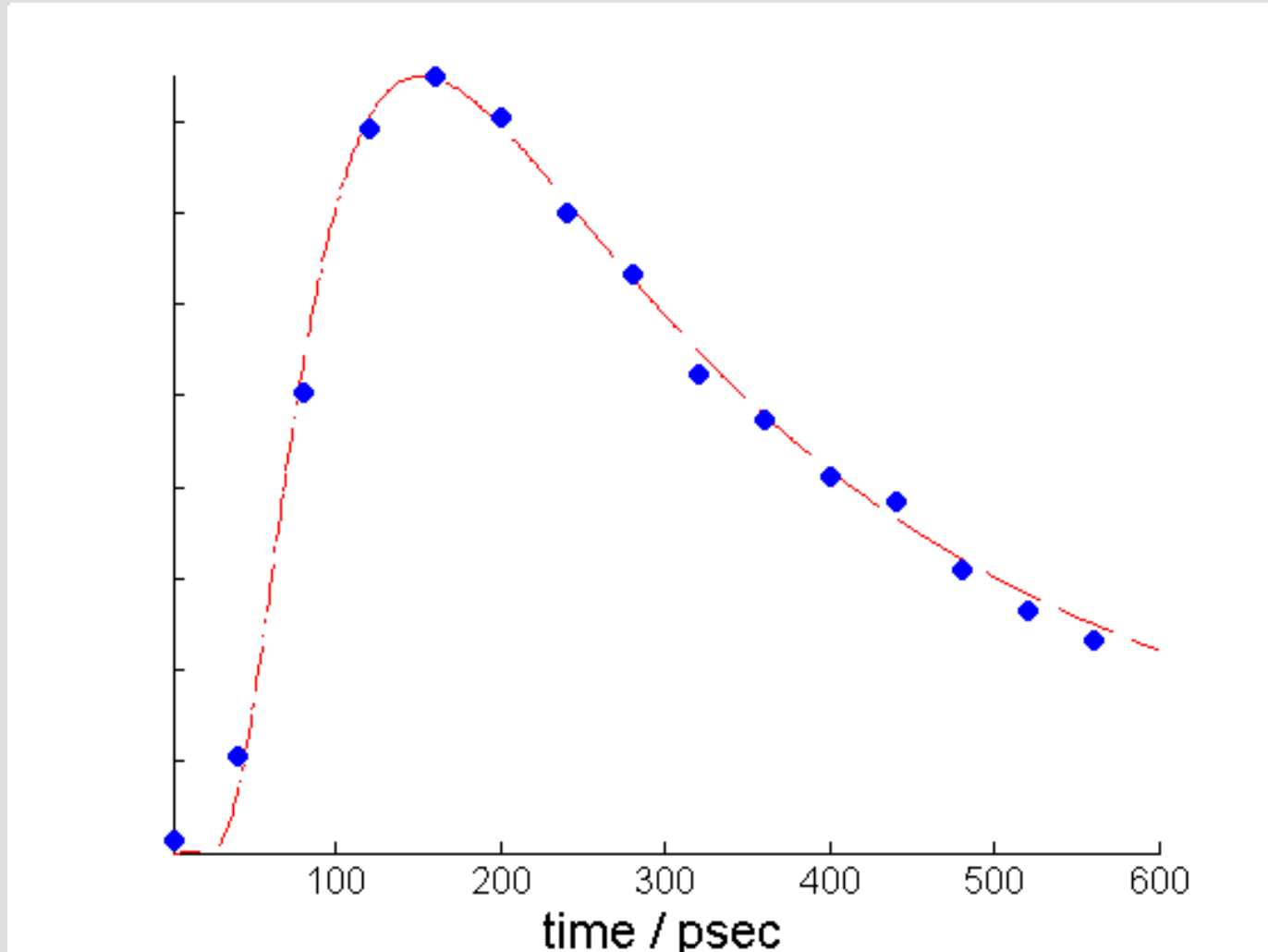


- Distinguish benign from malignant tumor tissue
- Map blood activity (hemodynamics) within brain

