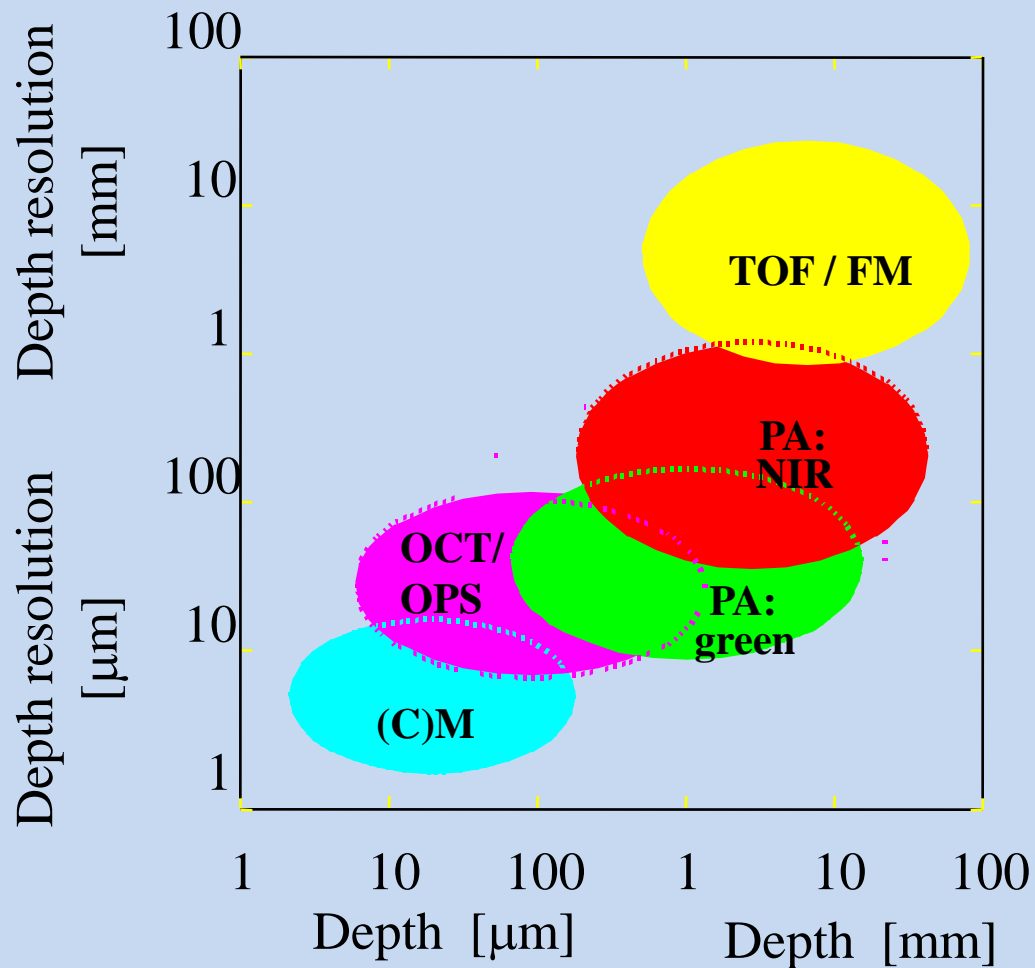


# Photoacoustic Imaging of Blood Vessels in Tissue

F.F.M. de Mul

(University of Twente, Enschede, the Netherlands)

# Imaging methods for hidden structures in turbid media (tissue)



C(M) : (confocal) microscopy

OCT: optical coherence tomography

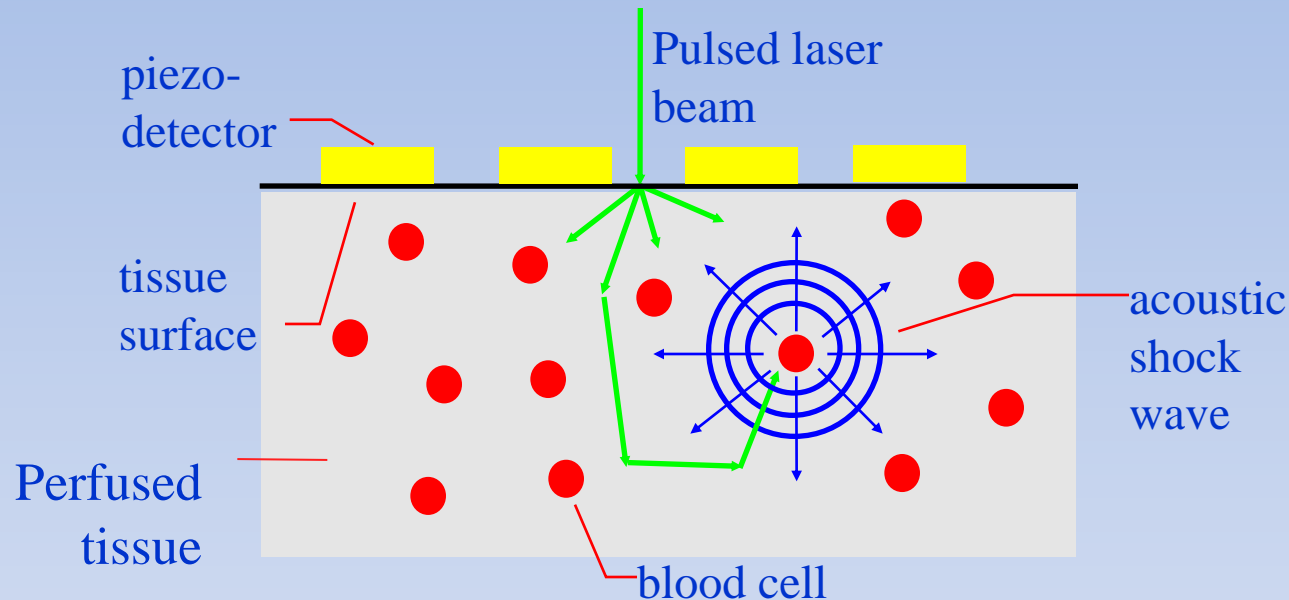
OPS: orthogonal polarization spectral imaging

PA: photoacoustics

TOF: time-of-flight tomography

FM: frequency-modulated tomography

# Photoacoustic Imaging



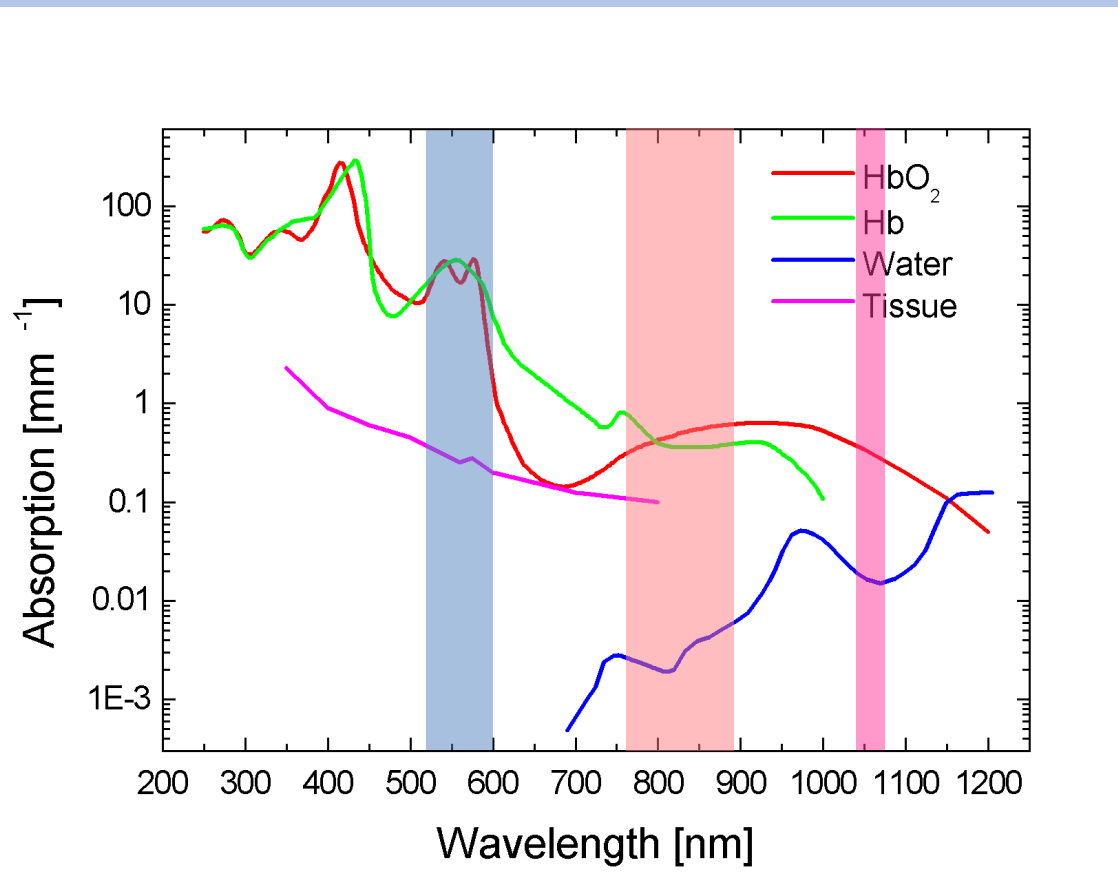
- light pulse is absorbed in blood cell
- adiabatic heating
- pressure pulse emerging ( $\approx 1500$  m/s)
- detection at tissue surface

Depth:

- **Green light:**  $\approx 0 - 9$  mm
- **Near-infrared:**  $\approx 0 - 30$  mm

Depth resolution:  $\approx 10$   $\mu$ m

## Scattering and absorption



(Reduced)  
Scattering coefficient:

- $\lambda = 580 \text{ nm}$ :

Dermis:  $3 \text{ mm}^{-1}$

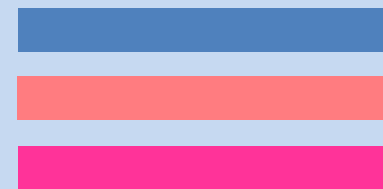
Blood: 1 ...

- $\lambda = 850 \text{ nm}$ :

Dermis: 1 ...

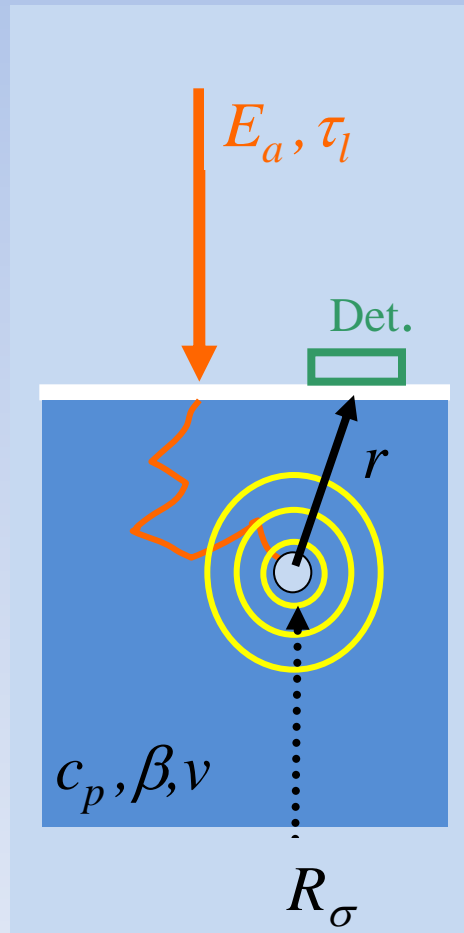
Blood: 0.5 ...

