

# Opto-acoustic Relative-Colorization of Blood Oxygen Saturation in Biologically Relevant Phantoms

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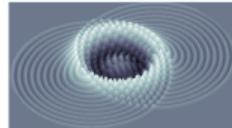
<sup>3</sup>Tomowave Laboratories, Houston TX, USA

SPIE Photonics West

February 5, 2019

The logo for Ryerson University, featuring the text "Ryerson University" in white on a blue rectangular background, with a yellow vertical bar to the right.

**Tomowave**



Laboratories, Inc.

# Opto-acoustic Imaging

**Opto-Acoustic (OA) Imaging Technology**  
Imagio™ Breast Imaging System

The diagram illustrates the Imagio™ Breast Imaging System. A probe is shown on the right, emitting acoustic waves (represented by red and blue concentric arcs) into a breast. Two lesions are identified: a malignant lesion and a benign lesion. The malignant lesion is characterized by irregular vascularity and reduced oxygen saturation, while the benign lesion has normal vascularity and normal oxygen saturation. The probe is connected to a cable with blue and white sections.

**malignant lesion**  
irregular vascularity  
reduced oxygen saturation

**benign lesion**  
normal vascularity  
normal oxygen saturation

- developed for **breast cancer diagnosis** to distinguish benign from malignant lesions and potentially reduce need for biopsy
- **blood oxygen saturation** and **vascularity** of breast tissue is detected with **acoustic waves** and **dual optical wavelengths**
- **safe to use** because there are no injected contrast agents and no ionizing radiation is required

# Pivotal Study

findings from clinical study of 2100 subjects



**OA improved specificity** to distinguish benign from malignant lesions

(source: Neuschler, E. et al. "A Pivotal Study of Optoacoustic Imaging to Diagnose Benign and Malignant Breast Masses: A New Evaluation Tool for Radiologists.". In: *Radiology* 287.2 (2018), pp. 398–412)

# Motivation

for today's talk

- **blood oxygen saturation ( $sO_2$ )** is an important parameter for assessing differences between benign and malignant lesions

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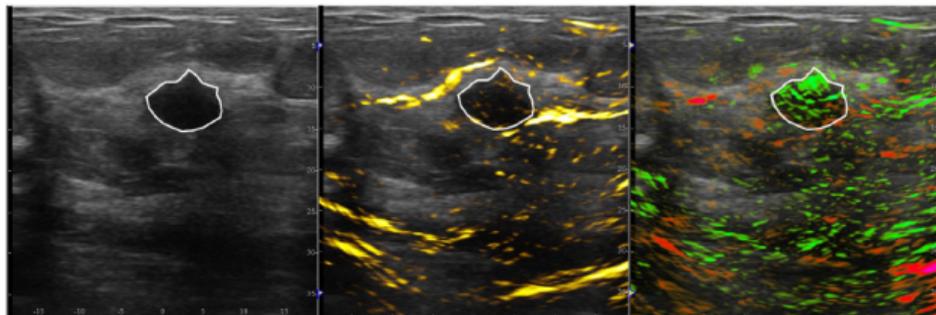
- **blood oxygen saturation ( $sO_2$ )** is an important parameter for assessing differences between benign and malignant lesions
- accurate diagnosis involves proper interpretation of **color mapped images** and an understanding of how images relate to tumor physiology

# Motivation

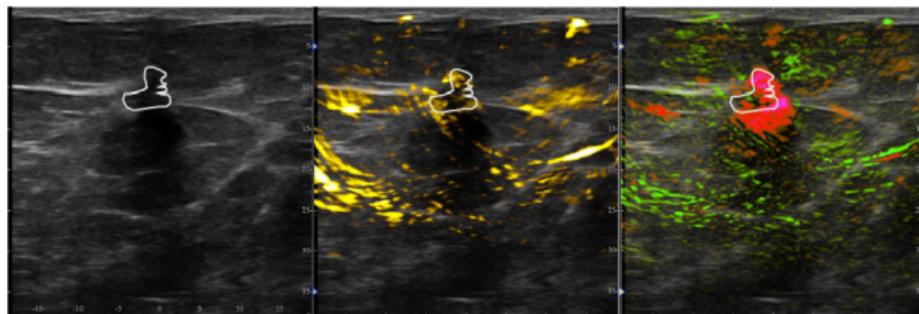
for today's talk

- **blood oxygen saturation (sO<sub>2</sub>)** is an important parameter for assessing differences between benign and malignant lesions
- accurate diagnosis involves proper interpretation of **color mapped images** and an understanding of how images relate to tumor physiology
- necessary to perform **system characterization** that relates RGB image colorization to known oxygen saturation values

# Opto-acoustic Images



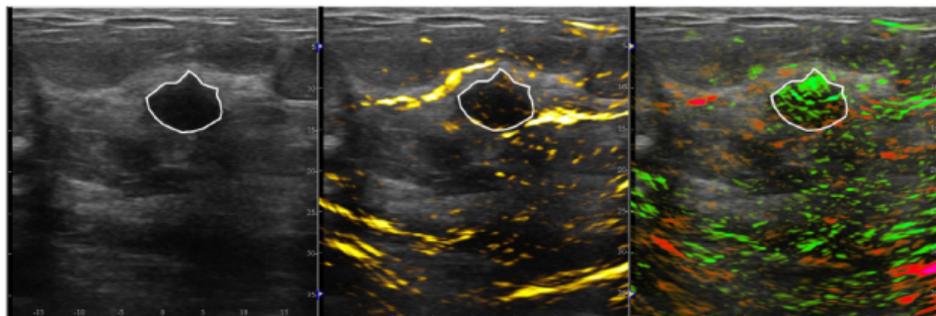
(a) **benign lesion** (fibroadenoma)



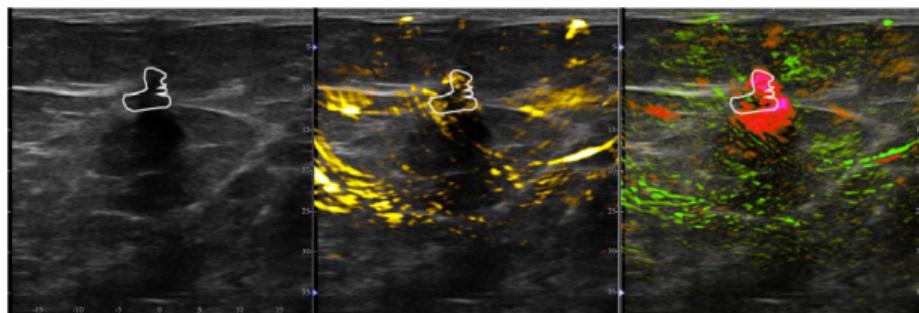
(b) **malignant lesion** (grade II invasive ductal carcinoma)

(source: Zalev, J. et al. "Opto-acoustic image fusion technology for diagnostic breast imaging in a feasibility study". In: *SPIE Medical Imaging*. International Society for Optics and Photonics. 2015, pp. 941909–941909)

# Opto-acoustic Images



(a) **benign lesion** (fibroadenoma)



(b) **malignant lesion** (grade II invasive ductal carcinoma)

total hemoglobin  
low high

relative oxygenation  
low high

(source: Zalev, J. et al. "Opto-acoustic image fusion technology for diagnostic breast imaging in a feasibility study". In: *SPIE Medical Imaging*. International Society for Optics and Photonics. 2015, pp. 941909–941909)

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  - perform quantitative measurement using **biologically relevant phantoms** and **bovine blood** at controlled oxygenation levels
  - assess the ability to distinguish **small differences in oxygenation** as the **oxygenation level is varied**