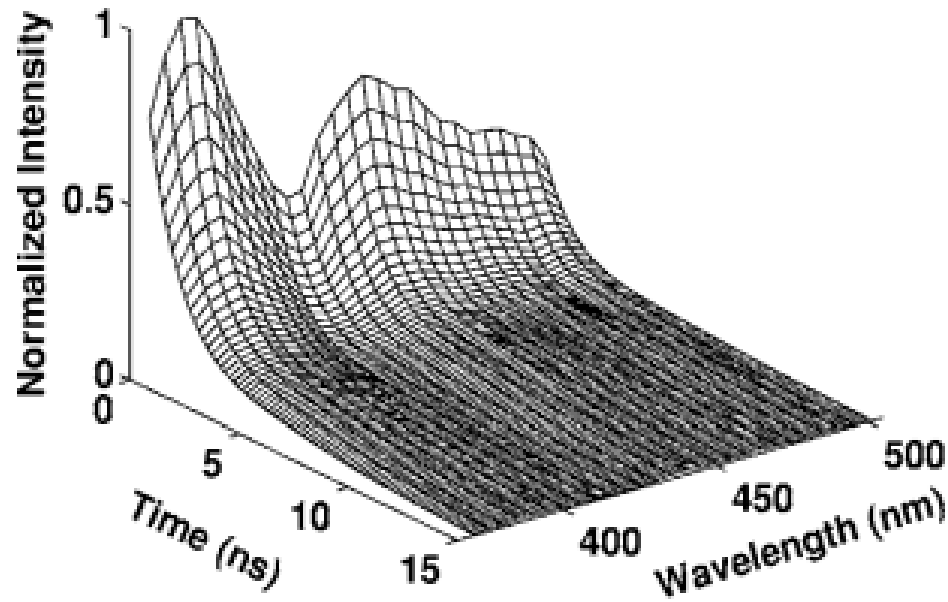
- Sense deep within tissue (cm)
- Record at many locations
- Record at many wavelengths
- Time resolution to few psec

Area #2: Fluorescence lifetime spectroscopy

Once again, psec-nsec timescale!



