

# La fluidodinamica valutata con mezzi di contrasto ecografici

ECOCARDIOGRAFIA 2015 – XVII CONGRESSO NAZIONALE  
SIEC Napoli, 16-18 aprile 2015



Prof Luciano Agati



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# **NEWS on Contrast Ultrasound 2015**

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- **World Imaging Agents Market to Exceed US\$14 Billion by 2015, According to New Report by Global Industry Analysts, Inc.**
- This is mainly due to the wide availability of existing equipment in key markets including Europe and the US and increasing focus on diagnostic examinations that facilitate early identification of the disease and its treatment.



# Contrast Ultrasound in Cardiology

- In the USA in 2010, there was 28 millions of echocardiographic studies and 1 million were with contrast (3.5%)
- In EU it is more difficult to get precise data. Estimation is 250.000 contrast procedures out of 20 millions (1.5%) echocardiographic studies.
- In the UK we have most precise data:
  - Stress echo procedures: 40.876
  - Contrast proc: 28.279 (69%)



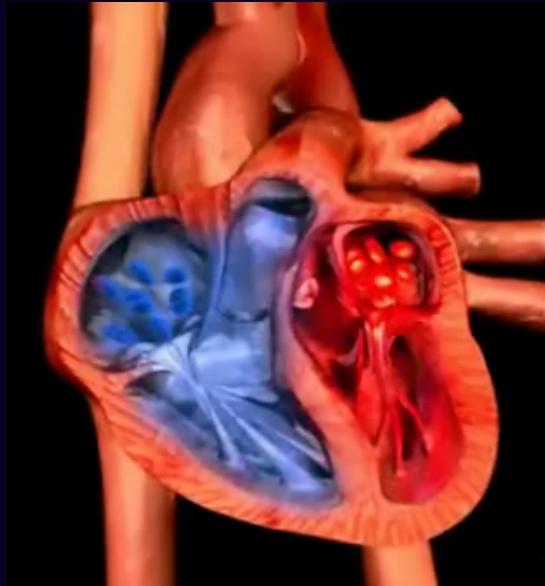
# ***Contrast echocardiography.***

## ***Main problems***

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- No reimbursement of contrast echo in several EU countries
- No approval for myocardial perfusion
- Need for “top level” echo instruments and dedicated software
- Need for learning curve
- Operator/ Acoustic window dependent





# *I.V. flow assessment*

Luciano Agati, MD

Gianni Tonti, MD

Gianni Pedrizzetti, PhD

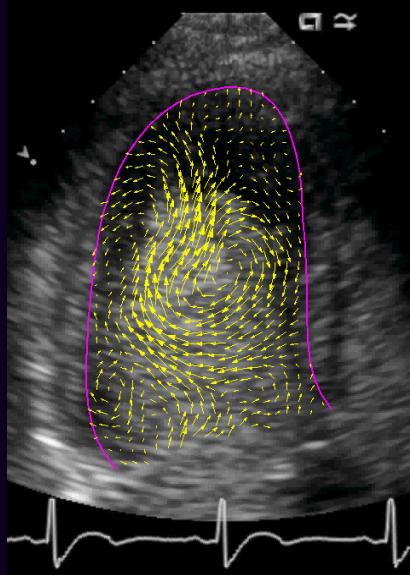
- **Dept of Cardiology, Sapienza University of Rome, Italy**
- **Dept of Engineering. University of Trieste, Italy**
- **AMID, Italy**



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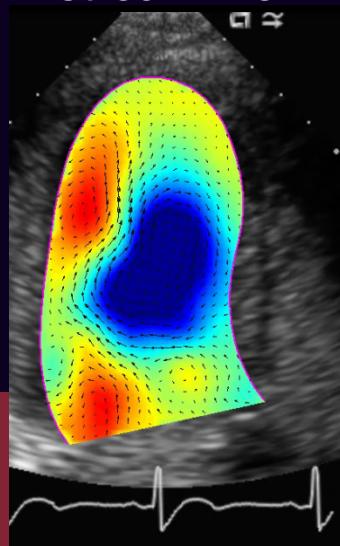
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# Vorticity & Energy Parameters

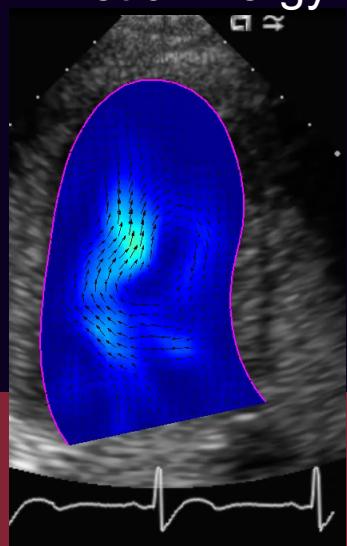
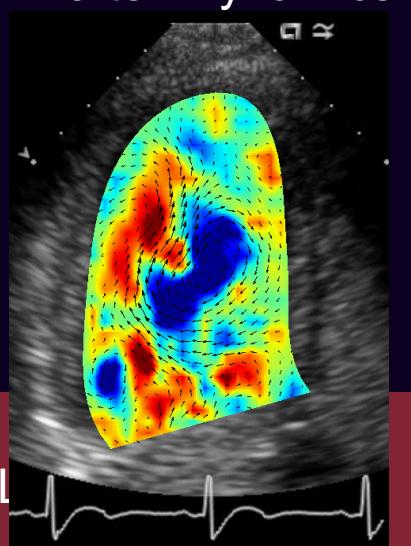
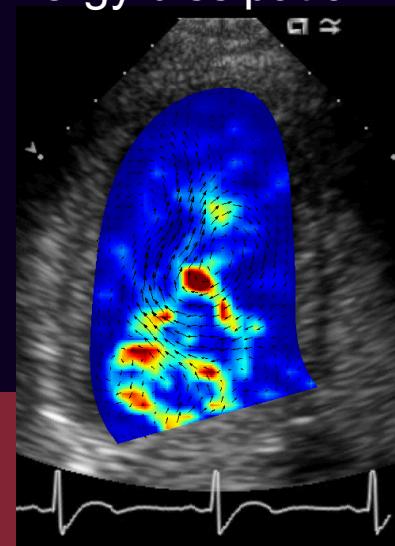


Parameter	Value
Vortex Area	0.360
Vortex Intensity	-0.392
Vortex Depth	0.473
Vortex Length	0.726
Energy Dissipation	0.633
Vorticity Fluctuation	0.838
Kinetic Energy Fluctuation	1.735
Shear Stress Fluctuation	0.185
Dominant Force Strength	12.397
CCW from Apex	-15.454

Stream function  
streamline



Energy dissipation

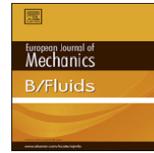


# 3D Echo flow



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*In vivo analysis of intraventricular fluid dynamics in healthy hearts*S. Cimino <sup>a</sup>, G. Pedrizzetti <sup>b</sup>, G. Tonti <sup>c</sup>, E. Canali <sup>a</sup>, V. Petronilli <sup>a</sup>, L. De Luca <sup>a</sup>, C. Iacoboni <sup>a</sup>, L. Agati <sup>a,\*</sup><sup>a</sup> Department of Cardiovascular Science, "Sapienza" University, Roma, Italy<sup>b</sup> DICAR, University of Trieste, Italy<sup>c</sup> SS. Annunziata Hospital, Sulmona, AQ, Italy

**Table 2**  
**Vortex and energetic dimensionless parameters.**

Parameter	Mean $\pm$ st. dev	CI
-		
Vortex area	$0.29 \pm 0.1$	0.12–0.46
Vortex circulation	$-0.33 \pm 0.28$	-0.56–0.17
Vortex length	$0.62 \pm 0.12$	0.4–0.75
Vortex depth	$0.36 \pm 0.12$	0.18–0.6
Energy dissipation	$0.37 \pm 0.13$	0.2–0.58
Vorticity fluctuation	$0.74 \pm 0.12$	0.61–0.93
Kinetic energy fluctuation	$1.38 \pm 0.3$	0.9–1.7
Shear stress fluctuation	$0.17 \pm 0.3$	-0.3–0.64

