



Multiscale Imaging in Cancer Research

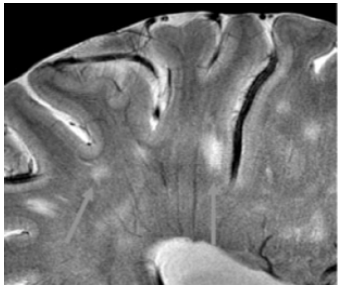
Sarah E Bohndiek PhD

10/5/2016



CAMBRIDGE
INSTITUTE





Radiowave
 $> 1 \text{ m}$



Visible/NIR
 $400 - 1,000 \text{ nm}$

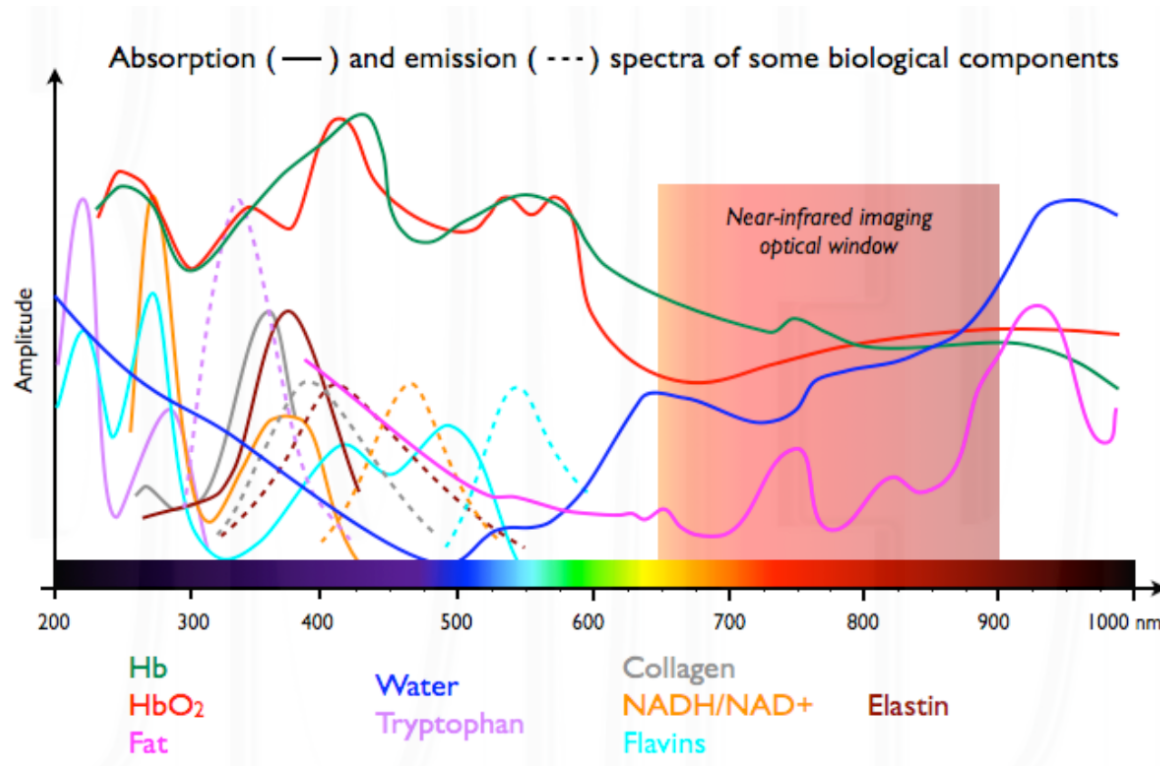


X-ray
 $10^{-8} - 10^{-11} \text{ m}$



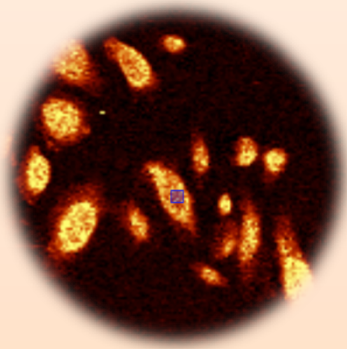
Gamma
 $< 10^{-11} \text{ m}$

Optical imaging gives high intrinsic contrast for understanding cancer metabolism



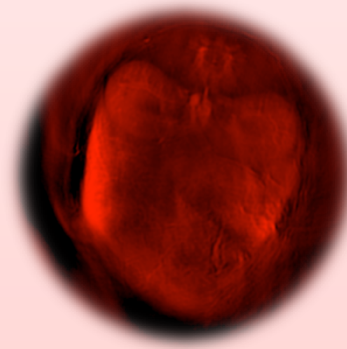
- **Penetration depth is limited**
- **Current imaging techniques lack multiplexing**

Microscopy



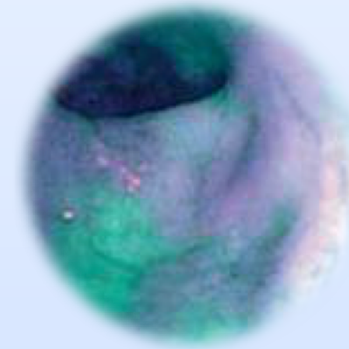
In vitro

Macroscopy



In vivo

Endoscopy



Translational

Interactions of light and tissue (experimental and theoretical)

Technology innovation and instrument development

Understanding the role of oxygen in cancer

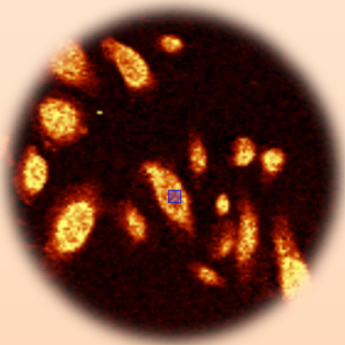
Technical Validation: Phantom trials

Biological Validation: Preclinical trials (cells & mice)

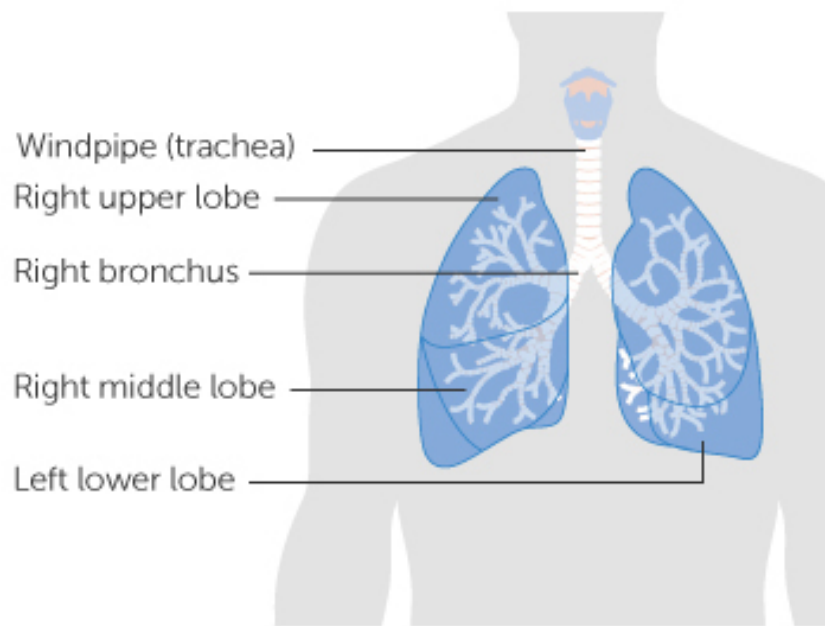
Biological Validation: Clinical trials (patients)

