

Multiple premature ventricular
contractions favouring heart
failure in candidates to CRT:
where do we start from?

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Background

Premature ventricular contractions (PVCs) are frequently found in patients both **with** and **without** structural disease.

The recommended treatment for PVCs has been evolving over the years.

The level of importance relegated to the presence of PVCs has also been a moving target

→ In the past, PVC only a negative prognostic factors after MI → moving to **PVC-induced CMP**

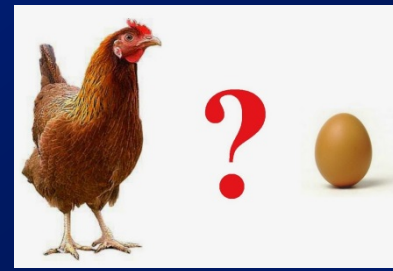
Vicious circle

HF



PVC

PVC and HF: who come first ?



PVCs usually precede cardiomyopathy

In the **absence** of **underlying cardiac** disease,

in the **absence** of **coronary artery** disease

- 1) PVCs >20,000 per day;
- 2) **one** or **two** primary PVC **morphologies**;
- 3) origin in RVOT, LVOT, or fascicular;

Provided that **preserved myocardial thickness** and **absence of scar** on echocardiogram are documented

Cardiomyopathy: how many PVCs are needed?

Impact of ventricular ectopic burden in a premature ventricular contraction–induced cardiomyopathy animal model

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(Heart Rhythm 2016;13:755–761)

Sort of threshold effect

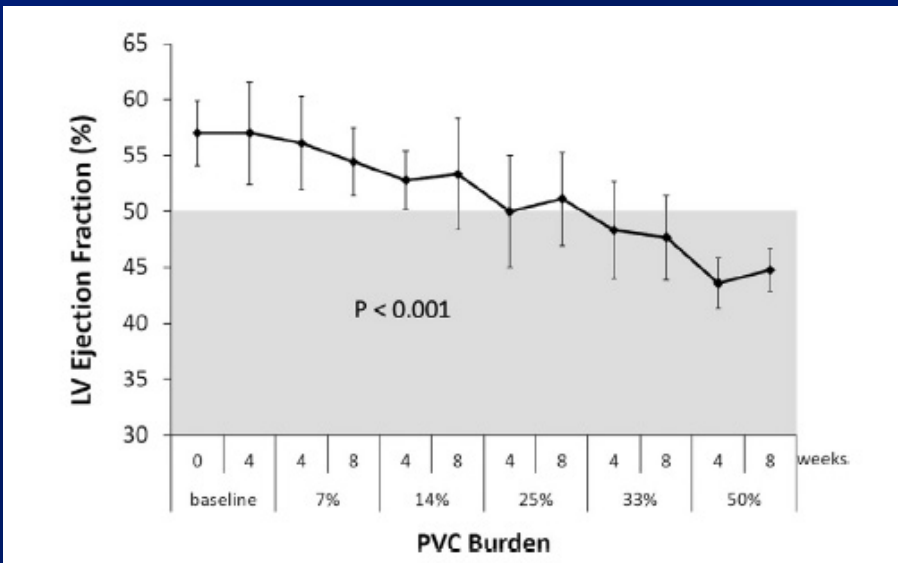


Figure 1 Progression of LV ejection fraction after 4 and 8 weeks of a progressive incremental PVC burden starting from 0% (baseline) to 7%, 14%, 24%, 33%, and 50%. *P* values were calculated using repeated-measures 1-way analysis of variance. LV = left ventricular; PVC = premature ventricular contraction.