European Heart Journal - Cardiovascular Imaging Advance Access published June 6, 2014



European Heart Journal – Cardiovascular Imaging  
doi:10.1093/eihci/jeu106

# **Quantitative analysis of intraventricular blood flow dynamics by echocardiographic particle image velocimetry in patients with acute myocardial infarction at different stages of left ventricular dysfunction**

**L. Agati<sup>1\*</sup>, S. Cimino<sup>1</sup>, G. Tonti<sup>2</sup>, F. Cicogna<sup>1</sup>, V. Petronilli<sup>1</sup>, L. De Luca<sup>1</sup>,  
C. Iacoboni<sup>1</sup>, and G. Pedrizzetti<sup>3</sup>**



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# Methods

## Echo - PIV

### Characterization and Quantification of Vortex Flow in the Human Left Ventricle by Contrast Echocardiography Using Vector Particle Image Velocimetry

Geu-Ru Hong, MD, PhD,\*† Gianni Pedrizzetti, PhD,‡ Giovanni Toni, MD,§ Peng Li, MD, PhD,\* Zhao Wei, MD, PhD,\* Jin Kyung Kim, MD, PhD,|| Abinav Baweja,|| Shizhen Liu, MD, PhD,\* Namsik Chung, MD, PhD,¶ Helene Houle, RDCS,# Jagat Narula, MD, PhD, FACC,|| Mani A. Vannan, MBBS, FACC\*

JACC Card Imag 2008

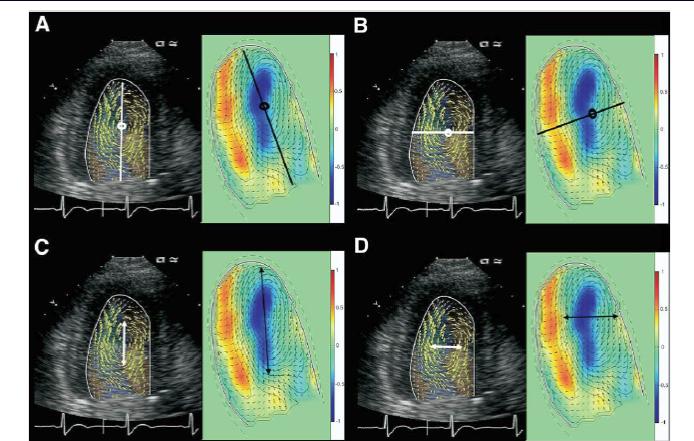
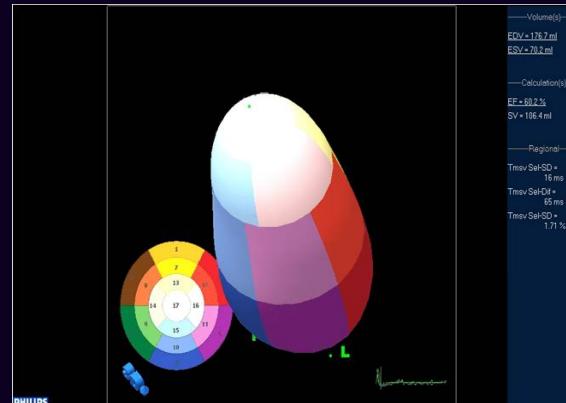
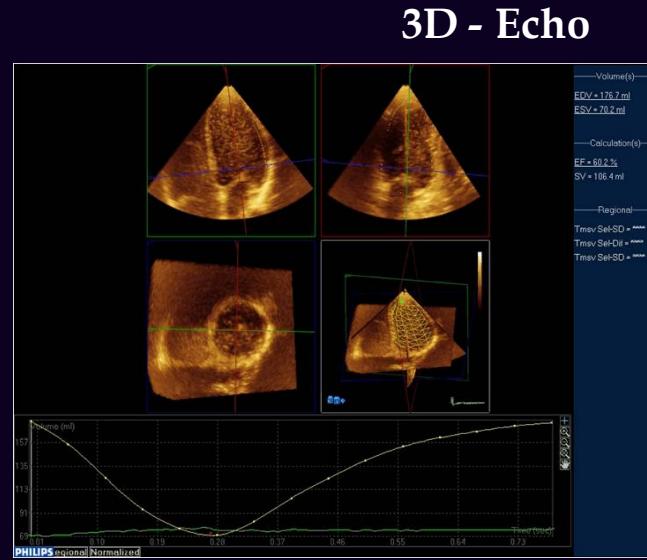


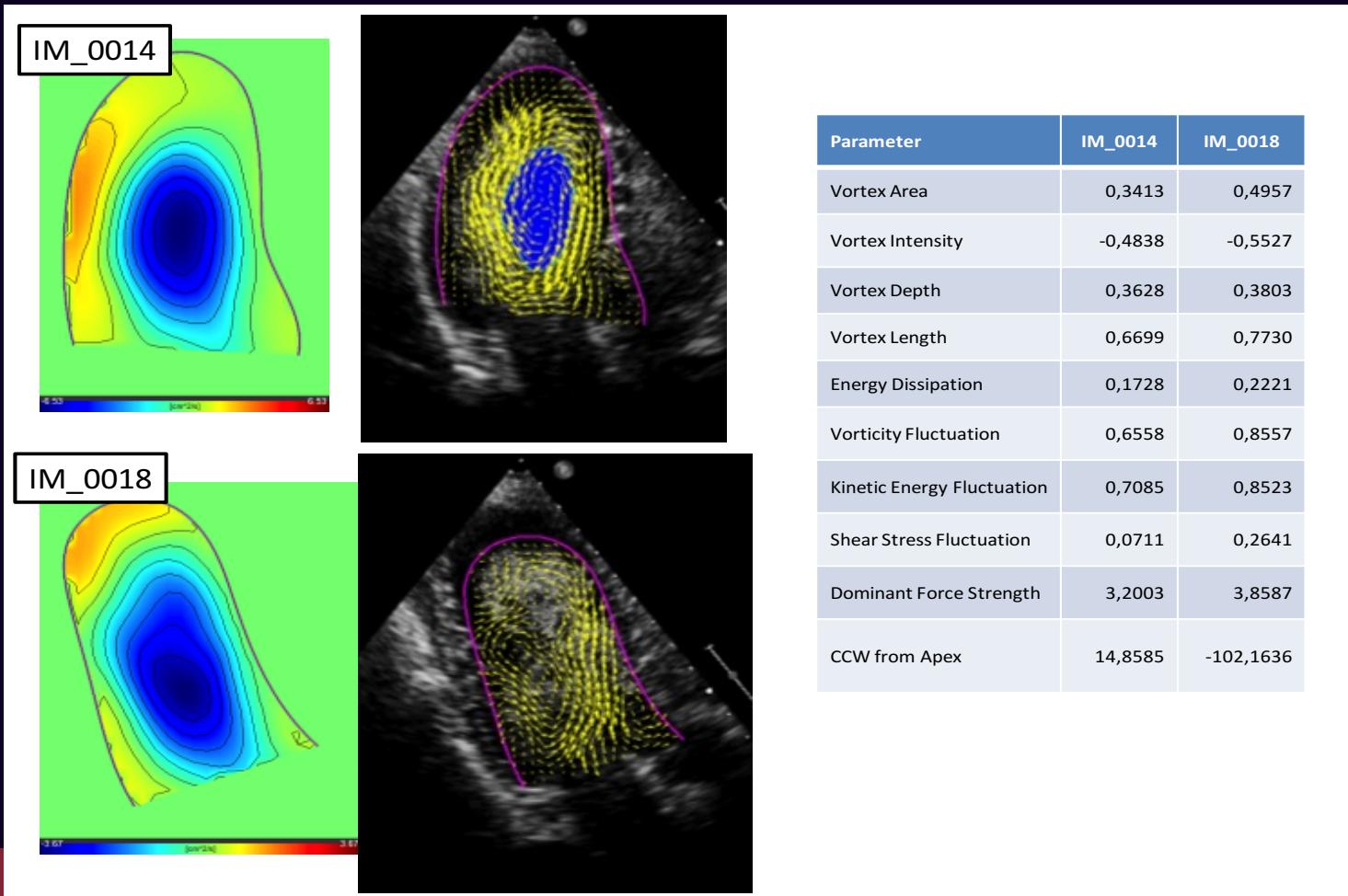
Figure 2. Description of How to Measure Quantitative Average Vortex Parameters That Represent Vortex Location and Shape

Vortex depth represents vertical position of center of vortex relative to left ventricular long axis (A, white line), and vortex transverse position represents transverse position relative to posteroseptal axis (B, white line). Vortex length was measured by longitudinal length of vortex relative to left ventricular length (C, white arrow), and vortex width was measured by horizontal length of vortex relative to left ventricular length (D, white arrow). A vortex sphericity index was calculated by vortex length and vortex width.