Lung cancer is a major cause of cancer death related to smoking

Microscopy



In vitro



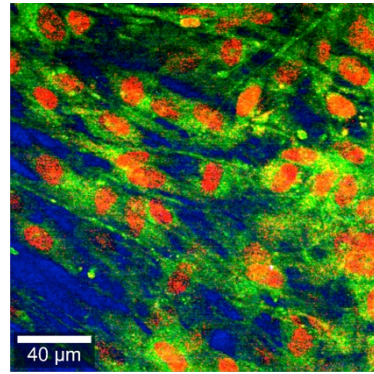
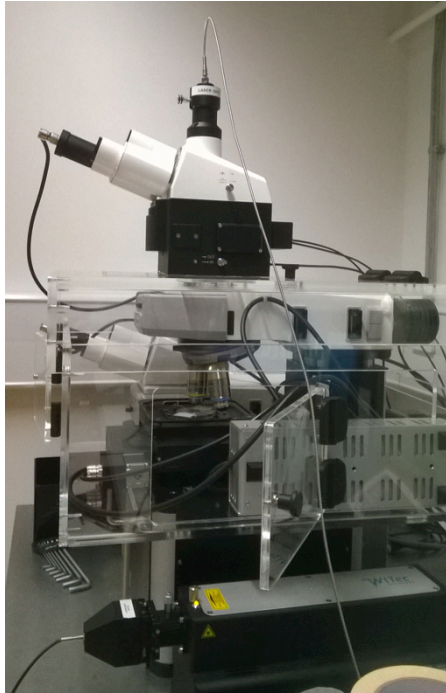
Cancer Research UK

Number of new cases per year



Number surviving after 10 years



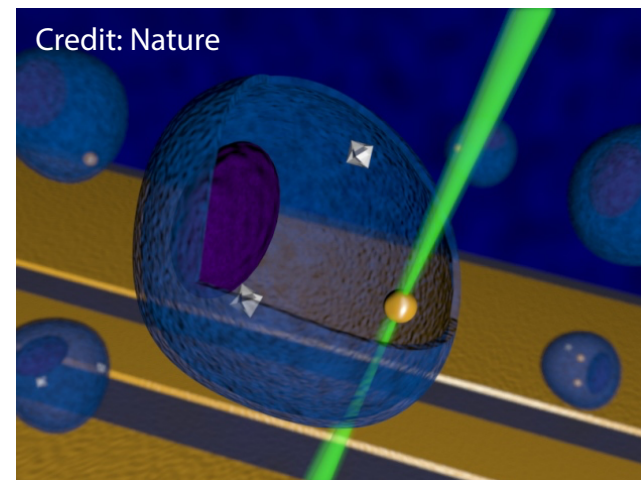


*Label free live
cell imaging of
oxidative stress
with Raman
spectroscopy*

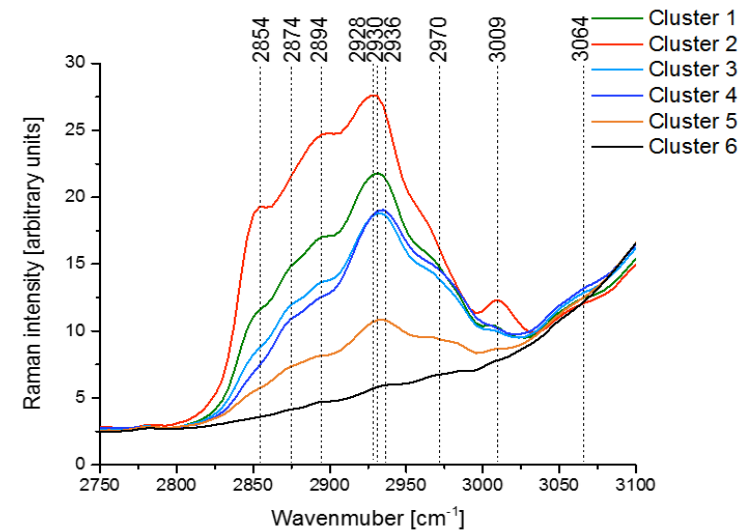
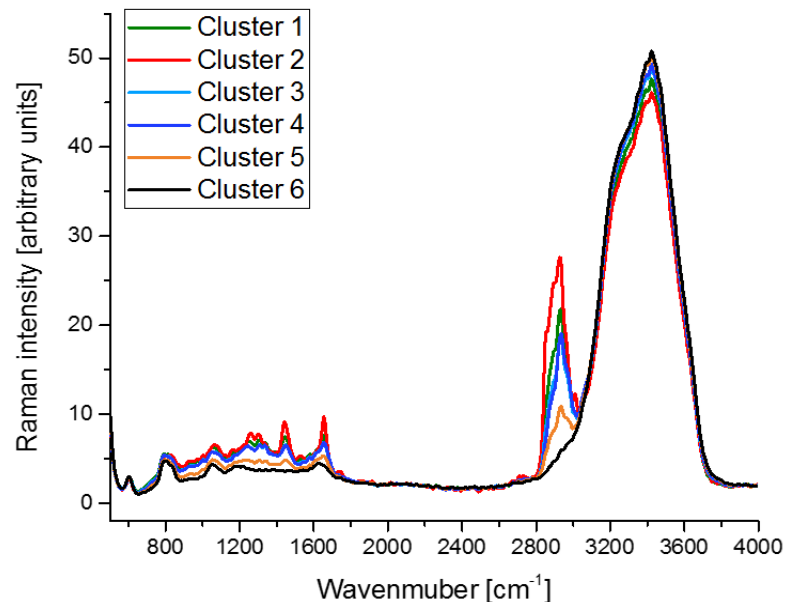
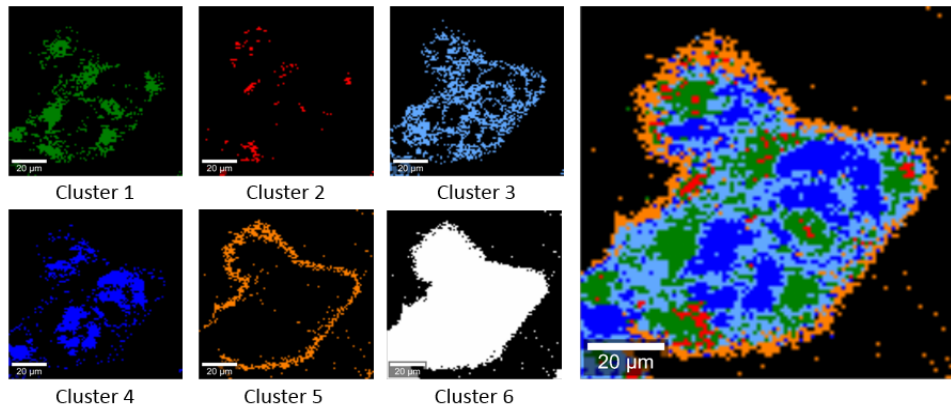
Collaboration with:

Prof. Bruce Ponder (Oncology)
and
Dr Robert Rintoul (Papworth)

*Intracellular
sensing*

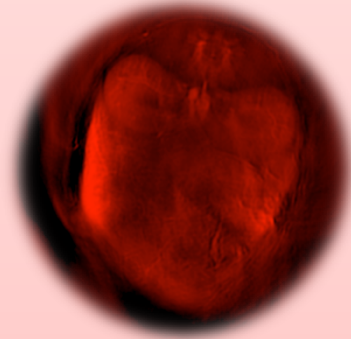


Raman spectroscopy enables spectral identification of cancer cells

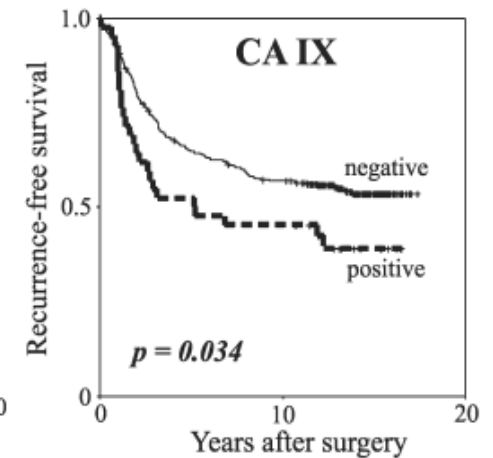
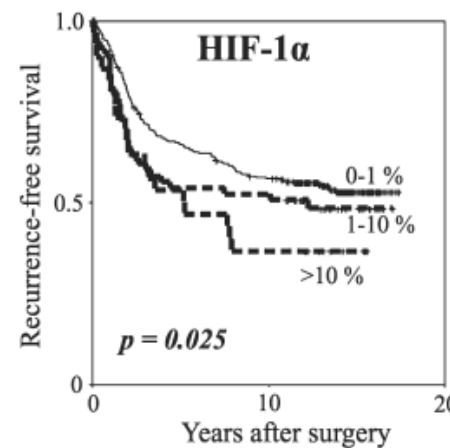
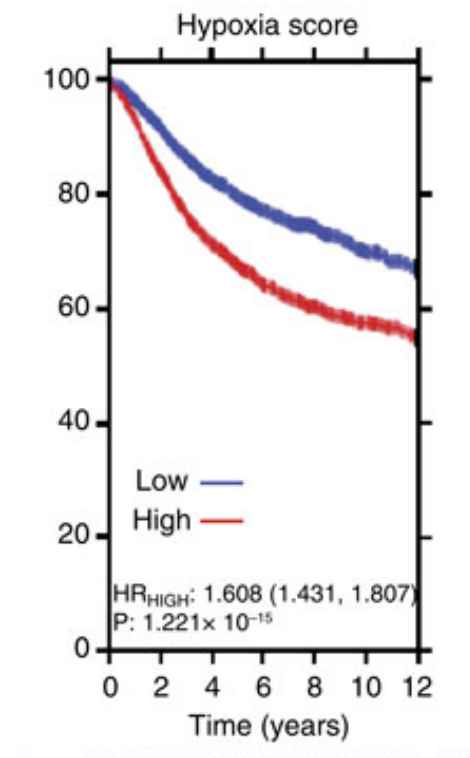


Prognosis of cancers in hormone sensitive tissues is linked to hypoxia

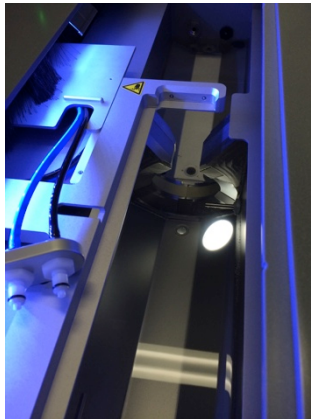
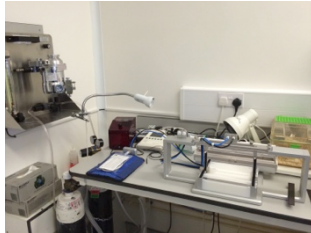
Macroscopy



In vivo

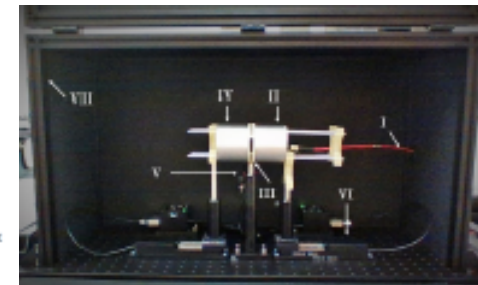
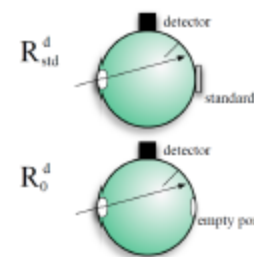
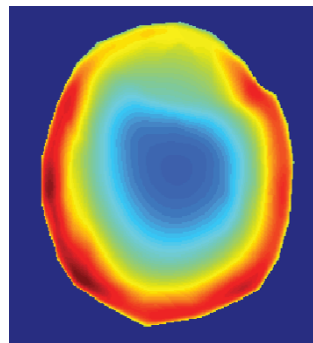