Available windows:



# Photoacoustic Signal Generation

Bipolar PA-signal generated by a spherical Gaussian Source



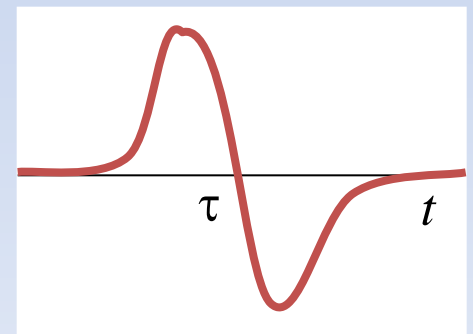
$$\tau_a = \frac{R_\sigma}{\nu}$$

$$\tau_e = \sqrt{\tau_a^2 + \tau_l^2}$$

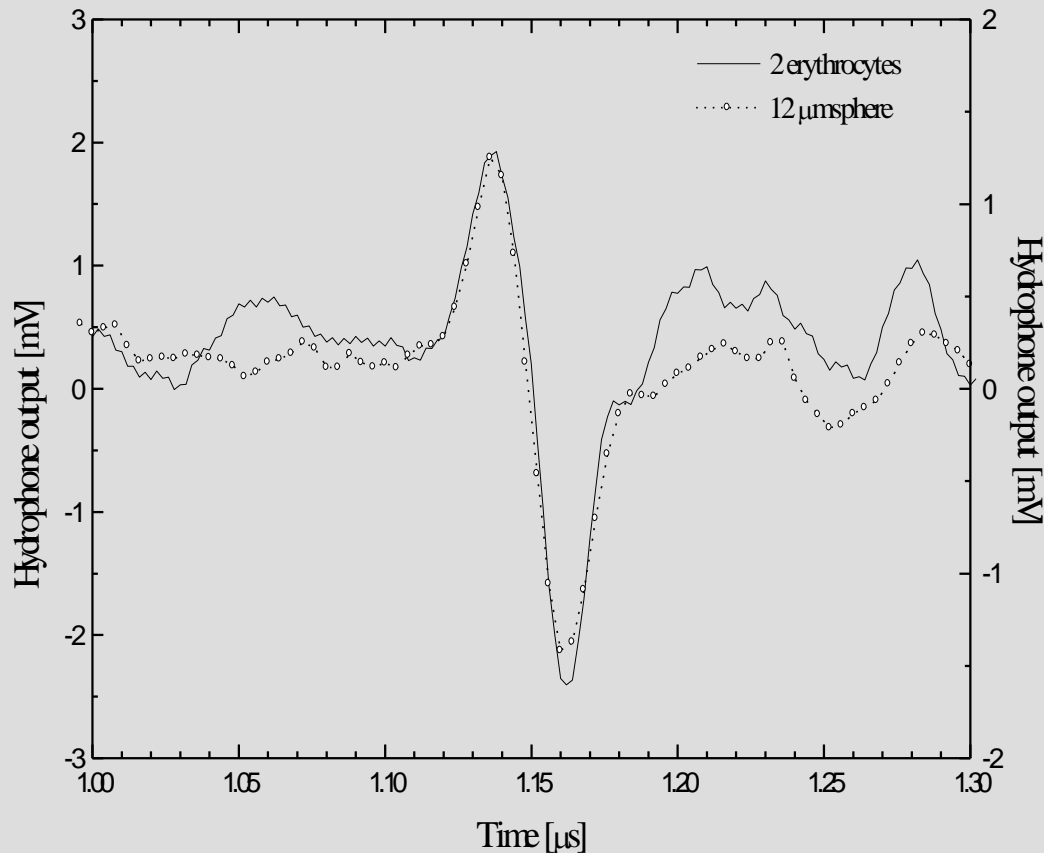
$$P(r, t) = -P_{\max}(r) \sqrt{e} \frac{t - \tau}{\tau_e} \exp\left\{-\frac{1}{2} \frac{(t - \tau)^2}{\tau_e^2}\right\}$$

$$\tau = \frac{r}{\nu}$$

$$P_{\max}(r) = \frac{\beta E_a}{2\sqrt{e}(2\pi)^{3/2} c_p \tau_e^2 r}$$



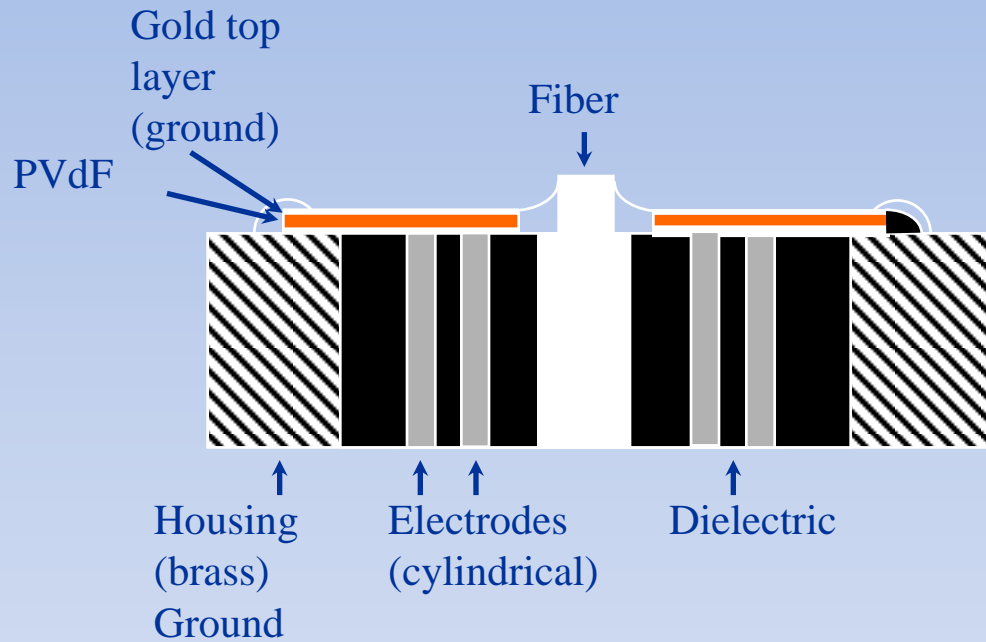
# Photoacoustic Imaging



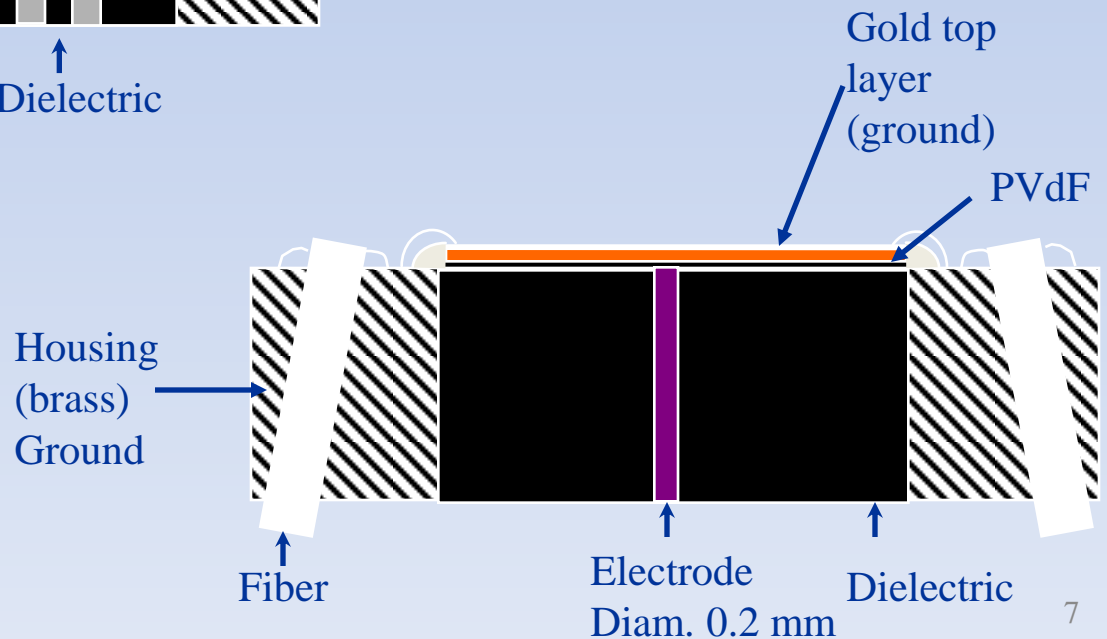
Two erythrocytes

Diameter:  $\approx 10 \mu\text{m}$   
 (compared with a  $12 \mu\text{m}$  blue polystyrene sphere)  
 detection distance:  $\approx 1.7 \text{ mm}$  ( $= 1.15 \mu\text{s} \times 1500 \text{ m/s}$ )  
 medium: water/PBS

# Photoacoustic Imaging

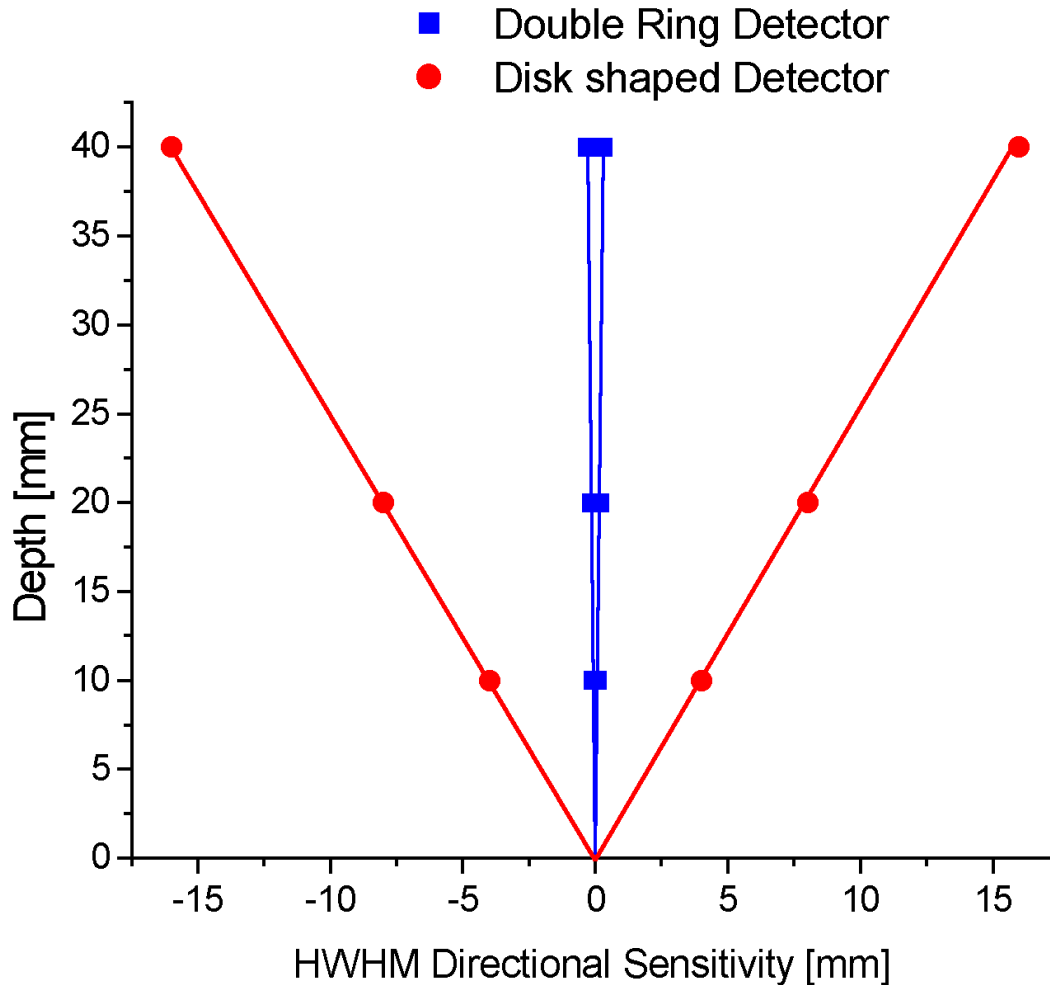


## Double-Ring Detector One-fiber illumination



## Disk-shaped Detector Ring illuminator

# Photoacoustic Imaging



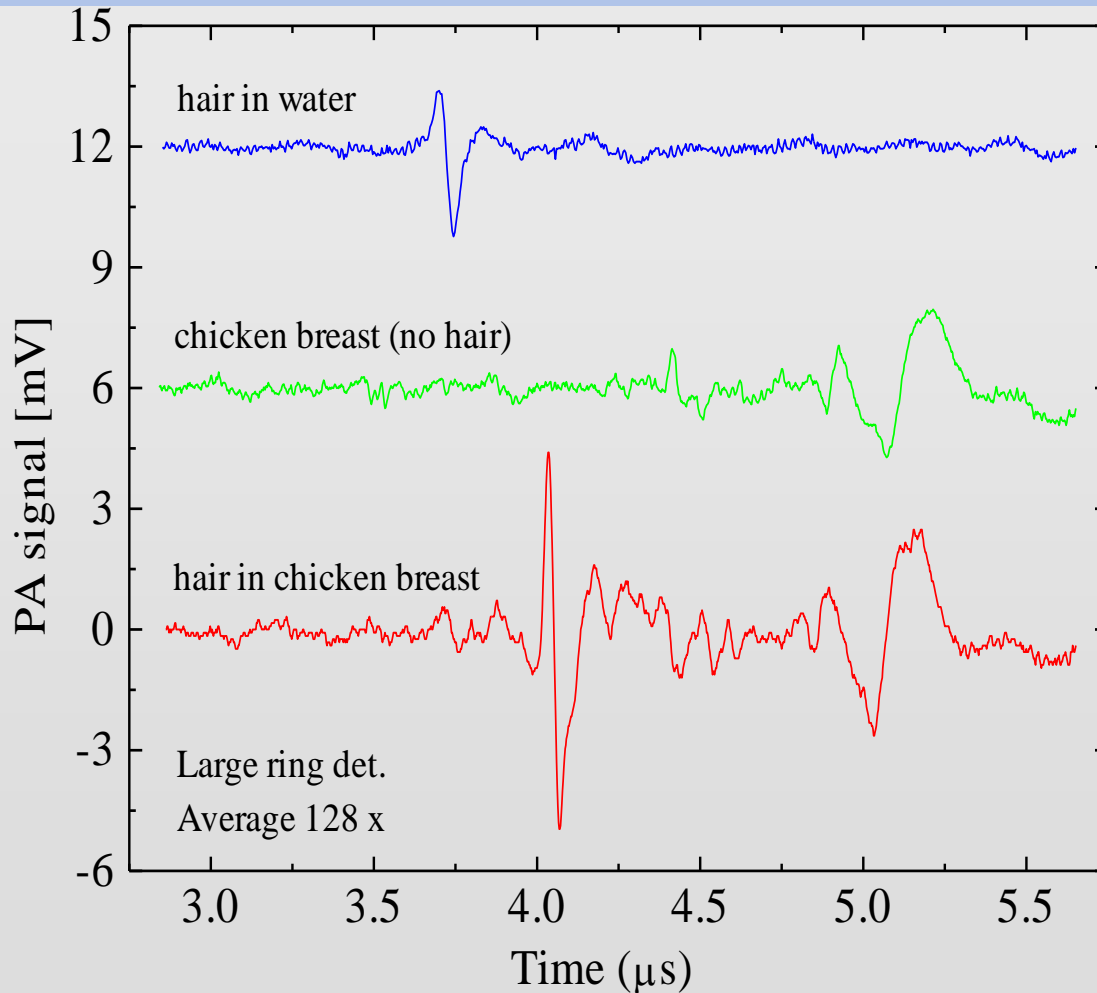
Directional Sensitivity:

Disk Detector:  
FWHM : Depth  
1 : 5

Ring Detector:  
FWHM : Depth  
1 : 70



# Photoacoustic Imaging

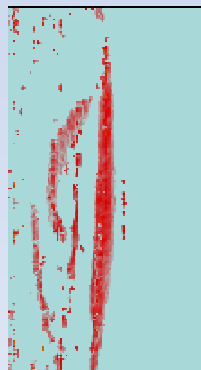
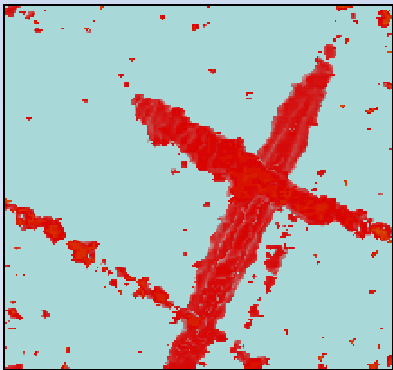
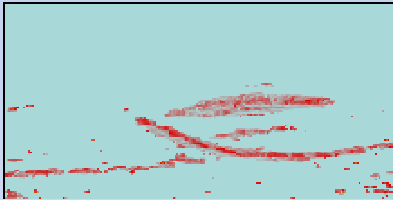
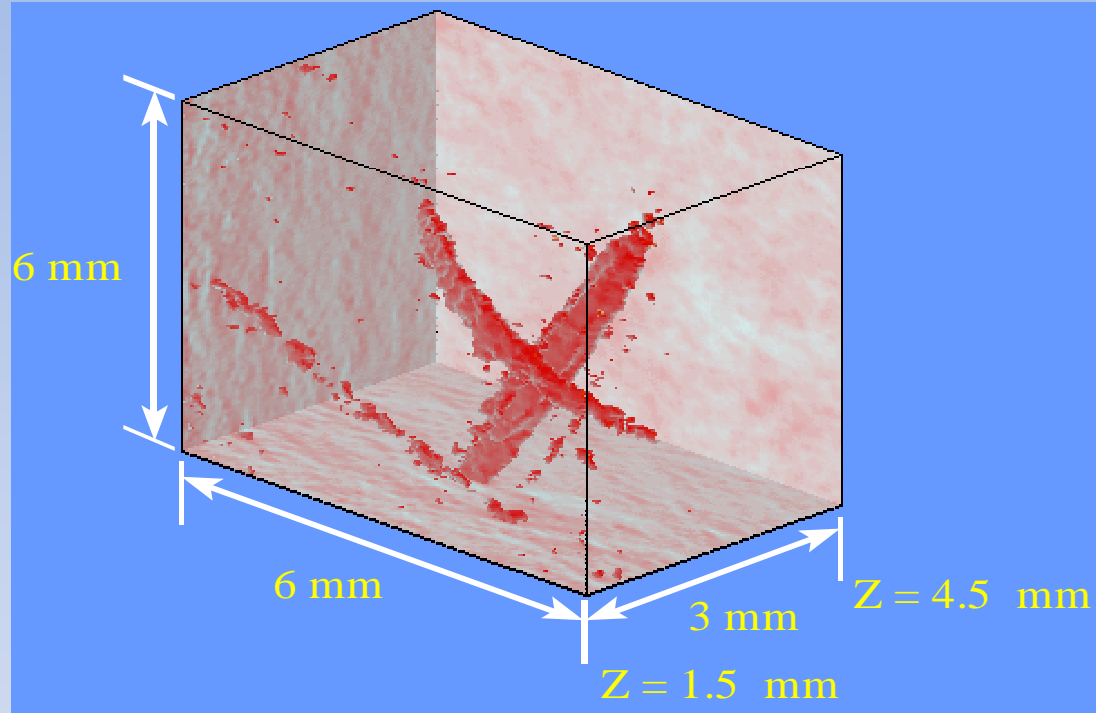


A human  
hair in  
chicken  
breast  
tissue.

Depth:  $\approx 6$  mm  
( $\approx 4 \mu\text{s}$  x  
1500 m/s)

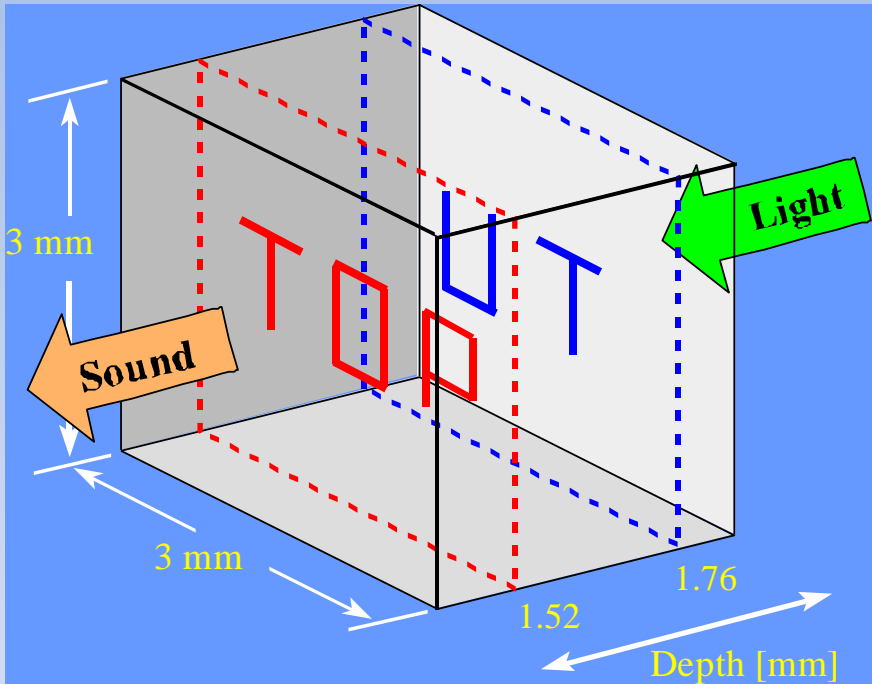
# Photoacoustic Imaging

Vessels in  
chicken  
breast tissue

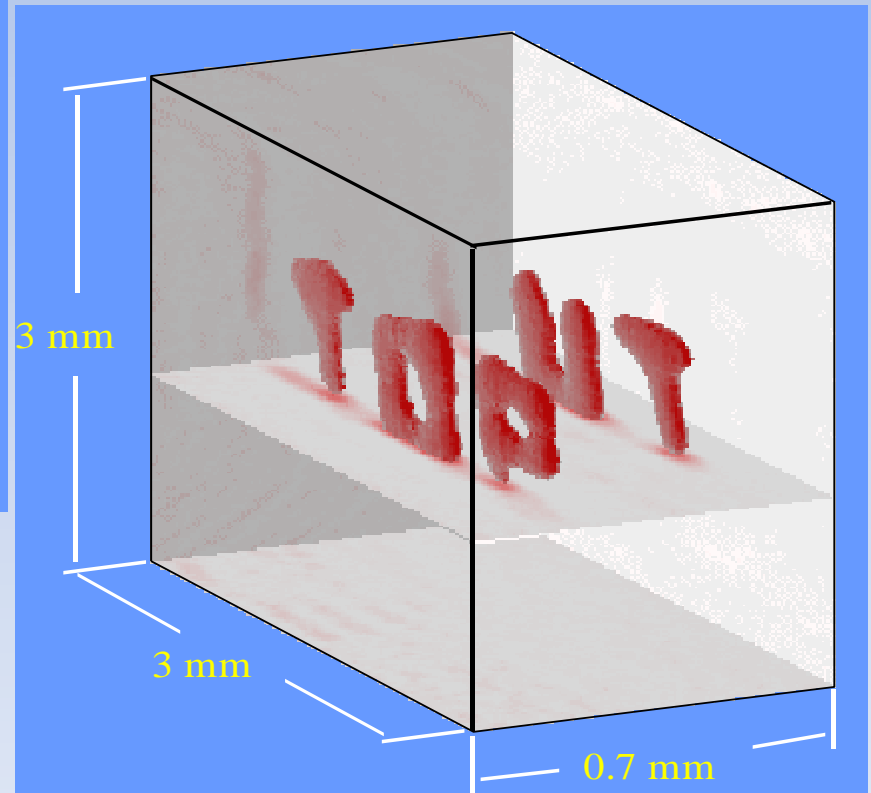


Sample : 5 mm thick chicken breast tissue in water  
 Image : 663 nm, inside sample, 35 % isosurface threshold  
 Vessels : 3 Nylon capillaries, 0.28 - 0.40 mm diameter  
 Absorber : Evans Blue, flowing,  $a = 300 \text{ cm}^{-1}$   
 Detection : at Z = 0 mm, 51 x 51 points, 0.15 mm spacing