# Statistical Color Mapping

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- **analyzes statistics** of reference region of each image to automatically determine color scaling and offset parameters
- performs color mapping to display opto-acoustic contrast using **relative colorization**

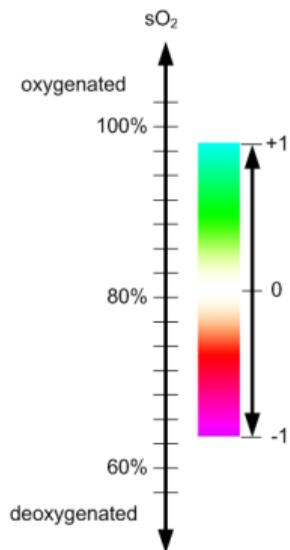
# Statistical Color Mapping

how it works

- **analyzes statistics** of reference region of each image to automatically determine color scaling and offset parameters
- performs color mapping to display opto-acoustic contrast using **relative colorization**
- visualizes structural and functional information from areas of strong blood oxygenation and total hemoglobin **relative to background tissue**

# Statistical Color Mapping

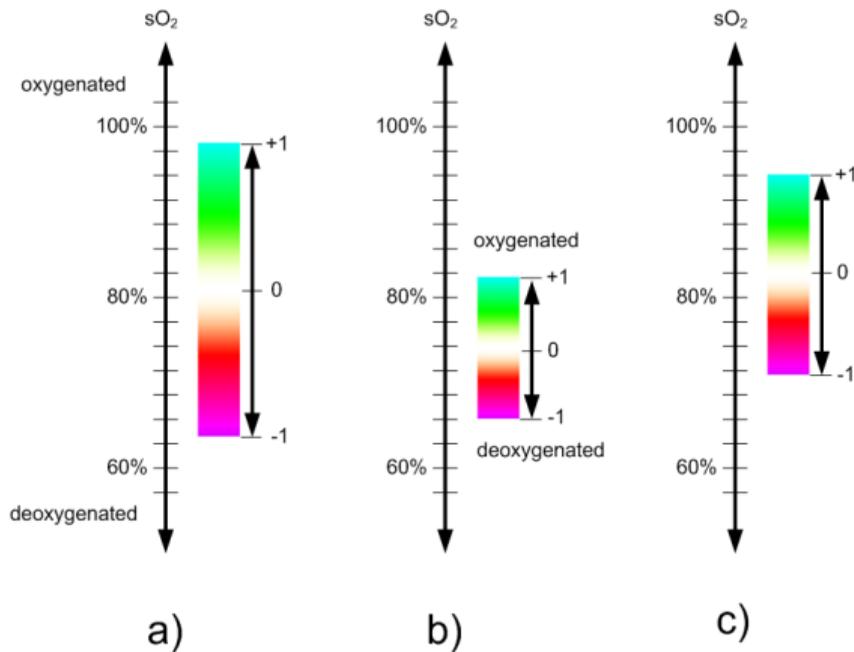
**relative colorization** of blood oxygen saturation



a)

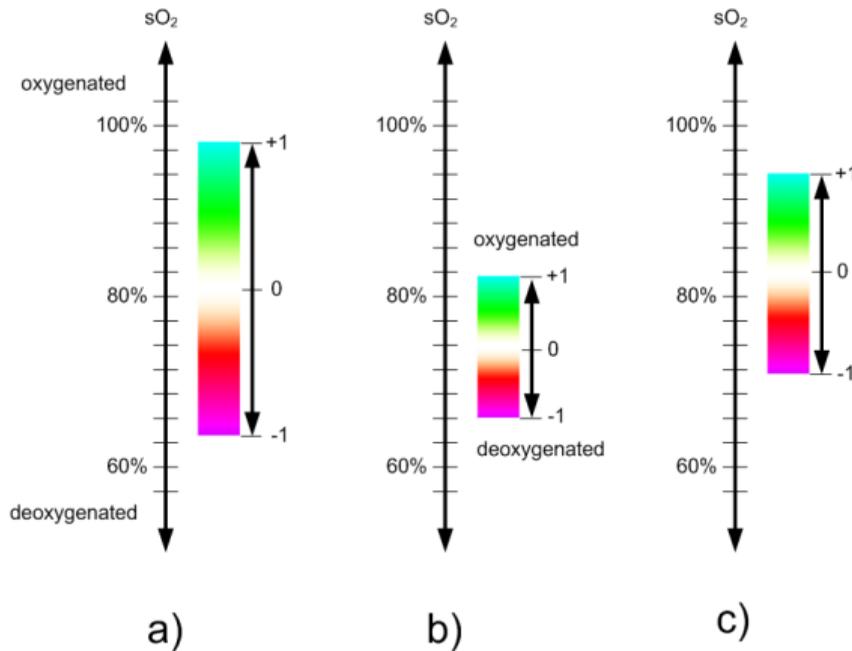
# Statistical Color Mapping

relative colorization of blood oxygen saturation



# Statistical Color Mapping

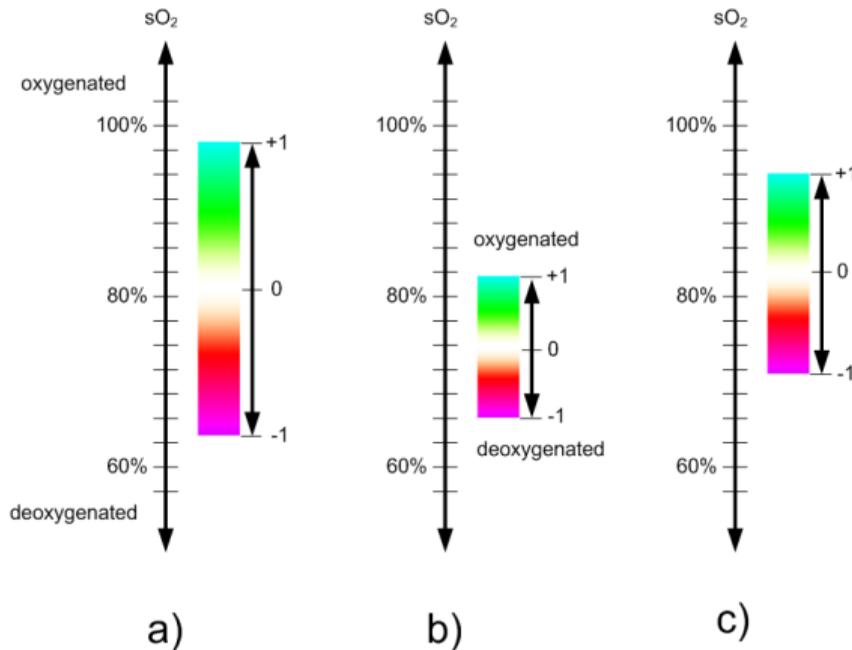
relative colorization of blood oxygen saturation