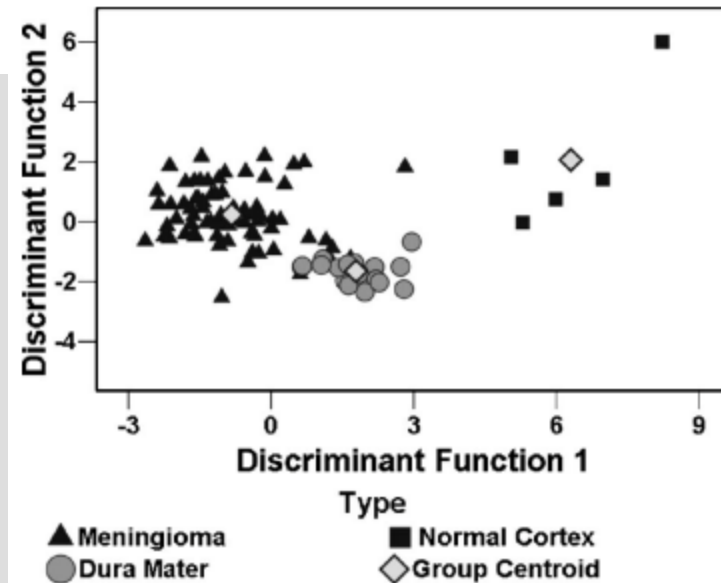
Fluorescence lifetime spectroscopy



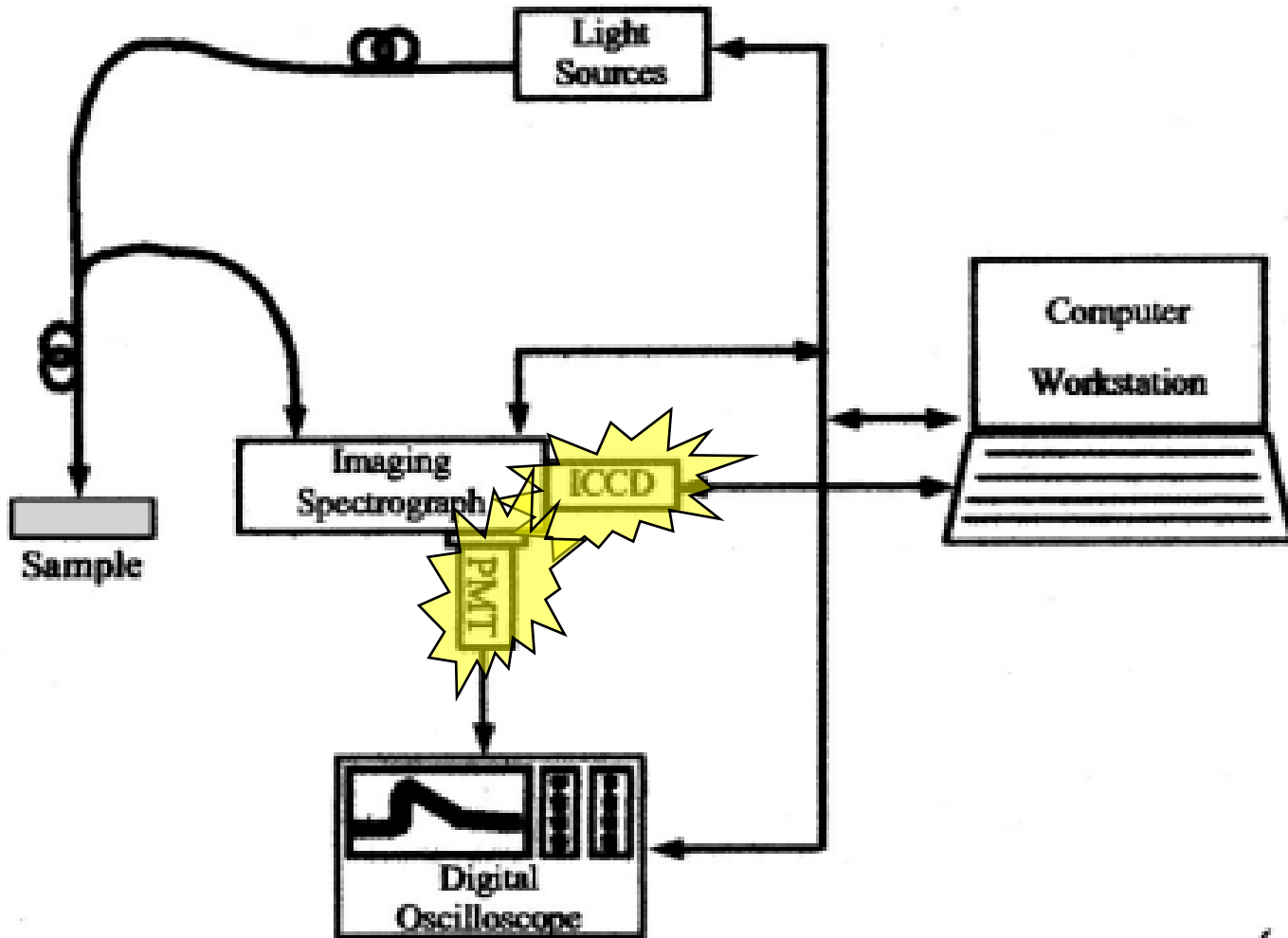
(b)

brain tissue

Butte et al., "Diagnosis of meningioma by time-resolved fluorescence spectroscopy," Journal of Biomedical Optics 10(6), 064026 (November/December 2005).

