



Electrical
Engineering,
Cardiology
and Physics

Cardiac
Imaging
Research



KU Leuven
UZ Gasthuisberg

Cardiovascular deformation imaging

Part II: cardiac

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Catholic University of Nijmegen, Nijmegen, THE NETHERLANDS

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October 18th, 2004



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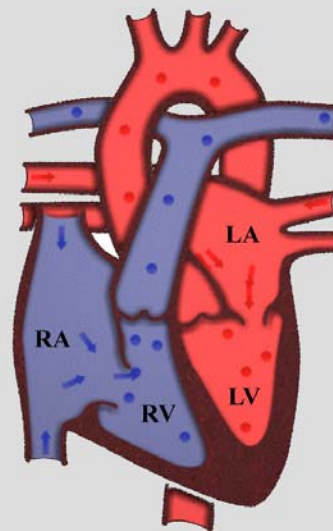
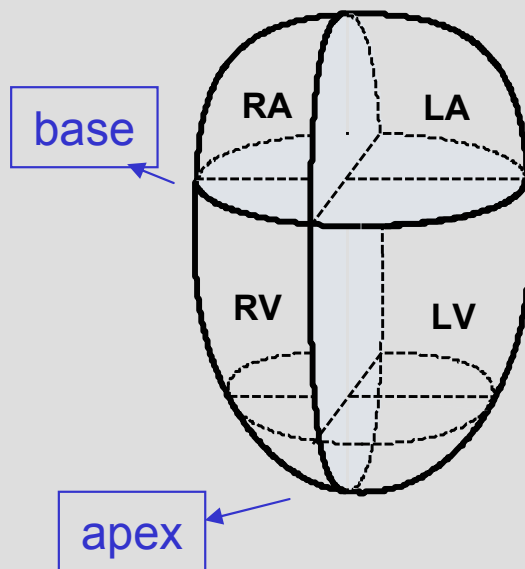
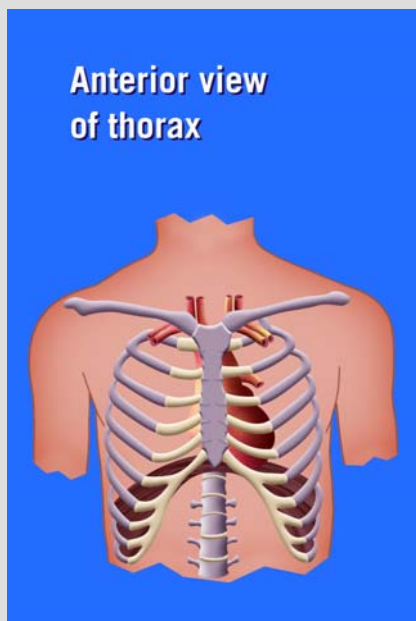
- Cardiac anatomy and function
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- Pitfalls of the current methodologies
- Estimating regional active stress development
- Estimating myocardial elasticity



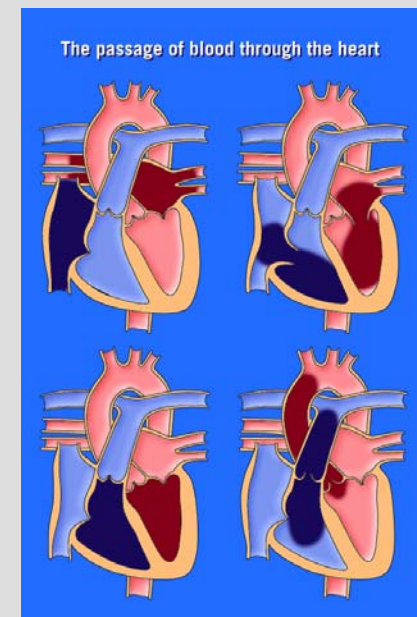
The heart: anatomy



Position of the heart in the human thorax



Circulation



4 compartment "box":

2 atria

2 ventricles

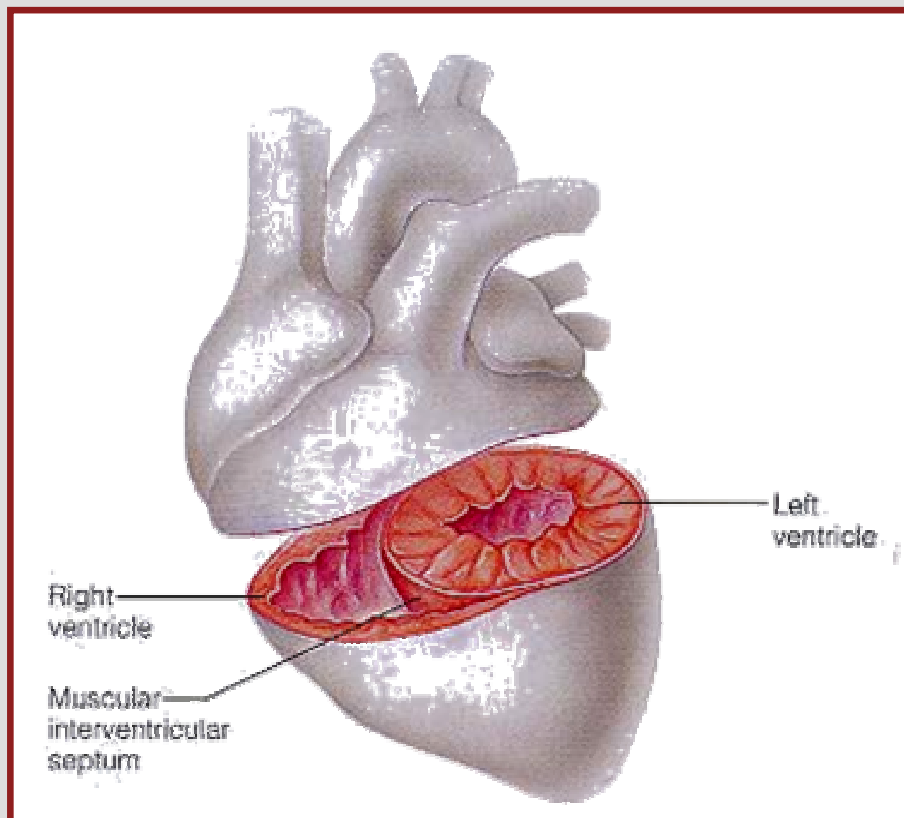
2 pumps in series:

Right heart: lung circulation

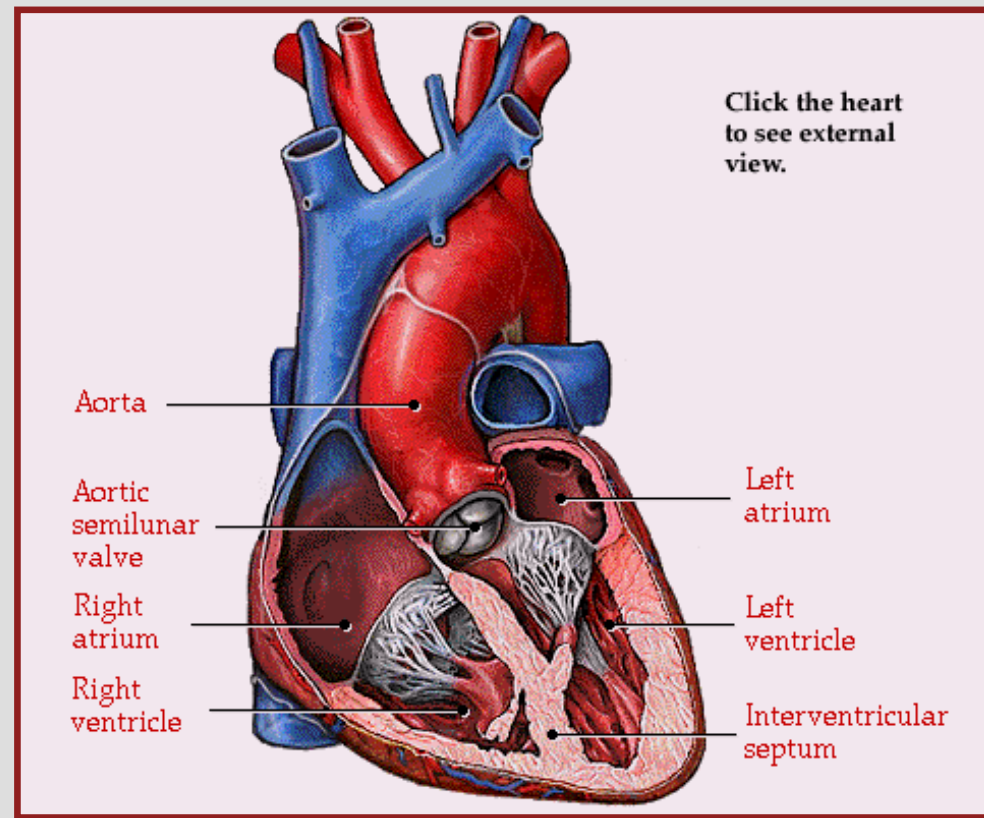
Left heart: systemic circulation



The cardiac muscle



Left ventricle =
Most important cavity of the heart
(systemic circulation)



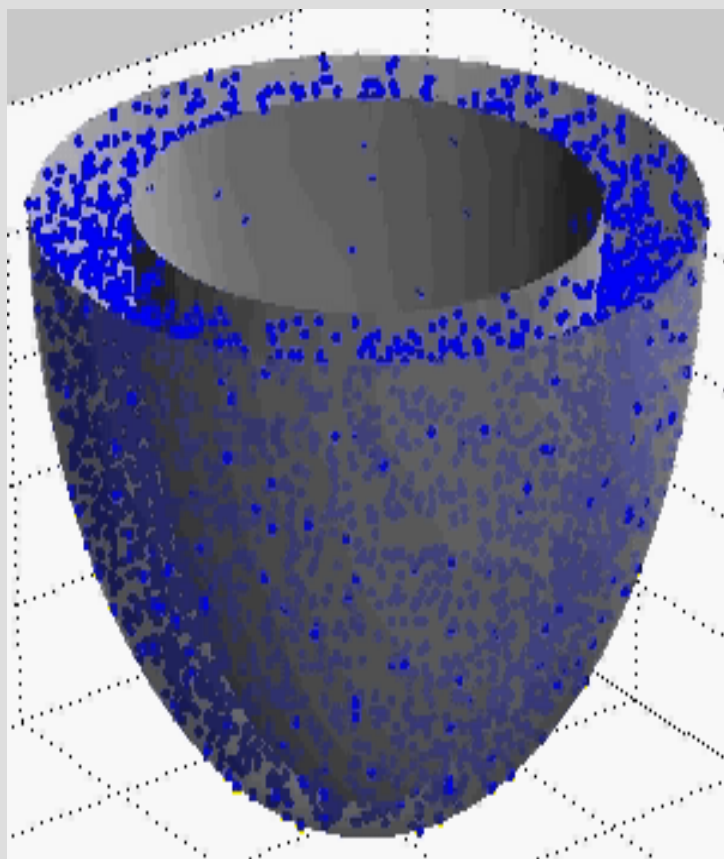
Left ventricle =
Most developed muscle)



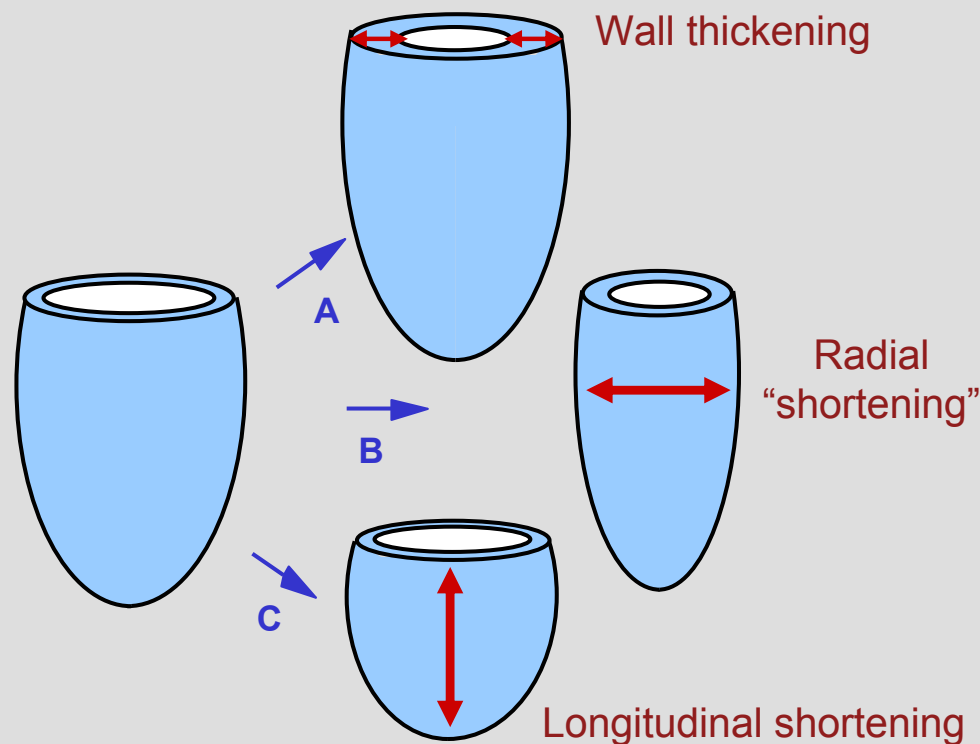
Cardiac function



Cardiac ejection/filling through myocardial deformation



Courtesy: S.I.Rabben, Univ. of Oslo, Norway





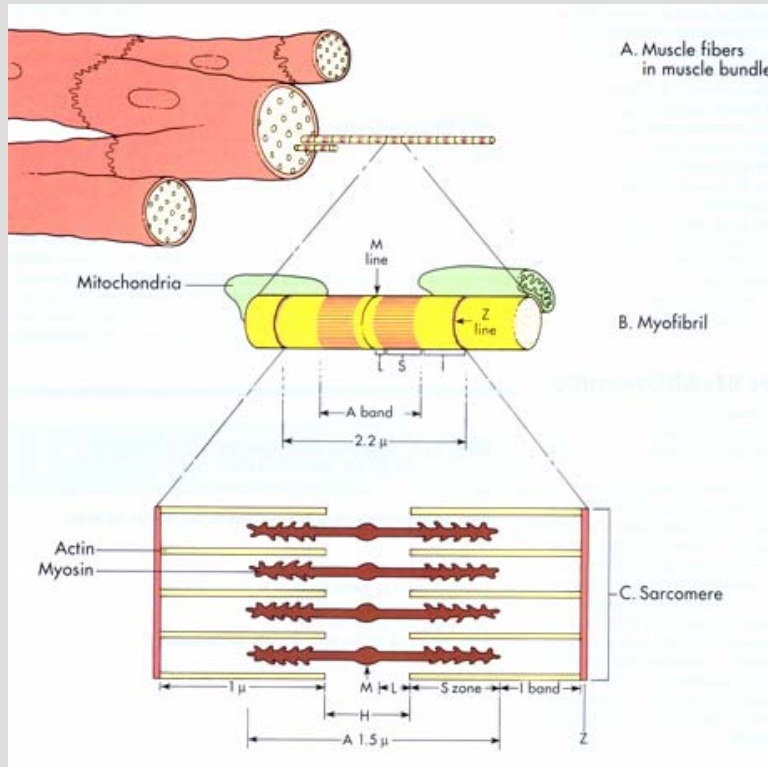
Ultrastructure



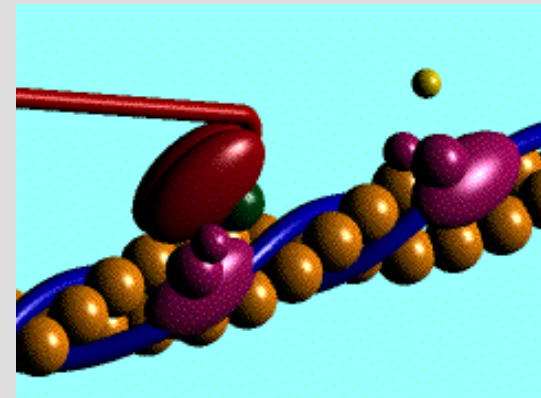
Fiber orientation



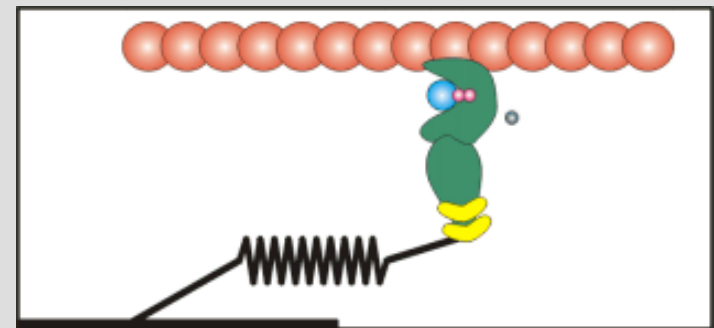
Ultra-structure



Actin-myosin interaction



Actin-myosin interaction



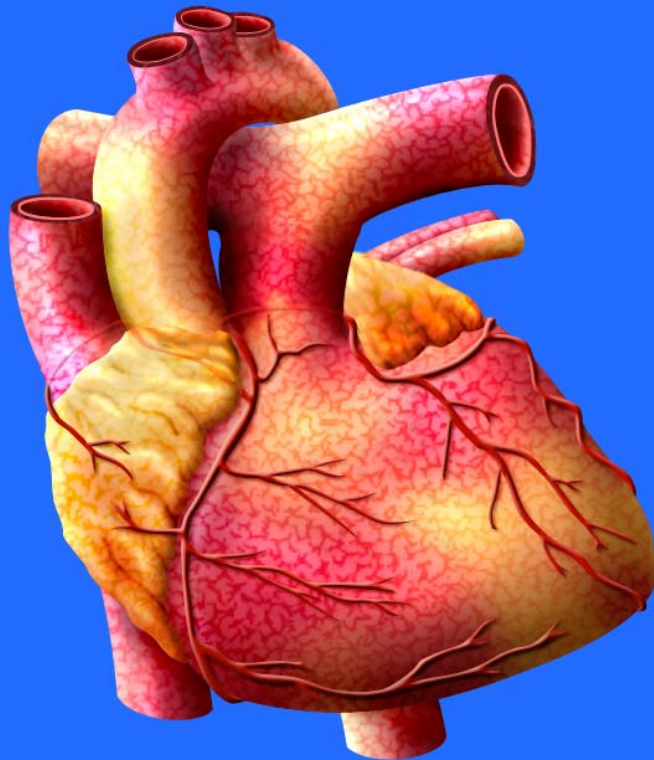


Coronary arteries



Coronary arteries

**Anterior heart showing
coronary arteries**



Most common
cardiac pathology:

Coronary Artery
Disease (CAD)

Therapy depends on:

- Severity
- Localization
- Period of time
-



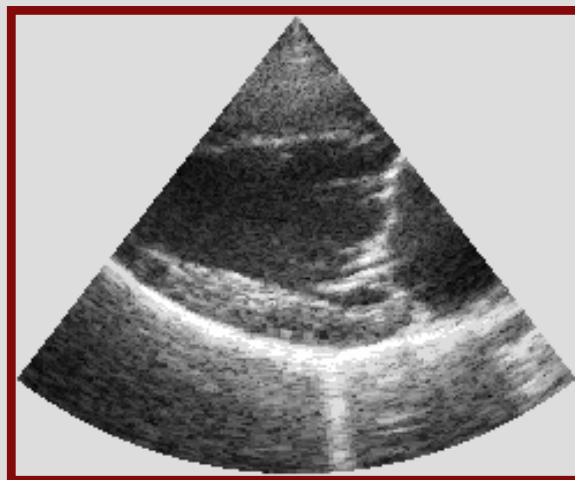
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Echocardiography: introduction



Strong points

- Real-time
- Non-invasive
- Relatively cheap
- Portable / bed-side
- Excellent temporal resolution

Clinical use

- Anatomy
- Blood flow
- Function
(= performance)



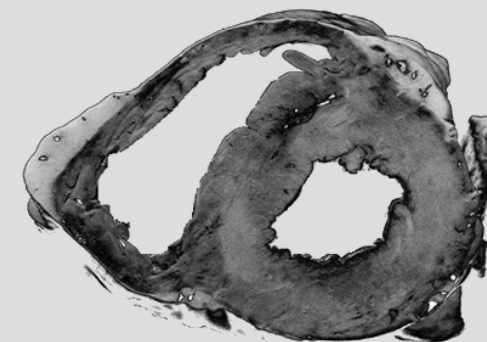
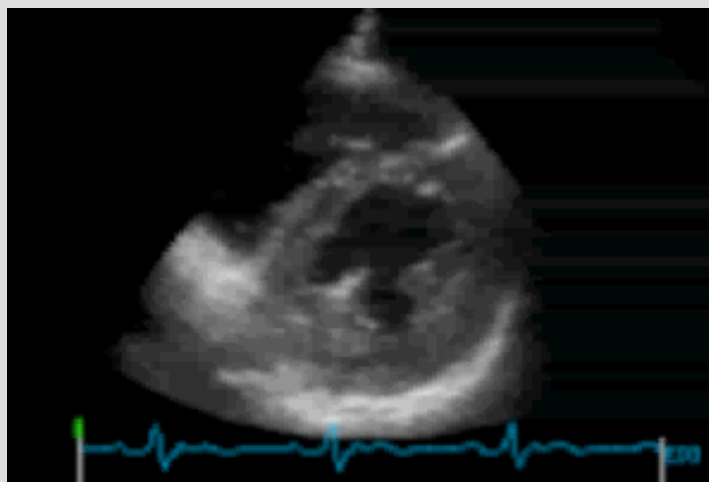
Standard echocardiographic views



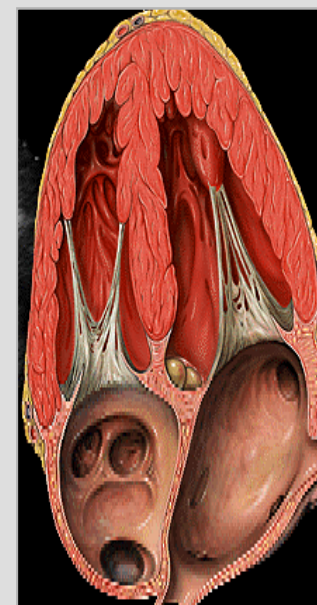
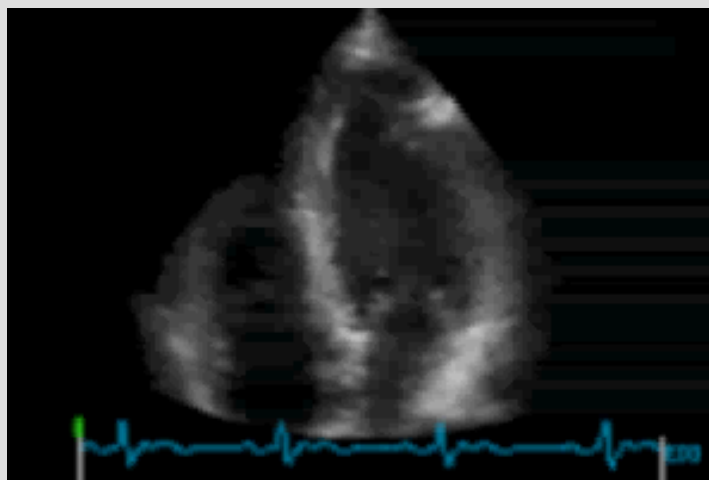
Parasternal long axis (M-mode)



Parasternal short axis

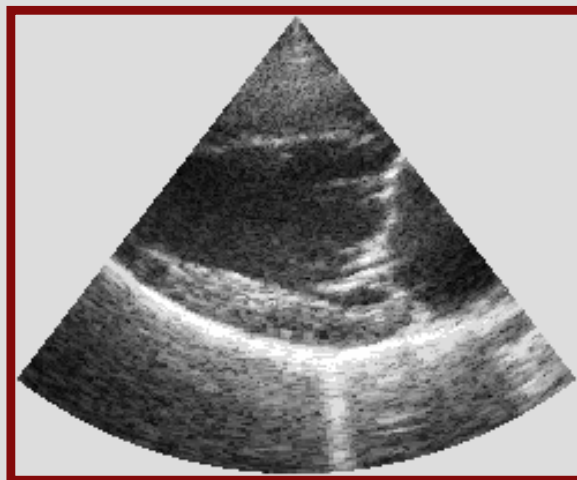


4 chamber view





Echocardiography: function



Qualitative

Visual assessment
and appreciation of
regional wall motion

Quantitative

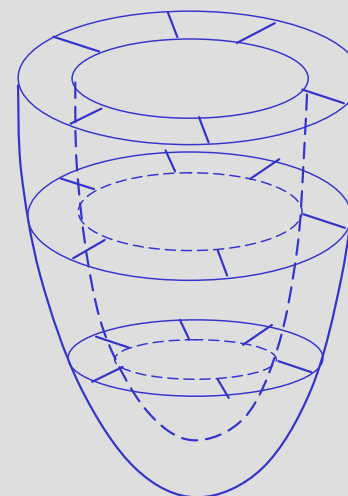
- Ejection fraction
 - Fractional shortening
 - Atrio-ventricular plane motion
 - E/A ratio blood Doppler
 - ...
- Global measures!



The quantitative assessment of *regional* myocardial function remains an important goal in clinical cardiology



17 segment model of the heart (LV)



basal

mid

apical

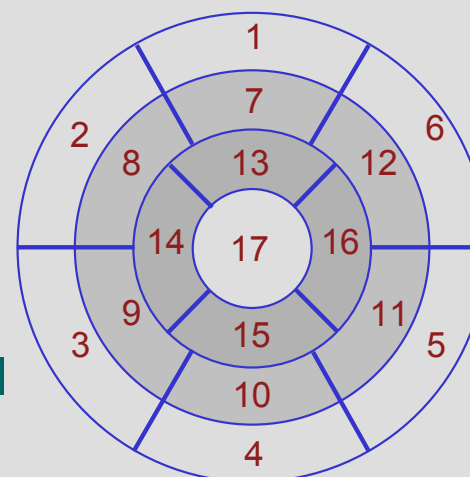
Anterior

Antero-septal

Antero-lateral

Infero-septal

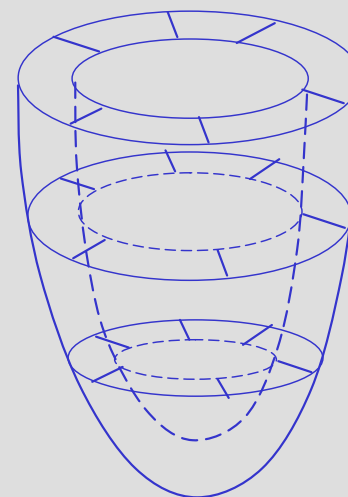
Infero-lateral



Inferior



17 segment model of the heart (LV)



basal

mid

apical

Anterior

A2C

Antero-septal

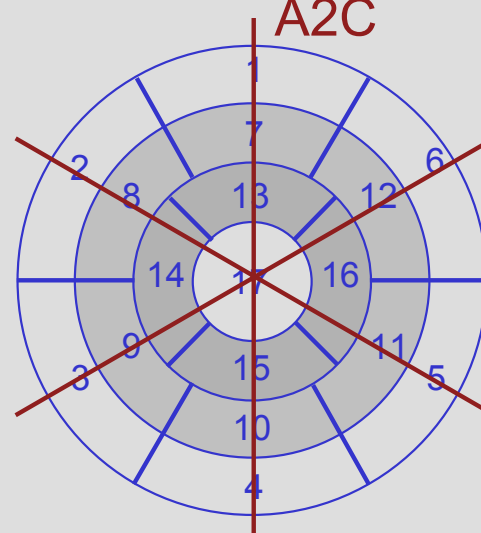
A4C

Antero-lateral

Infero-septal

A3C

Infero-lateral
(posterior)



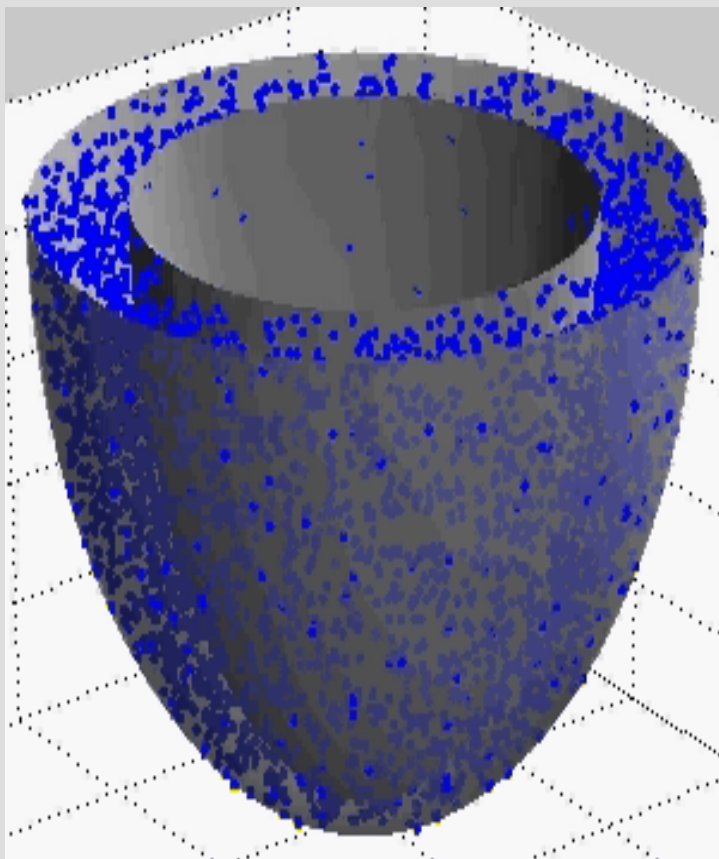
Inferior



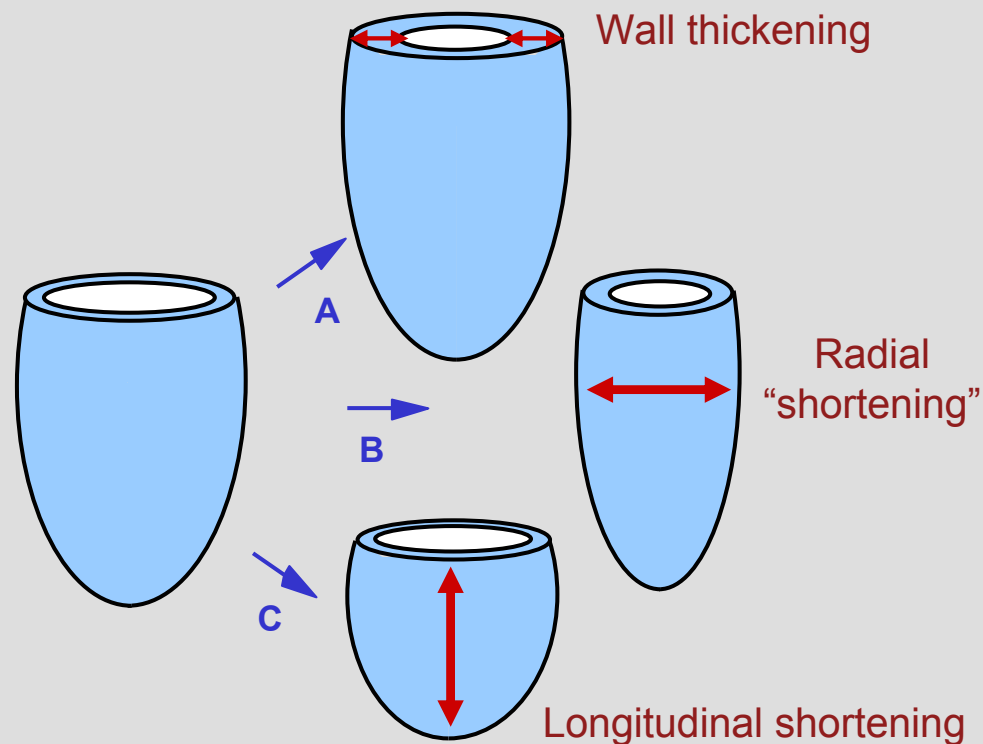
Regional myocardial function



Cardiac ejection/filling through myocardial deformation

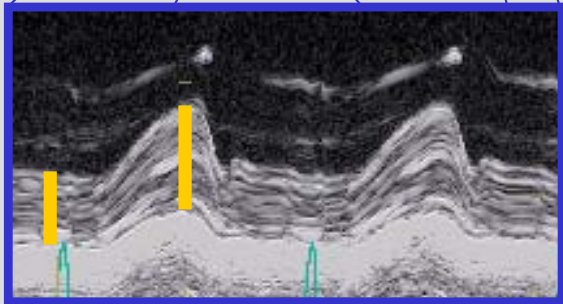
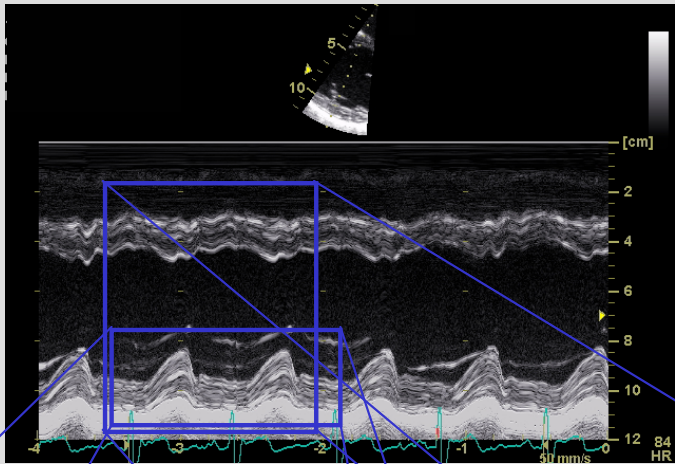


Courtesy: S.I.Rabben, Univ. of Oslo, Norway

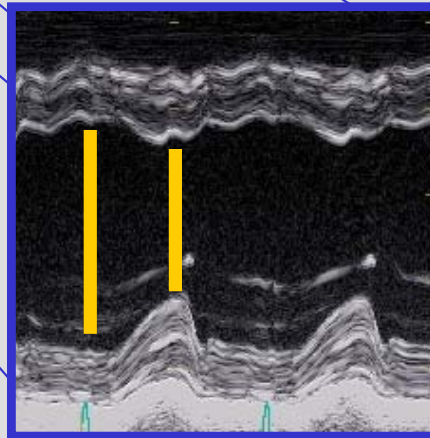


→ Measure local deformation to study the local contribution to ejection/filling (i.e. local myocardial function)

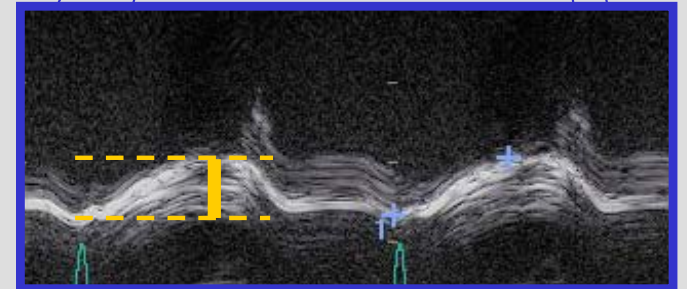
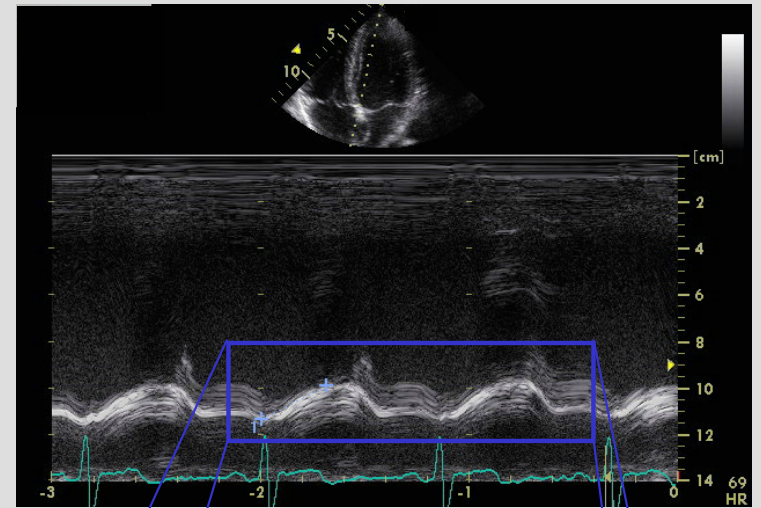
M-mode measurements



$$WT = \frac{ESWT - EDWT}{EDWT}$$



$$FS = \frac{ESD - EDD}{EDD}$$



$$D = EDL - ESL$$



Disadvantages of the M mode approach



- Dependent on endo- and epicardial border definition
 - ✓ Image quality dependent (cf. endocardial border)
 - ✓ Often difficult
 - ✓ User dependent and time consuming
- Relatively low spatial resolution
 - ✓ Use of envelope data (rather than RF)
 - ✓ Especially true for anatomic M-mode images
- Only *global* deformation characteristics can be measured
 - ✓ Global longitudinal shortening
 - ✓ Transmural (radial) wall thickening