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# *Non invasive assessment of LV flow dynamics and mechanical function; a new approach for the global evaluation of LV efficiency*



# Study population n = 41

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- Group I, LVEF >50%, (n= 14)
- Group II, LVEF 50-30%, (n=10)
- Group III, LVEF <30%, (n=10)
- Healthy controls (n=10)

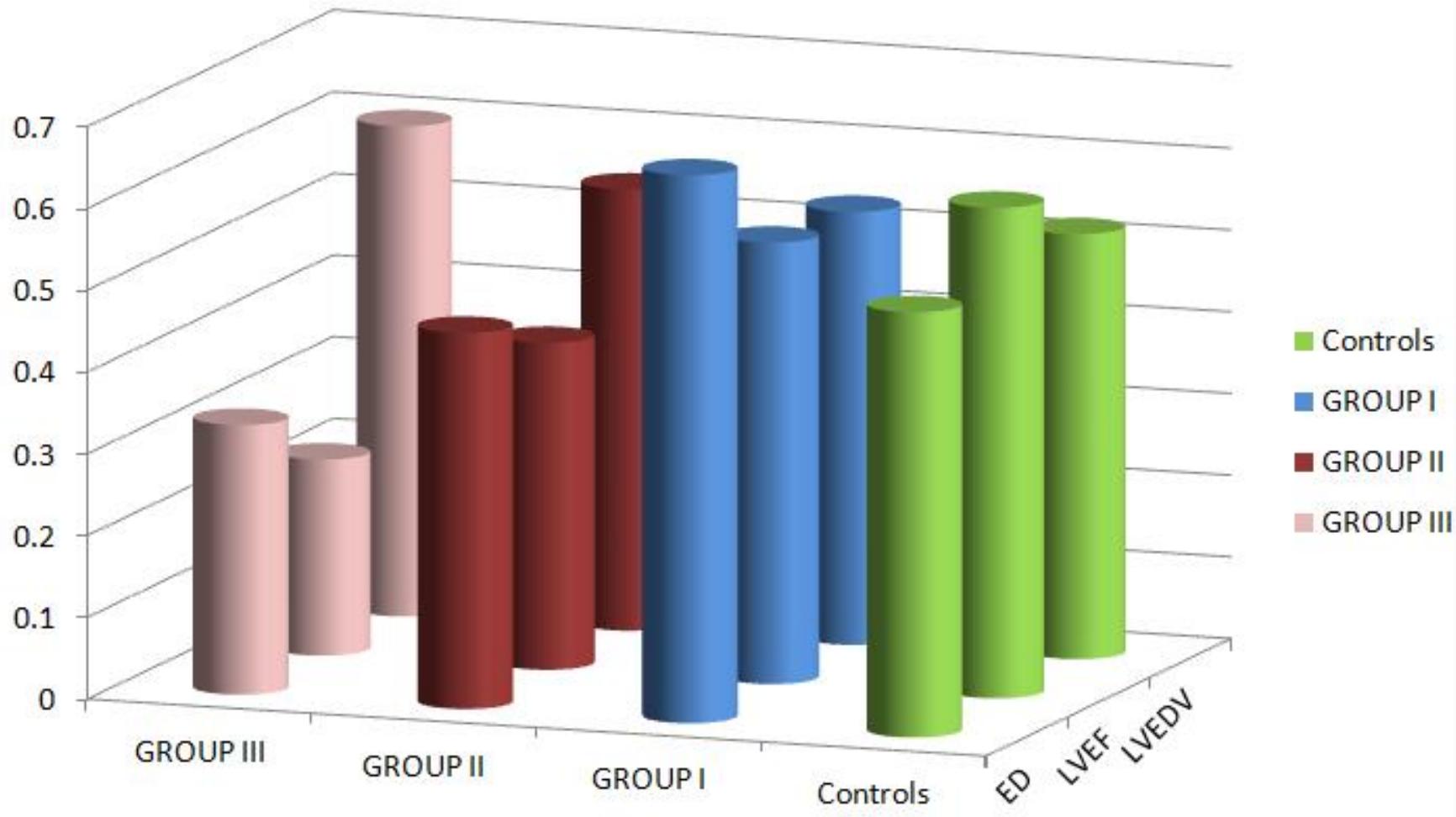


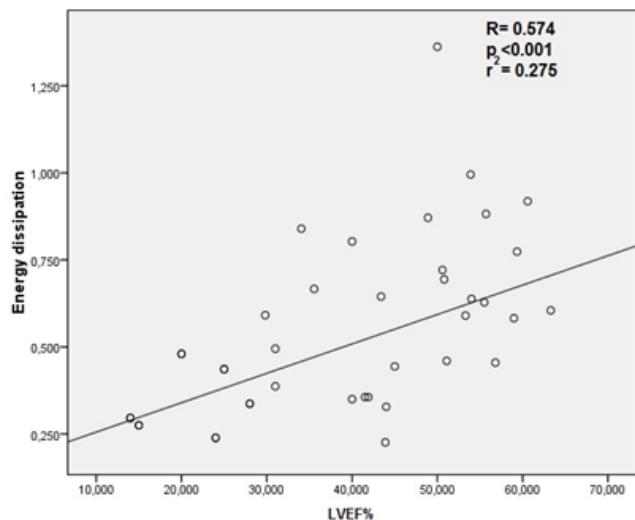
# *Echo Data*

	<b>Group I (n= 14)</b>	<b>Group II (n=10)</b>	<b>Group III (n= 10)</b>	<b>P value</b>
<b>LVEF, %</b>	<b>54.6±3.2</b>	<b>40.72±5.7</b>	<b>24.3±5.1</b>	<b>&lt;0.001</b>
<b>GLS</b>	<b>-16±5.3</b>	<b>-13±6.7</b>	<b>-8±4.2</b>	<b>&lt;0.001</b>
<b>LVEDV/i, ml/m<sup>2</sup></b>	<b>53±15</b>	<b>54±26</b>	<b>60±12</b>	<b>0.045</b>
<b>LVESV/i, ml/m<sup>2</sup></b>	<b>24±7.7</b>	<b>30±18</b>	<b>46±9.2</b>	<b>0.032</b>
<b>GWMSI</b>	<b>1.7±0.6</b>	<b>2.2±0.9</b>	<b>3.3±1.8</b>	<b>0.008</b>