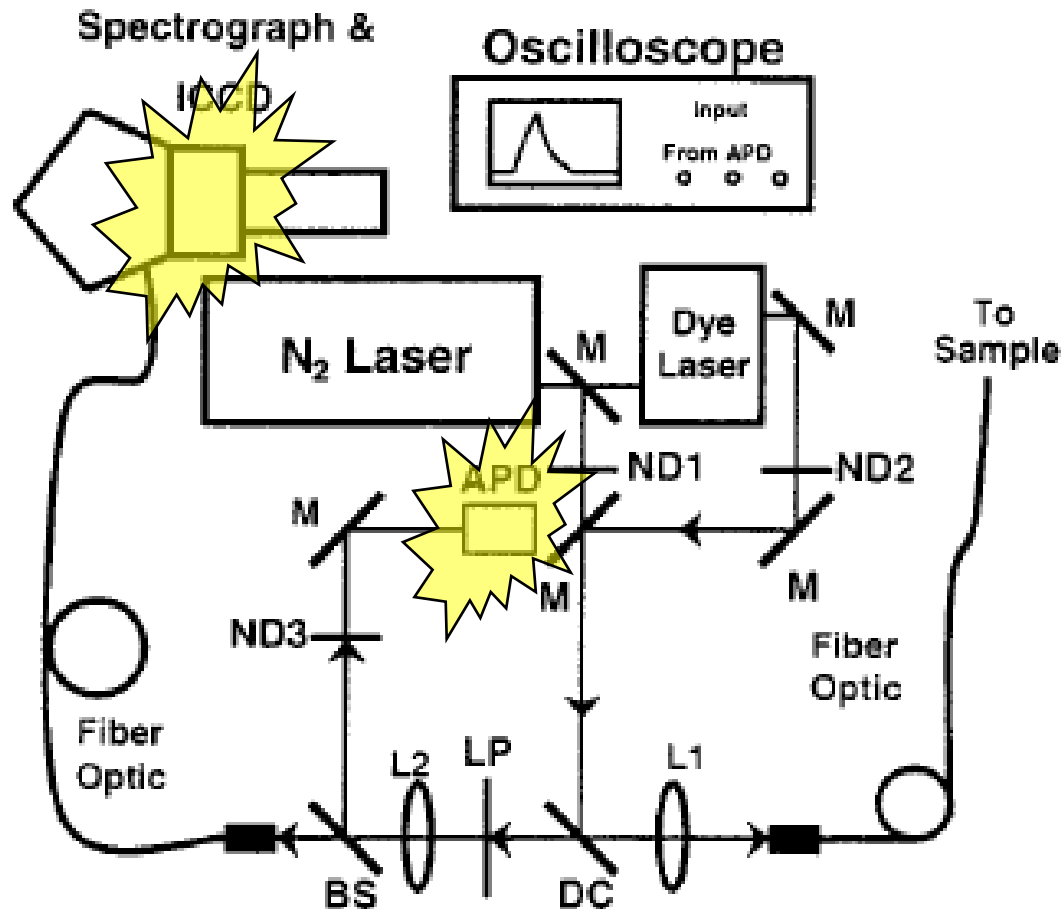


Instrumentation for temporal fluorescence



(a)

Same idea, different group!



Design and development of a rapid acquisition laser-based fluorometer with simultaneous spectral and temporal resolution

Jonathan D. Pitts and Mary-Ann Mycek^{a)}

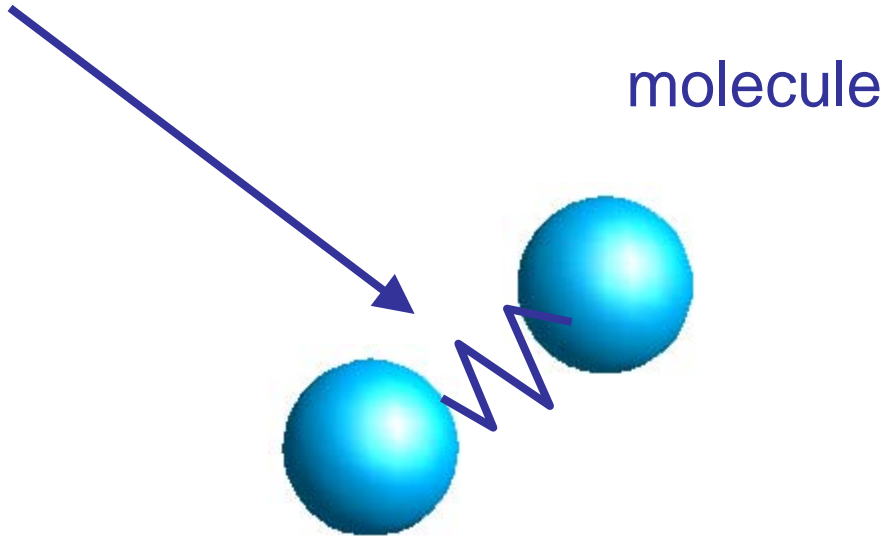
Department of Physics and Astronomy, Dartmouth College, 6127 Wilder Laboratory, Hanover, New Hampshire 03755