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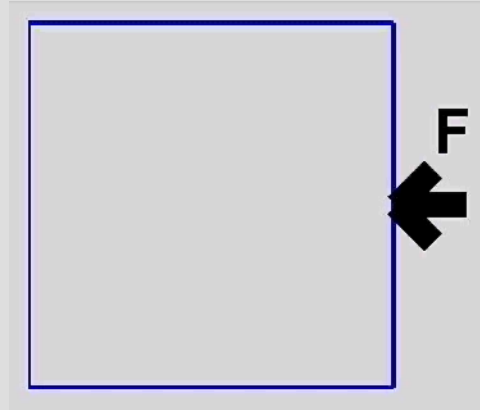
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Regional myocardial function



Passive elastic body



$$\sigma_{\text{load}}(t) = E \times \epsilon(t)$$



Wall stress
(total force on a segment)



segmental
elasticity



resulting
deformation

cf. elastography



Strain imaging \approx elasticity imaging



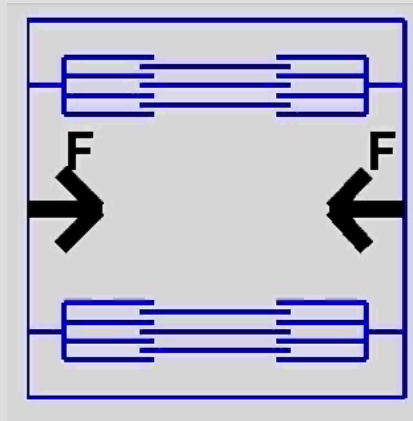
Strain image = elastogram



Regional myocardial function



Active elastic body



Active contractile stress

$C(t)$

+

$\sigma_{load}(t)$

=

E

x

$\epsilon(t)$

Regional myocardial function
(contractility)

Wall stress
(total force on a segment)
cavity pressure,
geometry,
neighbouring segment interaction

segmental elasticity

resulting deformation



If the assumption is made that external stresses (loading) can be neglected, local strain directly relates to local contractile stress



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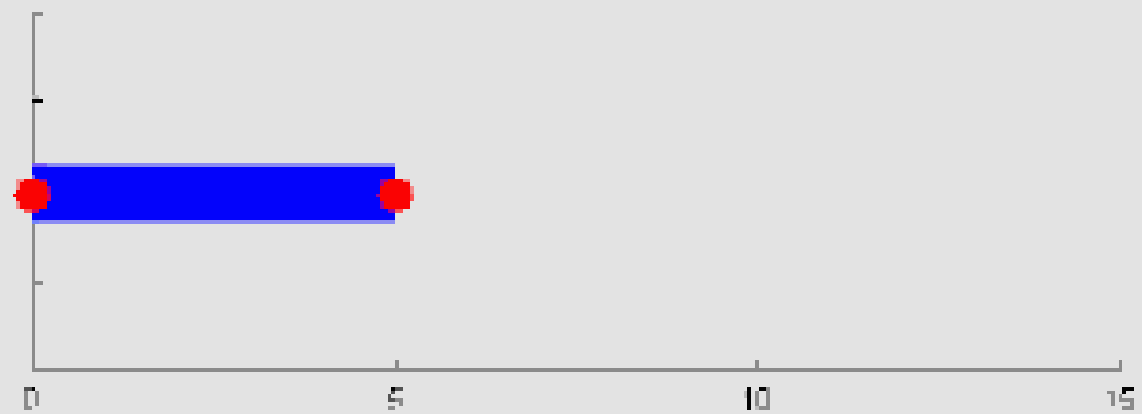
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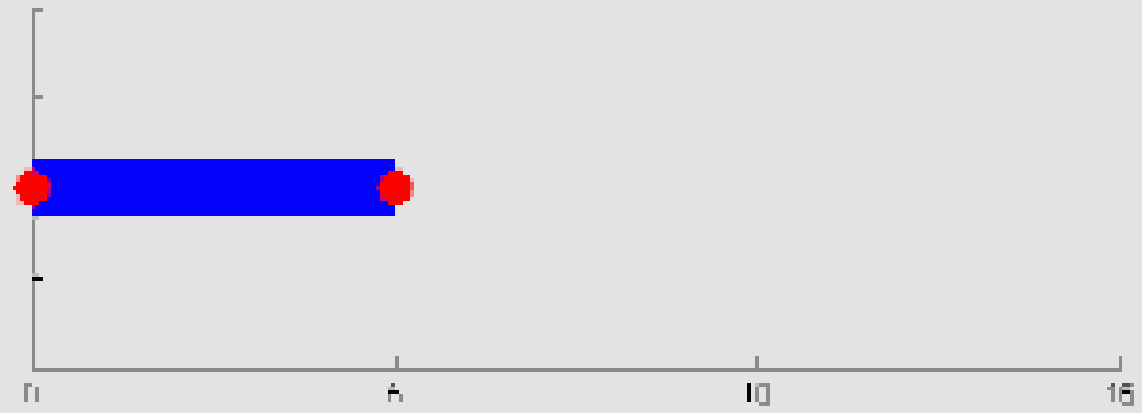
Velocity Gradient ~ Strain



Global motion
 $v_1 = v_2$



Lengthening
 $v_1 \neq v_2$



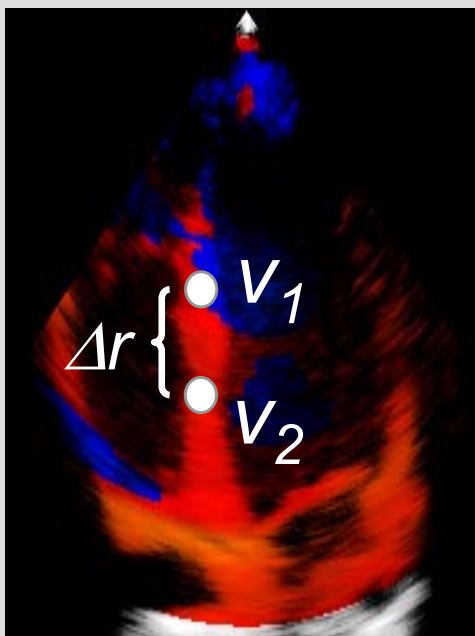
The spatial gradient in velocities (instantaneous motion) = strain rate (instantaneous strain)

$$\longrightarrow \frac{V_2 - V_1}{\Delta x} = \dot{\epsilon}$$

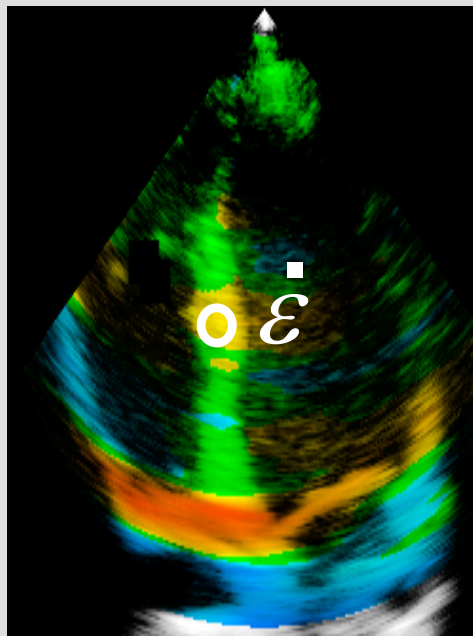


Cardiac Deformation Imaging

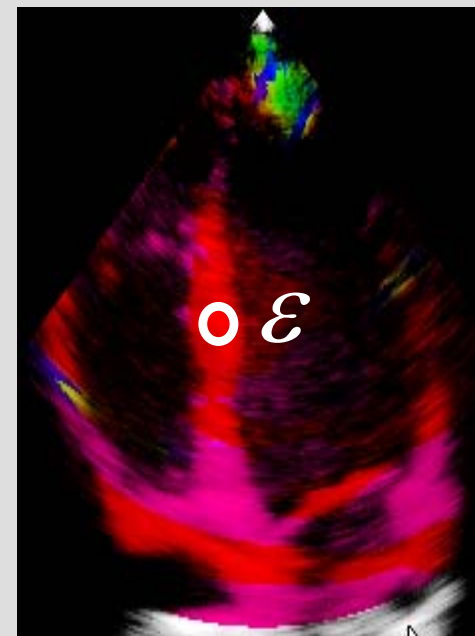
Velocities



Natural strain rate



Natural strain



Calculate spatial
gradient

Integrate
temporally



Strain and strain rate estimation =
motion/velocity estimation + post-processing

Note:

<i>Elastography:</i>	motion	→	(instantaneous) strain	→	cumulative strain
<i>SRI:</i>	velocity	→ ^{grad}	strain rate	→ ^{int}	strain



Measuring myocardial motion and deformation I

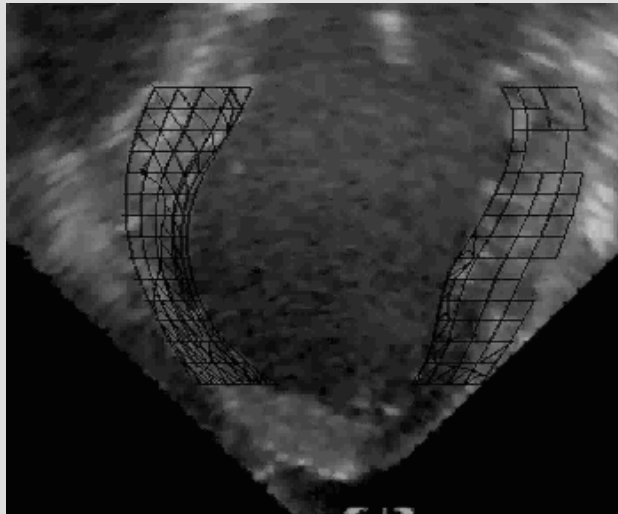


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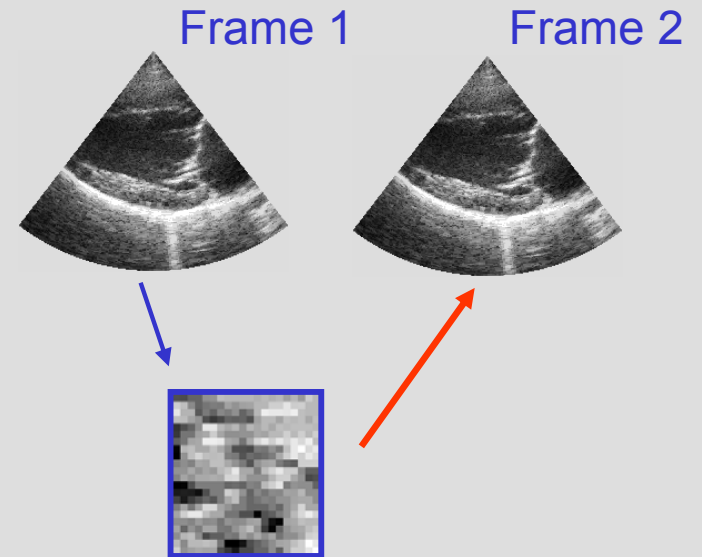
B-mode (envelope) data

Segmentation –
mechanical model based

Speckle tracking



Papademetris *et al.*,
Medical Image Analysis 5(1):17-28, 2001
IEEE-TMI 21(7):786-800, 2002



Optical flow ($E_x u + E_y v = -E_t$)

J. Meunier *et al.*, 1984
C. Jansen *et al.*, *EuroEcho*, 2002

Cross correlation based (block-matching)

G. Trahey *et al.*, *IEEE-TBME*, 34:965-967
E. Konofagou *et al.*, *IEEE Symposium* 2002

Behar *et al.*,
Ultrasonics,
43:57-65, 2004



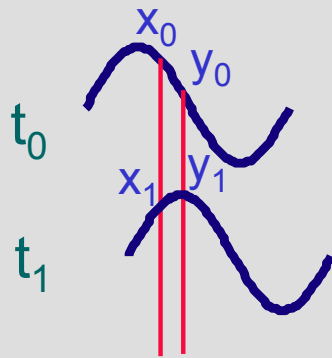
Measuring myocardial motion and deformation II



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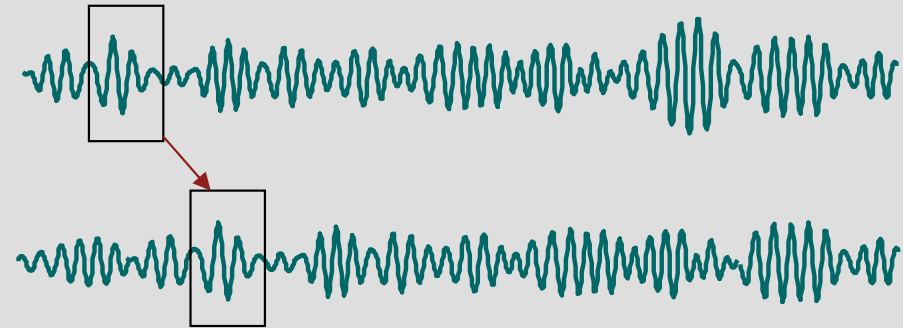
RF data

Auto-correlation



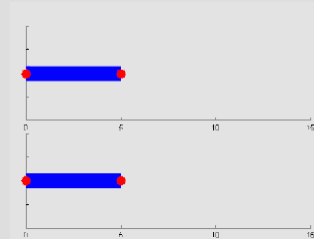
M-mode /
B-mode
+
Off-line /
Real-time

Cross-correlation



Motion → Velocity

Spatial
gradient



Instantaneous strain → Strain rate

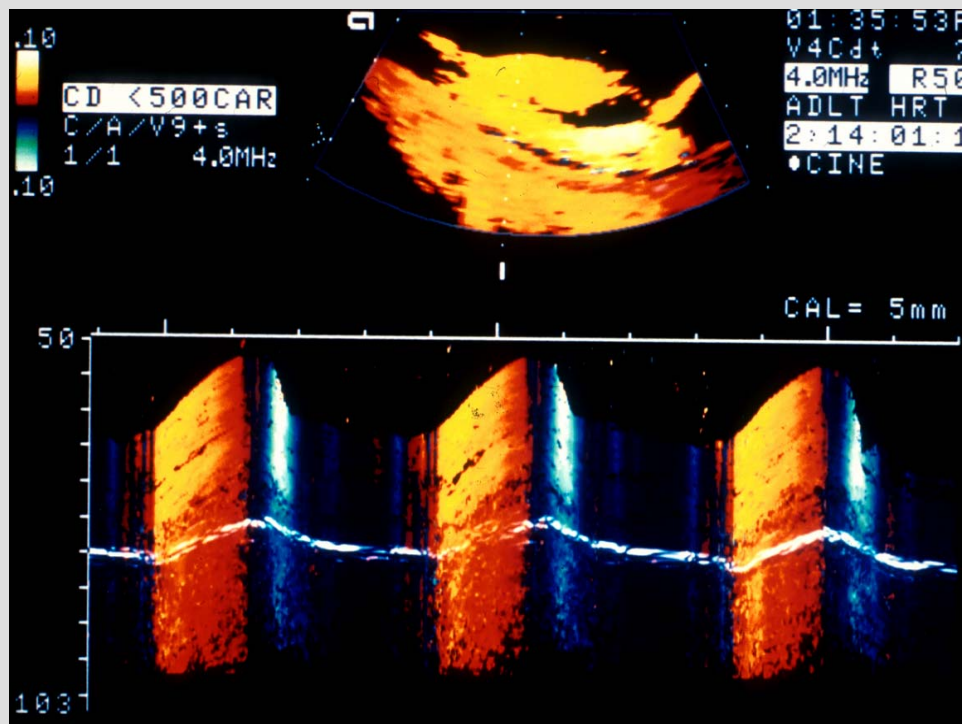
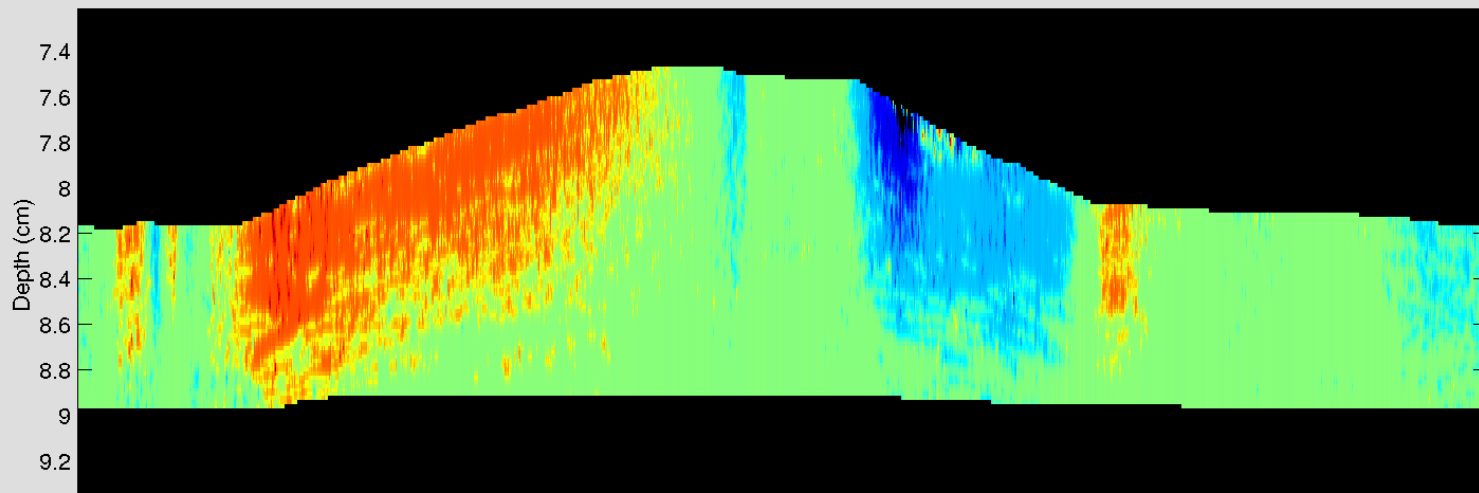