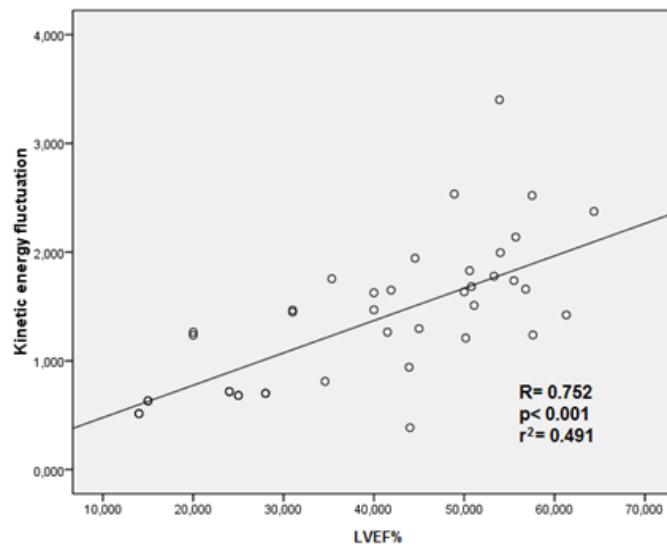


	Controls	Group I	Group II	Group III	P
	(n=10)	(n= 14)	(n=10)	(n= 10)	value
Vortex Area	0.34±0.08	0.35±0.07	0.35±0.09	0.33±0.09	0.42
Vortex Length	0.77±.5	0.7±0.1	0.68±0.14	0.64±0.09	0.38
Vortex Depth	0.4±0.1	0.37±0.1	0.45±0.11	0.46±0.054	0.25
Vortex Intensity	-0.44±0.1	-0.43±0.2	-0.43±0.23	-0.37±0.26	0.58
Energy Dissipation*	0.52±0.22	0.67±0.29	0.46±0.21	0.33±0.09	0.008
Vortex Fluctuation**	0.80±0.03	0.83±0.06	0.75±0.16	0.60±0.07	0.024
Kinetic Energy Fluctuation***	1.55±0.23	1.83±0.54	1.40±0.54	0.74±0.22	0.004