- color midpoint corresponds to average  $sO_2$  value of reference region

# Statistical Color Mapping

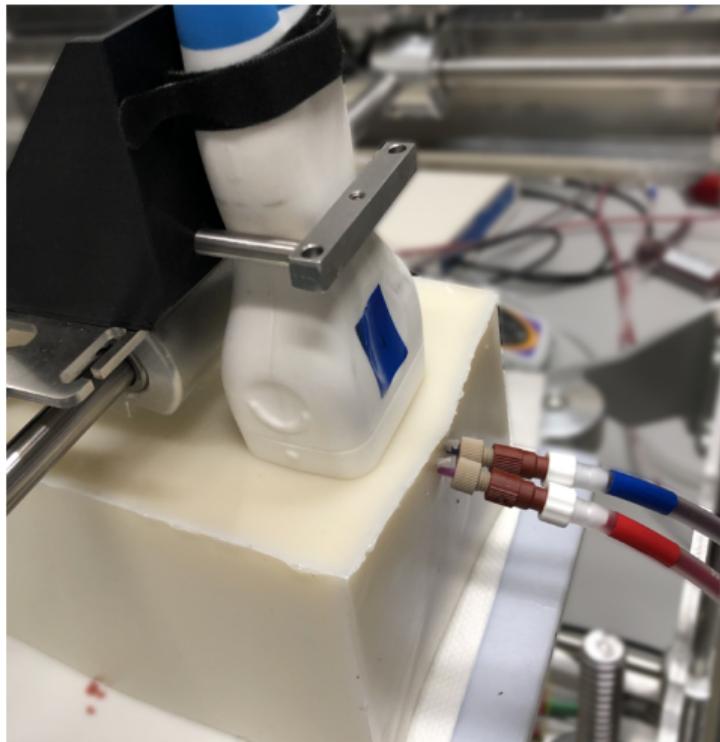
relative colorization of blood oxygen saturation



- color midpoint corresponds to average  $sO_2$  value of reference region
- color limits determined by  $sO_2$  variability of reference region

# Biologically Relevant Phantoms

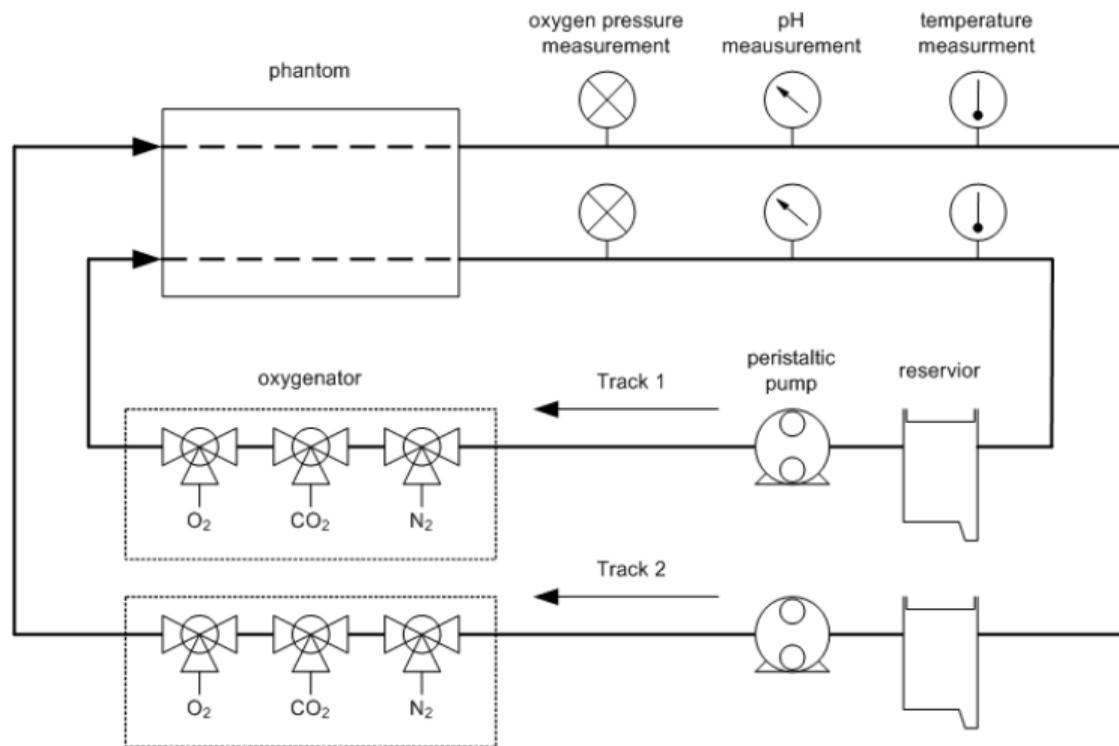
optical and acoustic properties emulate breast tissue



## Phantom properties

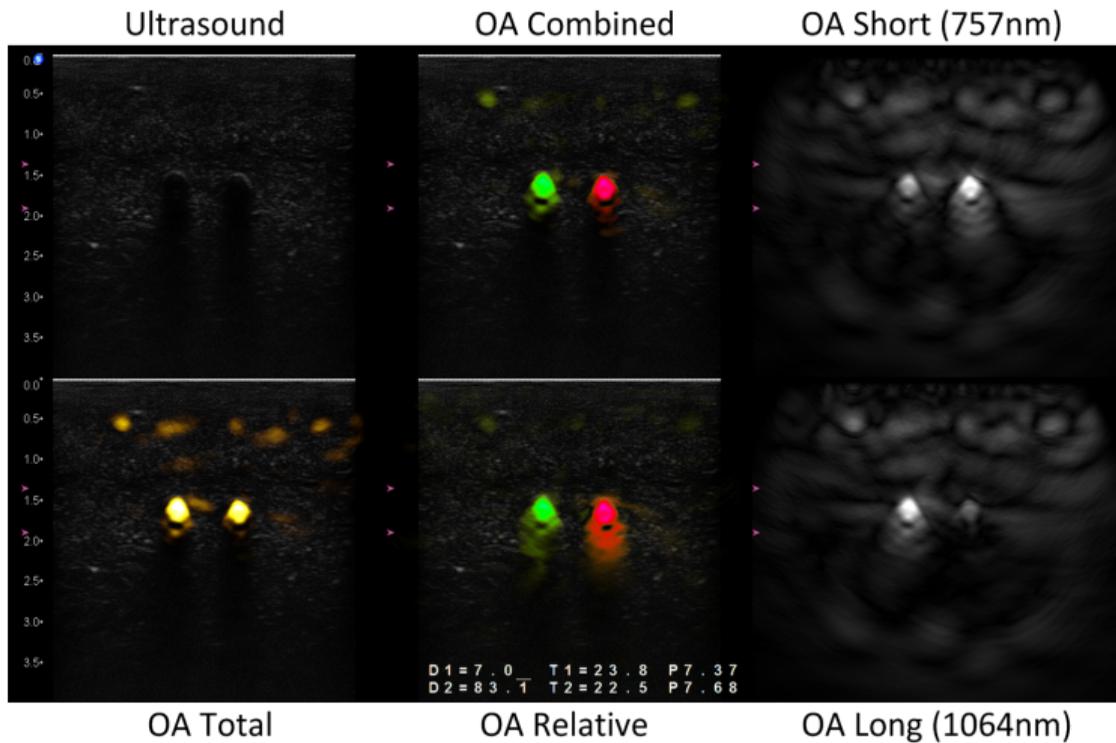
Material	PVCP (plastisol)	
<b>Optical Properties</b>	757nm	1064nm
Absorption $\mu_a$ (1/cm)	$0.47 \pm 0.10$	$0.82 \pm 0.38$
Scattering $\mu'_s$ (1/cm)	$9.18 \pm 0.06$	$5.89 \pm 1.35$
<b>Acoustic Properties</b>		
Speed of sound (m/s)	1477	
Attenuation (dB/cm)	0.8 – 8.2	
Vessel depth	15mm	
Vessel diameter	1.6mm	