Auto- vs. cross-correlation



Cross-correlation

Broad bandwidth
data



Auto-correlation

“Classic” velocity (gradient) method

Similar but narrow band



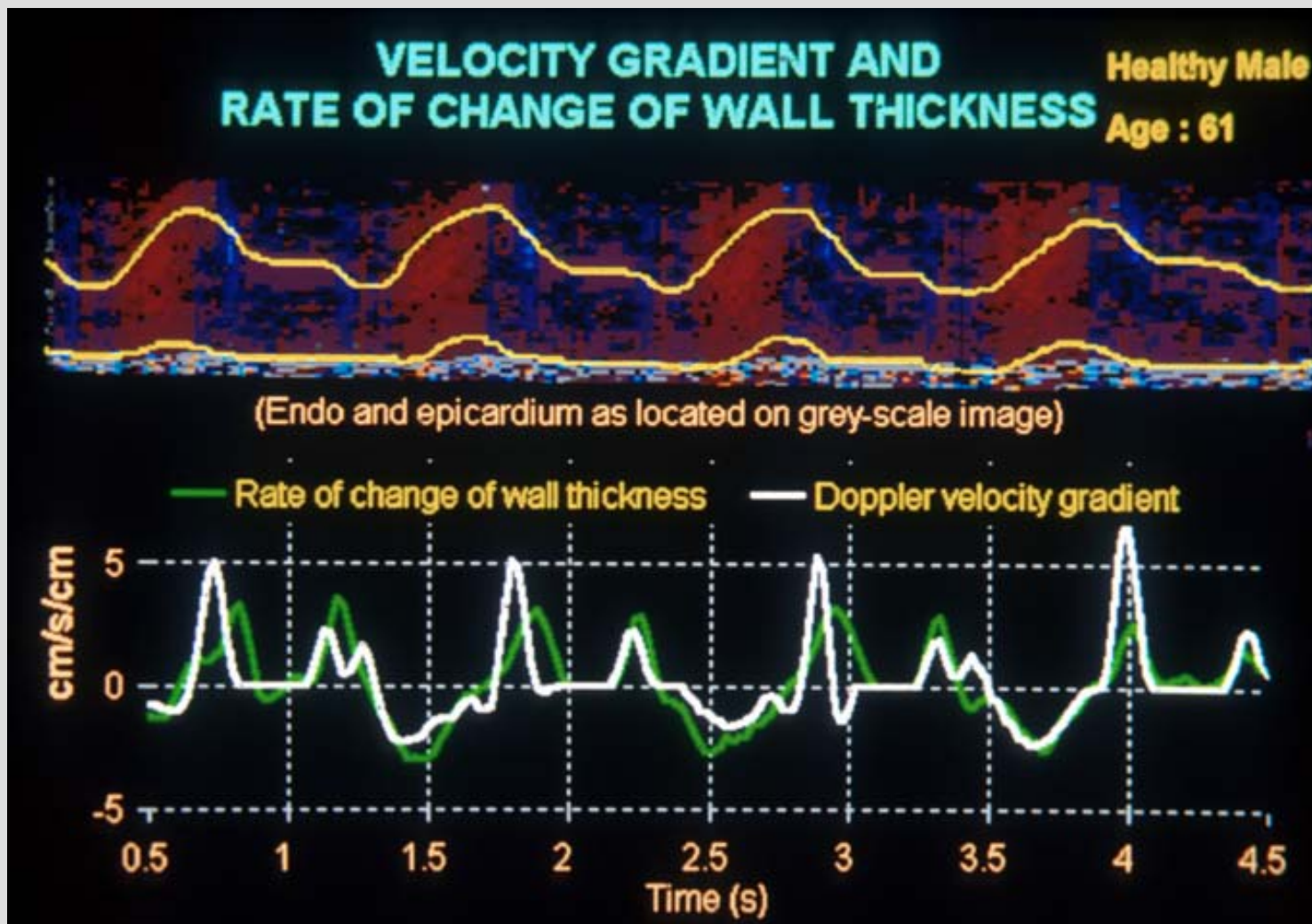
Spatial resolution intrinsically worse
but *real-time*



Auto correlation strain (rate) imaging



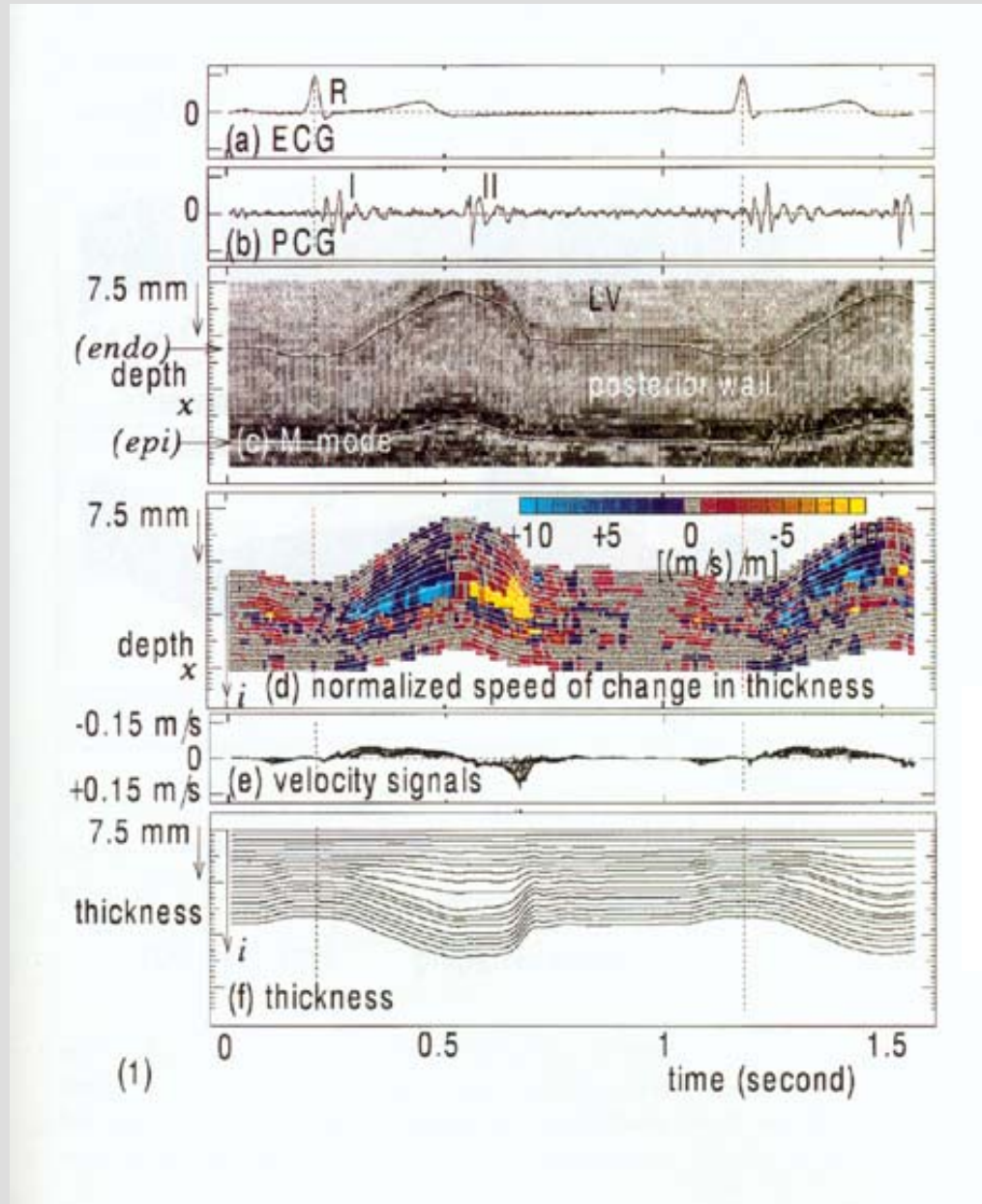
Conventional DMI imaging + offline post processing



Courtesy: George Sutherland '92



X correlation strain (rate) imaging



- Indicate initial position endo- and epicardium
- Automatic:
 - Transmural division
 - RF-tracking
 - Strain rate estimation
- Real-time system ('99)

Courtesy: IEEE-UFFC '97
H.Kanai
Sendai University, Japan



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Cardiology

Cardiac Imaging Research

Principle of deformation imaging



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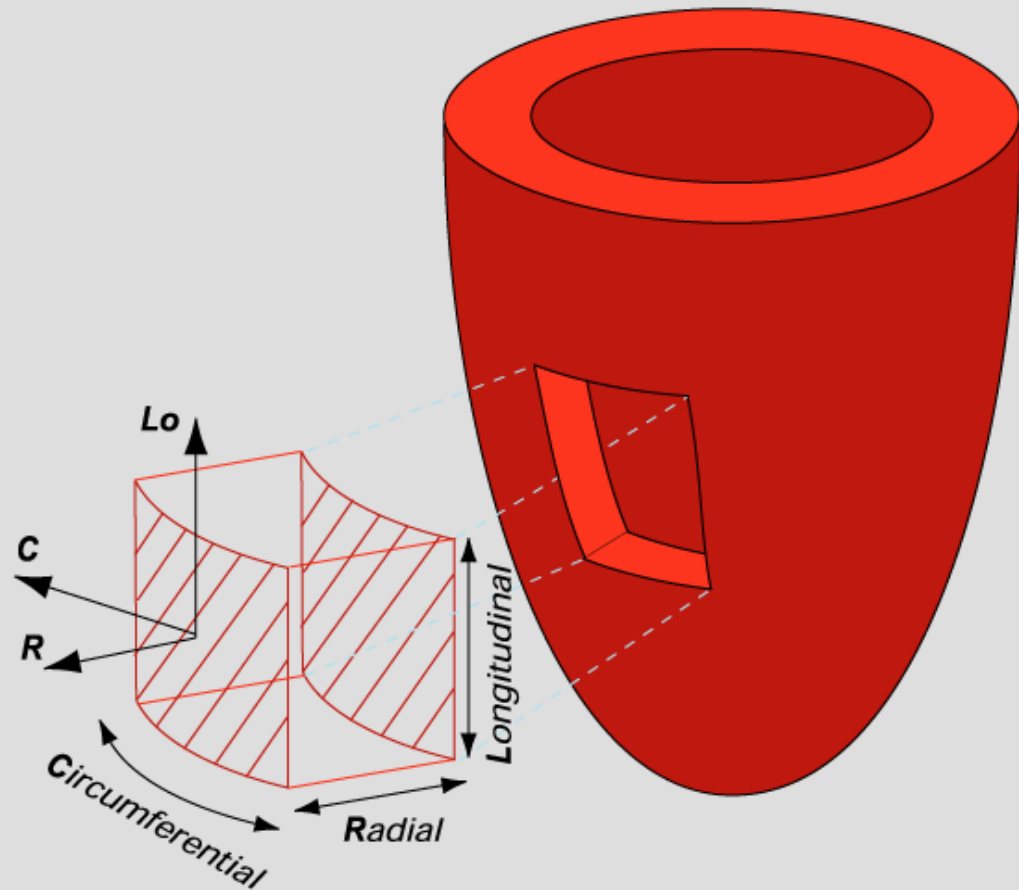
Local cardiac coordinates



- Radial
- Longitudinal
- Circumferential

↓

$$\boldsymbol{\varepsilon} = \begin{bmatrix} \varepsilon_{RR} & \varepsilon_{RC} & \varepsilon_{RL} \\ \varepsilon_{CR} & \varepsilon_{CC} & \varepsilon_{CL} \\ \varepsilon_{LR} & \varepsilon_{LC} & \varepsilon_{LL} \end{bmatrix}$$



Facilitates physical interpretation and mathematics of the strain values
(e.g. RR = wall thickening; CC/LL = circumferential/longitudinal shortening)

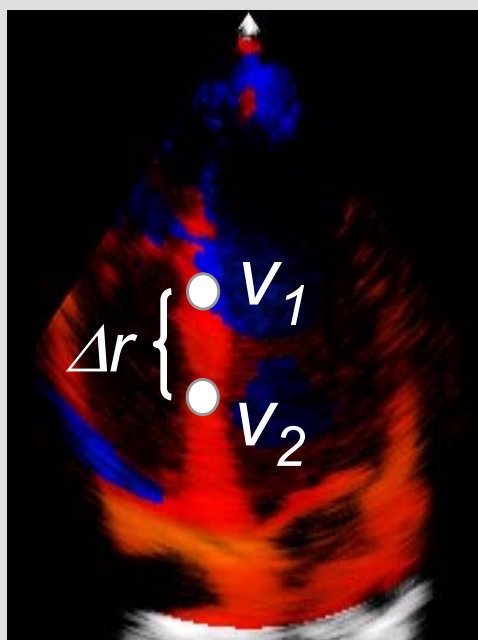


Strain and strain rate imaging

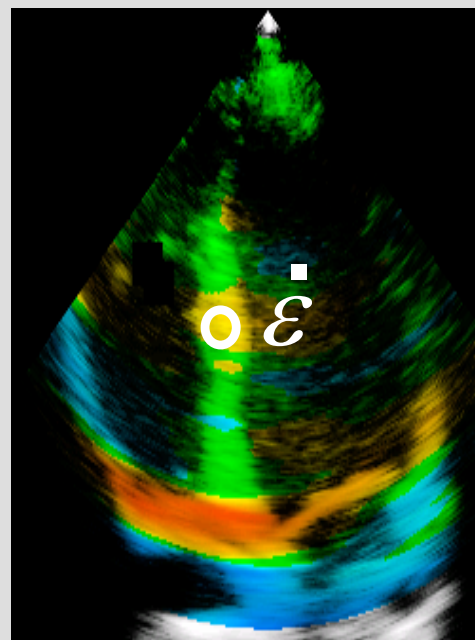


Most *animal and clinical* studies have been based on MVI (myocardial velocity imaging) data based on the auto-correlation methodology

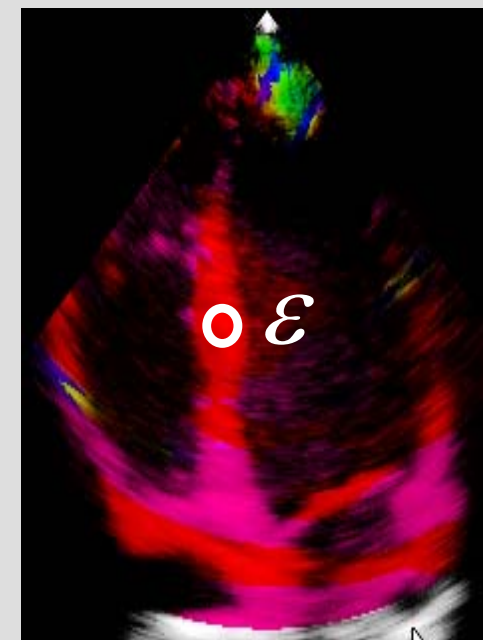
Velocities



Natural strain rate



Natural strain



Calculate spatial
gradient

Integrate
temporally



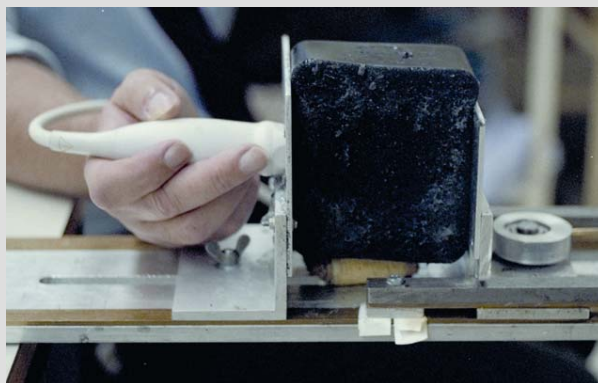
Strain and strain rate estimation =
motion/velocity estimation + post-processing



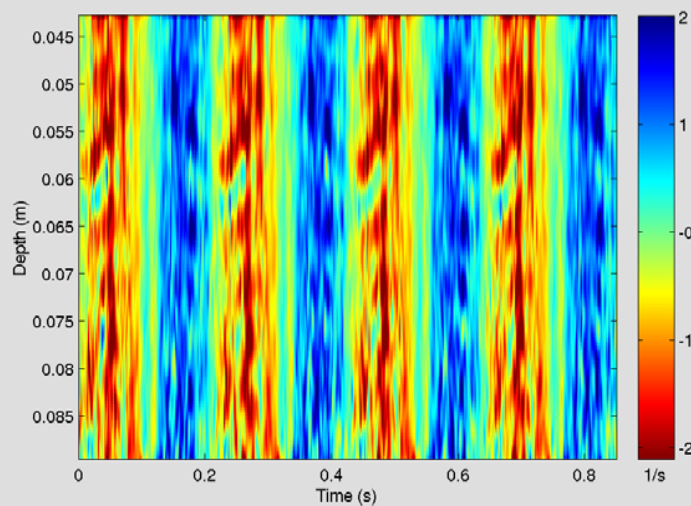
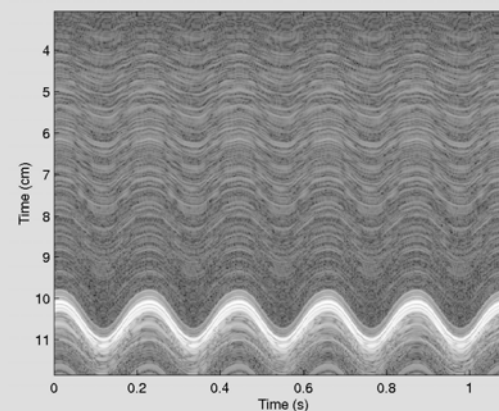
Validation in-vitro I



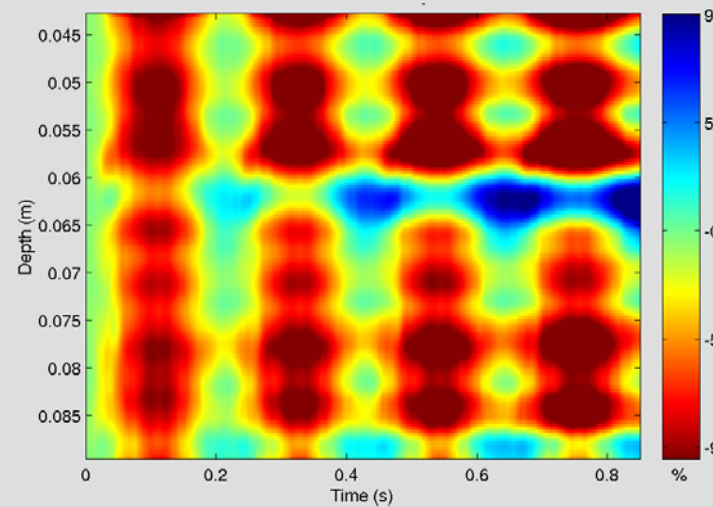
Cyclic
compression
of approx.
10% at 5Hz