Fluorescence lifetime: goals, requirements

- Distinguish benign from malignant tumor tissue
- Record at many wavelengths
- Time resolution *required* to few psec
- *Desirable* to record at many locations (imaging)

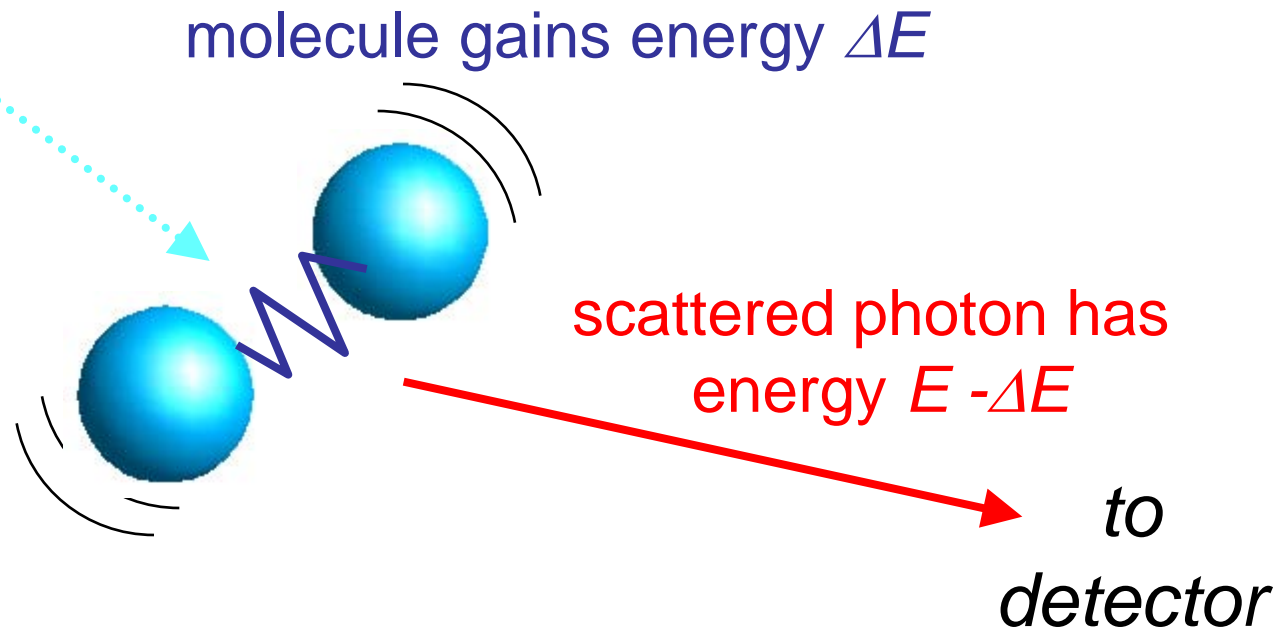
Area #3: Raman spectroscopy

incident photon
with energy E