# Photoacoustic Imaging



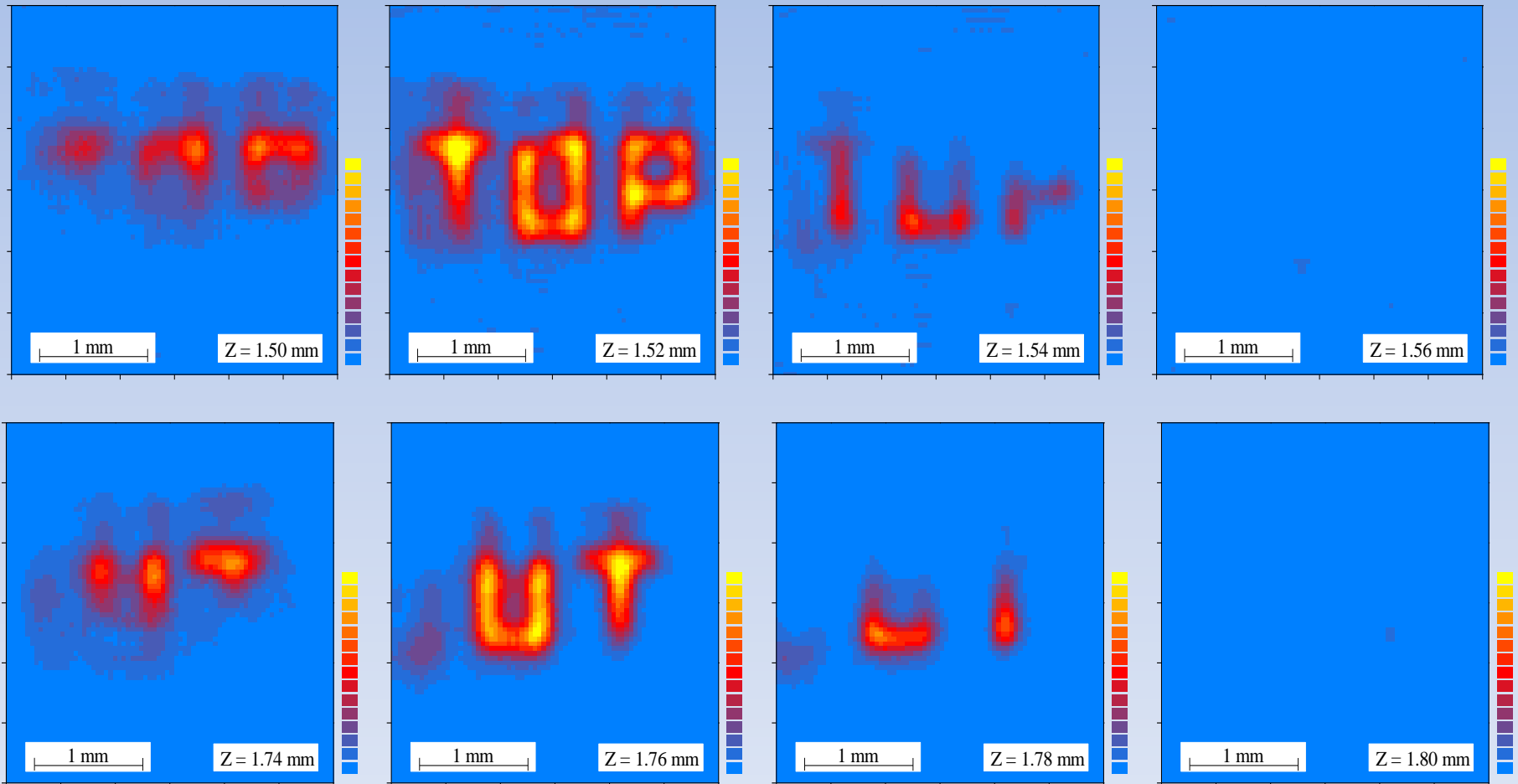
## Reconstruction of hidden objects



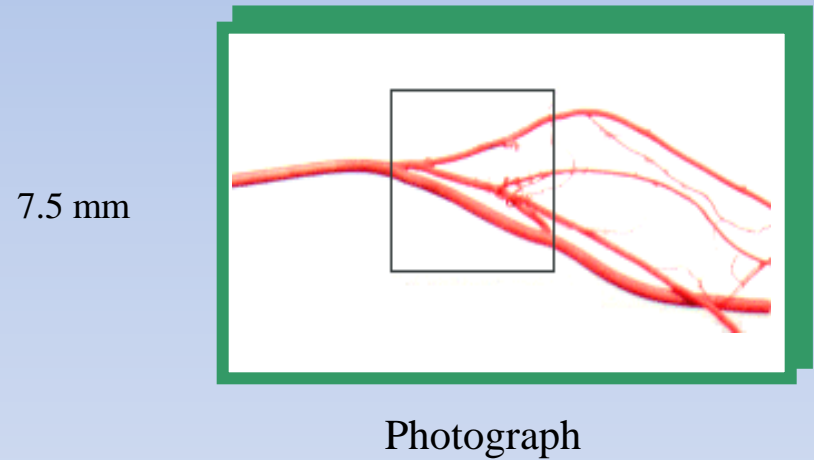
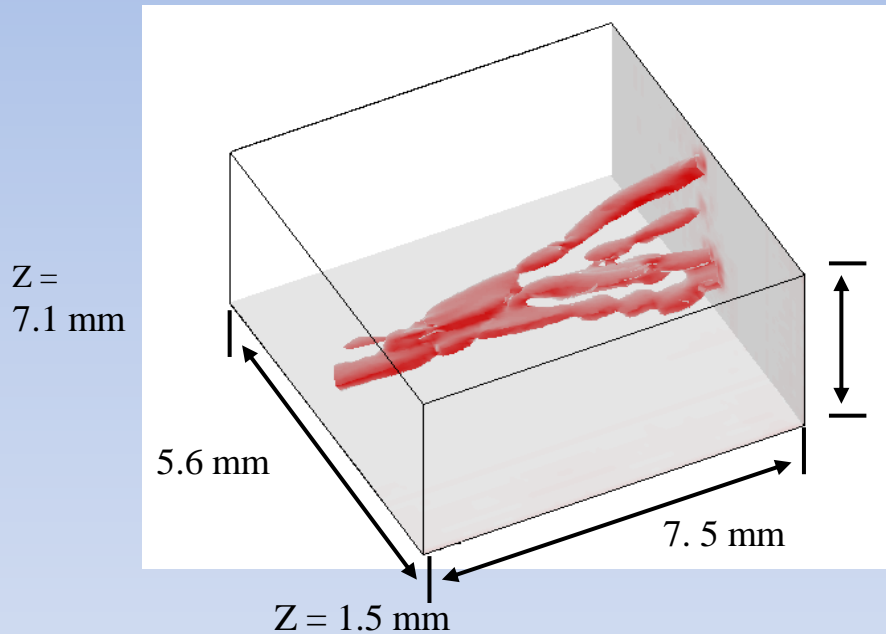
Material: carbon threads ( $10\ \mu\text{m}$ ) on transparent sheets, in 10 % Intralipid-10% (resembles human tissue scattering)

# Photoacoustic Imaging

Depth resolution:  $\approx 10 \mu\text{m}$



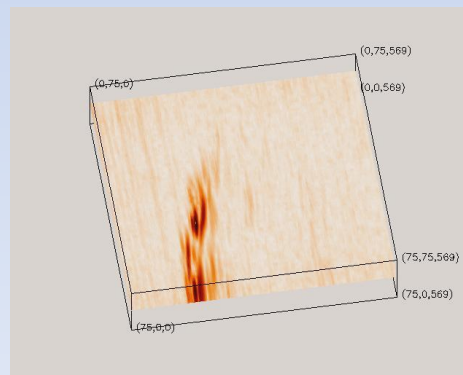
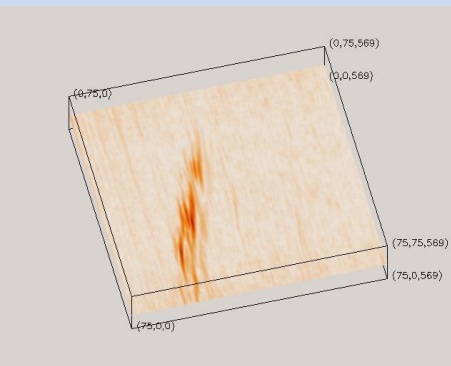
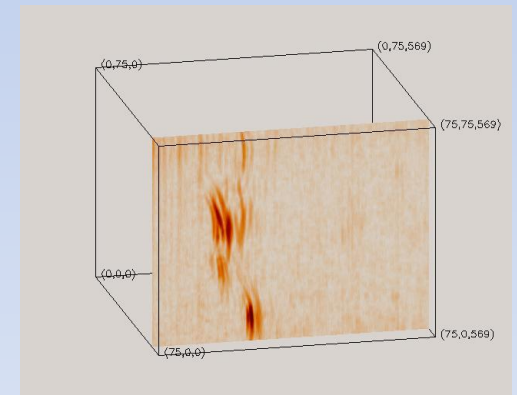
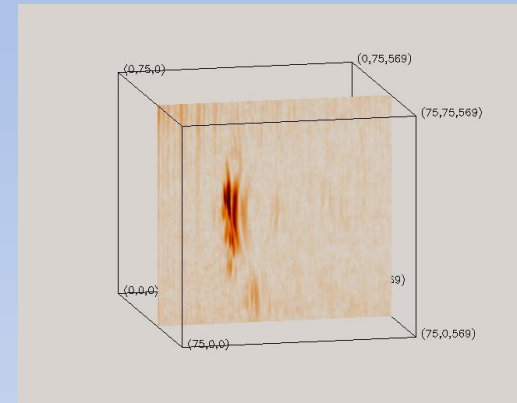
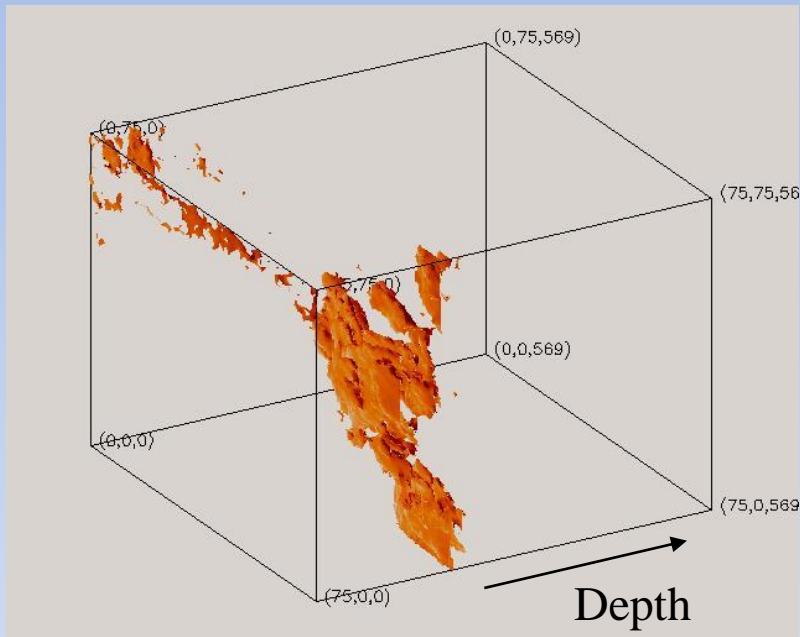
# Photoacoustic Imaging



Vascular tree from a branching epigastric artery of a rat.  
 Ex-vivo; medium: intralipid 1 % ( $\approx$  tissue).  
 Depth (Z-coord.)  $\approx 5 \text{ mm}$  ; indicated in figure.  
 Laser power 532 nm, 2mJ/pulse through fiber  $\varnothing 600 \mu\text{m}$ .  
 Depth resolution / lateral resolution: 10 / 100  $\mu\text{m}$  respectively.

# PA-imaging of blood vessels in tissue

## Live Albino Wistar Rat: Epigastric artery branching



$7.5 \times 7.5 \times 7.5 \text{ mm}^3$

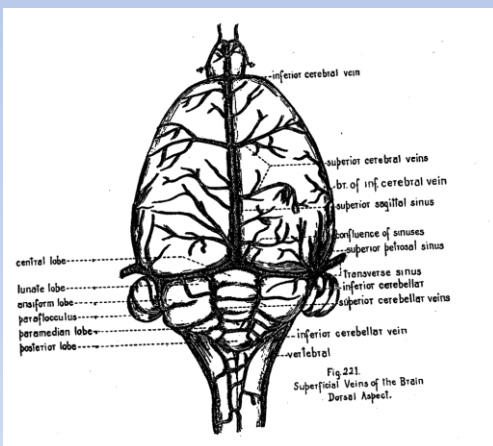
# PA-imaging of blood vessels in tissue

## Albino Wistar Rat: Brain Perfusion

**Alive**

Area: 4 mm x 4 mm

Depth 5mm



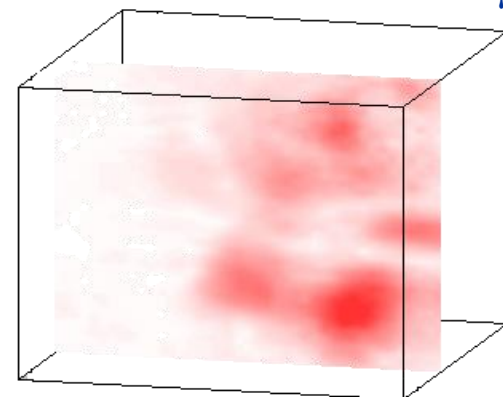
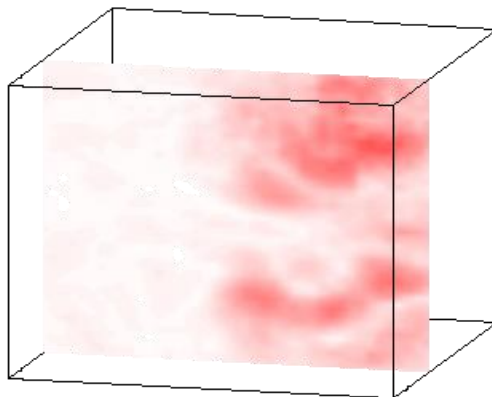
**Dead**

Area: 5 mm x 5 mm

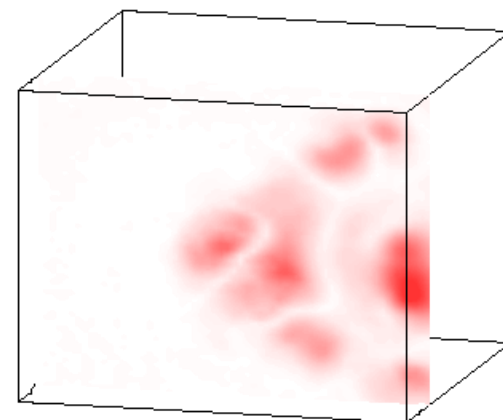
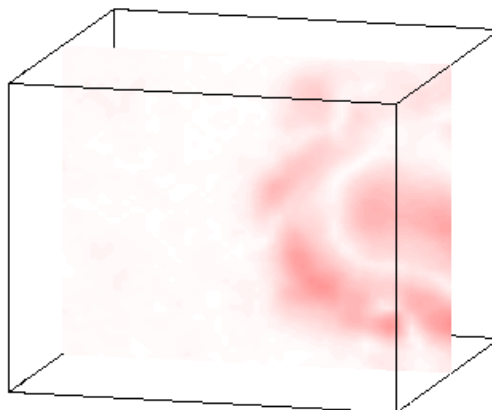
6 mm x 6 mm