M-mode



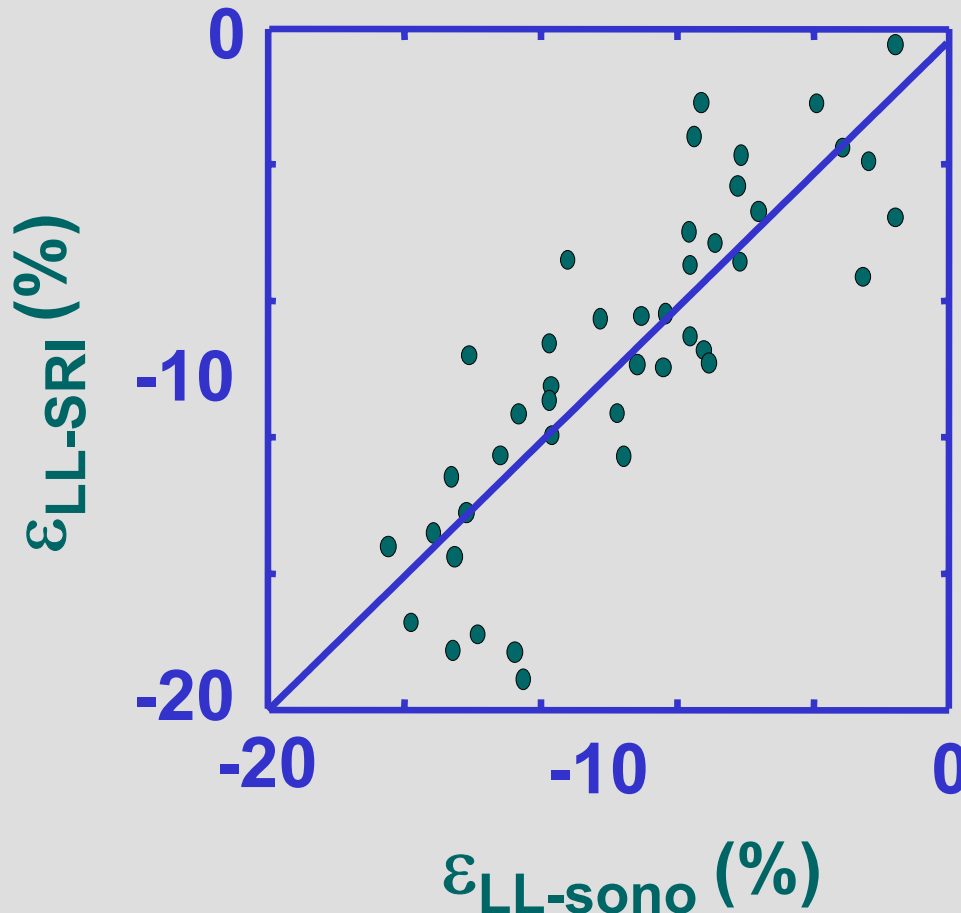
Strain rate



Strain



Validation in vivo I: SRI vs sonomicrometry

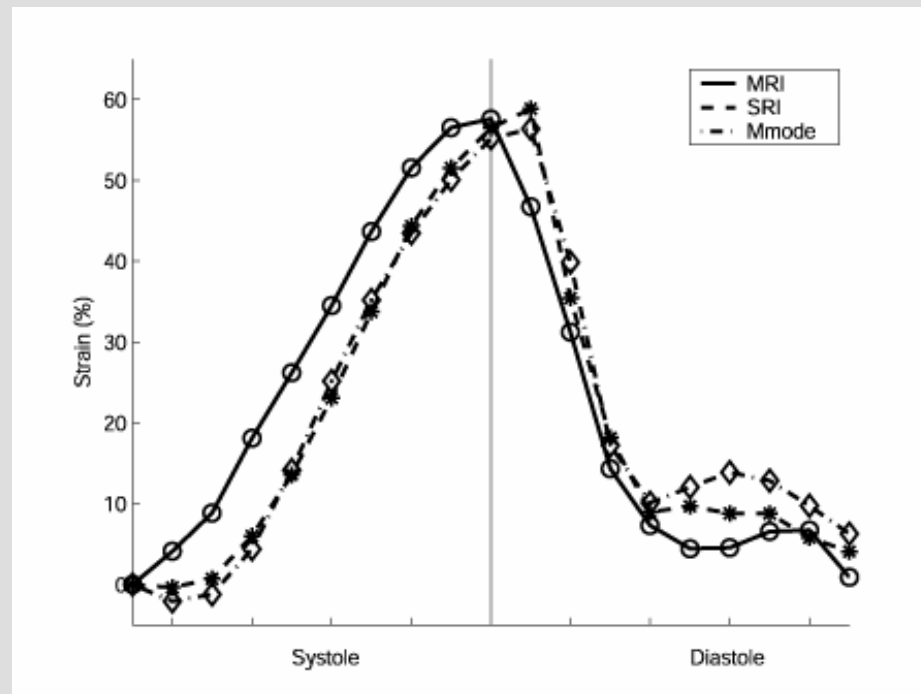
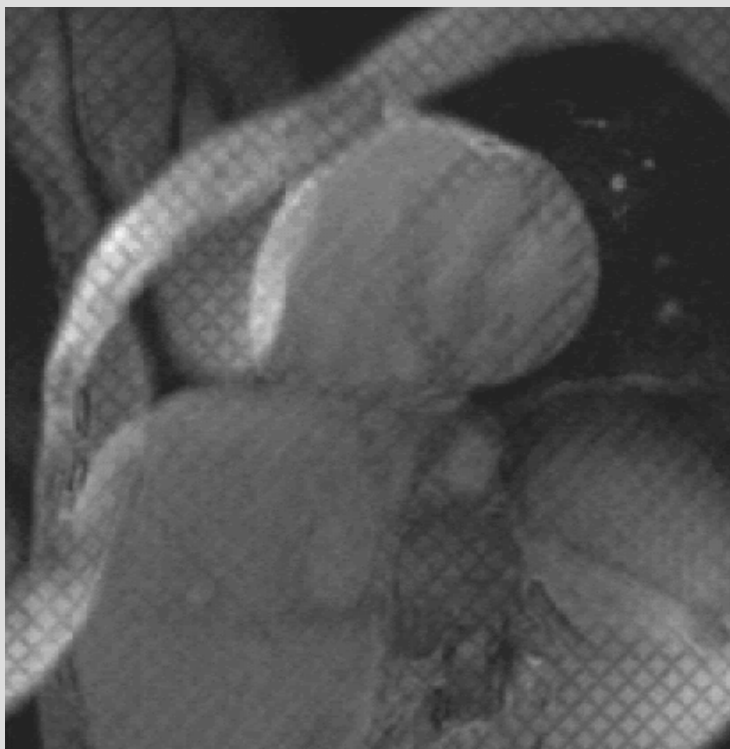


Linear correlation
 $\epsilon_{SRI} = 0.98 \cdot \epsilon_{sono} - 0.44$
 $R = 0.84$ - $N = 43$
 $p < 0.0001$

Courtesy: Fadi Jamal 2001
Lyon, France



Validation in vivo II: SRI vs MRI (tagging) / Mmode



Intra-class Correlation Coefficient: 0.69



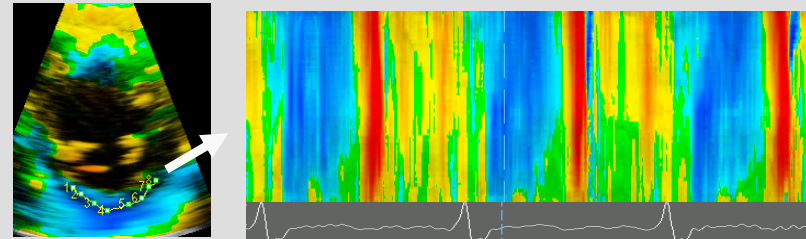
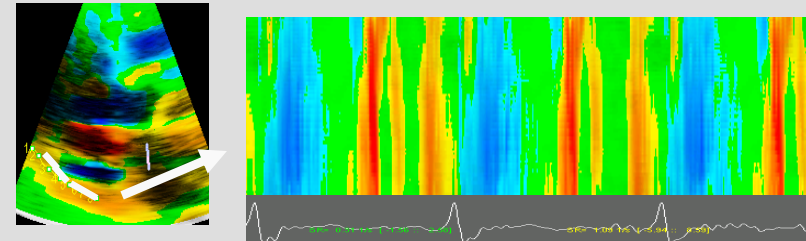
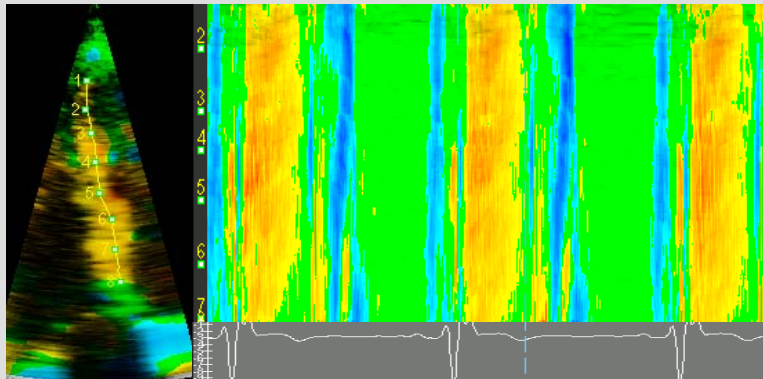
Normal Strain Rate



Longitudinal



Radial



Systolic shortening → **compression**

Systolic thickening → **expansion**

Diastolic lengthening → **expansion**

Diastolic thinning → **compression**

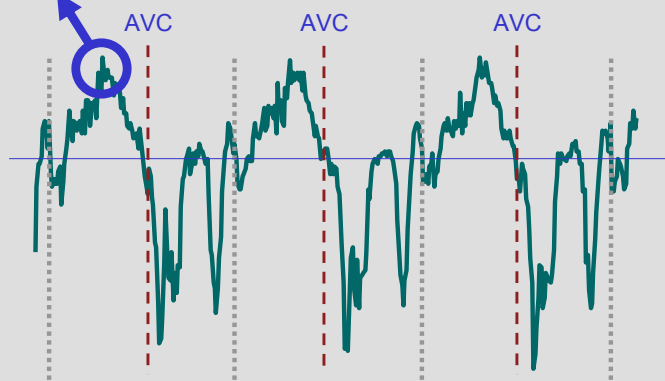
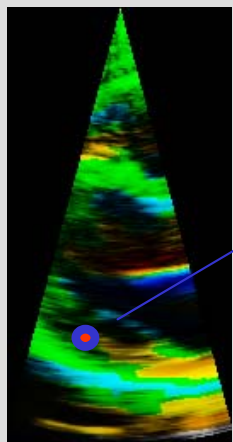


In-vivo normal examples



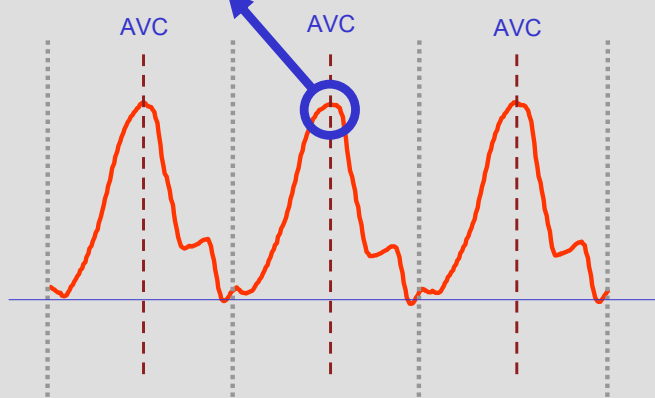
Peak systolic SR

Strain rate

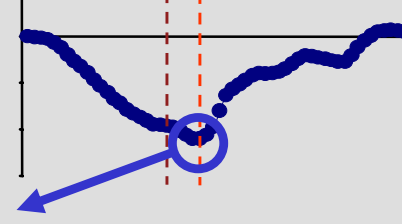
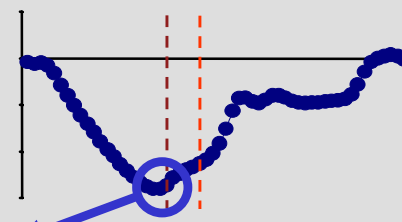
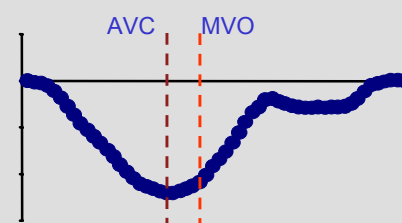
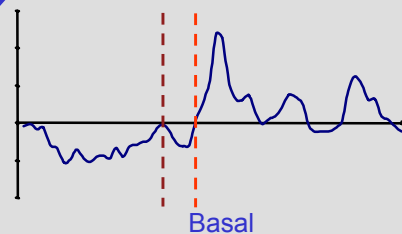
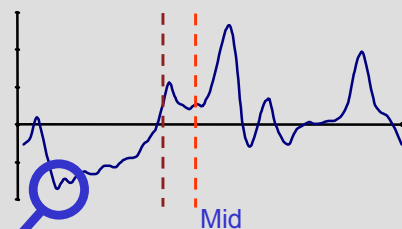
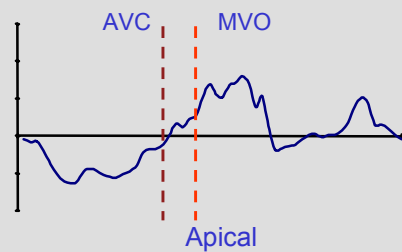
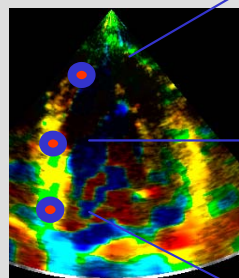