# Experimental Setup

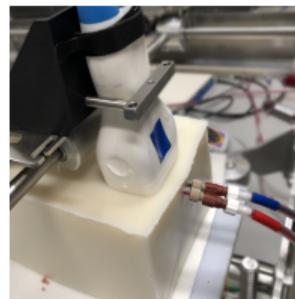
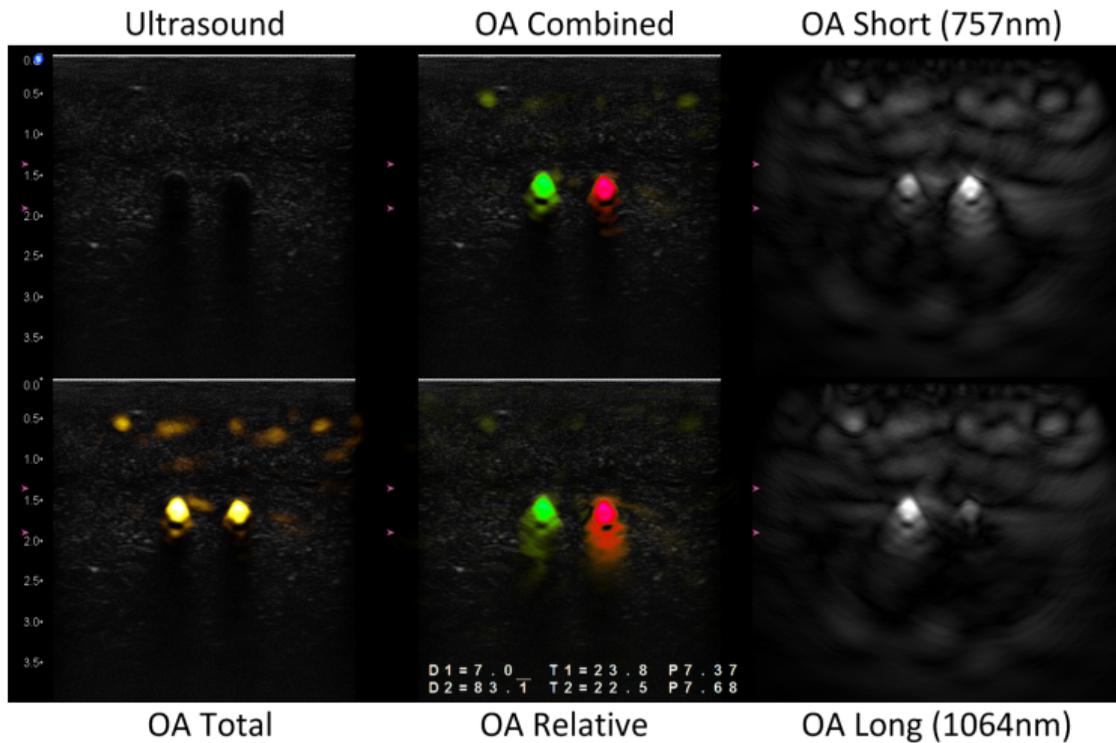


two independent tracks of bovine blood at controlled oxygenation levels

# Imagio Output



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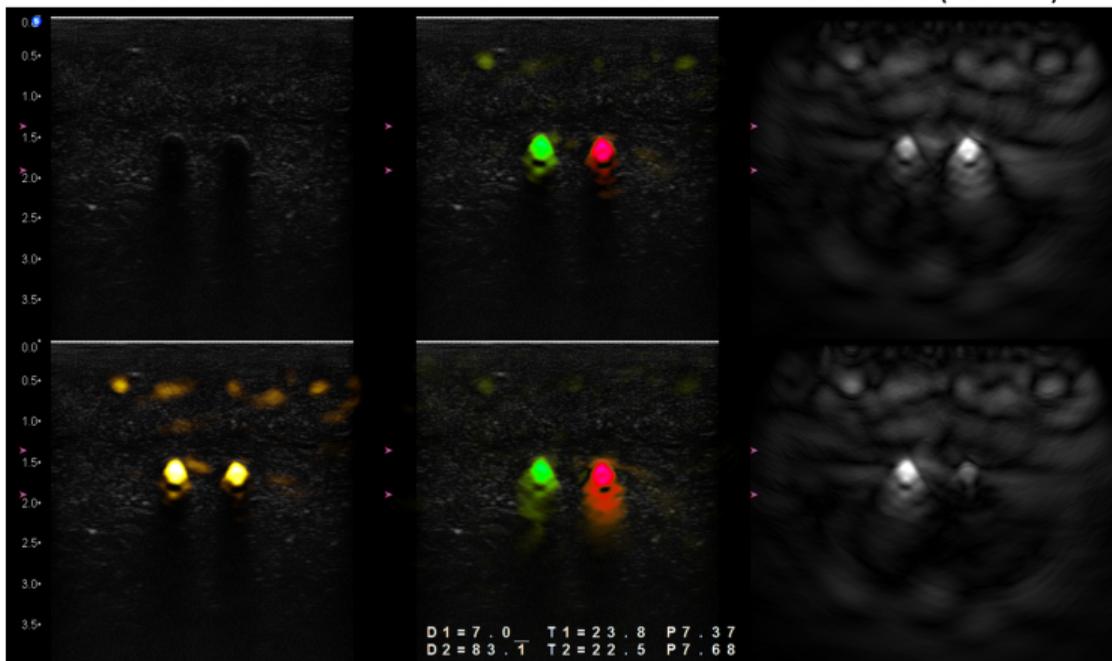
Ultrasound

OA Combined

OA Short (757nm)

total hemoglobin  
low high

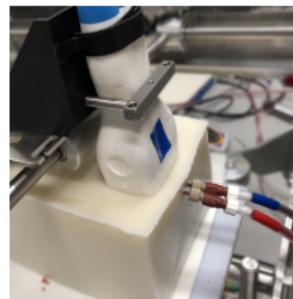
relative oxygenation  
low high



OA Total

OA Relative

OA Long (1064nm)



# Imagio Output

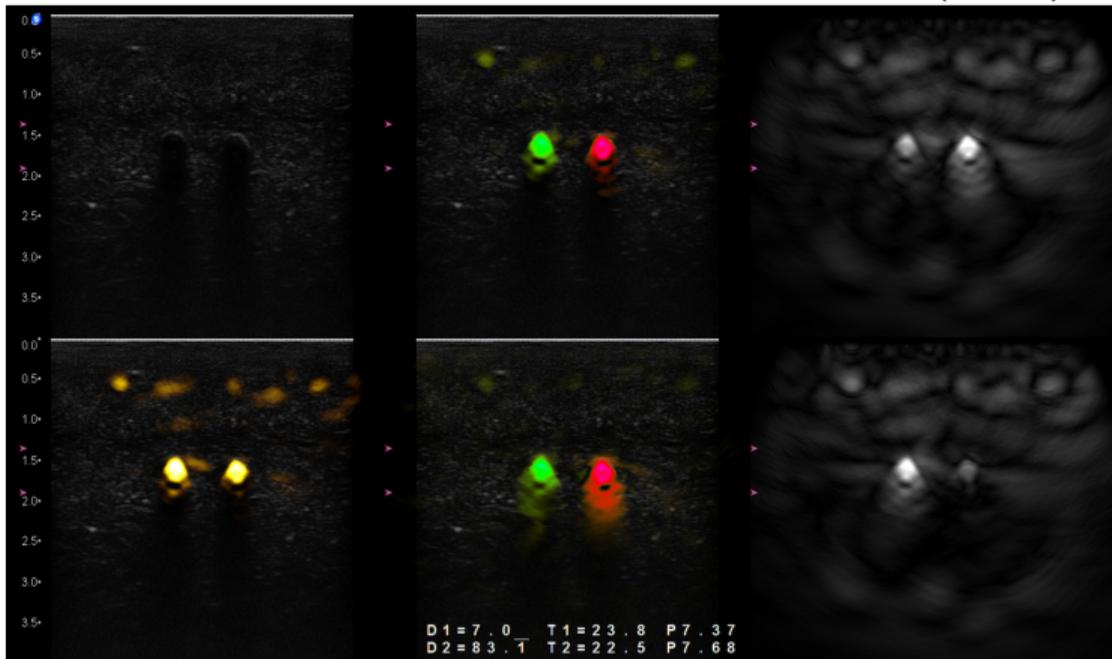
Ultrasound

OA Combined

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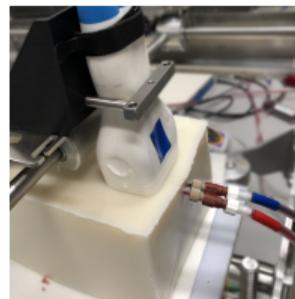
OA Total