Depth 5mm

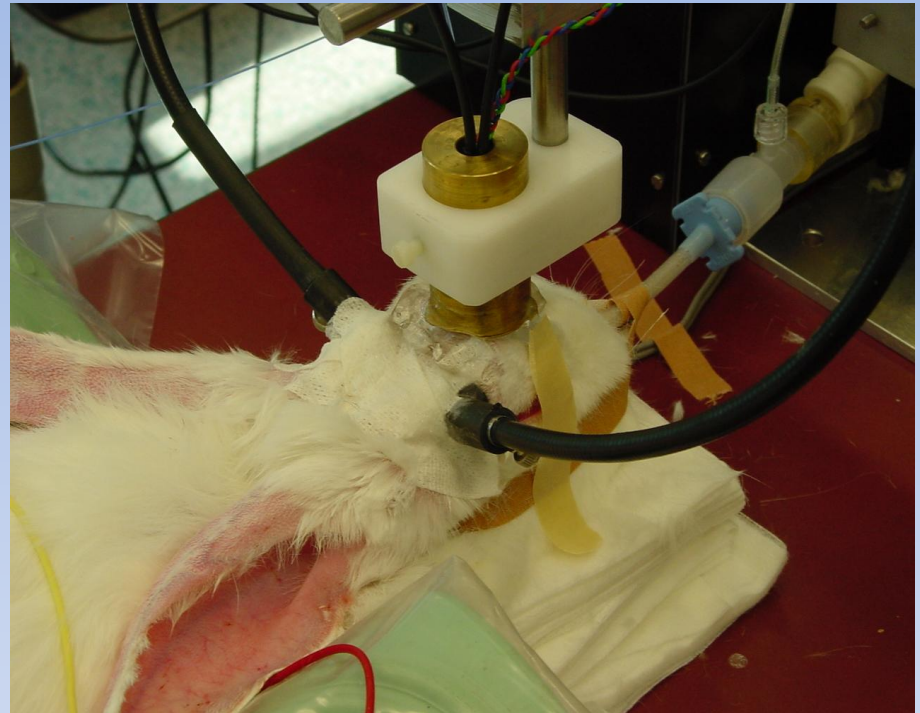
Slices of 100  $\mu$ m



Blood in the cartilage scalp, in the upper part of the occipital bone



## PA-image of Vessels in Rabbit Ear

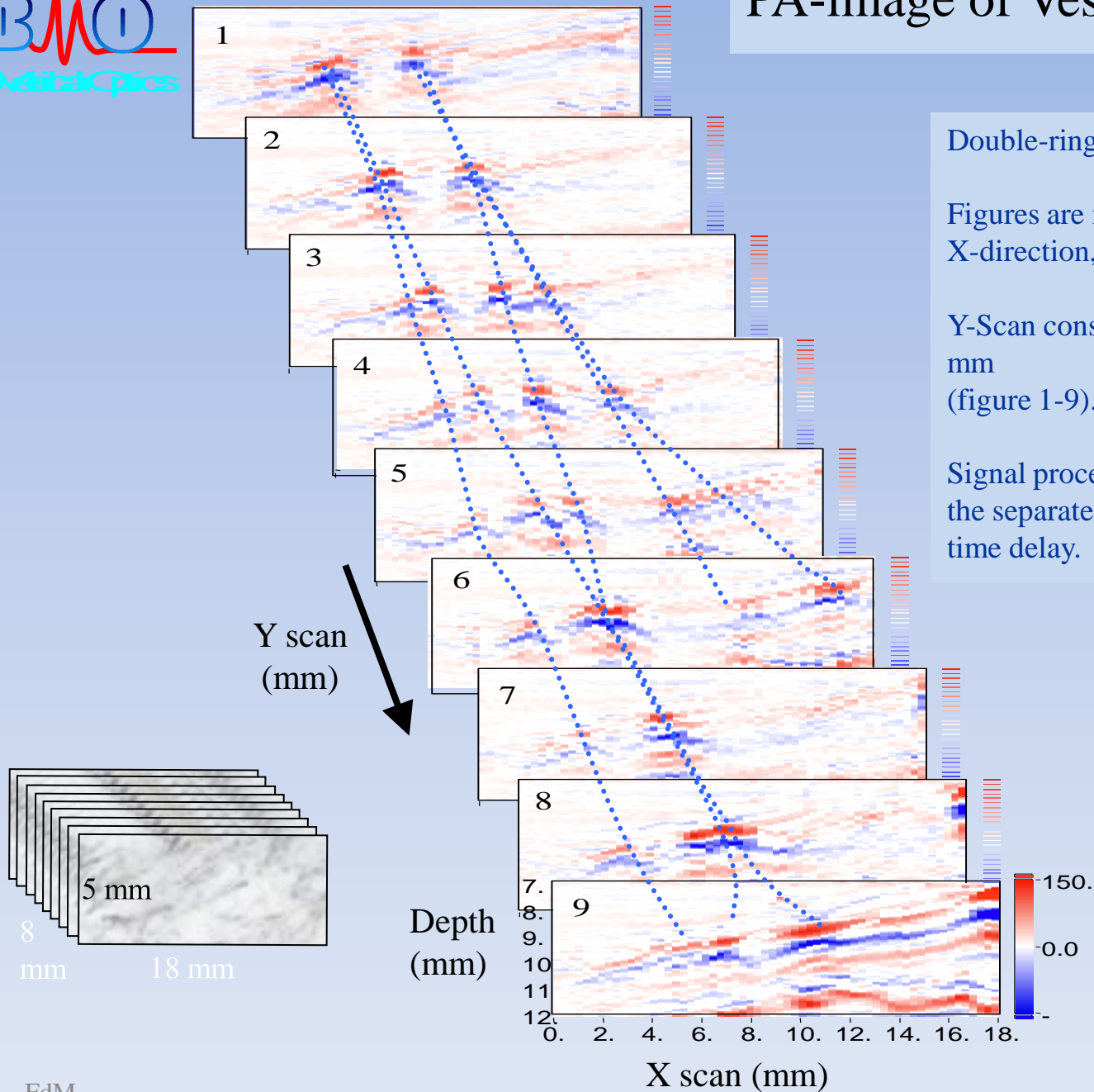


Double-ring detector.

Figures are made out of 61 scan steps in X-direction, with a step size of 0.3 mm.  
Y-Scan consists of 9 lines, separated 1.0 mm

Signal processing: summing the data of the separate rings,  
after correction for the time delay.



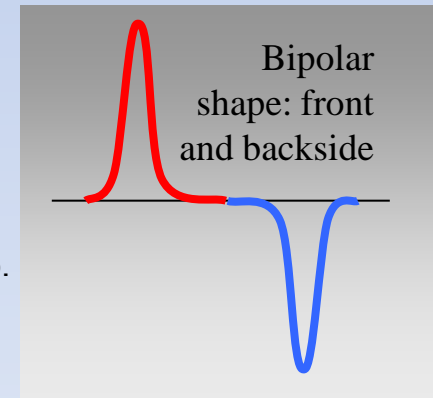


Double-ring detector.

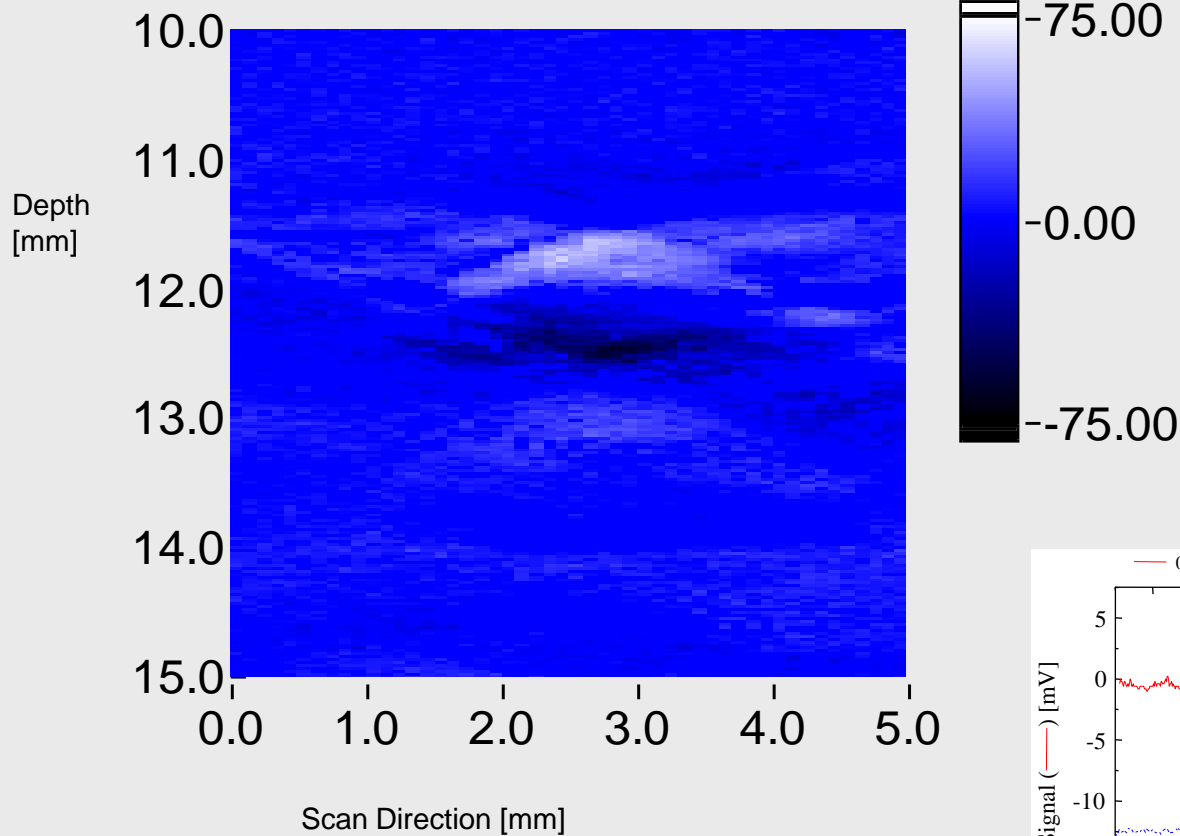
Figures are made out of 61 scan steps in X-direction, with a step size of 0.3 mm.

Y-Scan consists of 9 lines, separated 1.0 mm (figure 1-9).

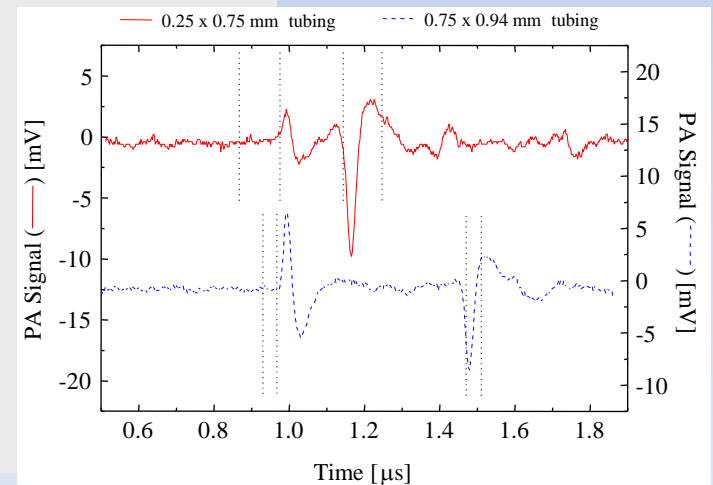
Signal processing: summing the data of the separate rings, after correction for the time delay.



# PA-imaging of blood vessels in rabbit ear

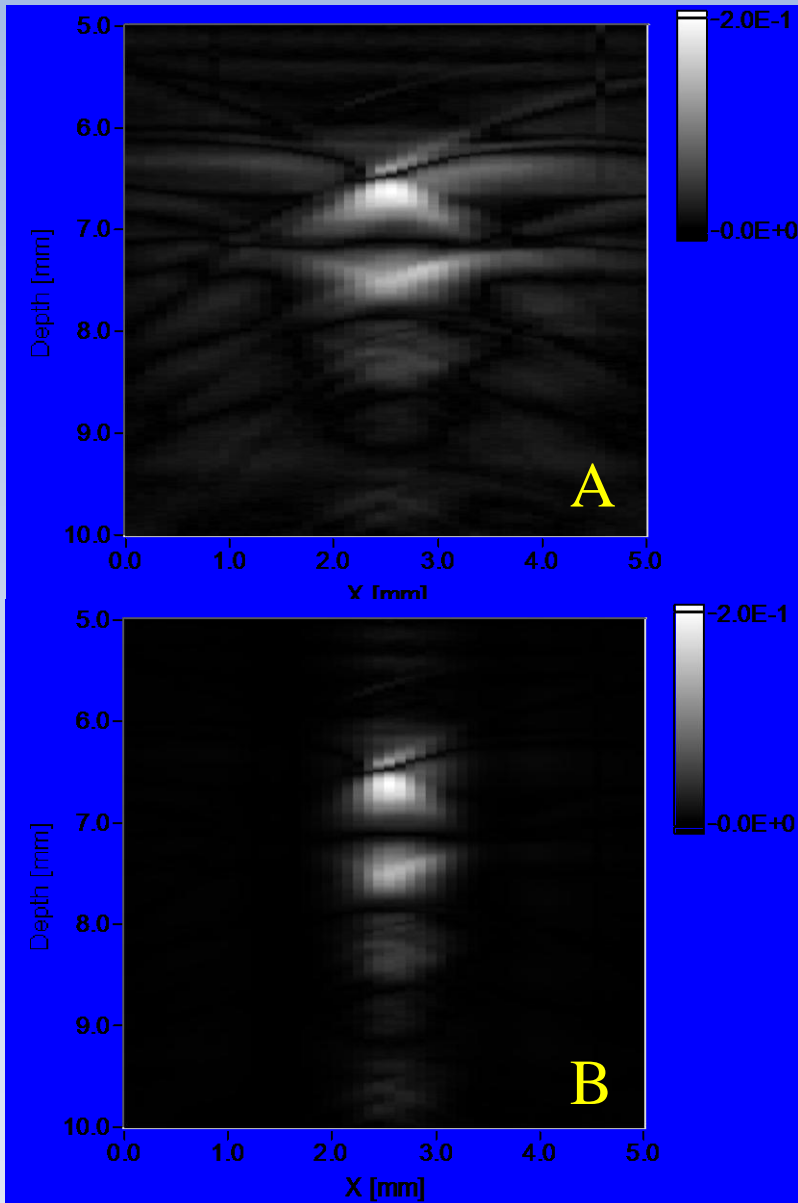


Front and  
backside  
of the vessel

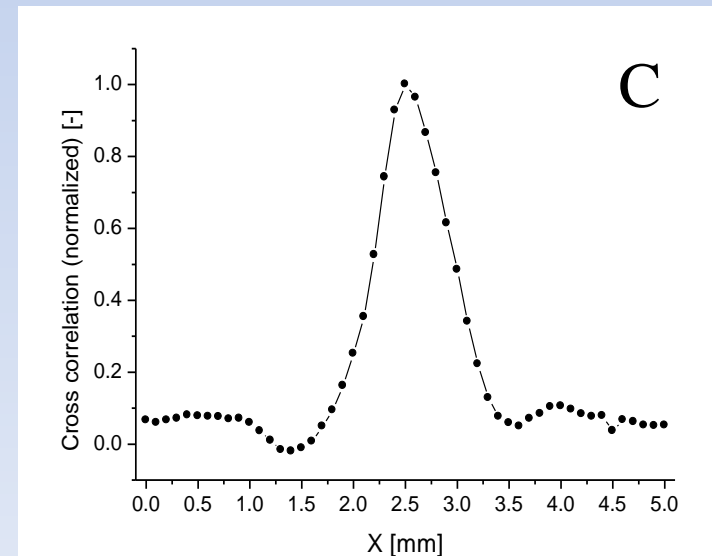


PA signals: capillaries in 10% IL-10%

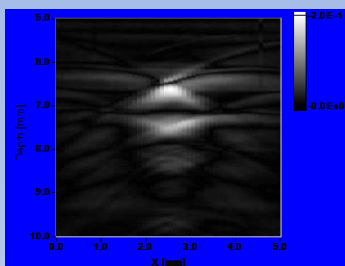
# PA-imaging of blood vessels: image correction