Imaging blood oxygenation and tissue hypoxia with optoacoustic tomography



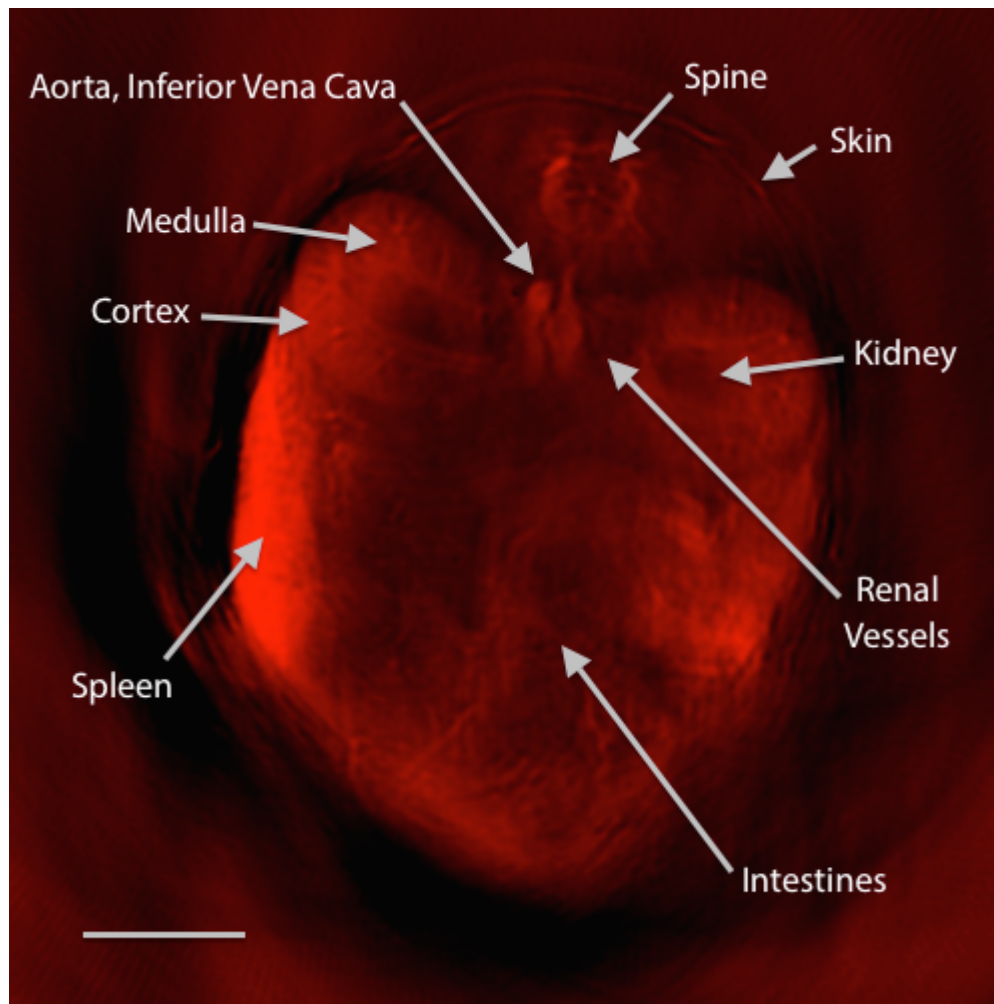
Collaboration with
Dept of Radiology:

Prof. Fiona Gilbert
and
Dr Oshaani Abeyakoon

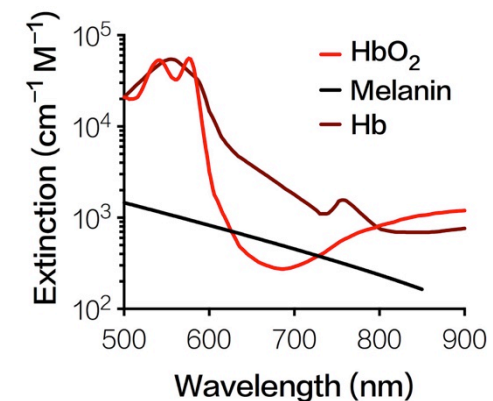


Understanding and compensating tissue optical properties

Optoacoustic tomography enables deep tissue imaging of optical absorption

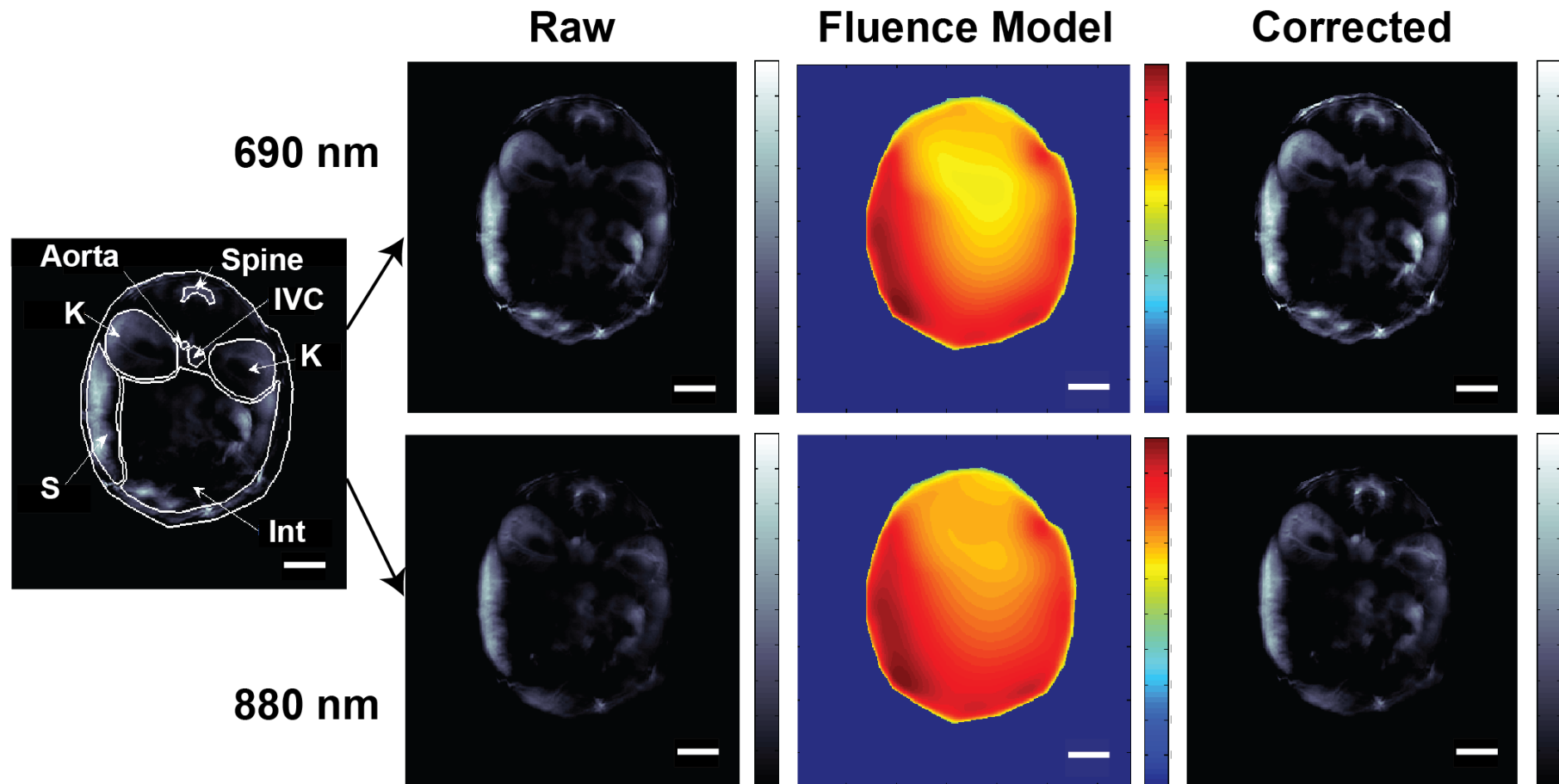


- Pulsed laser light excites tissue in NIR range
- Resulting ultrasound wave detected
- Increases penetration depth up to ~3 cm
- Spatial resolution ~ 150 μ m
- Temporal resolution ~ 10fps