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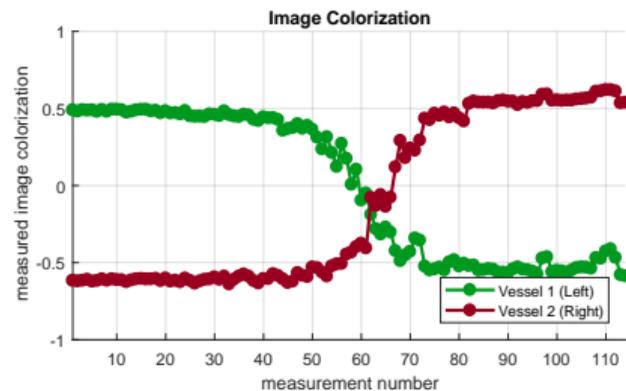
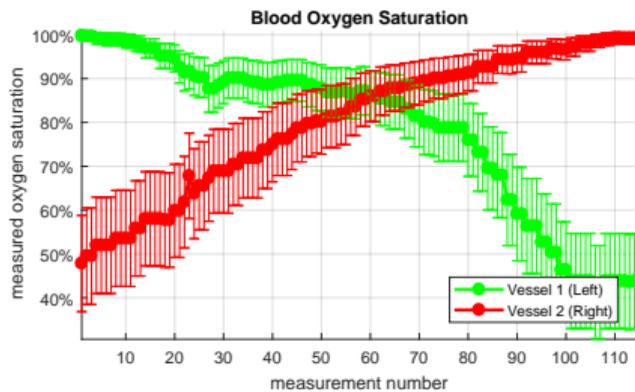
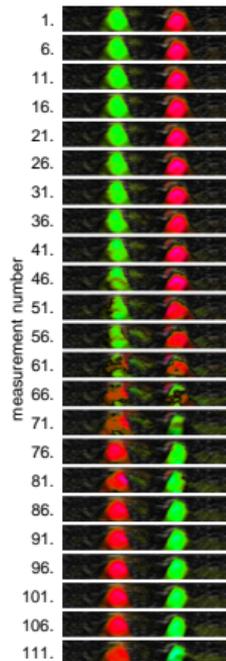
$[sO_2]_{\text{left}} = 99\%$ ,  $[sO_2]_{\text{right}} = 40\%$

arrow markers indicate reference region, which includes both vessels and phantom background



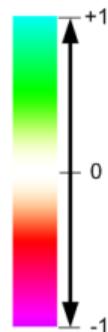
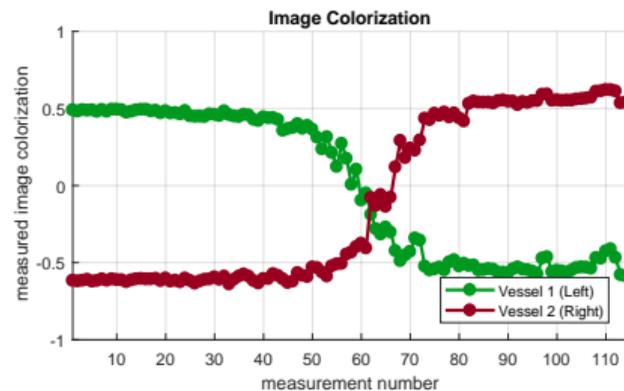
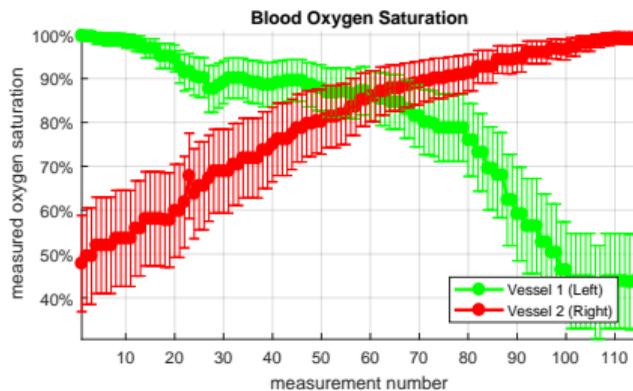
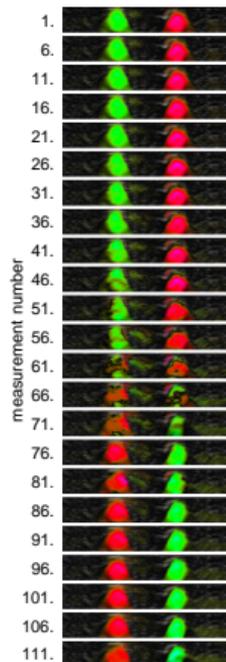
# Effect of Varying $sO_2$ on RGB Color

## Experiment 1



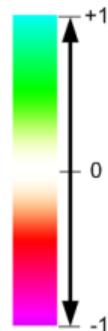
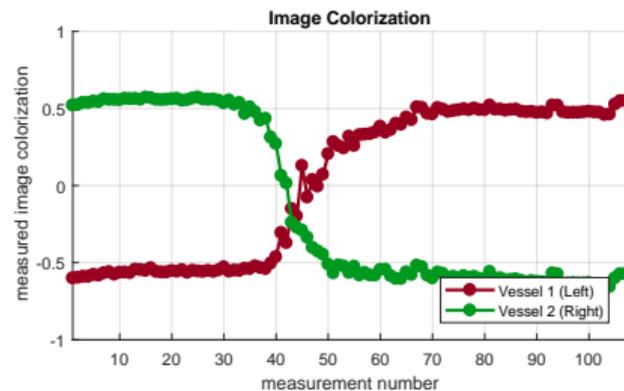
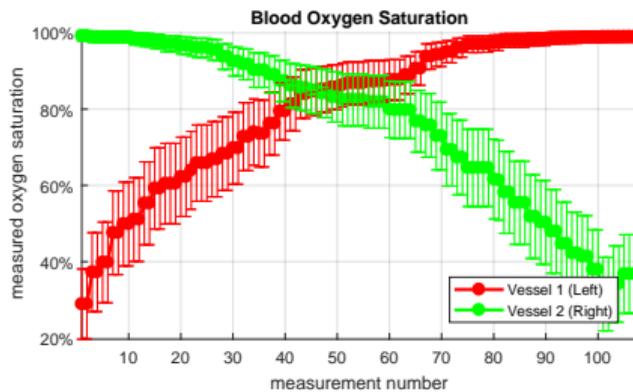
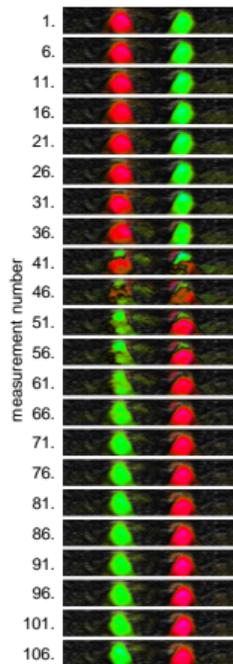
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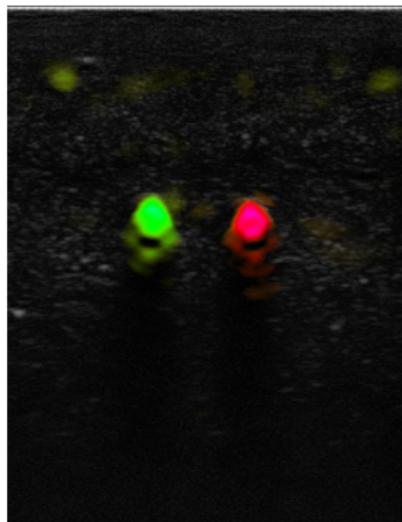
## Experiment 1 (repetition)