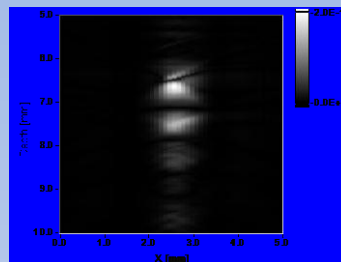
- 1mm diameter rubber tube, filled with human blood, in 7.5% Intralipid-10% dilution.
- **Image A** : 2D image, from measured time signals (1D-depth images), plotted as lines in the 2D image next to each other.
- **Image B** : 2D image, after correction by multiplication with the zero-time cross-correlation function, (C) : **STEP 1**



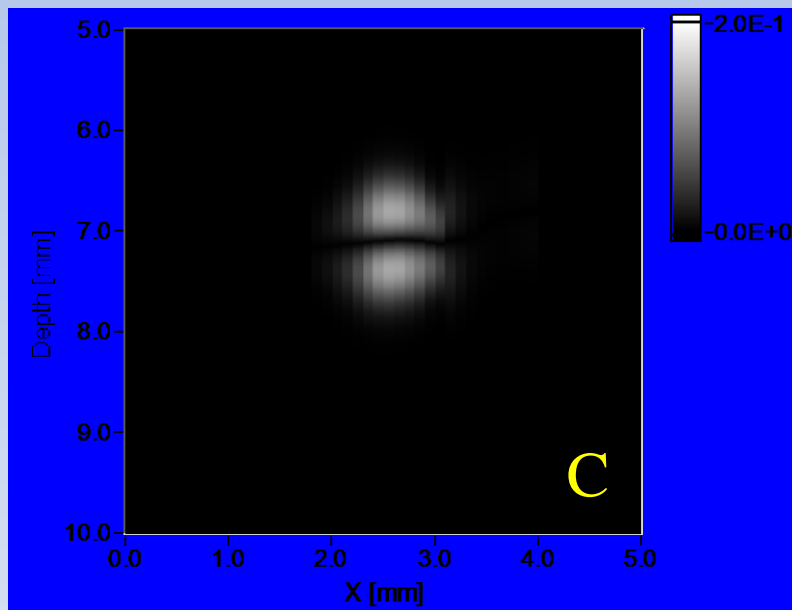
# PA-imaging of blood vessels: image correction



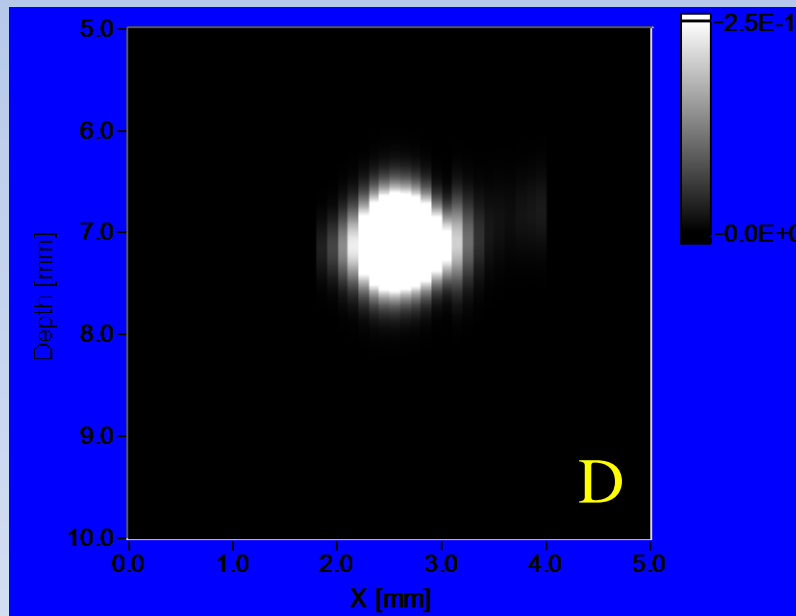
A: originally measured time signals



B: after multiplication with autocorrelation (STEP 1)

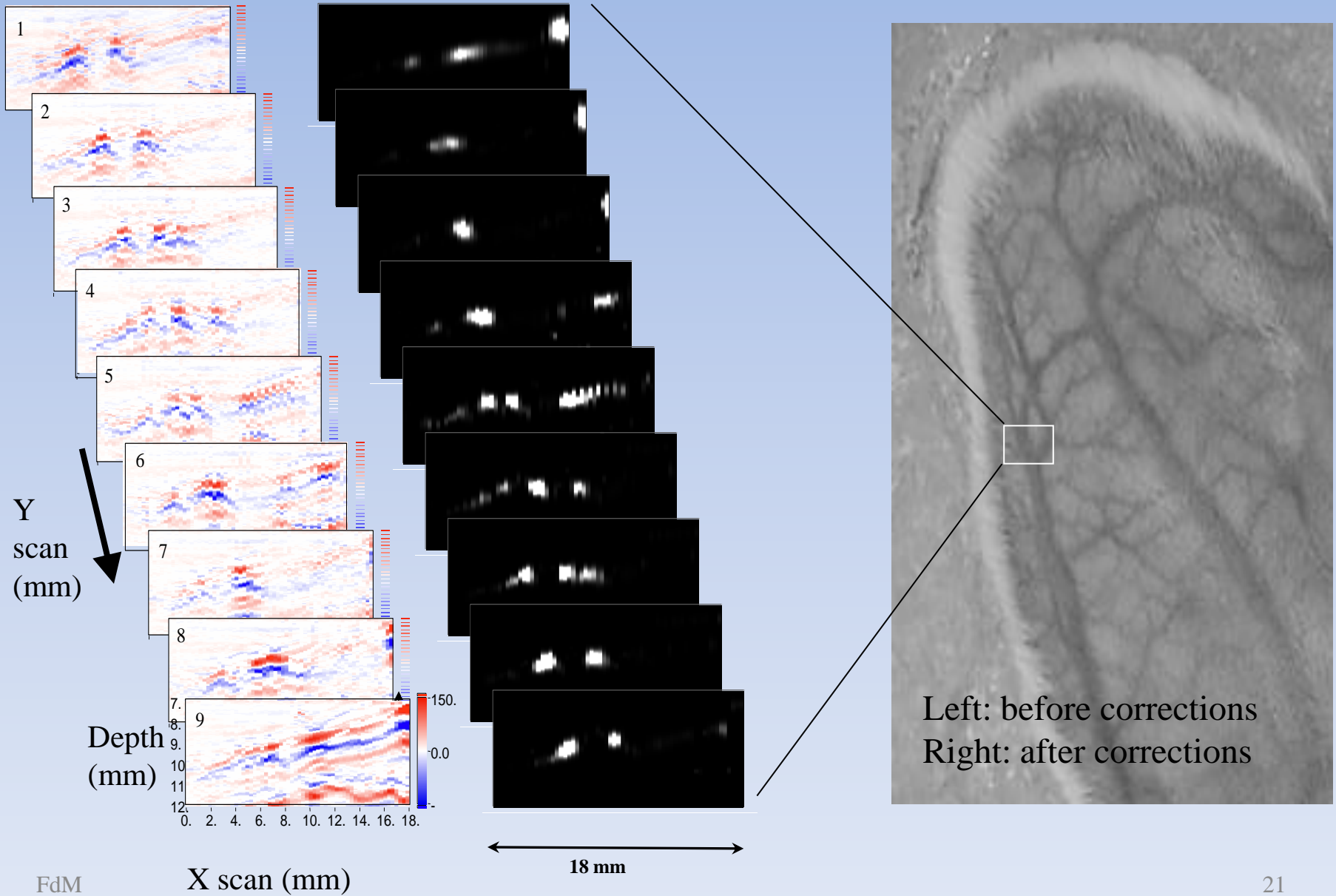


C: after fit to bipolar signal (STEP 2)  
Upper/lower image: pos./neg. peak signal (here absolute value plotted)

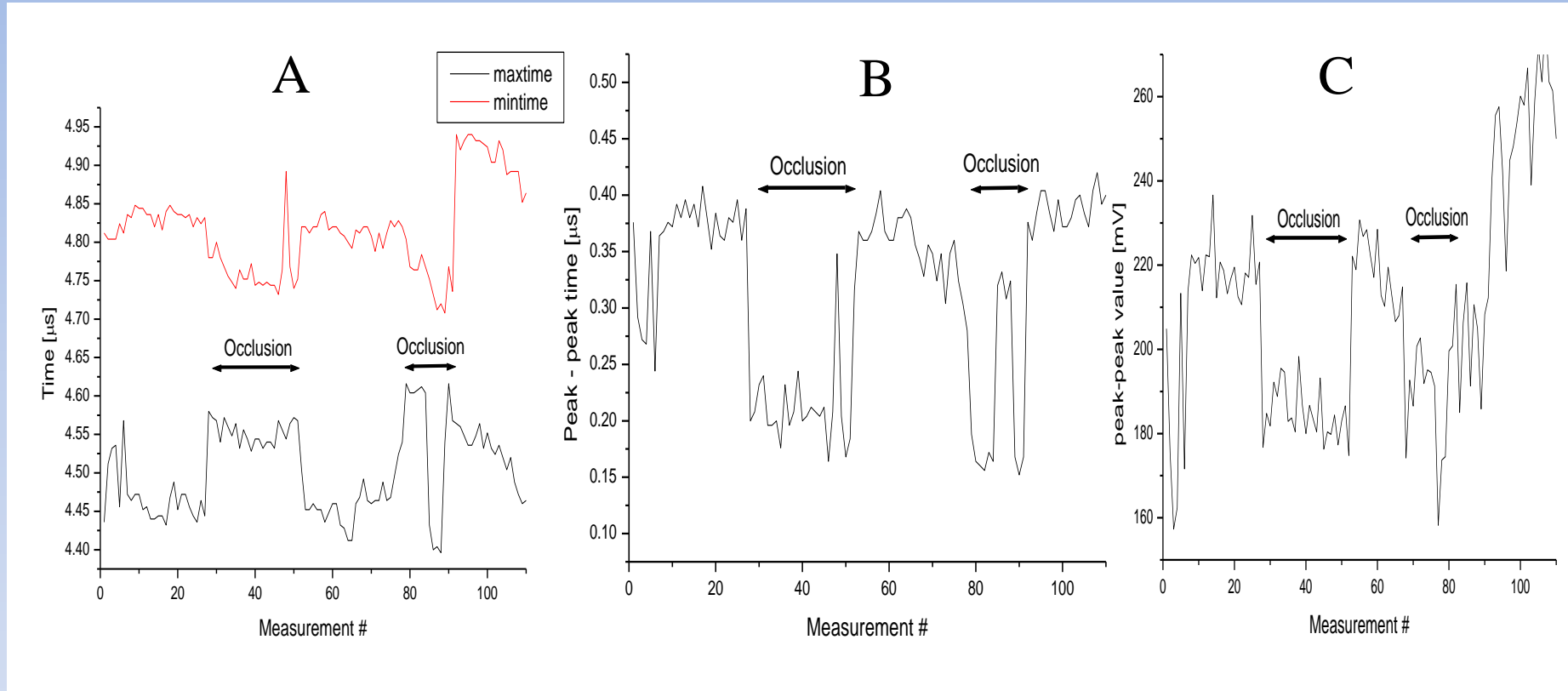


D: after running integration over depth (STEP 3)