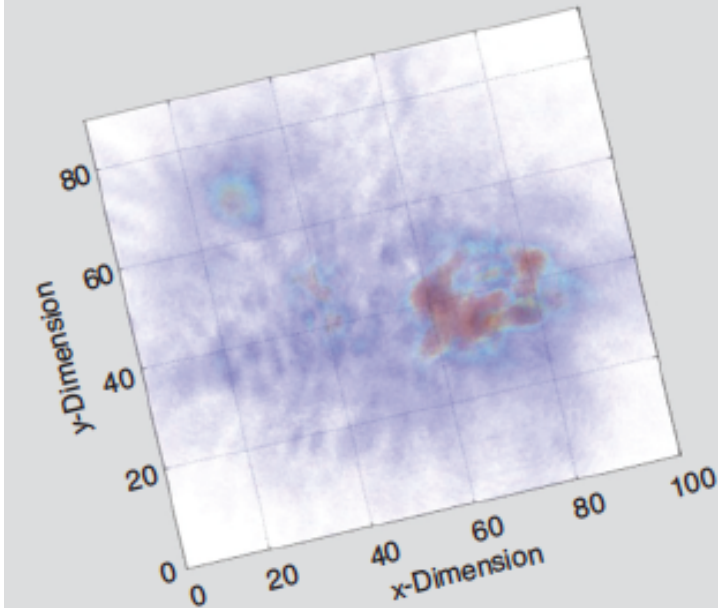
Oxyhaemoglobin weighted slice through the kidney. Scale 3 mm.

Applying light fluence corrections to *in vivo* improves the visibility of deep tissue structures

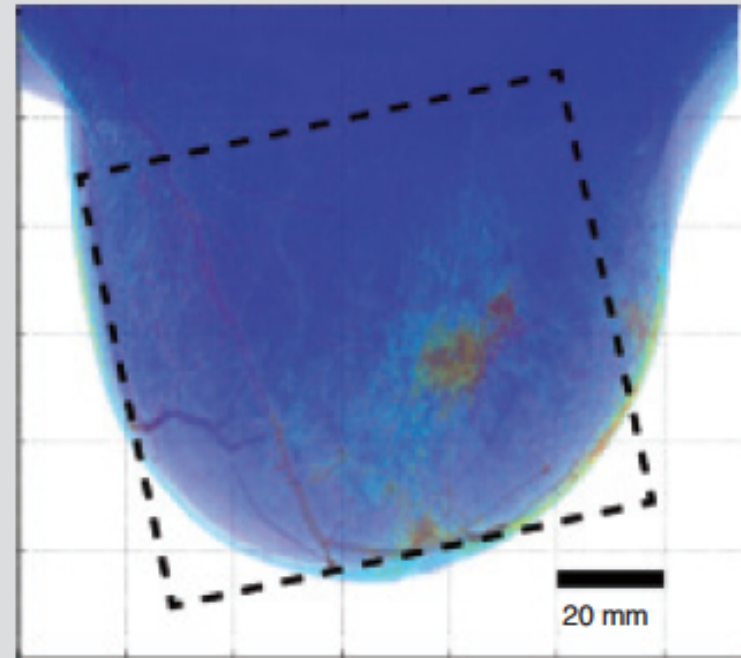


Clinical trials in breast cancer are underway worldwide and starting in Cambridge

Heijblom *et al* (2015) *IEEE Pulse* May/June 44



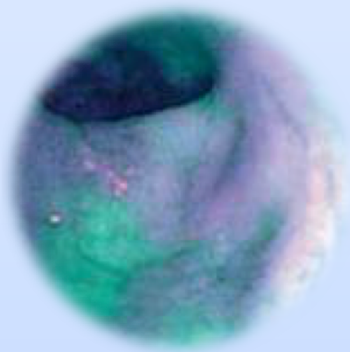
(a) PA Image



(b) MR Image

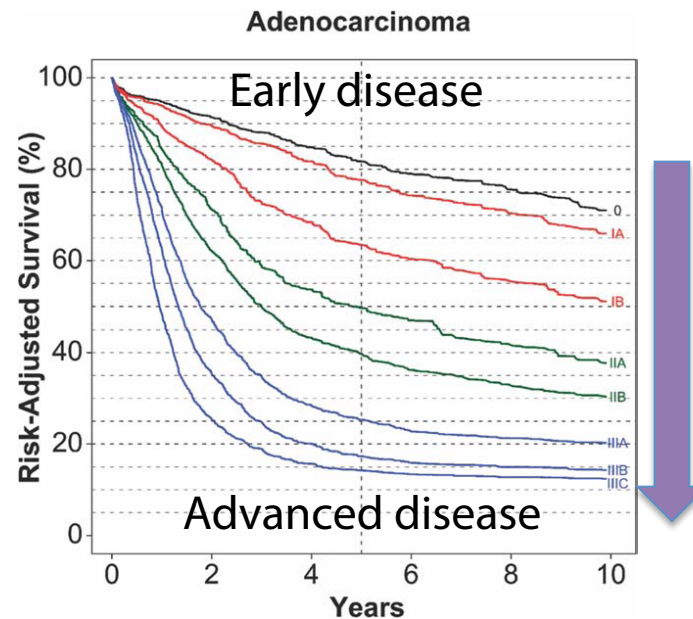
- Healthy volunteer study to confirm normal fluctuations in breast vascular density and oxygenation
- Breast cancer staging will assess prognostic potential compared to standard of care imaging

Endoscopy



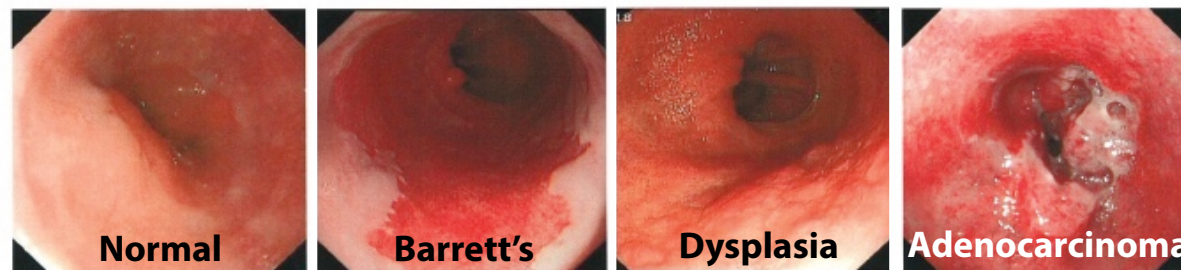
Translational

Barrett's patients undergo surveillance to improve early detection

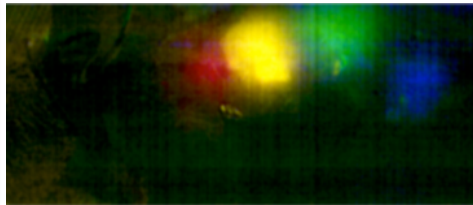


Standard of care:

- White Light Endoscopy
- Seattle Protocol Biopsy



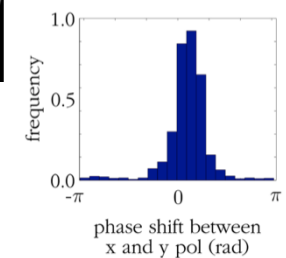
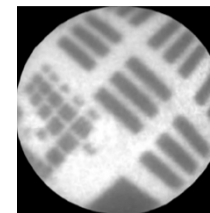
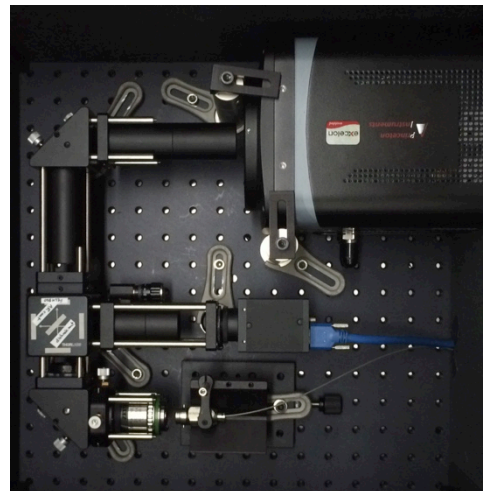
*Hyperspectral imaging
for multiplexed
fluorescence detection*



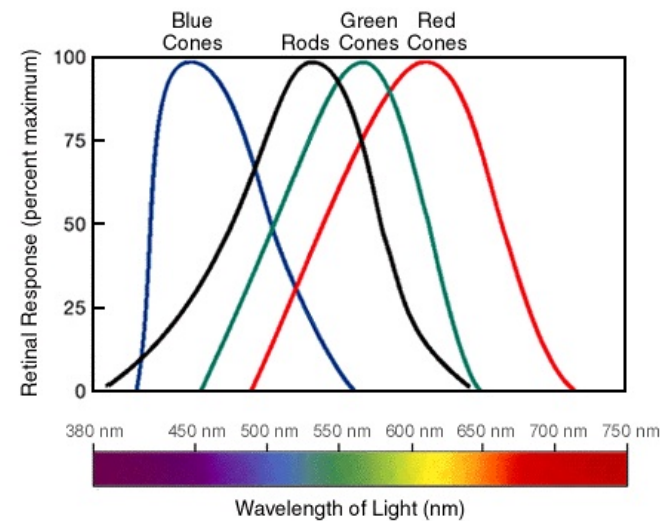
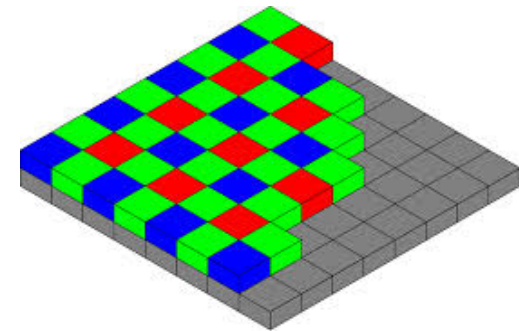
Collaboration with
MRC Cancer Unit:

Prof. Rebecca Fitzgerald
and
Dr Massi di Pietro

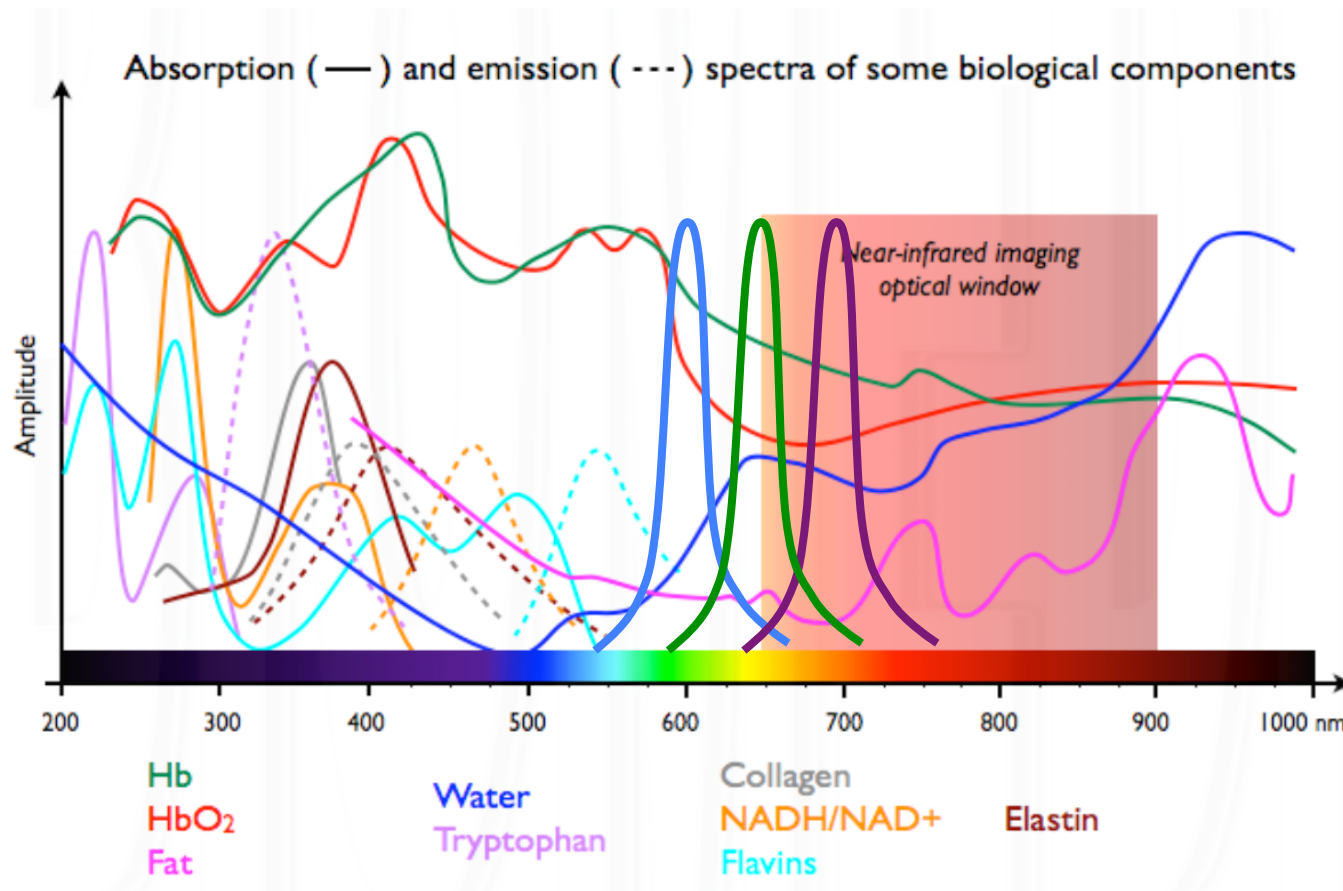
*Near infrared and coherent endoscopy for early
cancer detection*