## Measure RGB Color Distinction (Experiment 2)

Vessel 1 (left) is **held constant** as reference

Vessel 2 (right) is **varied**



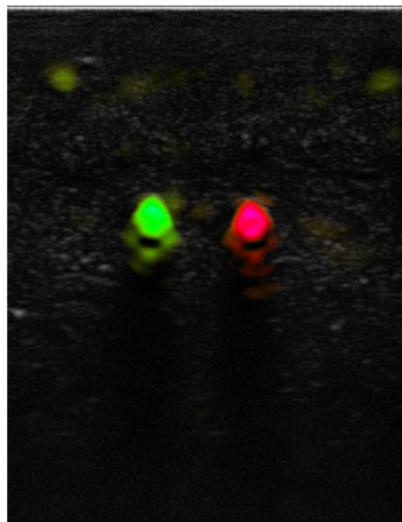
$$\Delta[sO_2] = 60\%$$

color distinction depends on **sO<sub>2</sub> difference** between the vessels

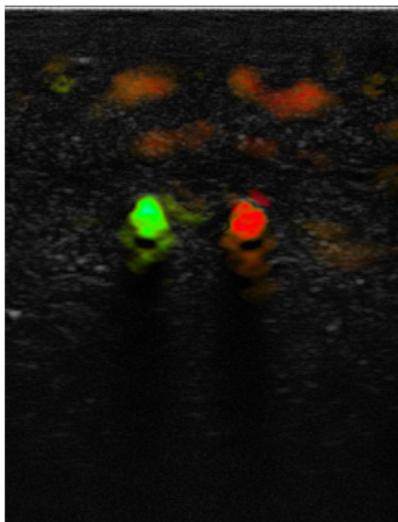
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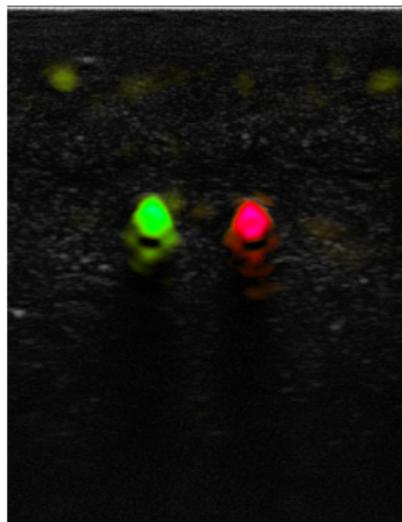
$\Delta[sO_2] = 14\%$

color distinction depends on **sO<sub>2</sub> difference** between the vessels

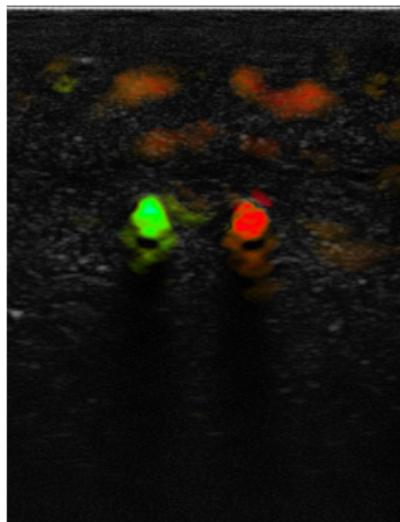
## Measure RGB Color Distinction (Experiment 2)

Vessel 1 (left) is **held constant** as reference

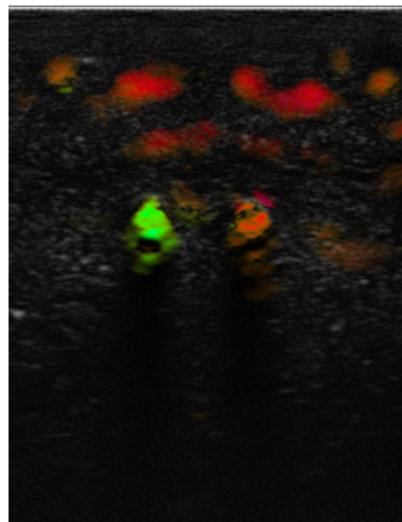
Vessel 2 (right) is **varied**



$\Delta[sO_2] = 60\%$



$\Delta[sO_2] = 14\%$



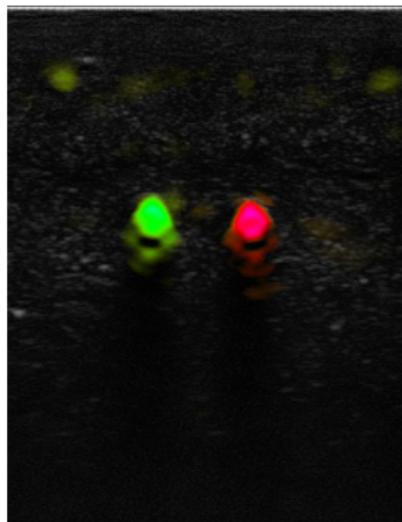
$\Delta[sO_2] = 6\%$

color distinction depends on **sO<sub>2</sub> difference** between the vessels

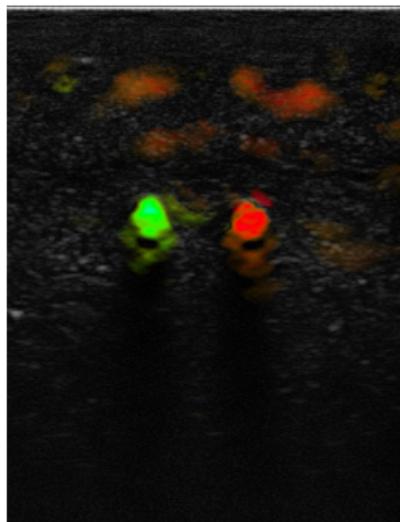
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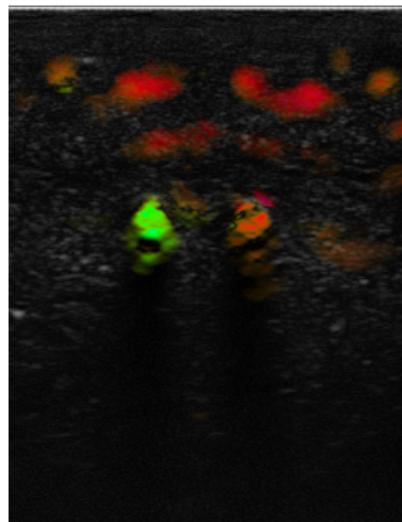
Vessel 2 (right) is **varied**