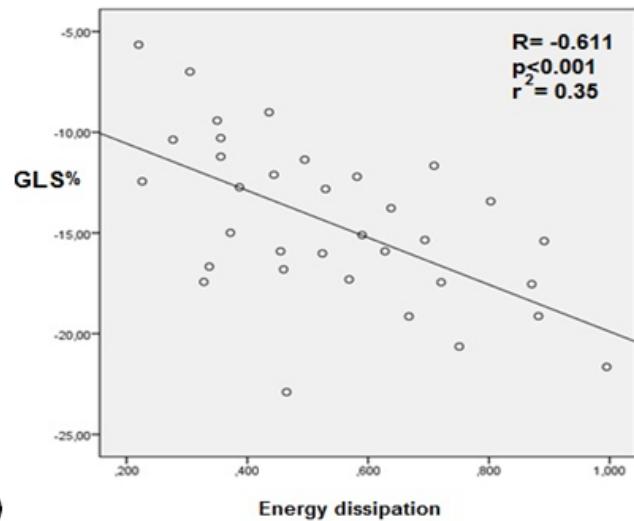




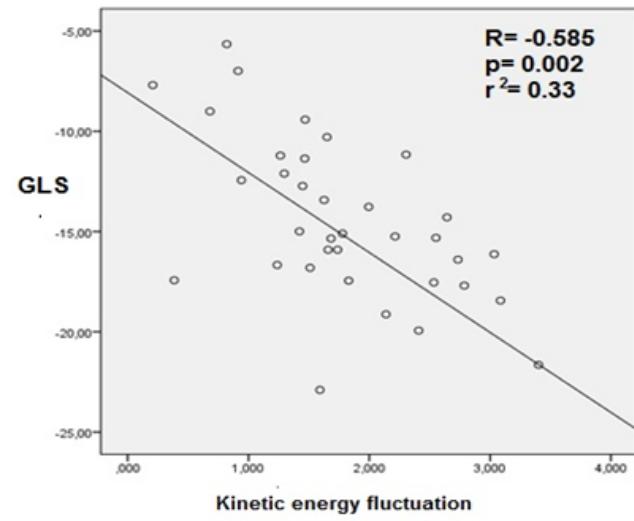
*a)*



*b)*



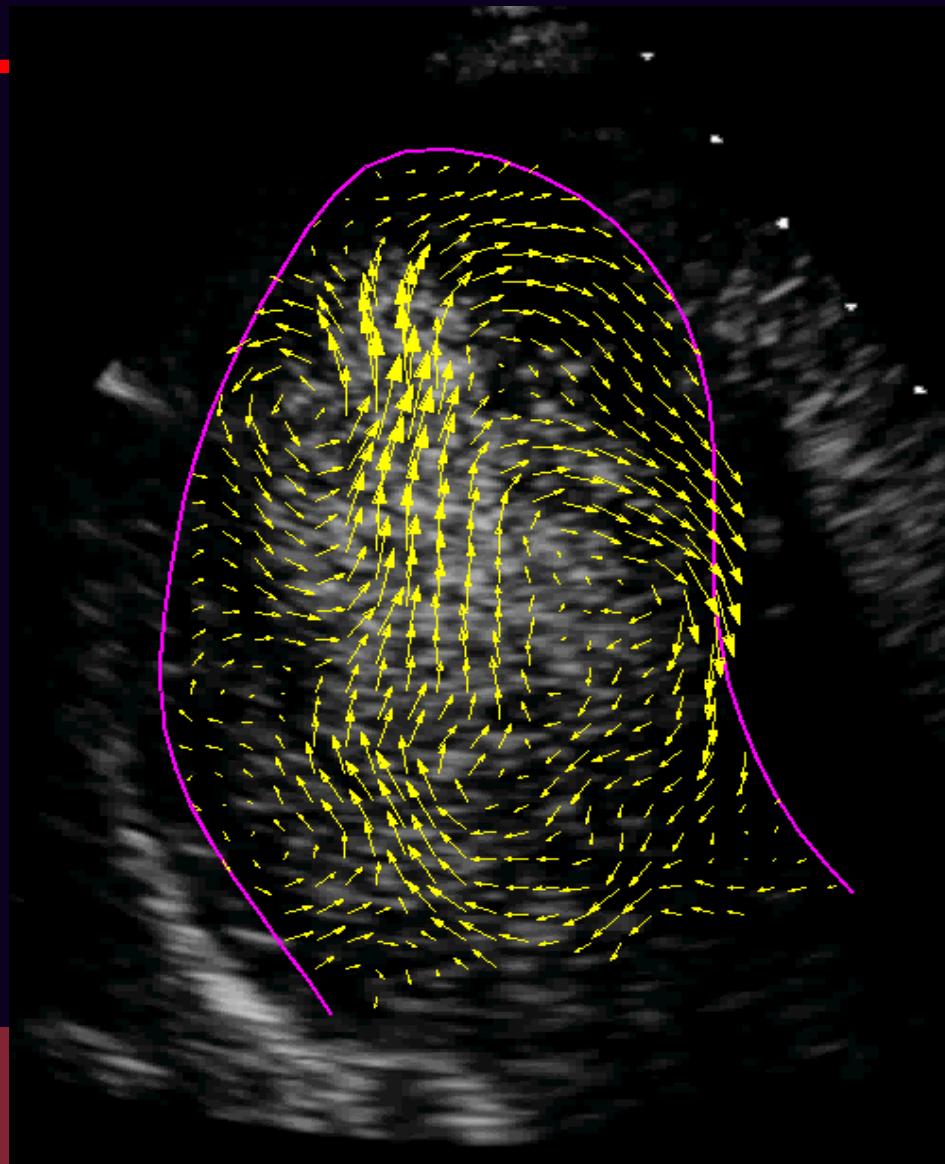
*c)*



*d)*



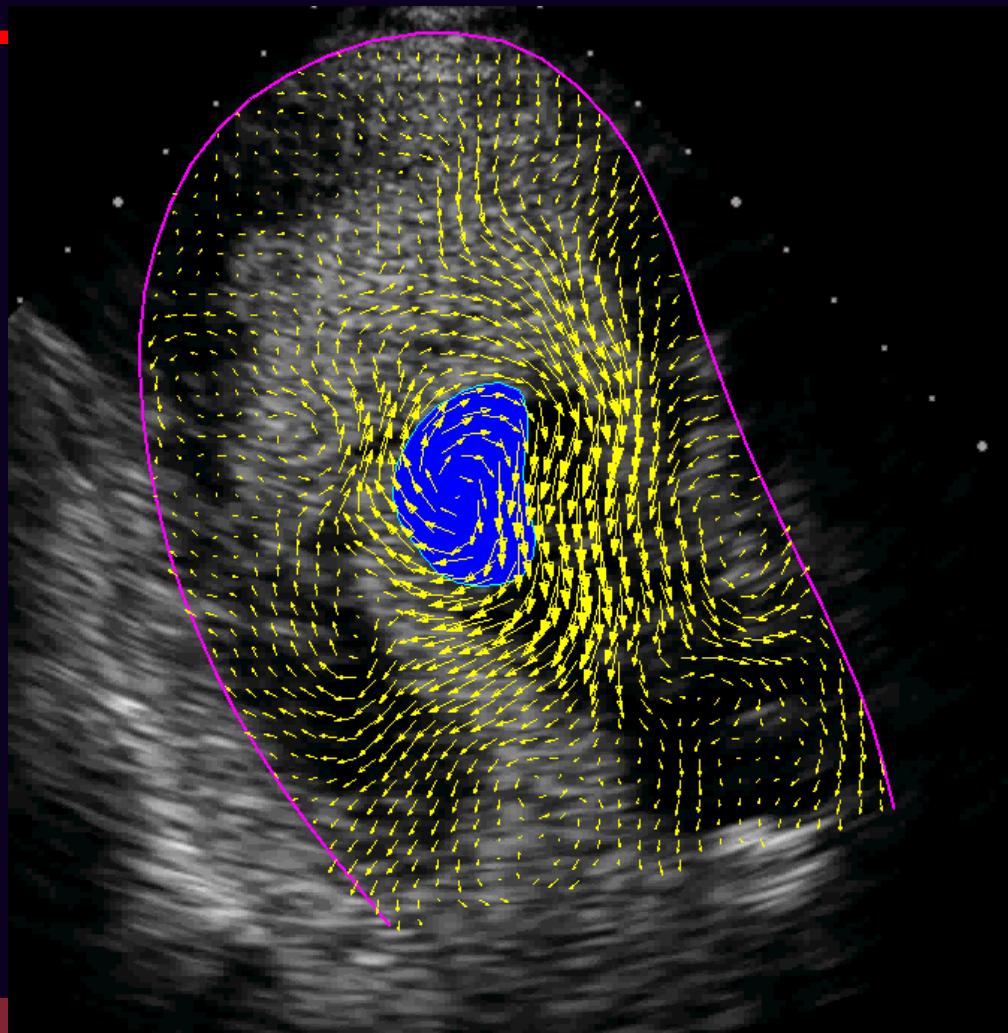
**LVEF 45%, Energy Diss. = 0.88**



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**LVEF= 25% , Energy Diss. = 0.27**



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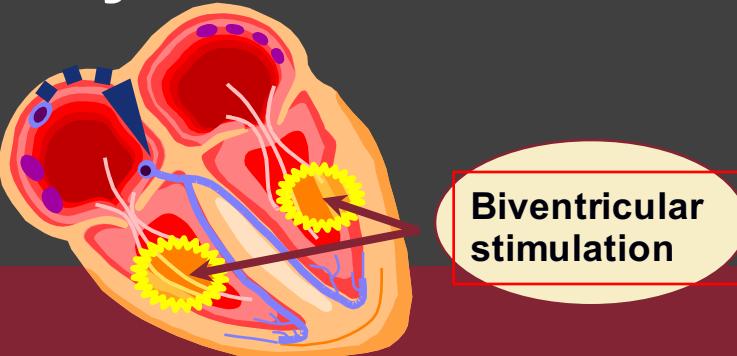
# Conclusions

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- Patients with preserved LV function exhibit a significant increase in energy dissipation suggesting the presence of a new fluid-tissue dynamical balance as compensatory mechanisms for maintaining an adequate LVEF.
- In the presence of a significant LV dysfunction, energy dissipation is markedly reduced as a consequence of a low flow kinetic energy.
- Serial monitoring of energy dissipation changes overtime allows quantitative information on LV pump efficiency.
- The role of these parameters in the development and maintenance of LV remodeling has to be clarified



# Significant increase of flow kinetic energy in non responders patients to Cardiac Resynchronization Therapy



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Luciano Agati, MD

# Guidelines Indication to CRT

Scientific Society	HFSA	ACCF/AHA/HRS	ESC/EHRA
CLASS I: Treatment should be performed (ACCF/AHA/HS) or is recommended (ESC/EHRA)	NYHA II – III LVEF≤35% Sinus rhythm QRS≥150ms Not due to RBBB	NYHA III and ambulatory NYHA IV LVEF≤35% Sinus rhythm QRS≥150ms LBBB	NYHA II, III, and ambulatory NYHA IV LVEF≤35% Sinus rhythm QRS≥150ms LBBB
<i>Level of Evidence A</i>			
CLASS I: Treatment should be performed (ACCF/AHA/HS) or is recommended (ESC/HFA)		NYHA II LVEF≤35% Sinus rhythm QRS≥150ms LBBB	NYHA II, III, and ambulatory NYHA IV LVEF≤35% Sinus rhythm QRS 120 - 150 ms LBBB
<i>Level of Evidence B</i>			NYHA III, and ambulatory NYHA IV LVEF≤35% Upgrade from IPG or ICD High percentage of ventricular pacing
CLASS IIa: Treatment is reasonable to be performed (ACCF/AHA/HRS) or should be considered (ESC/HFA)		NYHA III and ambulatory NYHA IV LVEF≤35% Sinus rhythm QRS≥150ms Non-LBBB morphology	



# Unsolved issues:

- Electrical dyssynchrony not necessary means mechanical dyssynchrony
- ~ 30% CRT “non-responders” (clinical and echo parameters)

*.... "no single echocardiographic measure of dyssynchrony may be recommended to improve patient selection for CRT beyond current guidelines..."*



European Heart Journal  
doi:10.1093/euroheartj/eht150

## 2013 ESC Guidelines on cardiac pacing and cardiac resynchronization therapy

The Task Force on cardiac pacing and resynchronization therapy of the European Society of Cardiology (ESC). Developed in collaboration with the European Heart Rhythm Association (EHRA).

ESC GUIDELINES

Circulation  
JOURNAL OF THE AMERICAN HEART ASSOCIATION



Results of the Predictors of Response to CRT (PROSPECT) Trial  
Eugene S. Chung, Angel R. Leon, Luigi Tavazzi, Jing-Ping Sun, Petros Nihoyannopoulos, John Merlino, William T. Abraham, Stefano Ghio, Christophe Leclercq, Jeroen J. Bax, Cheuk-Man Yu, John Gorcsan III, Martin St John Sutton, Johan De Sutter and Jaime Murillo



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## Aim of the study

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- To assess iv flow patterns in patients undergoing CRT
- To assess the correlation between reverse remodeling and iv flow dynamics in the attempt to find new non invasive predictive indexes of CRT responders