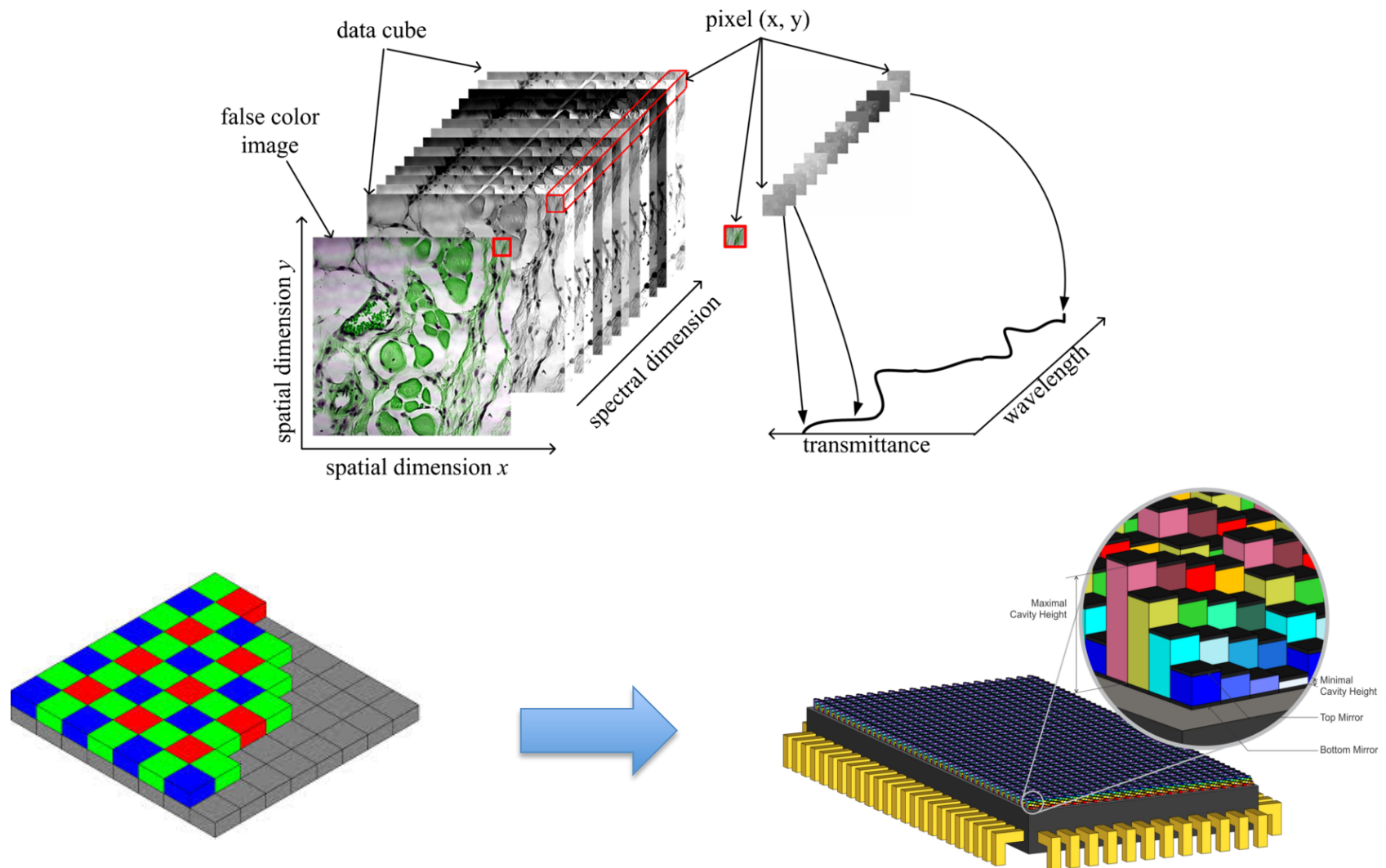
The use of light in endoscopy measures only amplitude



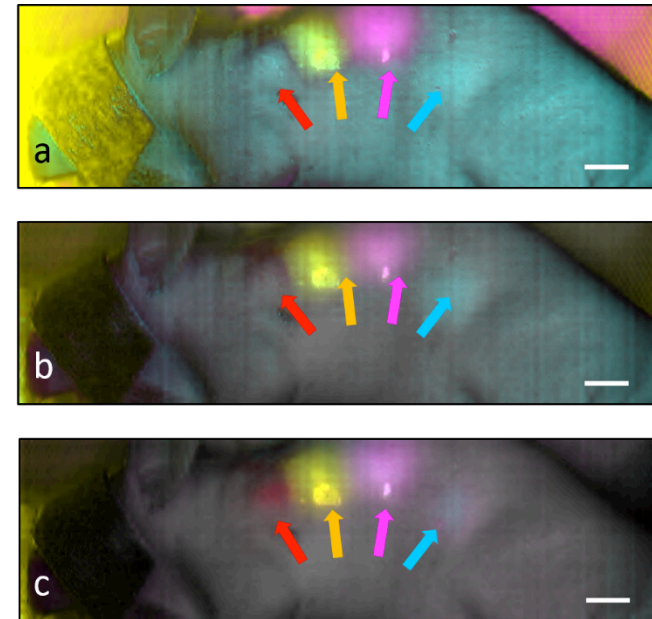
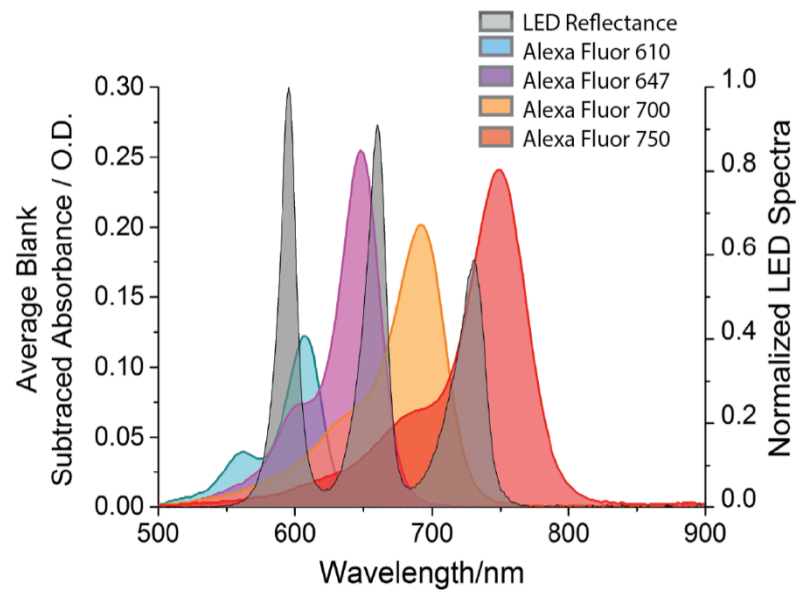
Spectral imaging can resolve information from absorption and fluorescence emission