End-systolic ϵ

Strain



Time
Integration
→



Peak systolic SR

End-systolic ϵ

Post-systolic ϵ

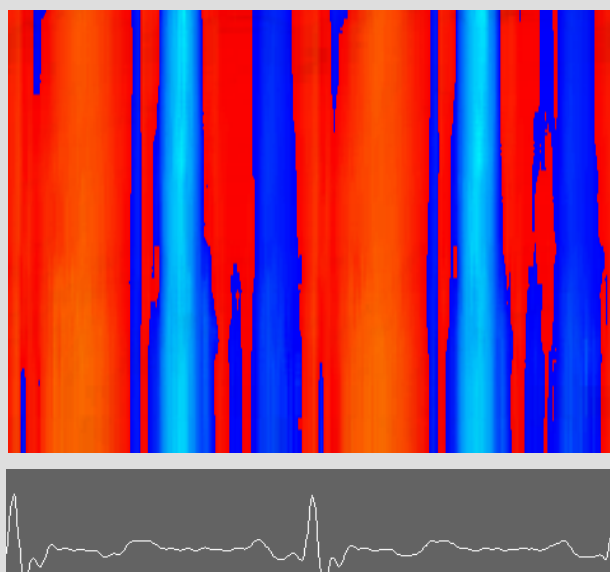


In-vivo abnormal example

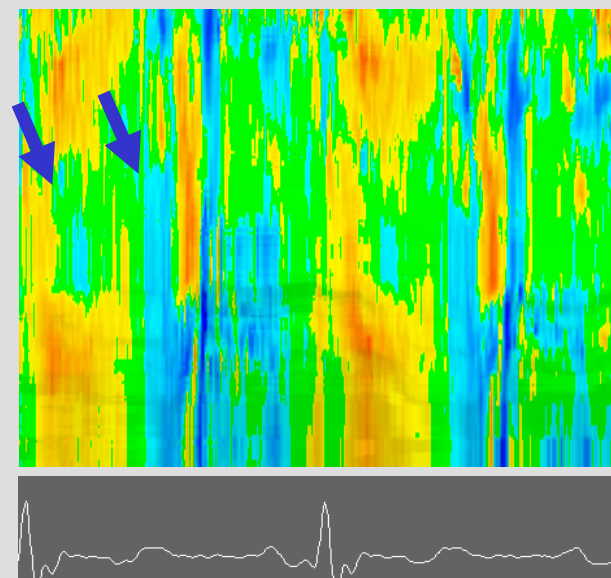


Hypertrophic non-obstructive cardiomyopathy

Velocity

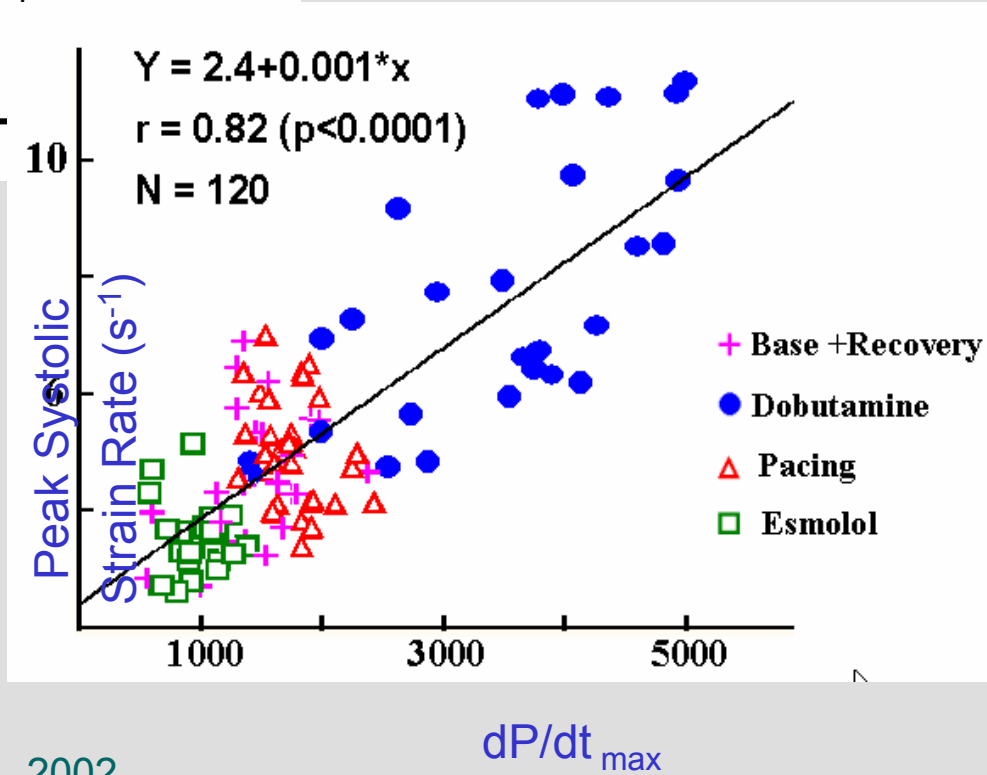
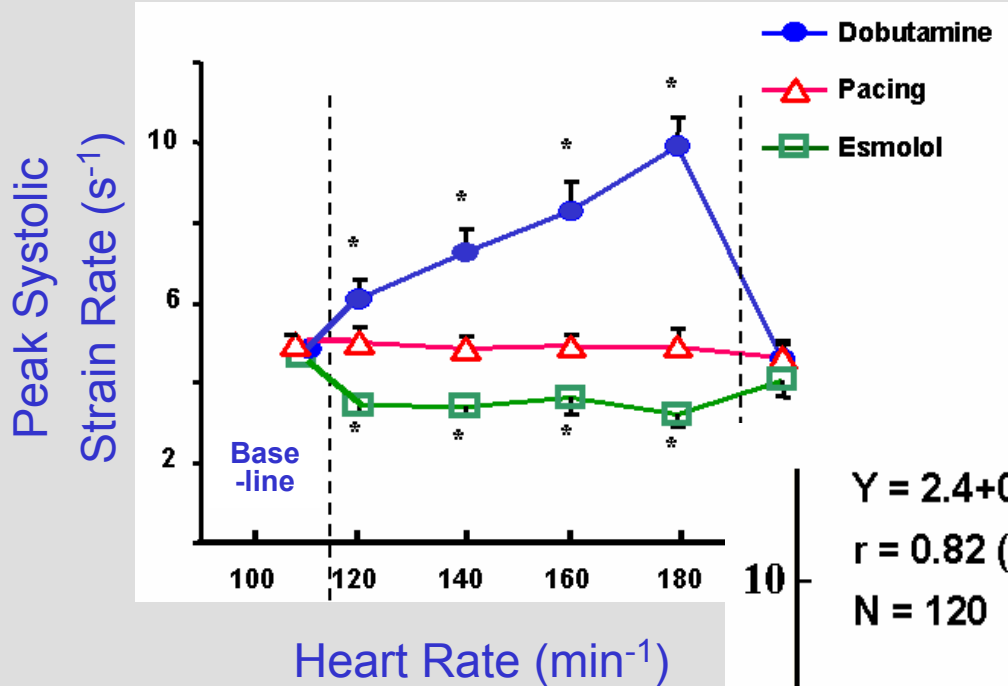


Strain Rate





Assessing myocardial function?



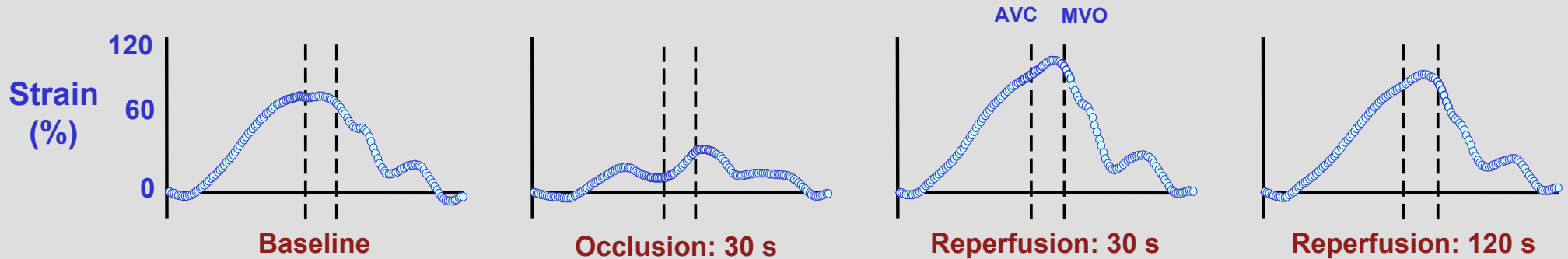
F. Weidemann *et al.*, *J. Am. Soc. Echocardiogr.*, 2002

F. Weidemann *et al.*, *Am. J. Physiology – Heart and Circ.*, 2002

Experimental ischemia

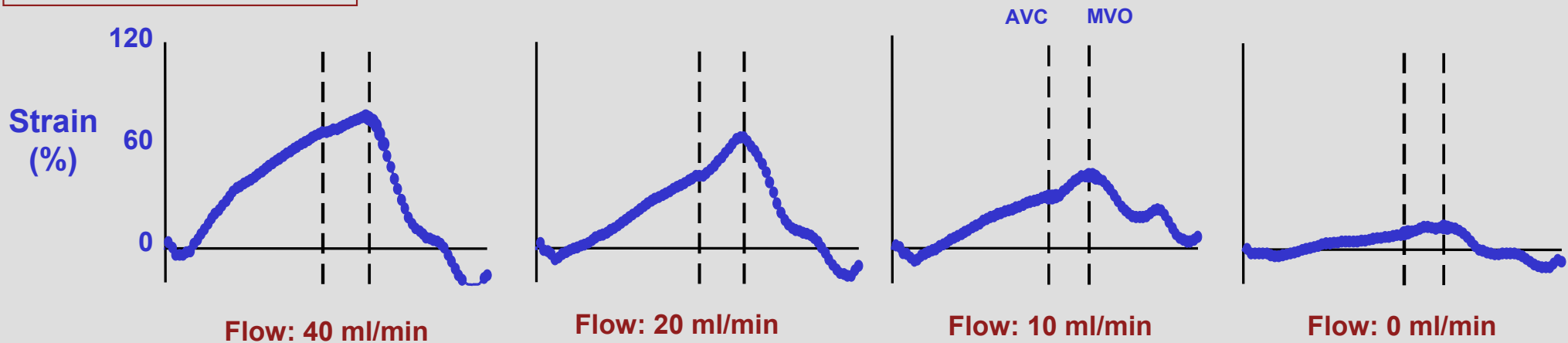
Acute total Ischemia

F. Jamal et al., *J.Am.Soc.Echocardiogr.*, 2001; C. Pislaru et al., *JACC*, 2001
M. Belohlavek et al., *JASE*, 2001, Pislaru et al., *JACC*, 2002



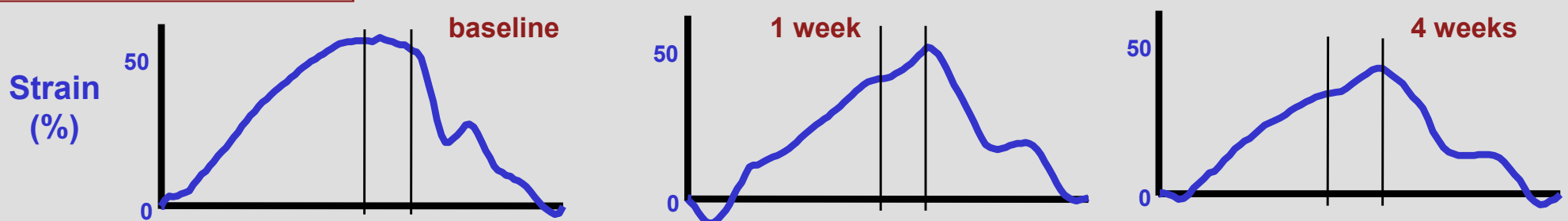
Graded Ischemia

Jamal et al., *J.Am.Soc.Echocardiogr.*, 2001; Weidemann et al., *Clin.Science*, 2003



Chronic Ischemia

F. Weidemann et al., *Circulation*, 2003





Dobutamine Response

	REST			DOBUTAMINE STRESS		
	SR _{SYS}	ϵ_{SYS}	PSI	SR _{SYS}	ϵ_{SYS}	PSI
Normal	5 /s	60 %	2 %	↗	↘	→
Stunning	↓	↓	↑	↗	↗	↘
Acute Ischemia	↓	↓	↑	↘	↘	↗
Nontransmural MI	↓	↓	↑	↘	→	↗
Transmural MI	↓	↓	↑	→	→	→

SRI can differentiate stunned from ischemic myocardium

F.Jamal *et al.*, *Circulation*, 2001

SRI can differentiate transmural from non-transmural MI

F.Weidemann *et al.*, *Circulation*, 2003

SRI + dobutamine stress can differentiate the different ischemic substrates

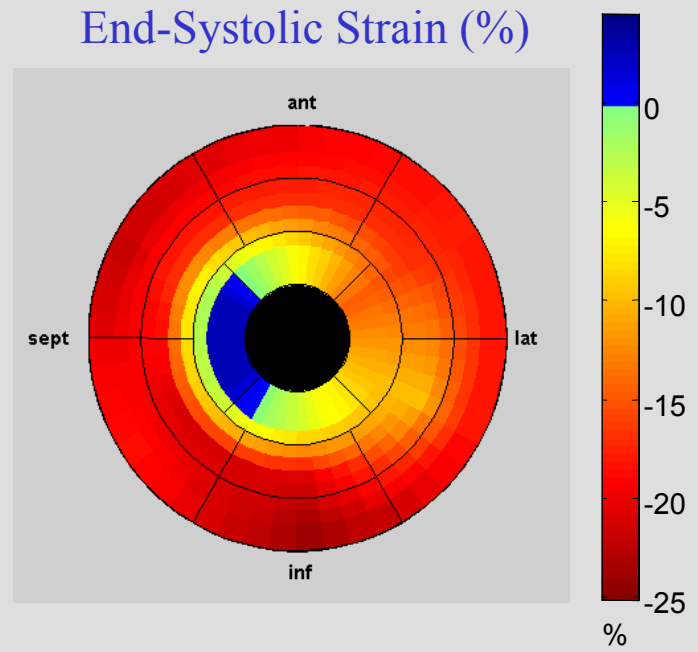
⇒ **Quantitative stress echo!**

M.Kowalski *et al.*, *Eur. J. Echocardiography*, 2003; J.Voigt *et al.*, *Circulation*, 2003

Bulls eye representation of ϵ_{LL} ?

15 patients with acute ischemia (stem-cell study)

- Baseline end-systolic ϵ_{LL} analyzed and represented as a 17 segment bulls eye
- Blinded (*non-medical!*) reader identified area at risk (per segment)
- Compared to angiographic data



⇒ Sensitivity = 81%; Specificity = 79%



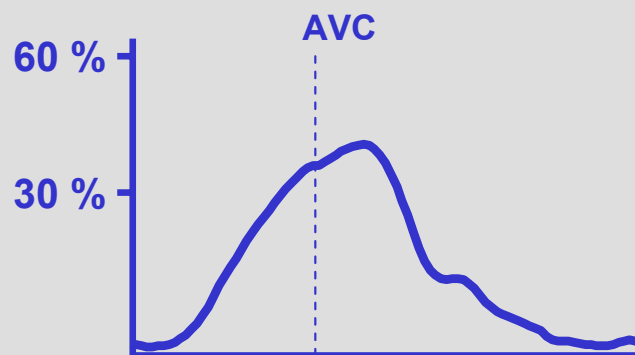
SRI for therapy follow-up



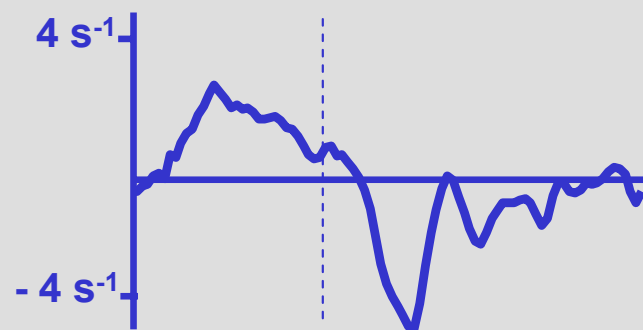
Baseline



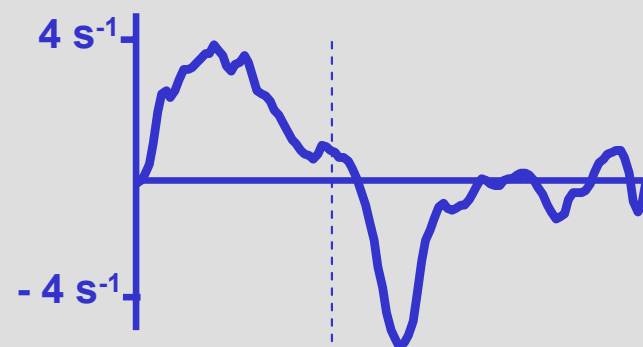
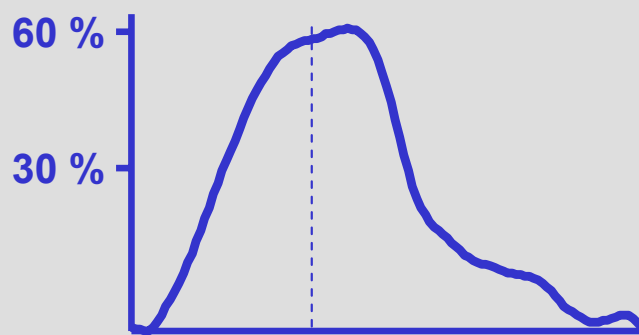
Strain



Strain Rate



8m follow-up



Study in patients with Friedreich's Ataxia using "Idebenone" as therapy
SRI could show early functional improvement

G. Buyse *et al.*, *Neurology*, 2003; G. Di Salvo *et al.*, *AJC*, 2003

F. Weidemann *et al.*, *Circulation*, 2003; F. Weidemann *et al.*, *Eur. Heart Journal*, 2004 (In press) (Fabry disease)



SRI for therapy: CRT



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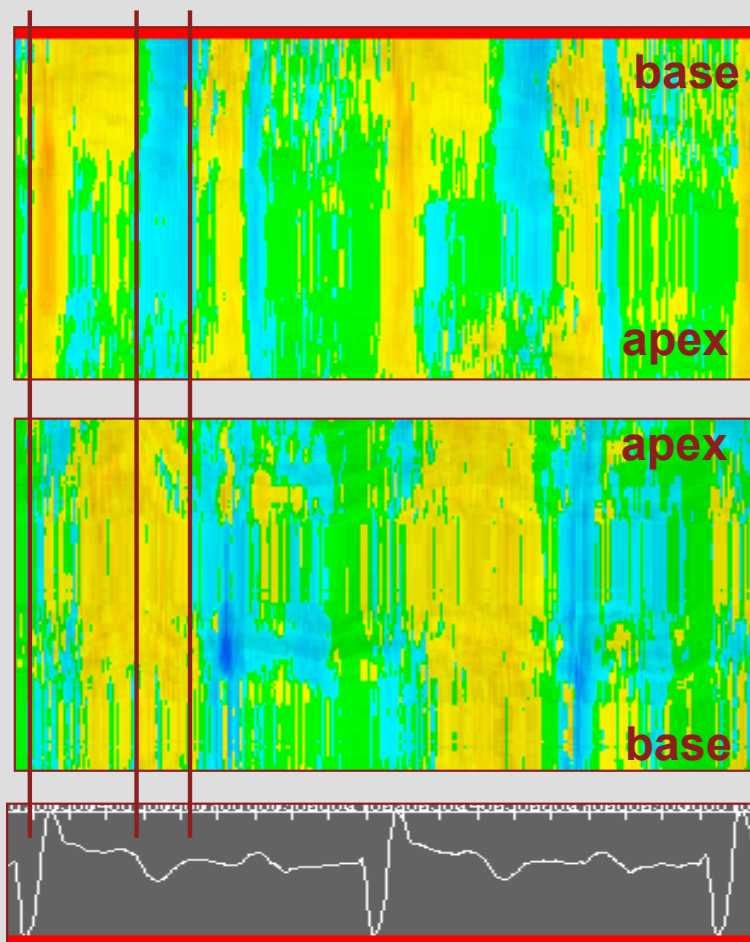
Cardiology
Cardiac Imaging Research

Strain Rate

Strain

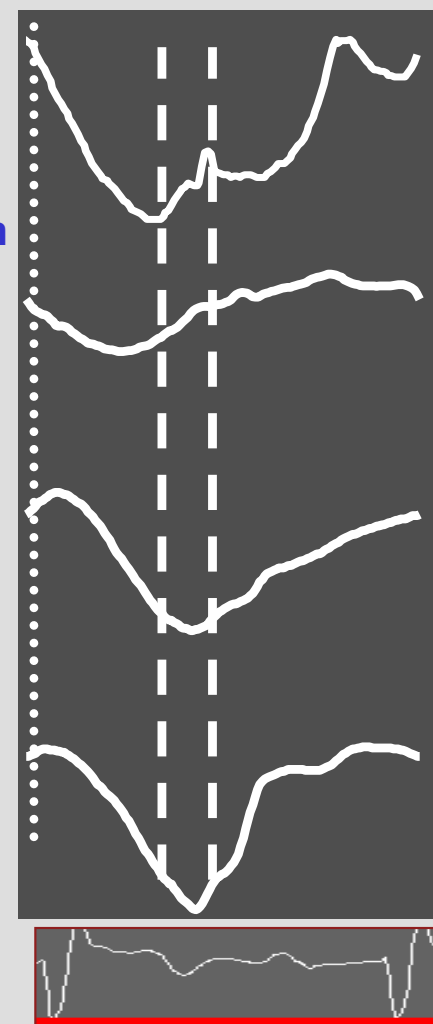


LBBB



Septum

Lateral wall





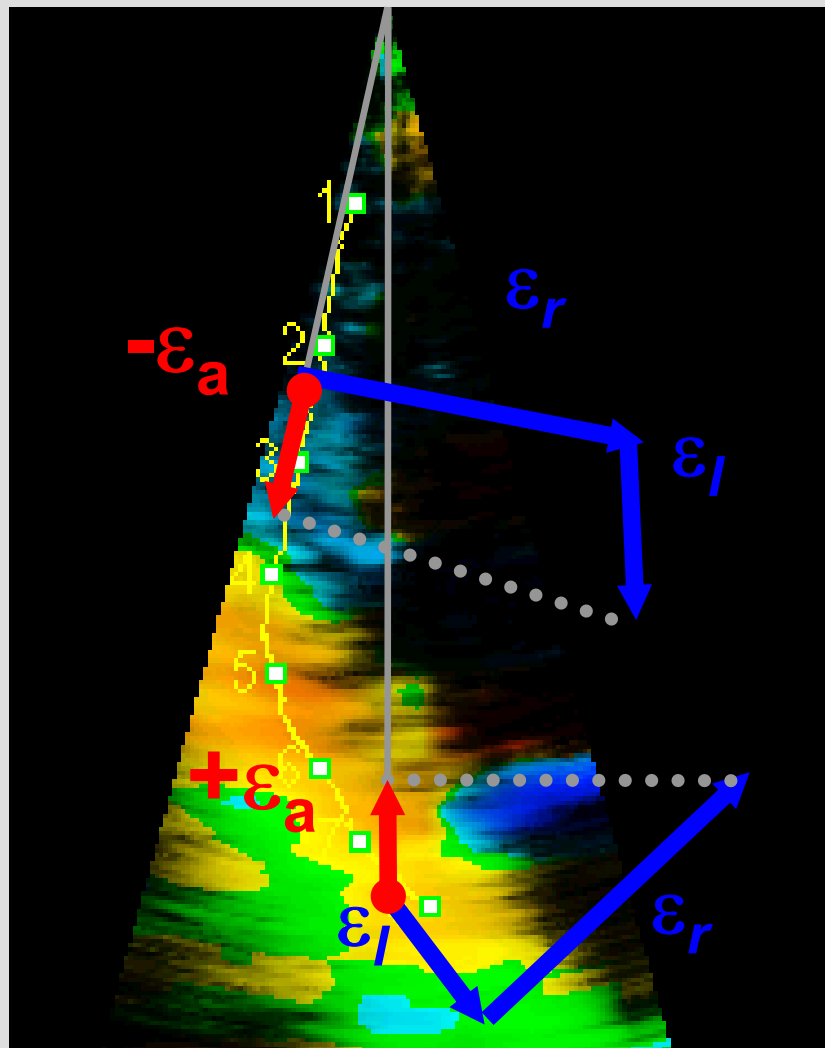
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- Clinical applications of cardiac deformation imaging
- **Pitfalls of the current methodologies**
- Estimating regional active stress development
- Estimating myocardial elasticity