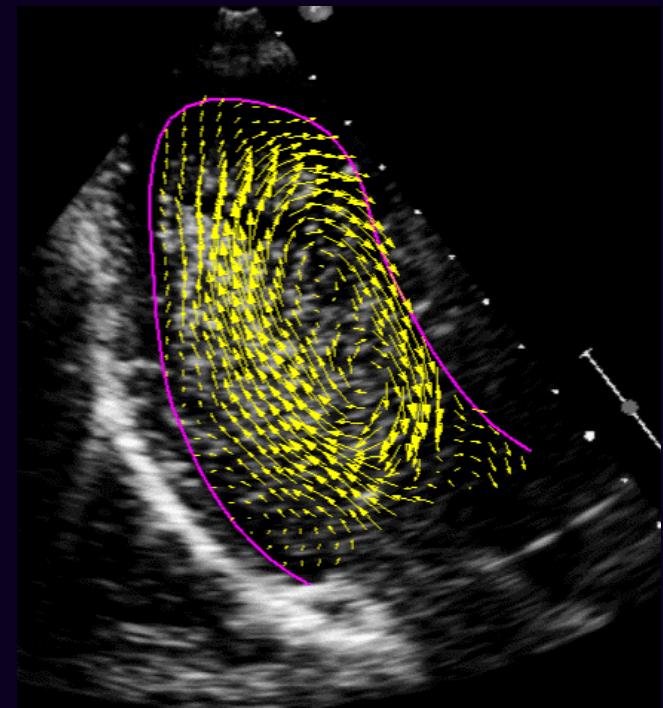
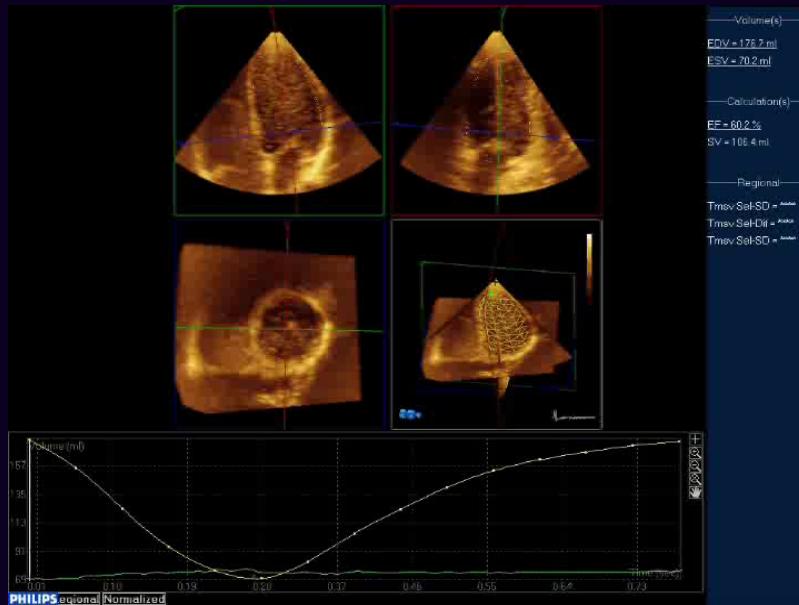
## Study population

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- 30 consecutive patients undergoing CRT following current guidelines.



# Study design



- Echo 2D and 3D at baseline and after 6 months (GLS- SDI)
- Echo-PIV - at baseline and after 6 months (PMK ON and OFF)

## CRITERIA TO DEFINE RESPONDERS

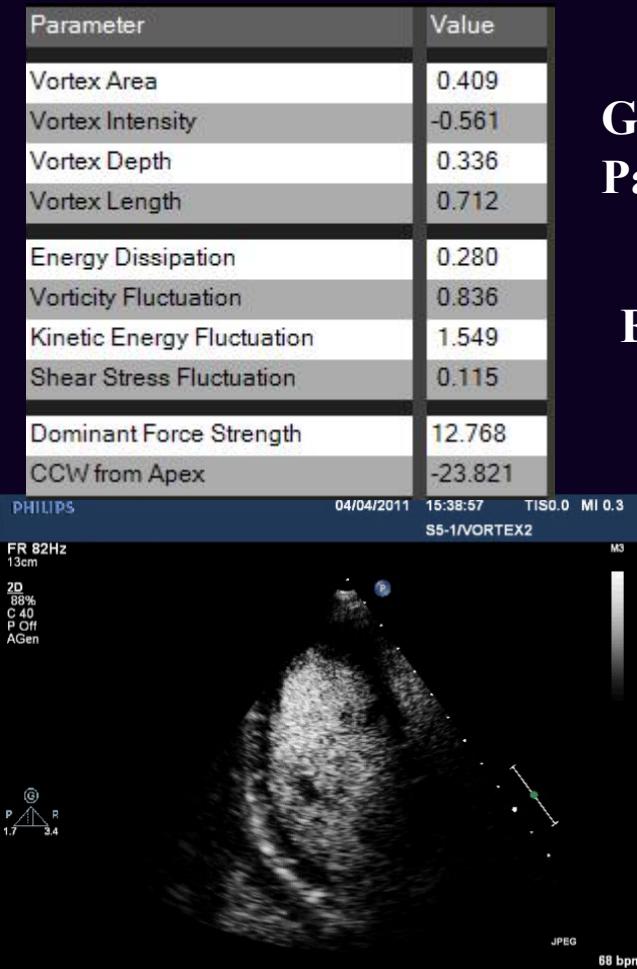
- Increase LVEF ( $> 20\%$ )
- NYHA class improvement
- LV end-systolic volume reduction



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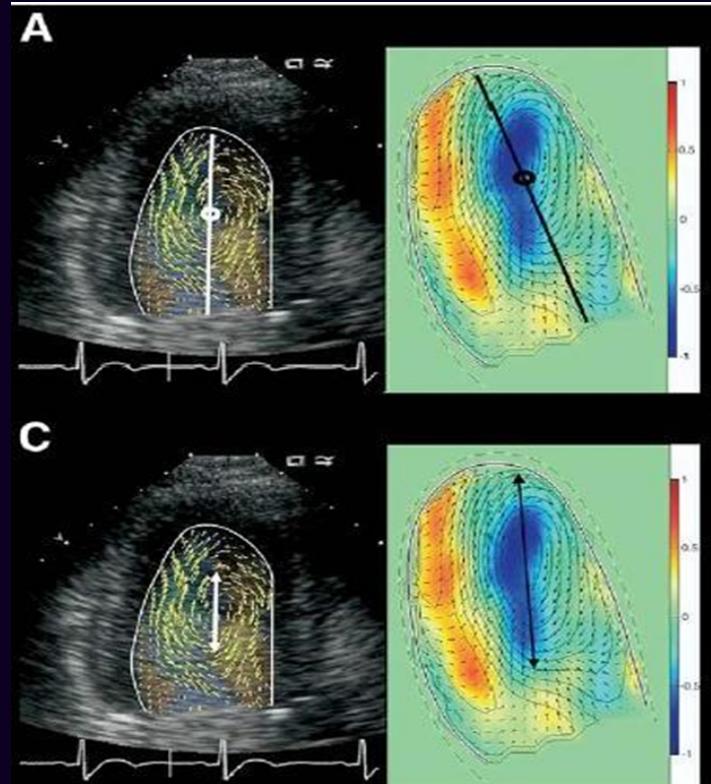
# Echo-PIV

Software Hyper Flow 6.0.0.3.



## Geometrical Parameters

## Energetics



European Heart Journal - Cardiovascular Imaging Advance Access published June 6, 2014



European Heart Journal – Cardiovascular Imaging  
doi:10.1093/eihci/jeu106



## Quantitative analysis of intraventricular blood flow dynamics by echocardiographic particle image velocimetry in patients with acute myocardial infarction at different stages of left ventricular dysfunction

L. Agati<sup>1\*</sup>, S. Cimino<sup>1</sup>, G. Tonti<sup>2</sup>, F. Cicogna<sup>1</sup>, V. Petronilli<sup>1</sup>, L. De Luca<sup>1</sup>, C. Iacoboni<sup>1</sup>, and G. Pedrizzetti<sup>3</sup>