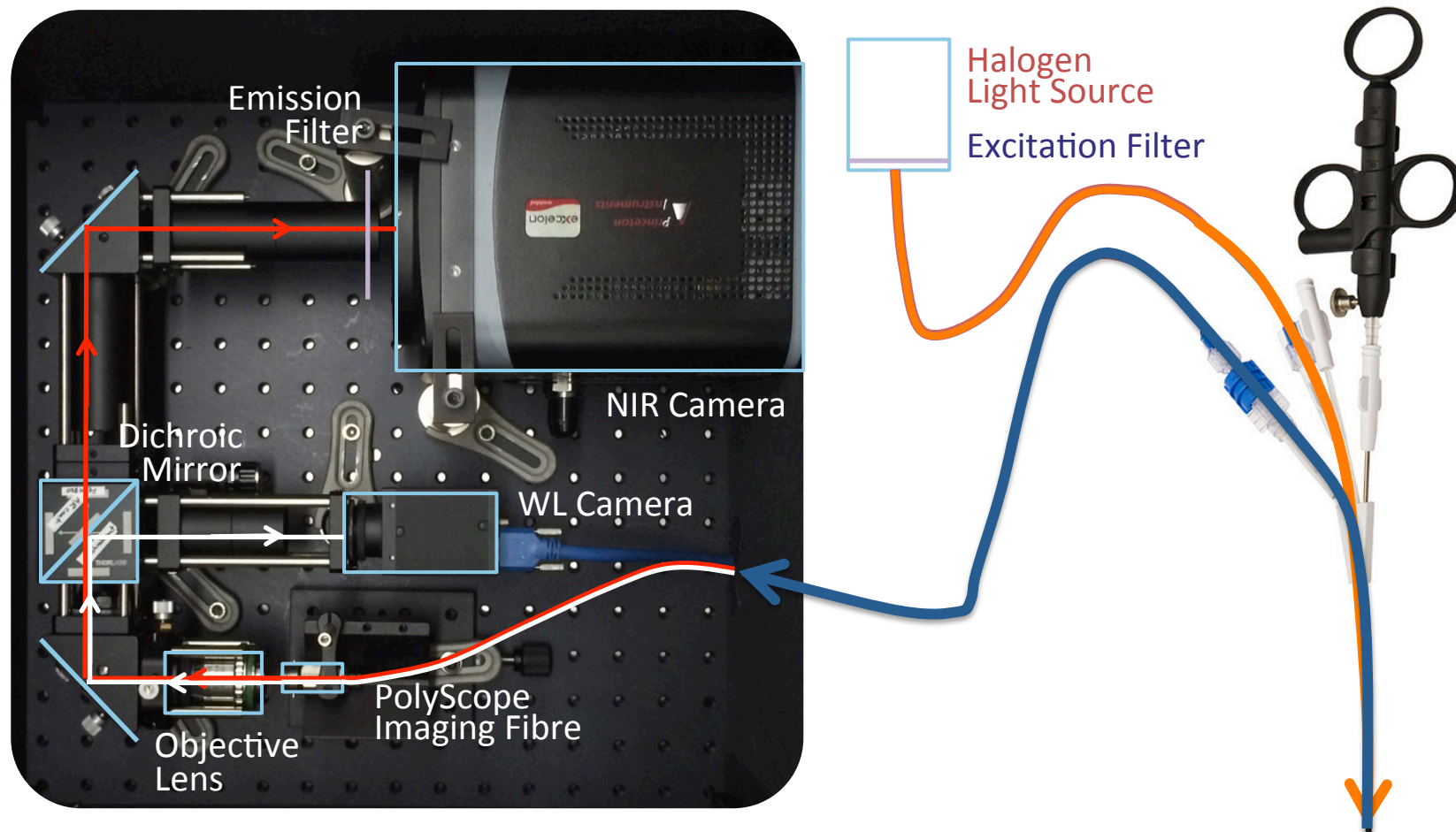
Hyperspectral imaging combines both spatial and spectral information



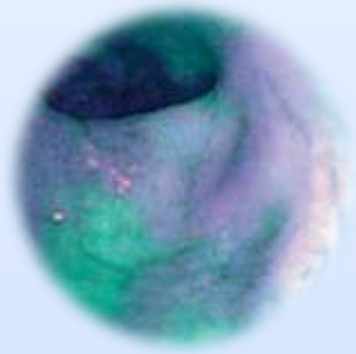
Initial data indicates that we can easily resolve at least four sources of fluorescence *in vivo*



We are using a generic platform for testing novel endoscopic technologies



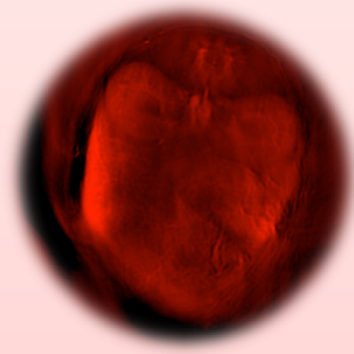
Endoscopy



Translational



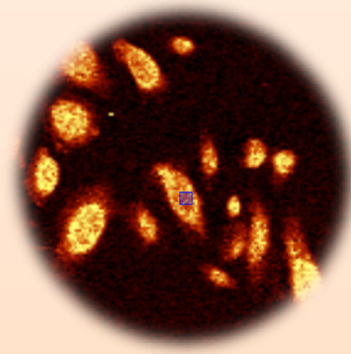
Macroscopy



In vivo



Microscopy



In vitro



Acknowledgements

The Bohndiek Lab

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Visitors

Dr Frederic Brochu (@HEP)

Lung Cancer

Prof. Bruce Ponder
Dr Robert Rintoul

Breast Cancer

Prof. Fiona Gilbert
Dr Oshi Abeyakoon

Oesophageal Cancer

Prof. Rebecca Fitzgerald
Dr Massi di Pietro

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Dr Andre Neves

Engineering @ Cam

Prof. Tim Wilkinson
Dr George Gordon

iThera Medical

Stefan Morscher
Christian Wiest



CRUK & EPSRC Cancer Imaging Centre
in Cambridge and Manchester



CANCER
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