SRI pitfall: angle dependency



Courtesy: Jens-Uwe Voigt, University of Erlangen, Germany

P.L. Castro *et al.*, *Biomed. Sci. Instrum.*, 2000

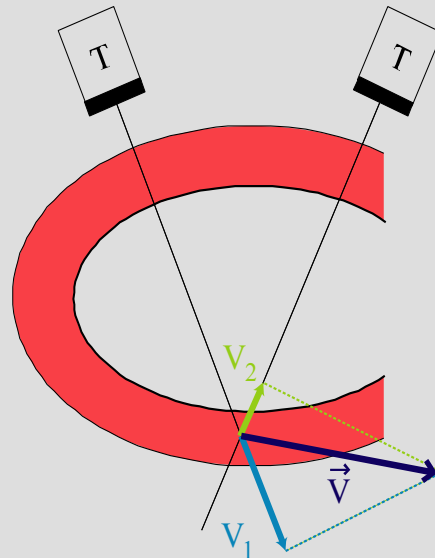
S.I. Rabben *et al.*, *IEEE Ultrasonics Symp.*, 2003 (Effect of angle on SRI values)



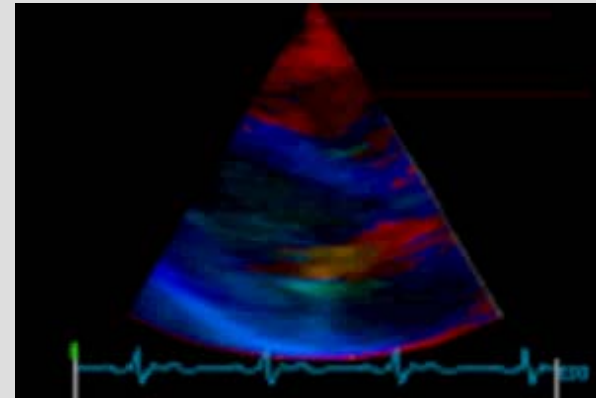
Possible solution



Origin problem: 1D velocities \rightarrow 1D strain rate

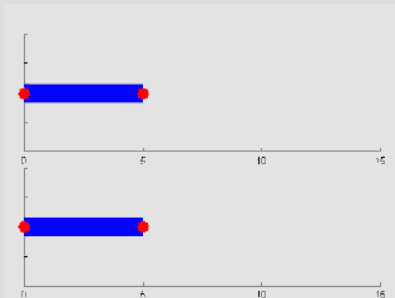


MVI data set



Contains only velocity info
ALONG THE IMAGE LINE

Assume: 2D velocities \rightarrow 2D strain (rate) tensor



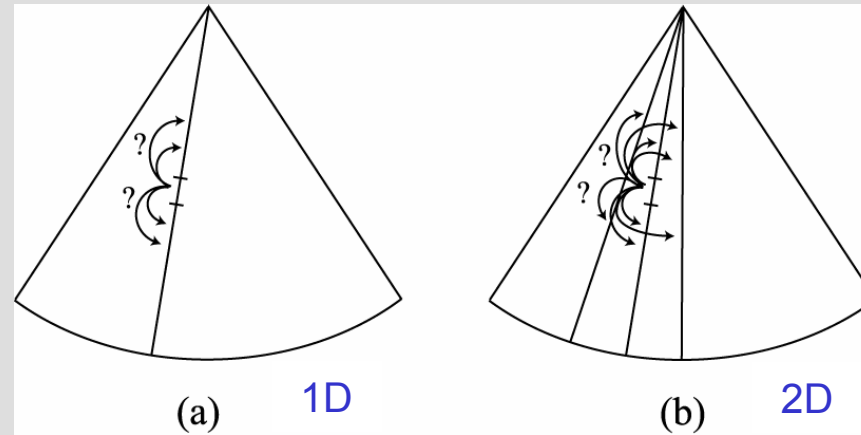
$$\begin{pmatrix} \frac{\partial v_x}{\partial x} & \frac{\partial v_x}{\partial y} \\ \frac{\partial v_y}{\partial x} & \frac{\partial v_y}{\partial y} \end{pmatrix} = \begin{pmatrix} \dot{\epsilon}_{xx} & \dot{\epsilon}_{xy} \\ \dot{\epsilon}_{yx} & \dot{\epsilon}_{yy} \end{pmatrix} = \dot{\epsilon}$$



RF based 2D motion/velocity



Generalized RF time-shift estimator (displacement estimator)



1D RF kernel

2D RF kernel

Optimal distance measure between RF
patterns for cardiac SRI?

Viola *et al.* IEEE-UFFC, 2003
S. Langeland *et al.*, UMB, 2003

K. Kaluzynski *et al.* IEEE-UFFC, 2001
X. Chen *et al.* IEEE Symposium, 2002
X. Chen *et al.* IEEE Symposium, 2003

Cross Correlation (XC)

Normalized Cross Correlation (NXC)

Sum of Absolute Differences (SAD)

Sum of Squared Differences (SSD)



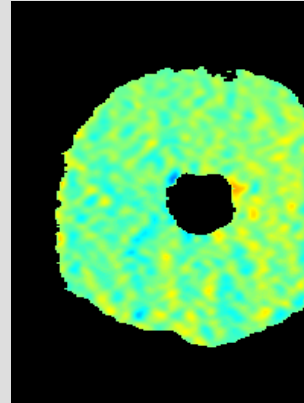
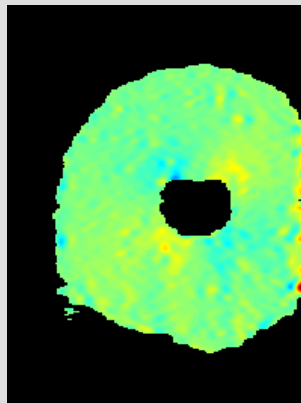
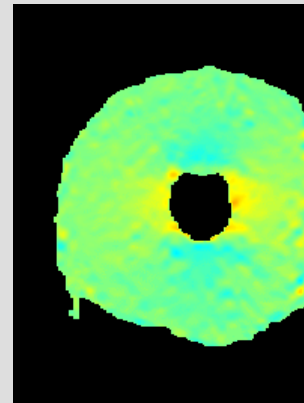
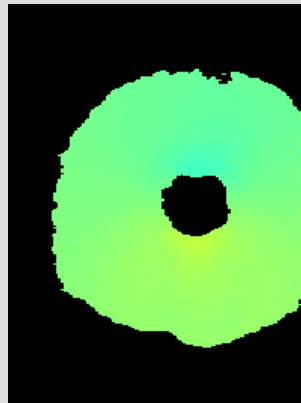
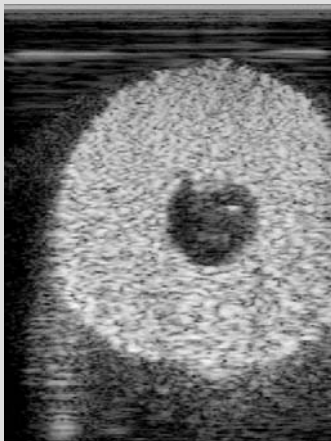
In vitro feasibility



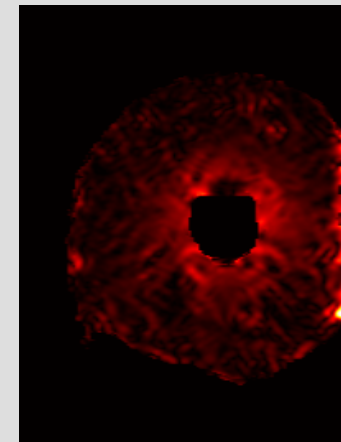
$$\begin{pmatrix} \frac{V_x}{x} & \frac{V_x}{y} \\ \frac{V_y}{y} & \frac{V_y}{x} \end{pmatrix} = \begin{pmatrix} \dot{\epsilon}_{xx} & \dot{\epsilon}_{xy} \\ \dot{\epsilon}_{yx} & \dot{\epsilon}_{yy} \end{pmatrix} = \dot{\epsilon}$$

2D kernel

B-mode image



ϵ intensity

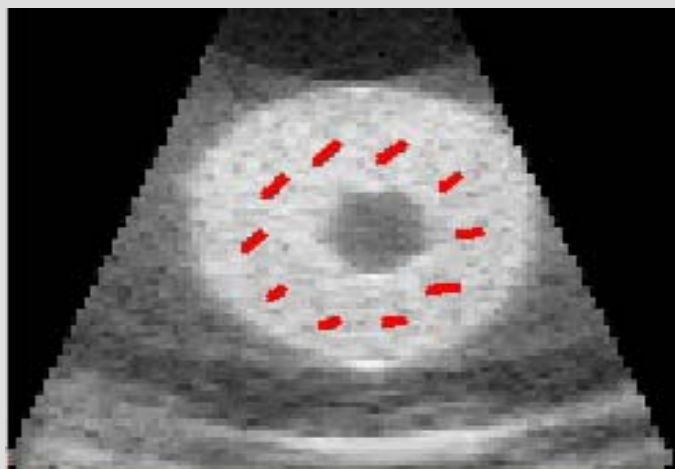




In vitro validation

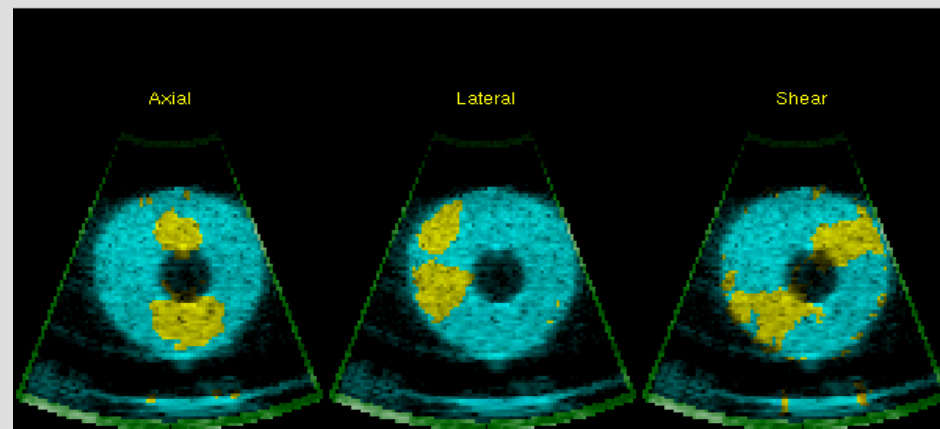
1D kernel

2D velocity vector

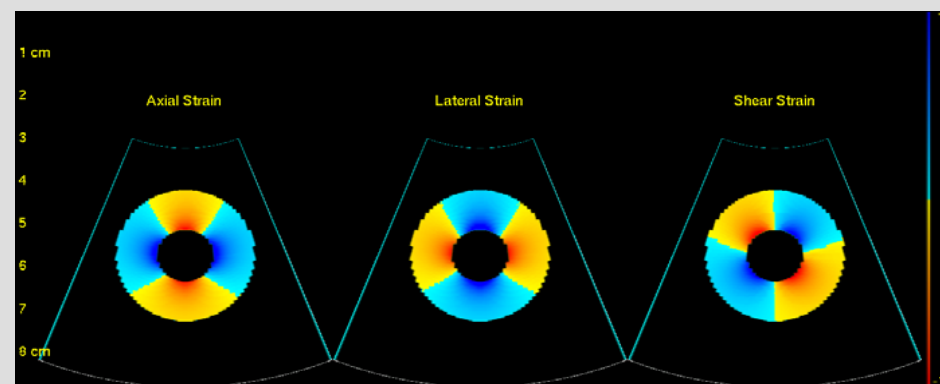


2D strain

Experiment



Theory



Axial

Lateral

Shear

Good correlation of both axial and lateral strain values
against micro-crystal data ($r \approx .90$)

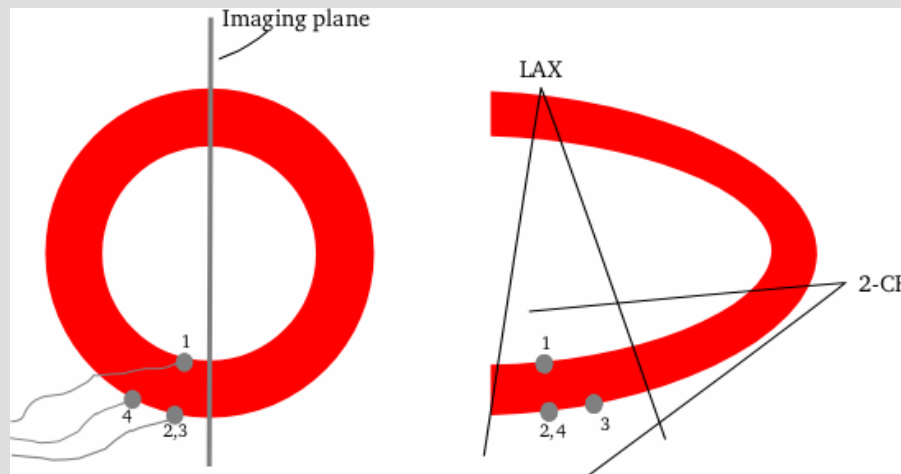
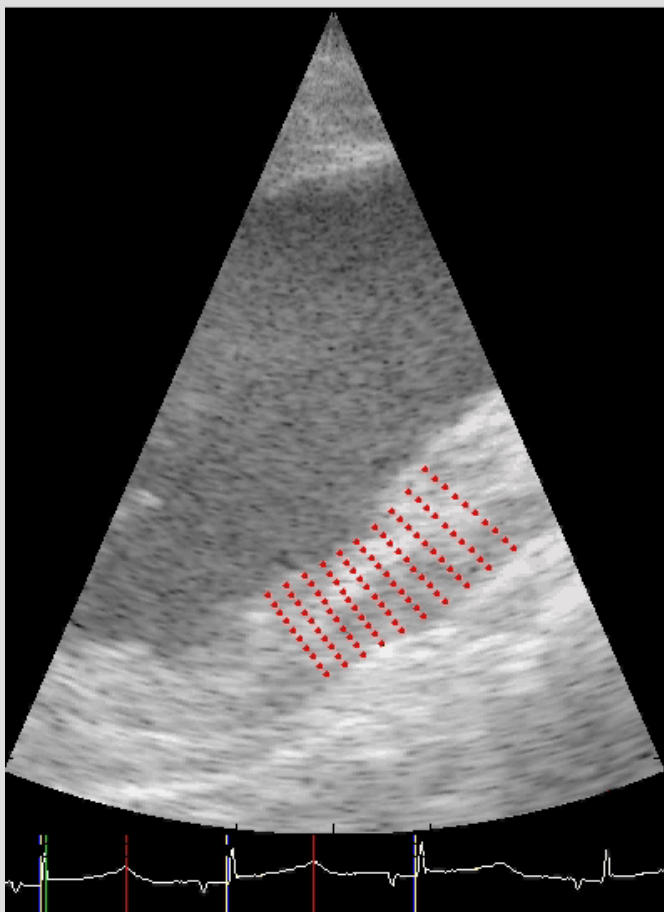