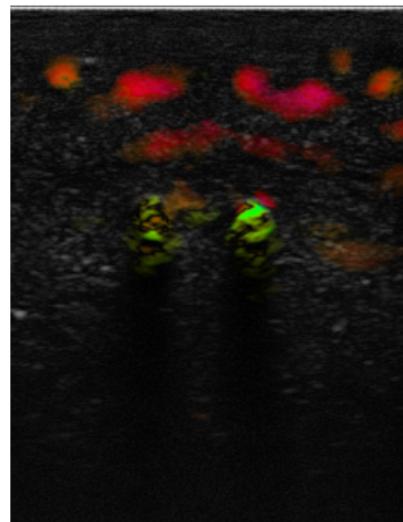
$\Delta[sO_2] = 60\%$



$\Delta[sO_2] = 14\%$



$\Delta[sO_2] = 6\%$

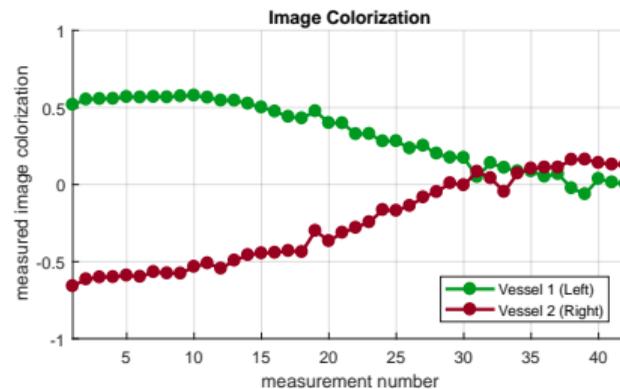
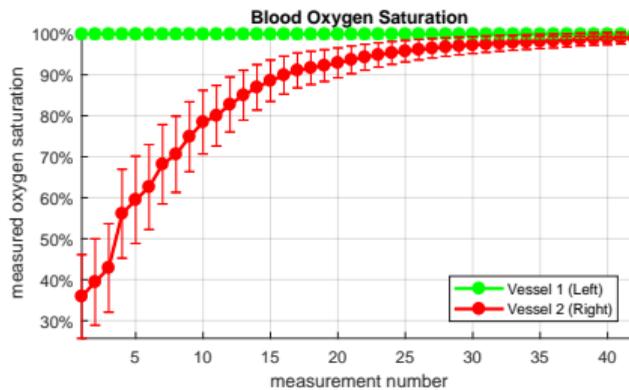
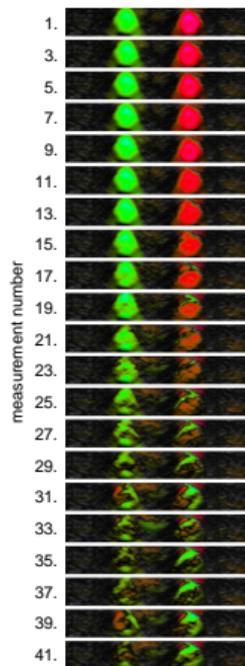


$\Delta[sO_2] = 2\%$

color distinction depends on **sO<sub>2</sub> difference** between the vessels

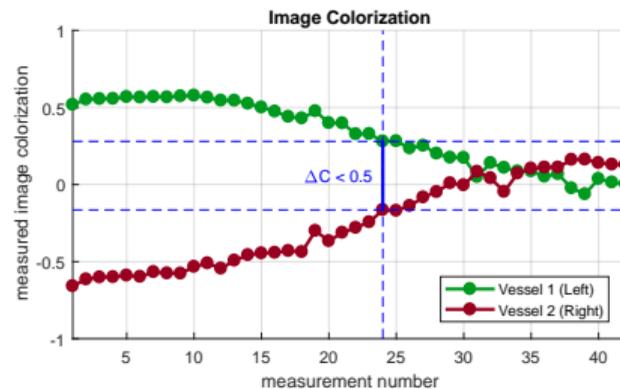
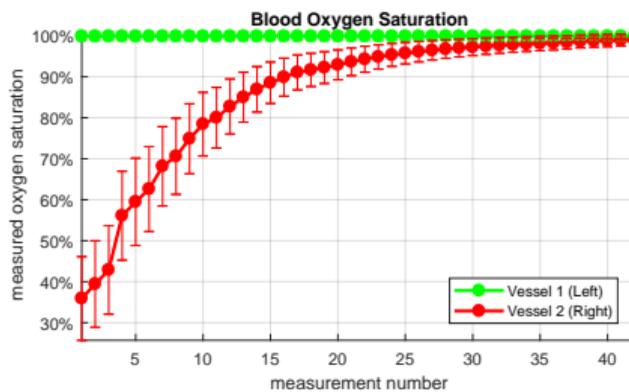
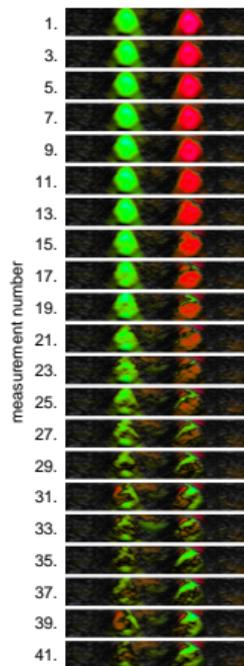
# Effect of Varying $sO_2$ on RGB Color Distinction

## Experiment 2



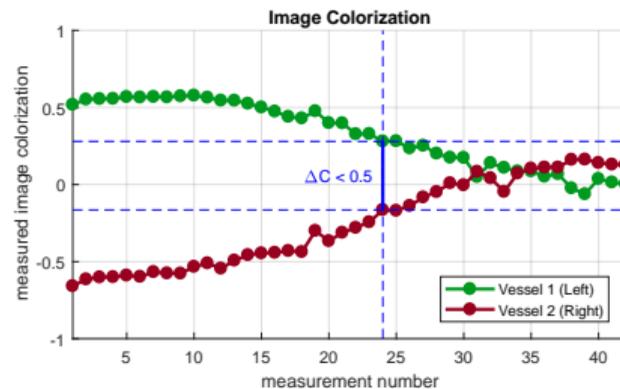
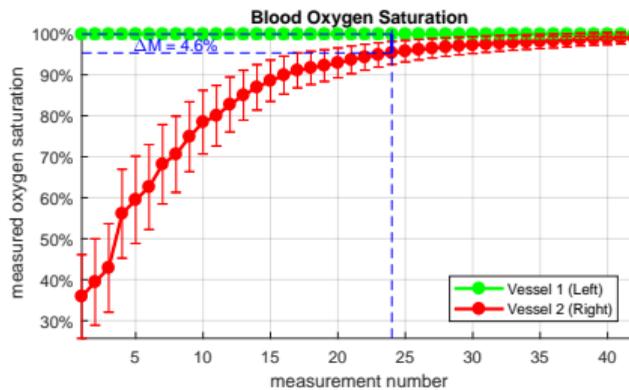
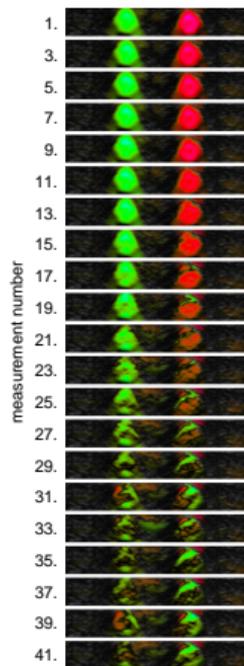
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## Experiment 2



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# Conclusion

- proposed a technique to characterize the ability to distinguish small differences in blood oxygen saturation when using relative colorization
- measuring effects of additional parameters (vessel depth, hematocrit, etc.) on  $\Delta M$  will be most effective if one reference vessel is held at constant  $sO_2$
- in PVCP phantom with reference vessel at 99%  $sO_2$ , the image colorization threshold  $\Delta C$  was reached at  $sO_2$  difference  $\Delta M$  of 4.6%