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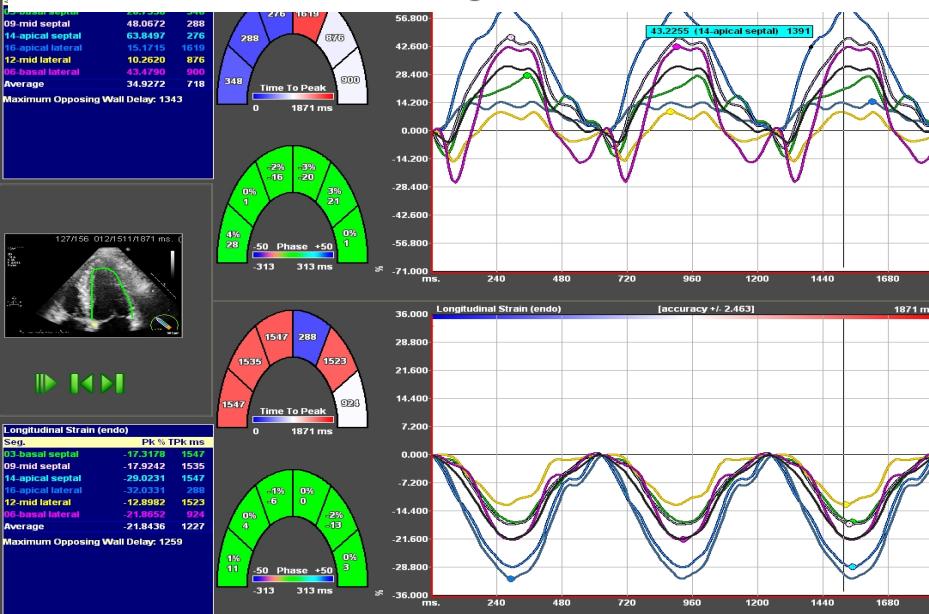
# Speckle Tracking Echocardiography



European Heart Journal – Cardiovascular Imaging  
doi:10.1093/ejci/jez295

**Global and regional longitudinal strain assessed by two-dimensional speckle tracking echocardiography identifies early myocardial dysfunction and transmural extent of myocardial scar in patients with acute ST elevation myocardial infarction and relatively preserved LV function**

S. Cimino<sup>1</sup>, E. Canali<sup>1</sup>, V. Petronilli<sup>1</sup>, F. Cicogna<sup>1</sup>, L. De Luca<sup>1</sup>, M. Francone<sup>2</sup>, G. Sardella<sup>1</sup>, C. Iacoboni<sup>1</sup>, and L. Agati<sup>1\*</sup>

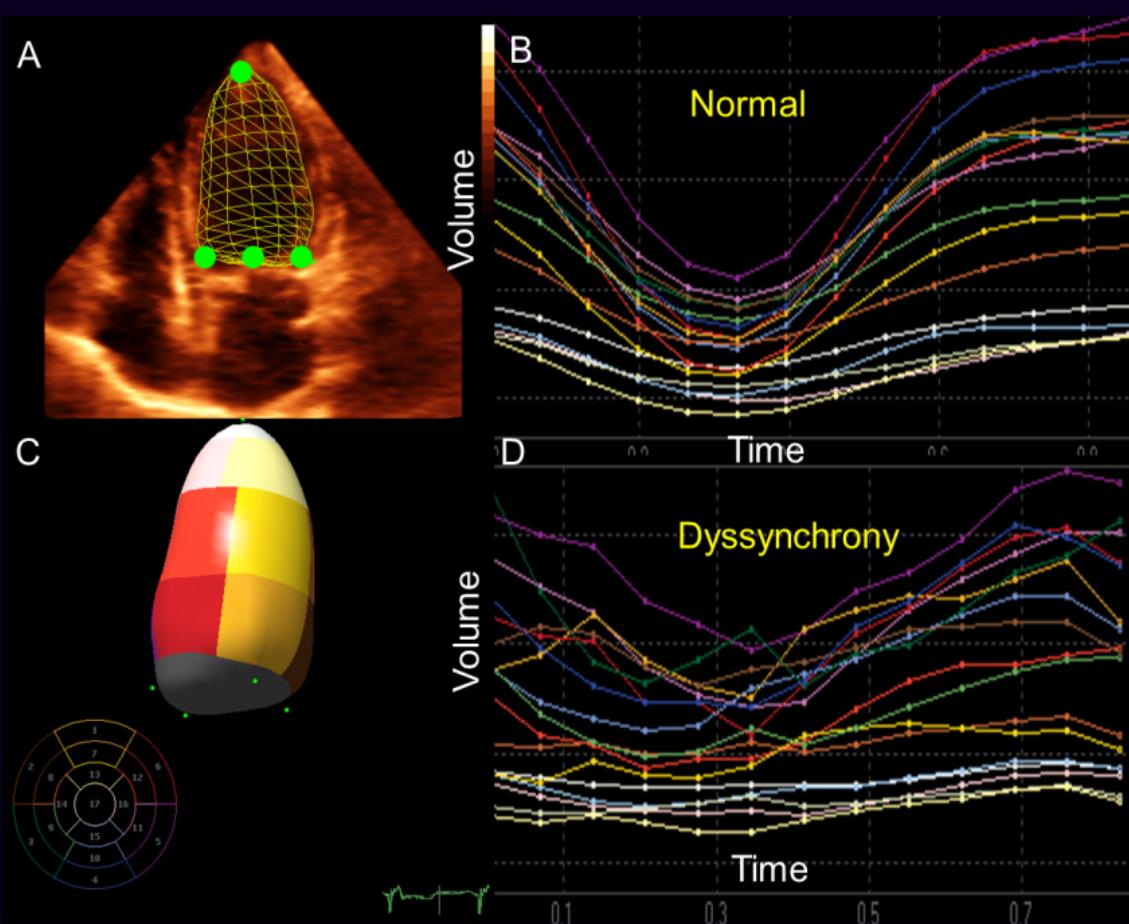


- Longitudinal Strain
- Strain rate
  - Velocity
  - Displacement
  - Torsion



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# ECO 3D/4D and CRT



## SYSTOLIC DYSSYNERGY INDEX (SDI):

Time necessary to reach minimum volume for each segment

$SDI < 3.5 \pm 1.8\%$  Normal

$SDI > 15 \pm 1\%$  Severe LV dysfunction

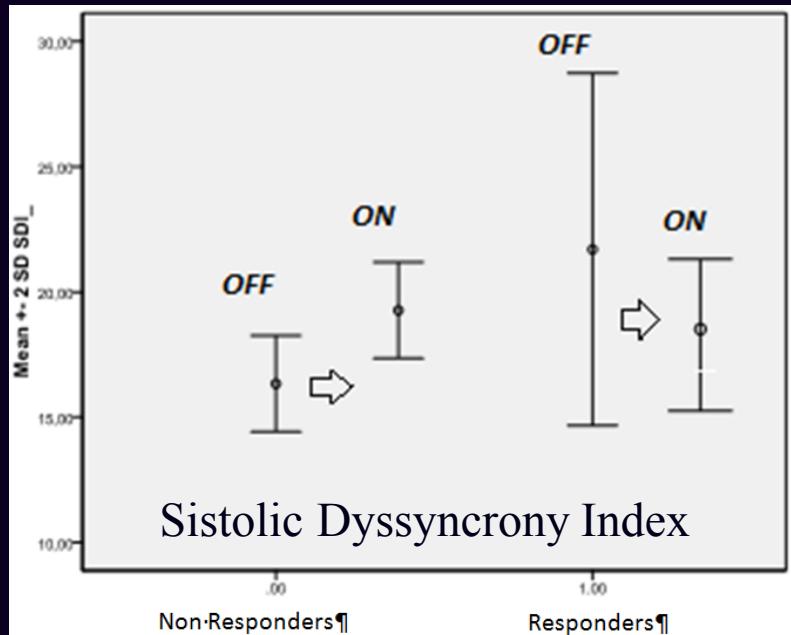
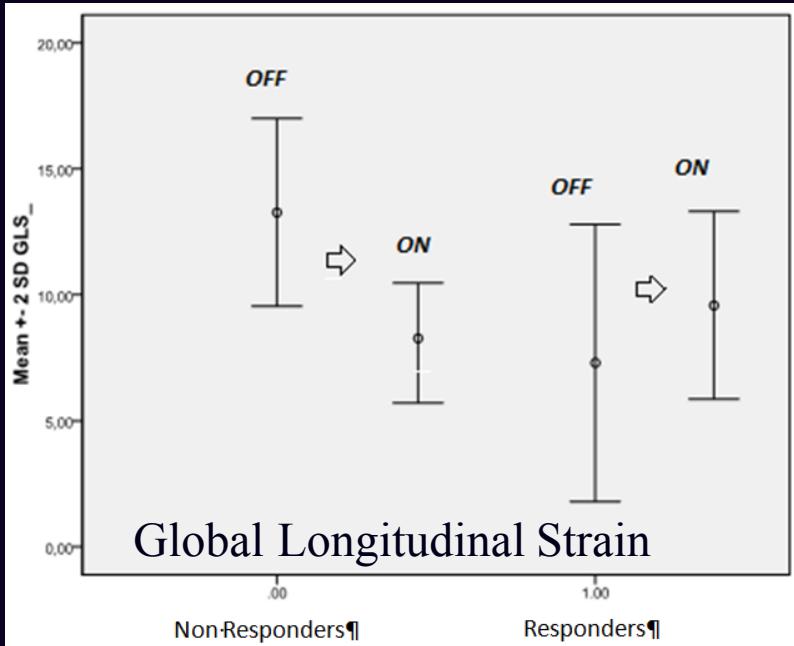