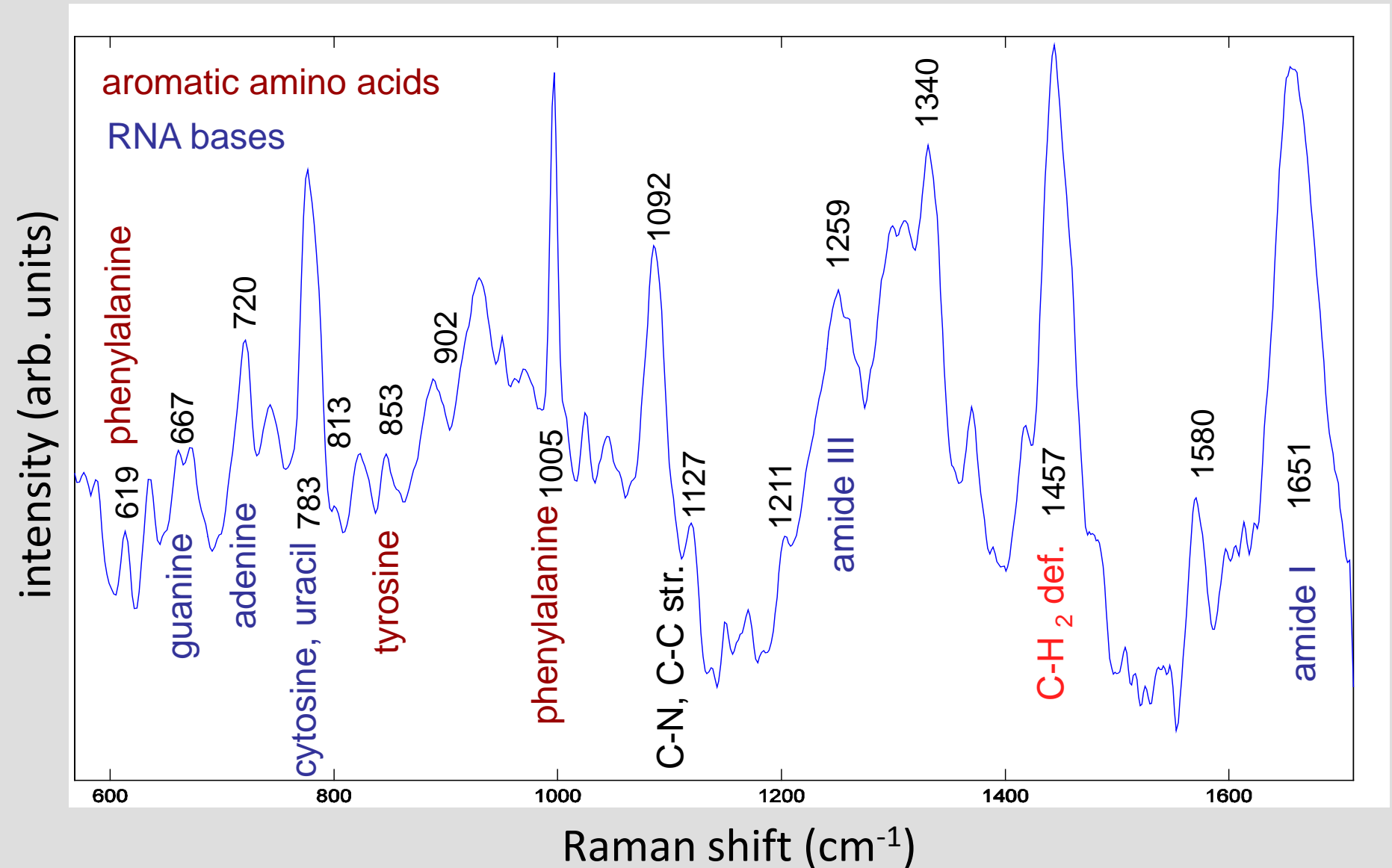


Raman spectroscopy

incident photon
with energy E



Raman spectrum of immune cell



Detectors for Raman spectroscopy

- Thermoelectrically-cooled CCD array detectors
- Sensitive out to ~ 1150 nm, limited by **Si bandgap**
- 25 micron square pixels
- typical dimensions, 256 x 1024 pixels