# Future Work

- perform additional measurements and characterize  $\Delta M$  while varying:
  - blood hematocrit level
  - vessel depth

# References I

- [1] Dash, R. K. and Bassingthwaite, J. B. "Erratum to: blood HbO<sub>2</sub> and HbCO<sub>2</sub> dissociation curves at varied O<sub>2</sub>, CO<sub>2</sub>, pH, 2, 3-DPG and temperature levels". In: *Annals of biomedical engineering* 38.4 (2010), pp. 1683–1701.
- [2] Neuschler, E. et al. "A Pivotal Study of Optoacoustic Imaging to Diagnose Benign and Malignant Breast Masses: A New Evaluation Tool for Radiologists." In: *Radiology* 287.2 (2018), pp. 398–412.
- [3] Vogt, W. C. et al. "Biologically relevant photoacoustic imaging phantoms with tunable optical and acoustic properties". In: *Journal of biomedical optics* 21.10 (2016), p. 101405.
- [4] Zalev, J. et al. "Opto-acoustic image fusion technology for diagnostic breast imaging in a feasibility study". In: *SPIE Medical Imaging*. International Society for Optics and Photonics. 2015, pp. 941909–941909.
- [5] Zalev, J. and Clingman, B. *Statistical mapping in an optoacoustic imaging system*. US Patent 9,330,452. 2016.

# Acknowledgement



Canada Research  
Chairs

Chaires de recherche  
du Canada



Canada Foundation  
for Innovation  
Fondation canadienne  
pour l'innovation

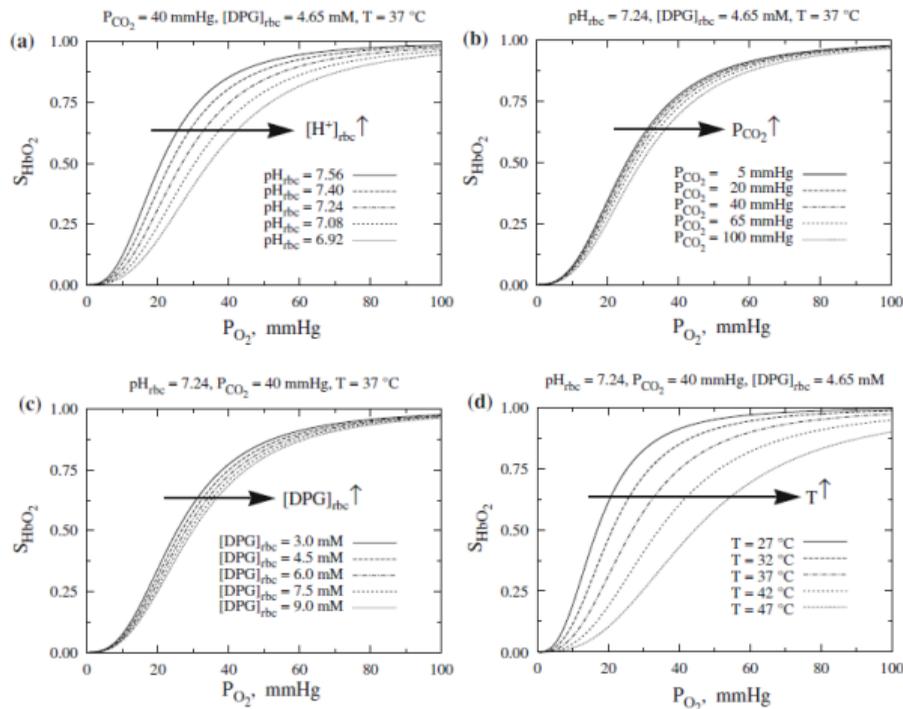
# Breast Tissue Properties

	Optical properties		Acoustic properties	
	Absorption coefficient ( $\text{cm}^{-1}$ )	Reduced scattering coefficient ( $\text{cm}^{-1}$ )	Speed of sound (m/s)	Acoustic attenuation coefficient (dB/cm)
General soft tissue	0.1 - 0.5	10 - 20	1450 - 1575	0.5 - 30
Breast fat	0.05 - 0.3	3 - 8	1430 - 1480	1 - 18
Breast parenchyma	0.1 - 0.3	5 - 15	1460 - 1520	2 - 25
Blood	2.0 - 10.0	10 - 15	$\sim$ 1560	0.1 - 2

Optical and acoustic properties of representative soft tissues. Optical properties cover a spectrum from 600 to 900 nm, while acoustic properties span 1 to 10 MHz. (source: Vogt, W. C. et al. "Biologically relevant photoacoustic imaging phantoms with tunable optical and acoustic properties". In: *Journal of biomedical optics* 21.10 (2016), p. 101405)

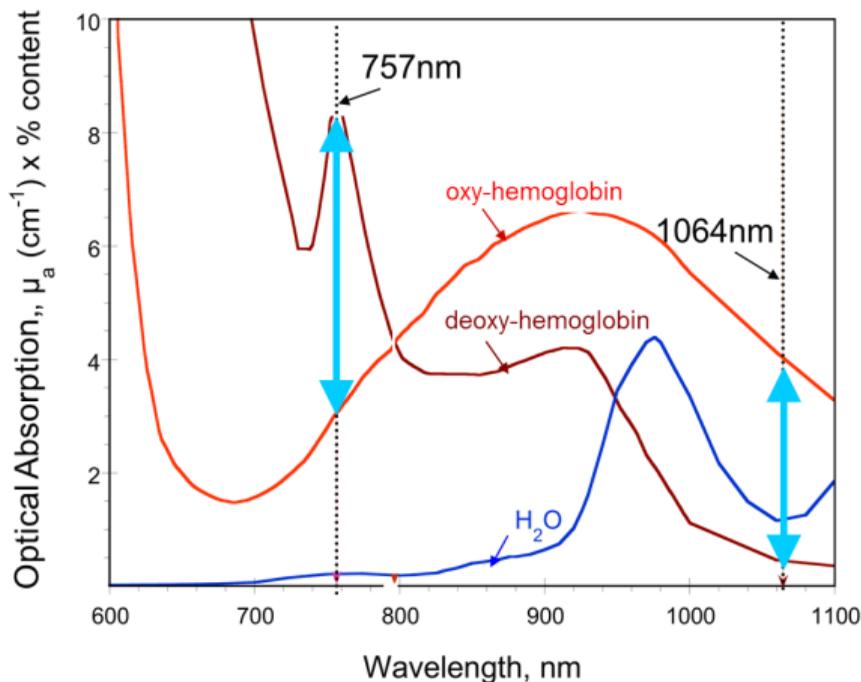
# Oxygen dissociation curves

Influence of Temperature, CO<sub>2</sub>, DPG and pH



source: Dash, R. K. et al. "Erratum to: blood HbO<sub>2</sub> and HbCO<sub>2</sub> dissociation curves at varied O<sub>2</sub>, CO<sub>2</sub>, pH, 2, 3-DPG and temperature levels". In: *Annals of biomedical engineering* 38.4 (2010), pp. 1683–1701

# Optical Absorption



Optical absorption coefficient for primary chromophores of tissue in near-infrared optical spectrum. It is indicated that at 757nm, deoxy-hemoglobin absorbs light more strongly than oxy-hemoglobin. At 1064nm, oxy-hemoglobin absorbs light more strongly than deoxy-hemoglobin (adapted from: [www.senomedical.com](http://www.senomedical.com)).