# BASELINE – Before CRT

Parameter	Responders (N=16)	Non responders (N=14)	p
Energy dissipation	<b>0.578<math>\pm</math>0.08</b>	<b>0.54<math>\pm</math>0.001</b>	NS
Vortex area	<b>0.23<math>\pm</math>0.02</b>	<b>0.19<math>\pm</math>0.05</b>	NS
Vortex intensity	<b>0.35<math>\pm</math>0.01</b>	<b>0.13<math>\pm</math>0.36</b>	NS
Vortex depth	<b>0.36<math>\pm</math>0.01</b>	<b>0.49<math>\pm</math>0.03</b>	<b>&lt;0.001</b>
Vortex length	<b>0.58<math>\pm</math>0.04</b>	<b>0.49<math>\pm</math>0.05</b>	<b>0.002</b>
Vorticity fluctuation	<b>0.81<math>\pm</math>0.001</b>	<b>0.82<math>\pm</math>0.06</b>	NS
Kinetic Energy fluctuation	<b>1.24<math>\pm</math>1.14</b>	<b>1.36<math>\pm</math>0.32</b>	NS
LVEF (%)	<b>17.5<math>\pm</math>2.6</b>	<b>25<math>\pm</math>7.3</b>	<b>0.014</b>
LVED vol(ml)	<b>223<math>\pm</math>11.2</b>	<b>159<math>\pm</math>28</b>	NS
LVES vol(ml)	<b>188<math>\pm</math>28</b>	<b>121<math>\pm</math>29</b>	NS
GLS (%) pre	<b>-7.39<math>\pm</math>3.55</b>	<b>-13.58<math>\pm</math>2.1</b>	<b>0.001</b>
SDI (%) pre	<b>22.1<math>\pm</math>4.5</b>	<b>16.5<math>\pm</math>0.7</b>	<b>0.048</b>
NYHA III-IV (%) pre	<b>80% (8)</b>	<b>100% (10)</b>	NS
Age (yrs)	<b>65<math>\pm</math>9</b>	<b>69<math>\pm</math>10</b>	NS
Male%	<b>60% (6)</b>	<b>50% (5)</b>	NS



# 6-Months Follow-up: GLS and SDI PMK ON vs PMK OFF



Parametro	Pmk-off	Pmk-on	P
GLS(%)	-7.39±3.55	-9.77±4.4	0.038
SDI(%)	22.1±4.5	15.5±3.7	0.02
GLS(%)	13.58±2.1	7.55±3.3	0.005
SDI(%)	16.5±0.7	19±1.4	NS

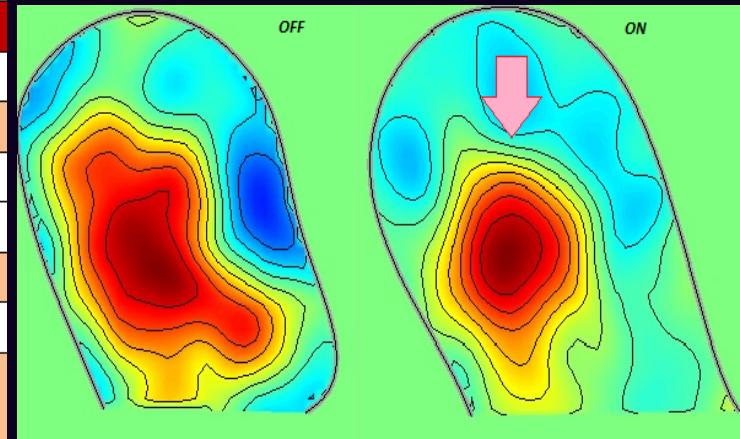
Responders

Non Responders



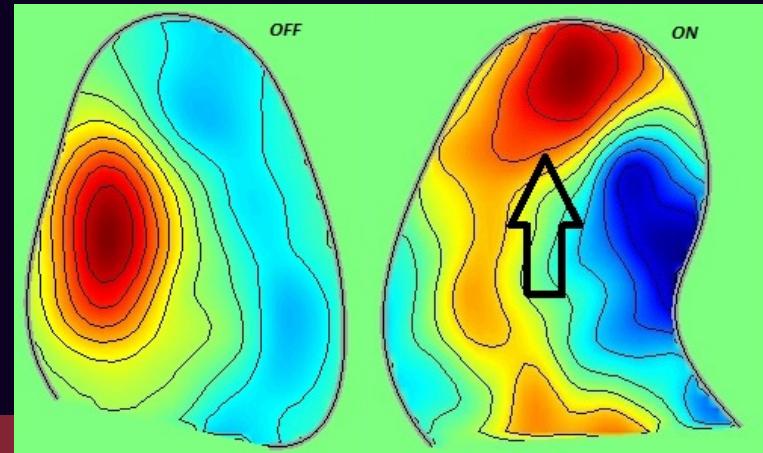
# 6-Months Follow-up: Flow Parameters Responders

Parametro	Pmk-off	Pmk-on	p
Energy-dissipation	0.578±0.08	0.523±0.09	NS
Vortex-area	0.23±0.02	0.19±0.01	0.02
Vortex-intensity	0.35±0.01	0.37±0.003	0.002
Vortex-depth	0.36±0.01	0.34±0.07	NS
Vortex-length	0.58±0.04	0.5±0.04	0.043
Vorticity-fluctuation	0.81±0.001	0.74±0.09	NS
Kinetic-Energy-fluctuation	1.24±1.14	1.07±0.34	0.039



## Non Responders

Parametro	Pmk-off	Pmk-on	p
Energy-dissipation	0.54±0.001	0.766±0.32	0.004
Vortex-area	0.19±0.05	0.21±0.05	NS
Vortex-intensity	0.13±0.36	0.08±0.03	NS
Vortex-depth	0.49±0.03	0.56±0.19	NS
Vortex-length	0.49±0.04	0.59±0.18	NS
Vorticity-fluctuation	0.82±0.05	0.87±0.05	<0.001
Kinetic-Energy-fluctuation	1.36±0.32	1.6±0.5	0.002



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# Before and After CRT

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# Conclusions

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- CRT in Responder significantly reduces energy dyssipation and increases flow uniformity.
- CRT in non-Responder further increase dyssynchronicity and iv flow turbulence likely responsible for adverse LV remodeling
- Fluid dynamics analysis before and after CRT may be useful to early identify non-Responders
- Prospective multicenter studies are needed to verify the additional value of fluid dynamics analysis in this setting



CRT optimization with quadripolar LV  
pacing leads through the novel  
Vortex® echocardiographic algorhythm

## The Vortex-CRT study