Princeton Instruments PIXIS CCD

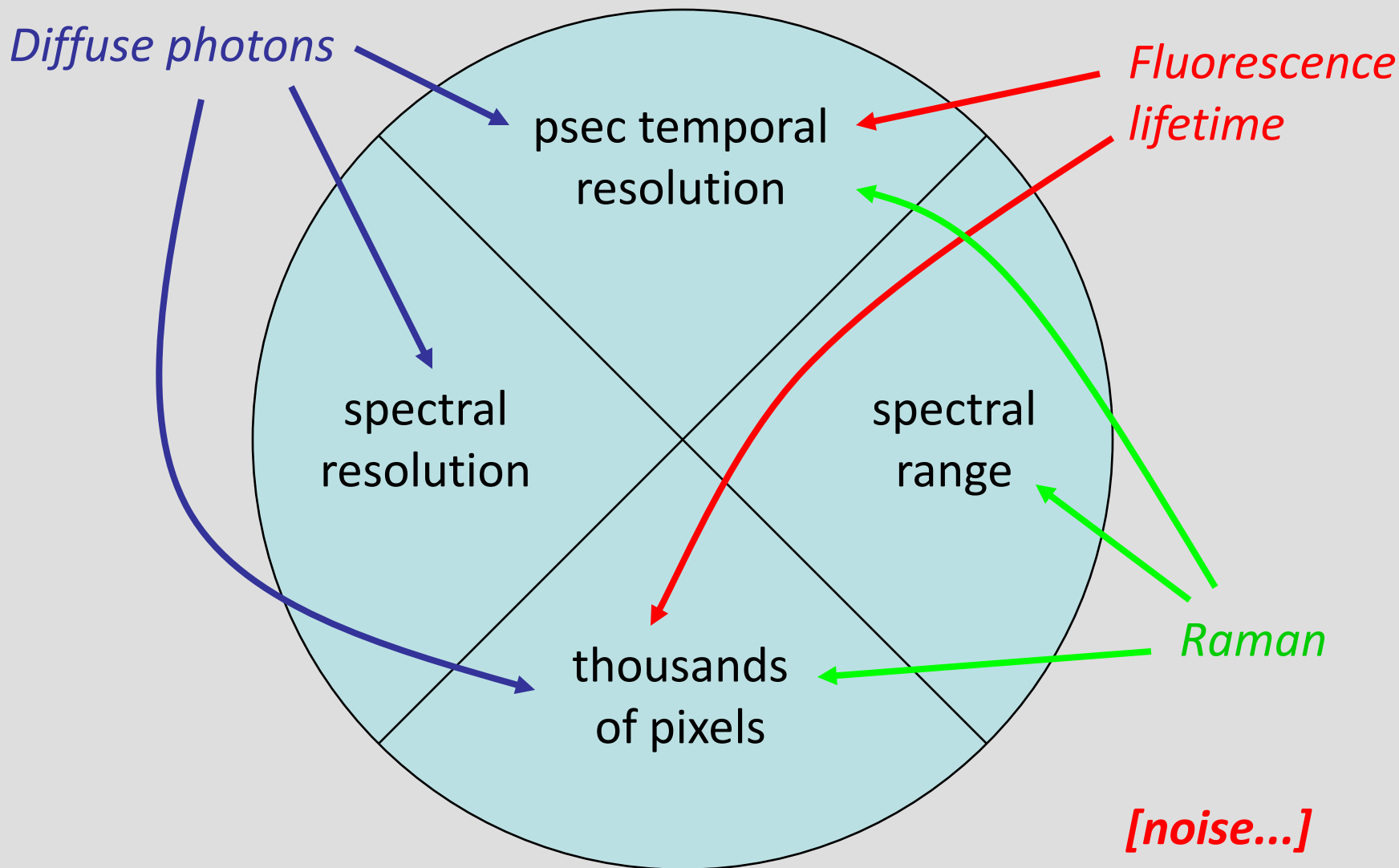
Raman spectroscopy: goals, requirements



- Distinguish one cell type/state from another
- Quantify chemical levels in biofluids (e.g. blood)
- Yes, distinguish cancer from non-cancer

- Record at many wavelengths
- Long acquisition times (sec-minutes)
- **Necessary** to wavelength-tune down the fluorescence
- **Desirable** to time-gate away the fluorescence (intensified CCD or more exotic gating)

Benefits of QLIDs for biomedical optics



Summary

- biomedical spectroscopy: characterize tissue, biofluids, cells
- frequently in near-IR
- multiple factors driving sub-nsec time resolution
- many-many-channel sensing: a game-changer
- get past the Si bandgap cutoff
- spectral resolution at each pixel: good for diffuse spectroscopy

Questions?