Results in-vivo validation

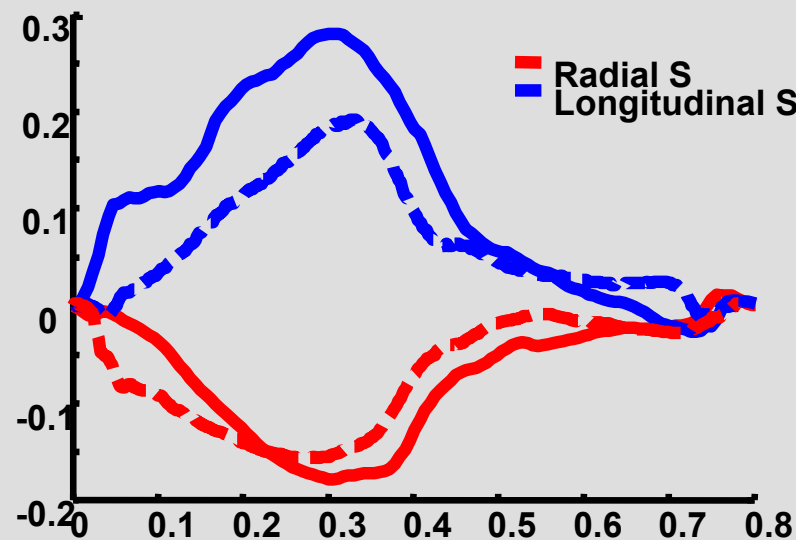


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2D strain in-vivo



Solid = ultrasound
Dashed = micro-crystals



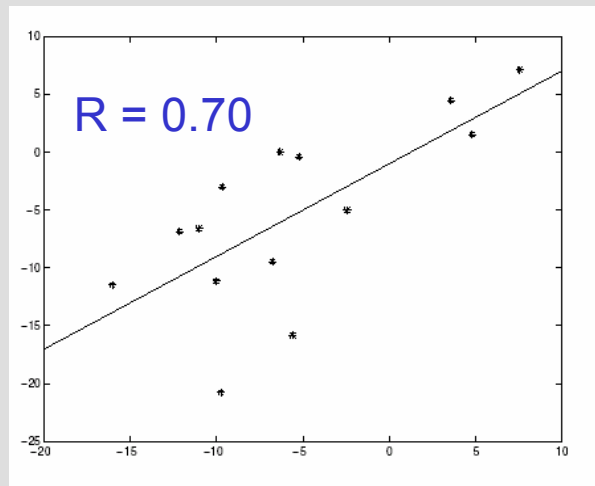
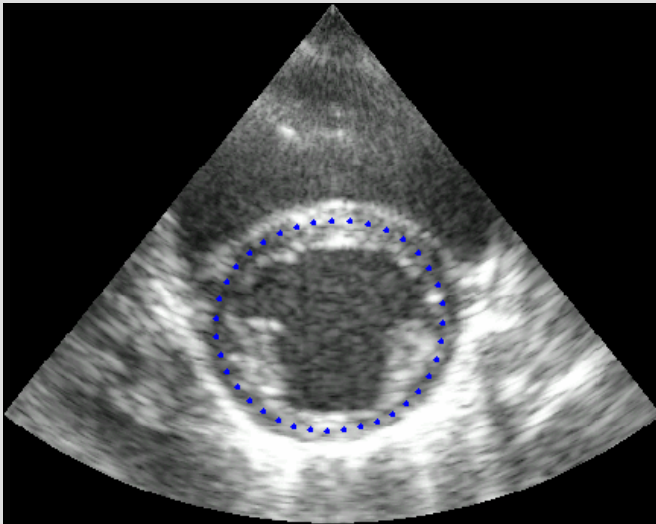


Envelope based 2D motion/velocity

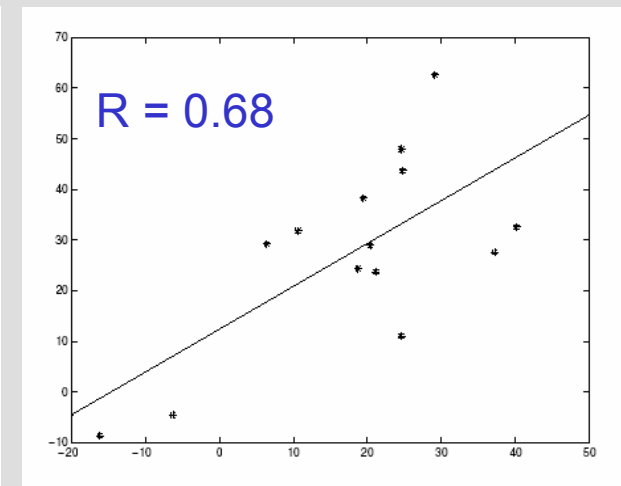


Optical Flow

C. Jansen *et al.* Eur. J. Echocardiogr., 2002 (abstract)
Behar *et al.* Ultrasonics, 2004



Longitudinal



Radial

Initial clinical results: M. Leitman *et al.*, *J. Am. Soc. Echocardiogr.* 17(10):1021-1030, 2004



Contents



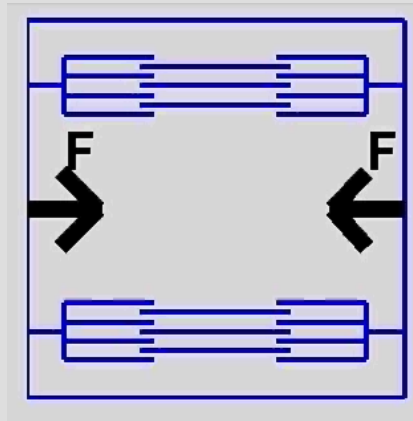
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- Estimating myocardial elasticity



Regional myocardial function



Active elastic body



Active contractile stress

$C(t)$

+

$\sigma_{load}(t)$

=

E

x

$\epsilon(t)$

Regional myocardial function
(contractility)

Wall stress
(total force on a segment)
cavity pressure,
geometry,
neighbouring segment interaction

segmental elasticity

resulting deformation



If the assumption is made that external stresses (loading) can be neglected, local strain directly relates to local contractile stress

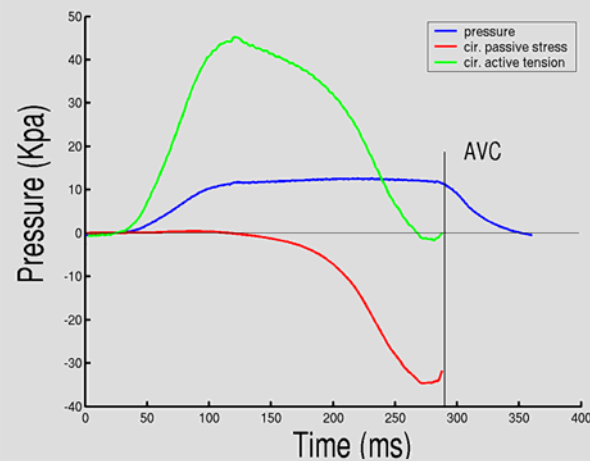
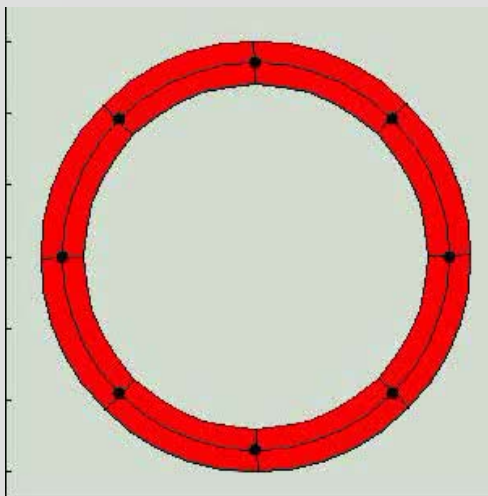
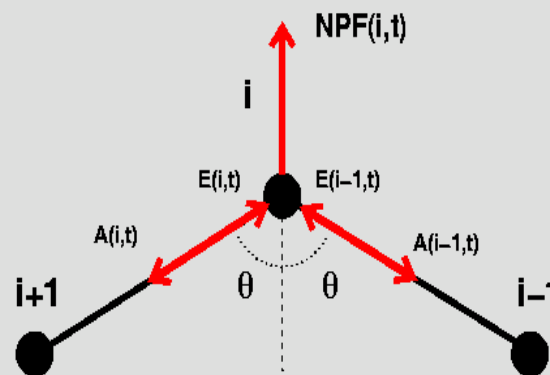
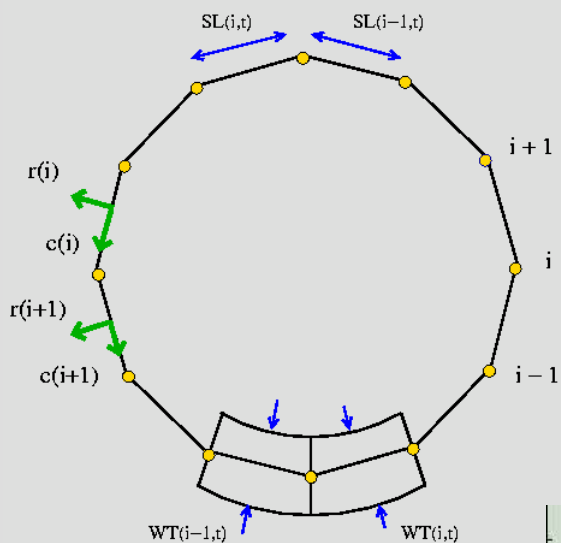


Towards active force development using SRI + mechanical models



The left ventricle was modeled as a 2D circular array of 400 nodes

In each node → balance of forces



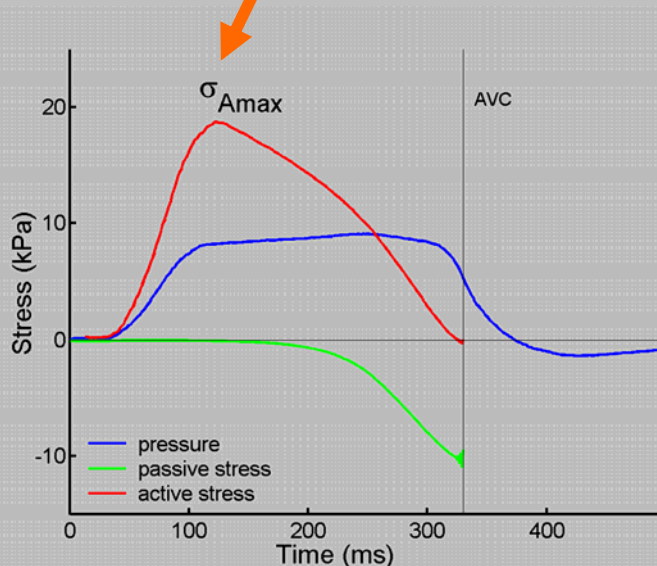
M. McLaughlin *et al.*,
IEEE EMBS Symposium, 2004



Results I

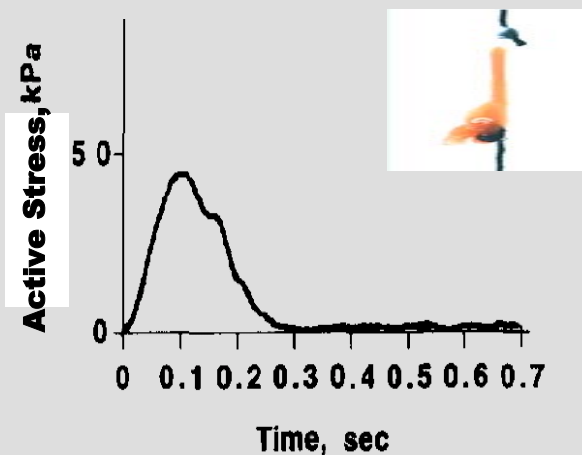


Normal Baseline Values



Parameter	Mean \pm SE
σ_{Amax}	20.5 \pm 1.3 kPa
$T_{\sigma_{Amax}}$	106 \pm 5.7 ms
AVC	317 \pm 9.8 ms
C_1	17 \pm 23 kPa
C_2	281 \pm 114

Active stress profiles measured
in isolated muscle strips



1. J.M. Guccione, et al. Measurements of active myocardial tension under a wide range of loading conditions. J. Biomechanics, 30(2): 189-192, 1997



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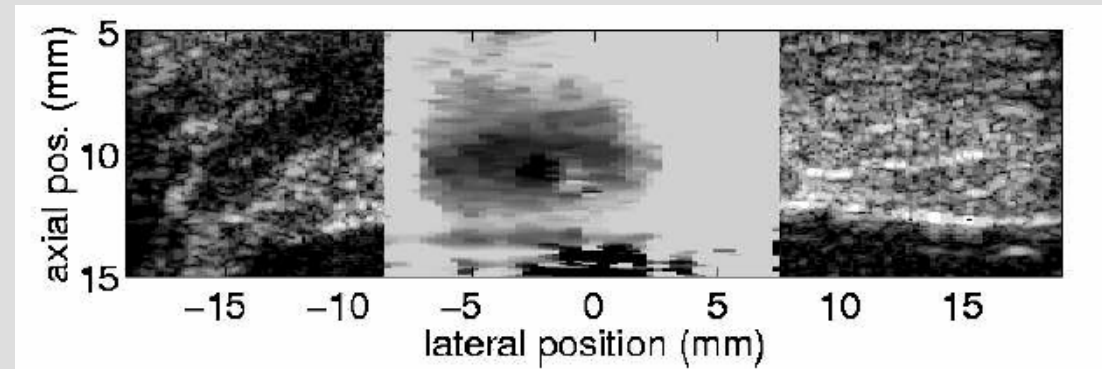
Non-invasive assessment of cardiac elasticity



KULeuven
UZ Gasthuisberg

Relative elastic properties

ARFI



B. Fahey *et al.*
IEEE Ultrasonics Symposium, 2003

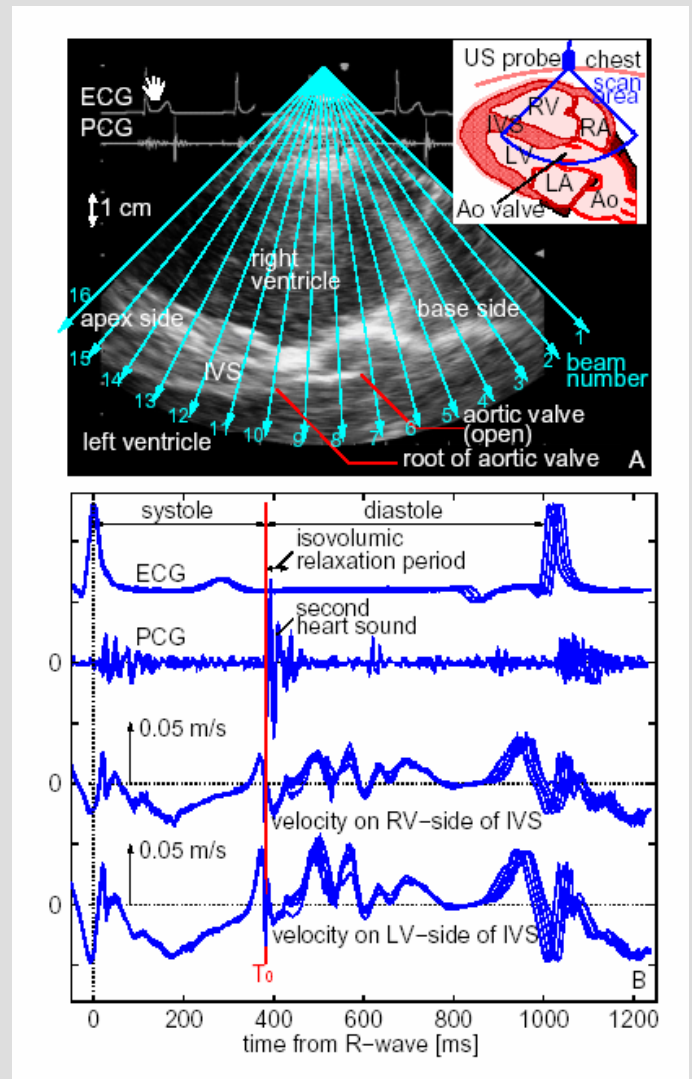
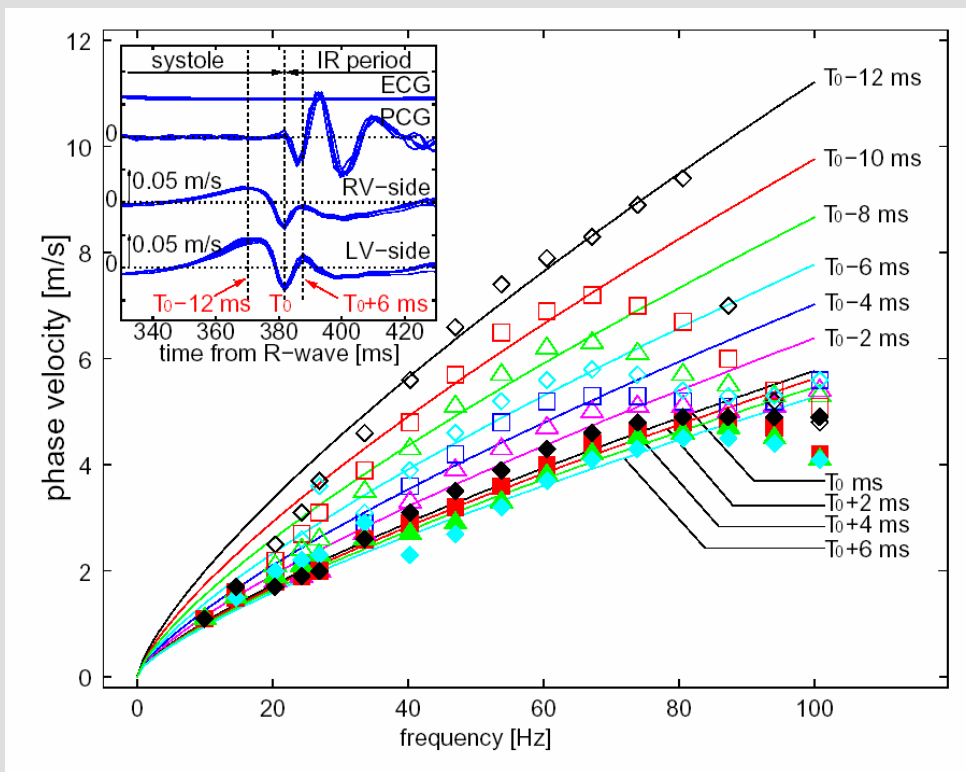


Non-invasive assessment of cardiac elasticity



Elastic properties

Using Lamb wave propagation?



H. Kanai *et al.*
IEEE Ultrasonics Symposium, 2003



Conclusions I



- Strain and strain rate imaging are new echocardiographic tools for the assessment of regional myocardial function
- Different approaches towards cardiac deformation estimation have been proposed both based on the envelope and RF signals
- Important clinical applications will be:
 - Quantitative stress echocardiography
 - Therapy guidance and follow-up
- Further research is required to optimize the use of the current technique and to further test its clinical applicability



Conclusions II



- Multi-dimensional strain (rate) estimation is becoming possible (angle dependency will thus be solved)
- Time-consuming analysis is being reduced through automated tracking of regions of interest
- Linking ultrasound deformation measurements to mathematical models of cardiac contraction can:
 1. Make the strain (rate) estimate more robust
 2. Estimate active stress development!
- New methods have been proposed for estimating myocardial relative and absolute elastic properties

www.strainrateimaging.org



- Teaching material

- Literature refs

- RF data sets

- SRI software

→ i.e. “Speqle”

- Latest

developments

Stay tuned!