# PA-imaging of blood vessels in rabbit ear



# PA-imaging of blood vessels in rabbit ear: occlusion

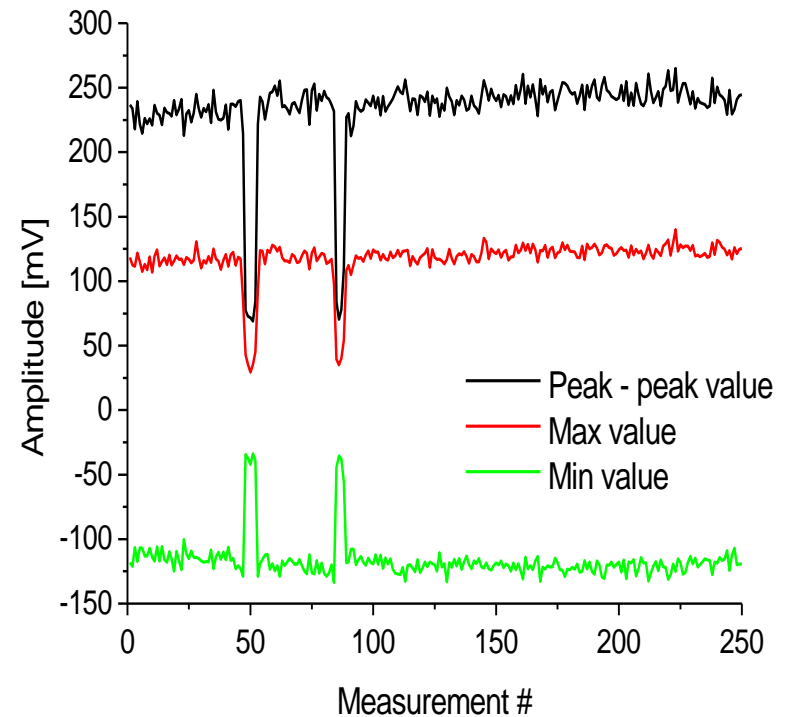
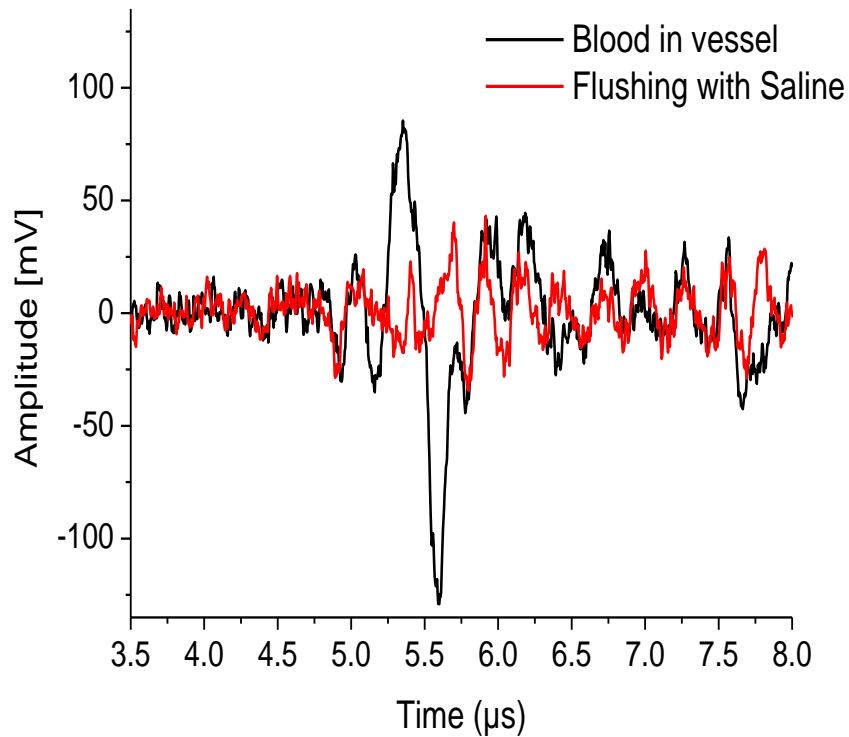


During occlusion: A: Time between max / min peak reduced ( $0.1 \mu\text{s} \approx 0.2 \text{ mm}$ )

B: Peak-peak time delay reduced  $\Rightarrow$  vessel thinner

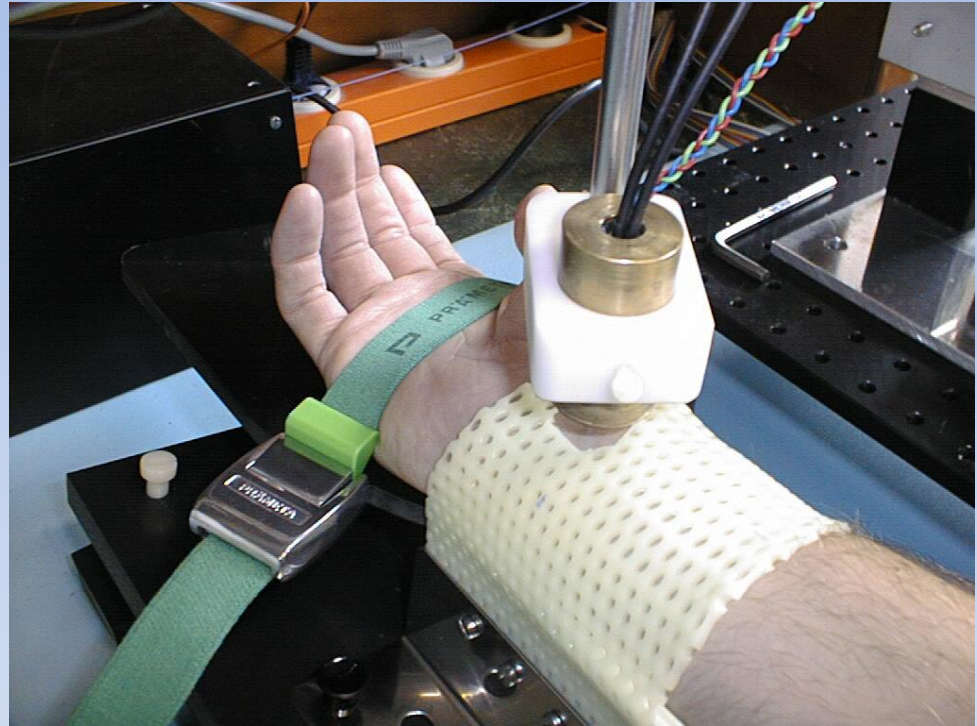
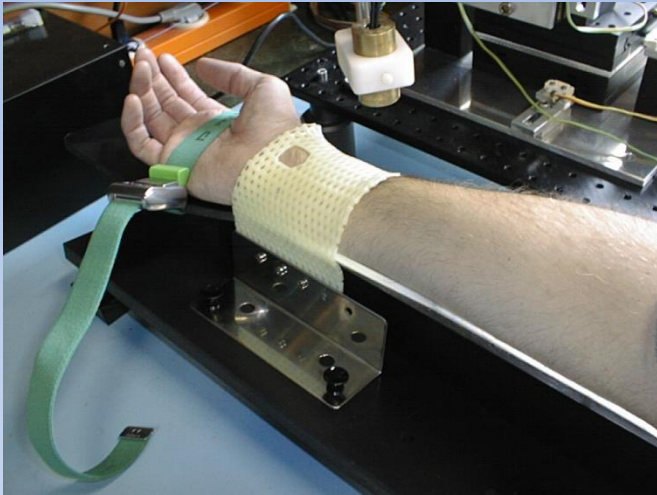
C: Peak-peak amplitude reduced  $\Rightarrow$  amount of blood lower

# PA-imaging of blood vessels in rabbit ear: flushing with saline





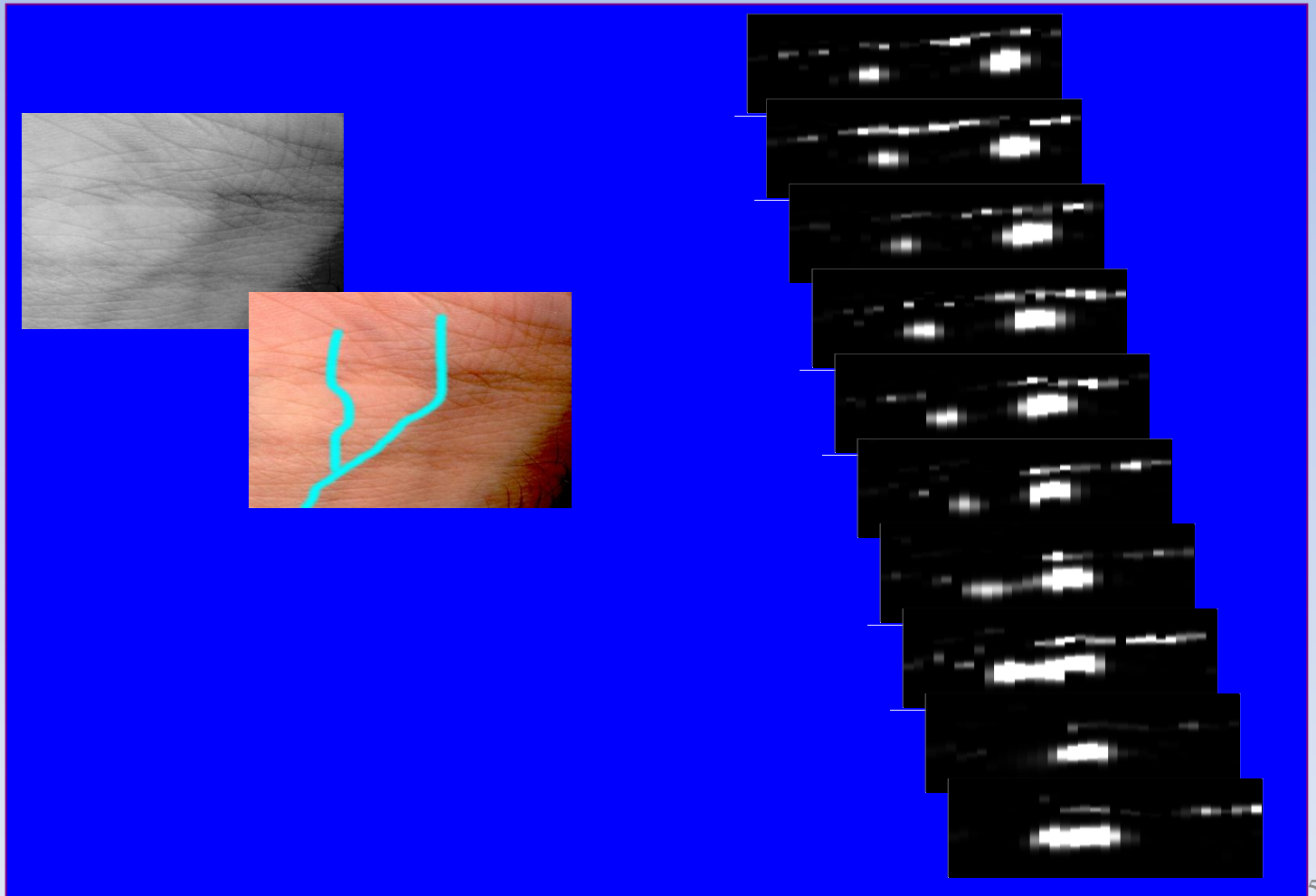
# In vivo Scan: Human Blood Vessels



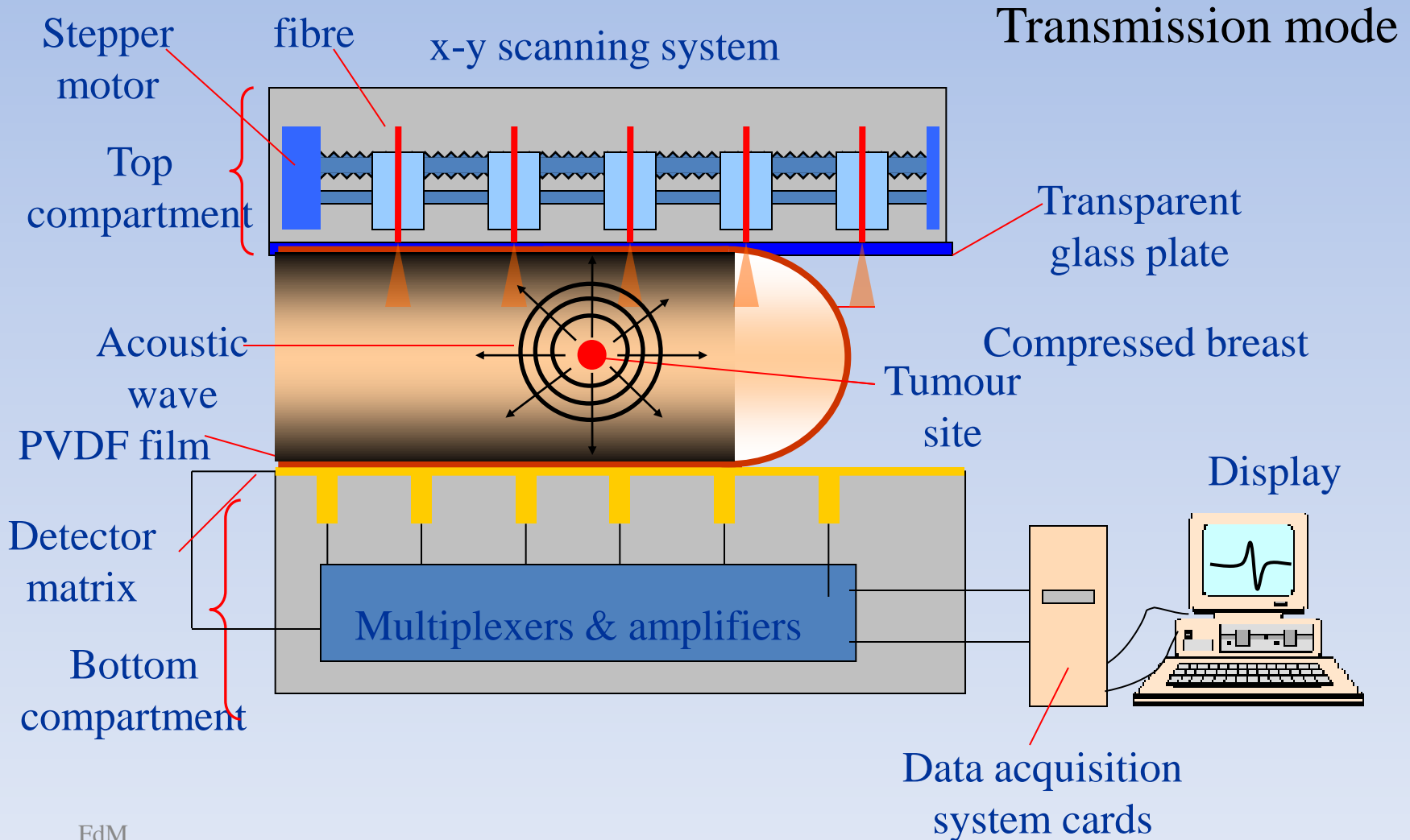


# PA-imaging of blood vessels in human arm

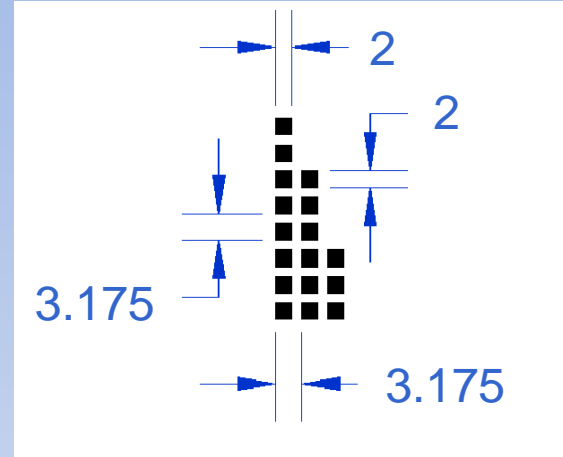
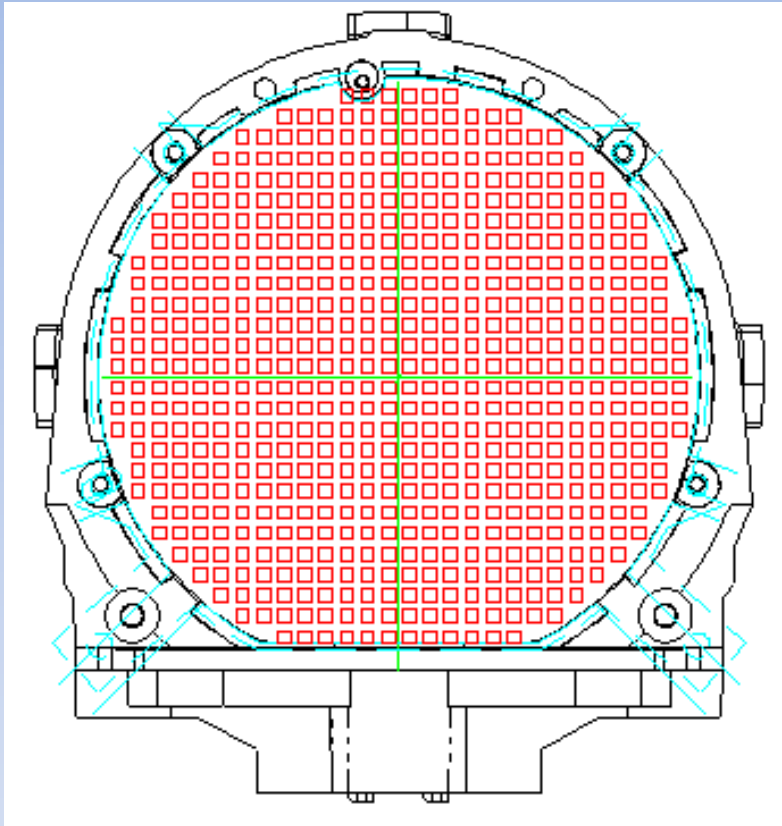
Depth  $\approx$  1-3 mm



# Photoacoustic Mammography



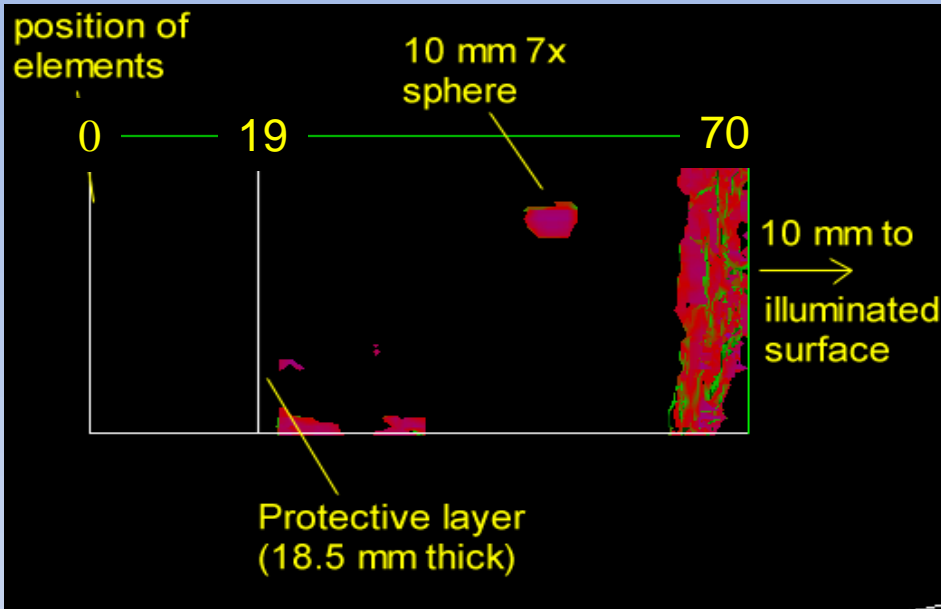
# The detector matrix



- 590 element microphone matrix
- active area: 90 x 85 mm

- arranged in a cartesian grid
- elements size: 2x2 mm
- separation: 3.175 mm
- 1 element accessed at a time

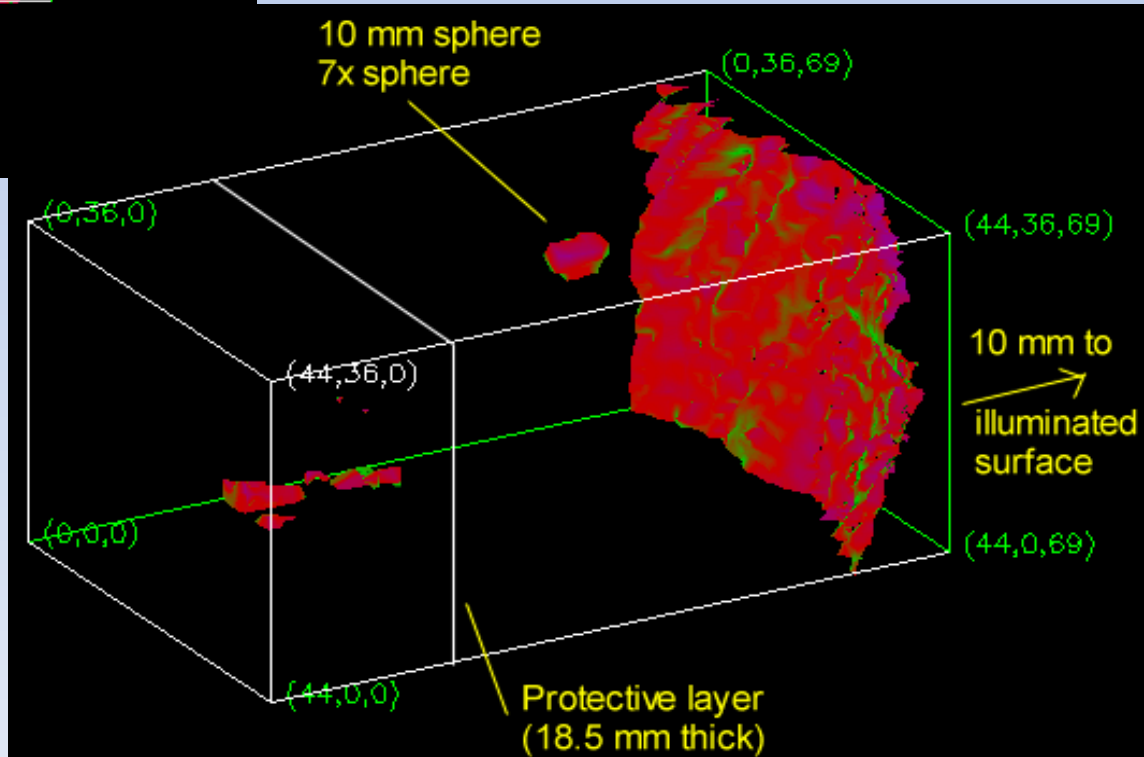
# Phantom: 10 mm, 7x sphere at depth 30 mm



Absorption contrast  
 Sphere : medium = 7 : 1

side view

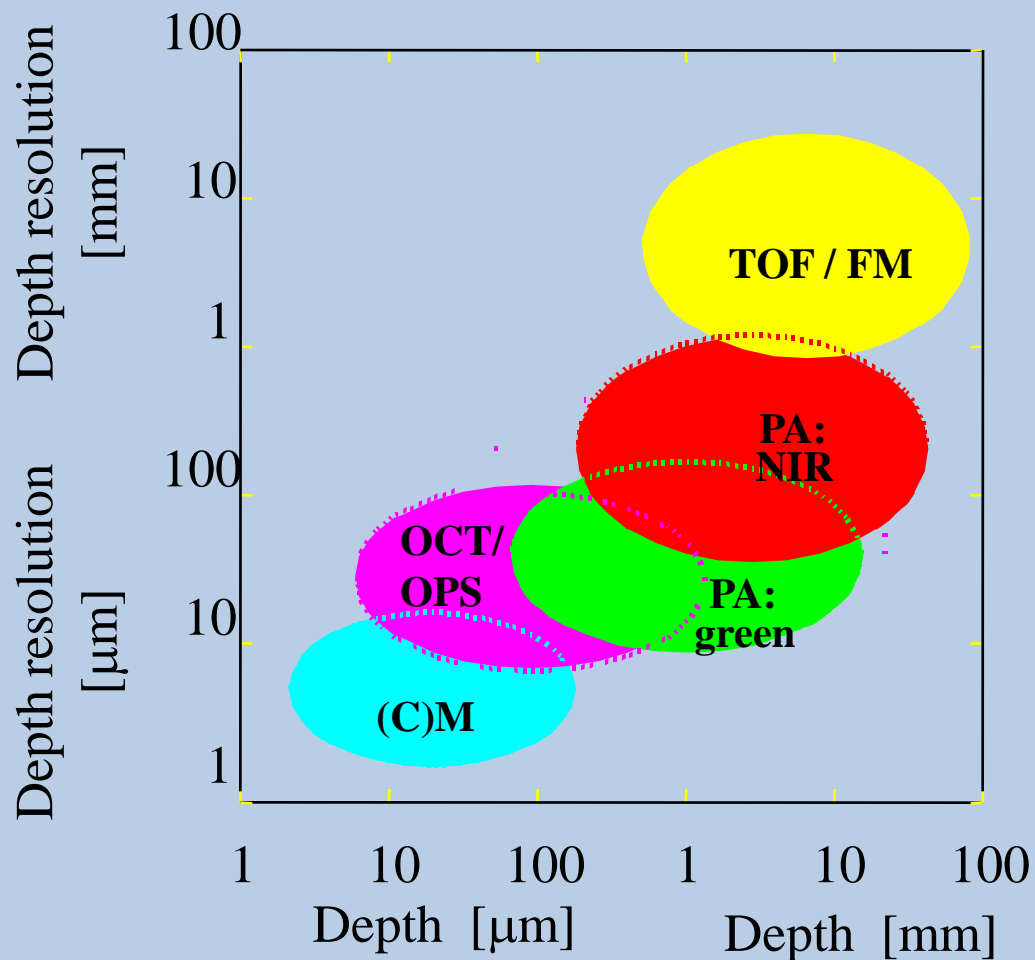
isometric view



# Photoacoustic Imaging

	Green 550 nm		NIR 800-1064 nm	
	dermis	blood	dermis	blood
• Absorption coefficient [1/mm]	0.03	32	0.01	1.2
• Scattering coefficient (reduced) [1/mm]	3	1	1	0.5
• Absorption Contrast	1000		100	
• Penetration into tissue [mm]	≈ 10		≈ 30	
• Applications	Cutaneous perfusion Wound healing Diabetes Vascular malformations Skin tumours		Cerebral perfusion Muscular perfusion Mammography Angiogenesis	

# Imaging methods for hidden structures in turbid media (tissue)



C(M) : (confocal) microscopy

OCT: optical coherence tomography

OPS: orthogonal polarization spectral imaging

PA: photoacoustics

TOF: time-of-flight tomography

FM: frequency-modulated tomography

# Photoacoustic Imaging Of Blood Vessels in Tissue

The end