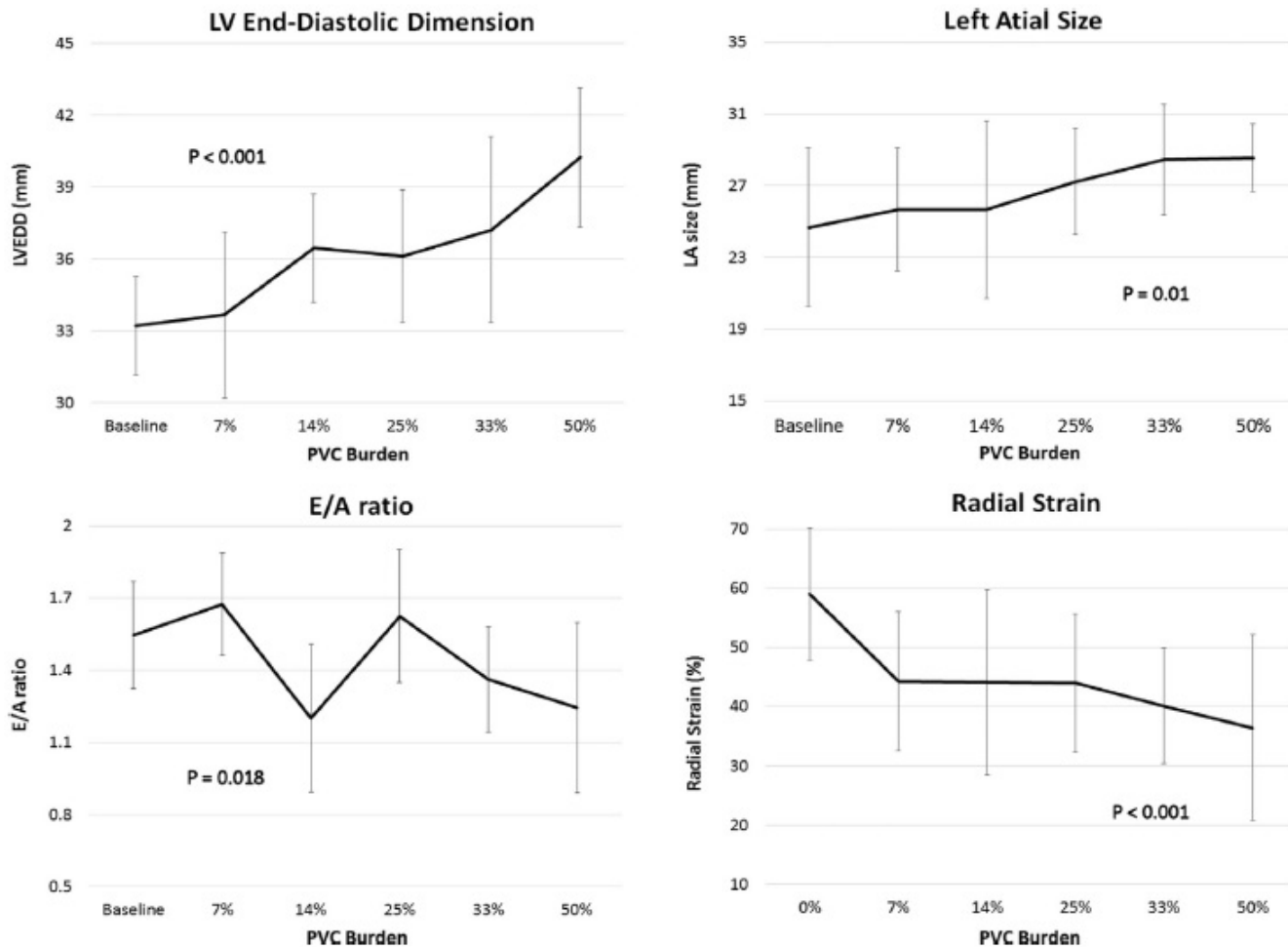


Figure 2 Estimated means of LVEDD, LA size, E/A ratio, and radial strain that showed a significant change between PVC burden levels (0%, 7%, 14%, 25%, 33%, and 50%). *P* values were calculated using repeated-measures 1-way analysis of variance. Details are given in Table 2. LA = left atrial; LV = left ventricular; LVEDD = left ventricular end-diastolic dimension; PVC = premature ventricular contraction.

25% 33% : minimum burden to induce CM
50% burden : 100 % CM

Yet, a high PVCs burden is not solely sufficient to induced CM ...

Predictors of PVC- induced CM

WIDER PVC QRS duration (epicardial ?)

NON OUTFLOW TRACT site of origin

~~PVC coupling interval~~

Mechanism of PVC- induced CM

→ ~~Tachycardia induced CM ?:~~
normal average rate !

→ ~~PVCs trigger a remodeling process →~~
structural abnormalities

no fibrosis on animal studies !

→ Dyssynchronous and ineffective
mechanical ventricular contraction:

Which therapy ?

→ AADs Class II (betablockers)


Class III (sotalol, amiodarone)

→ RF ablation





Heart Rhythm

Volume 11, Issue 2, February 2014, Pages 187–193



Relative efficacy of catheter ablation vs antiarrhythmic drugs in treating premature ventricular contractions: A single-center retrospective study ^{*}

Li Zhong, MD, PhD^{†,‡}, Ying-Hsiang Lee, MD^{†,‡}, Xin-Miao Huang, MD, PhD^{†,‡}, Samuel J. Asirvatham, MD, FHRSt, Win-Kuang Shen, MD, FHRSt, Paul A. Friedman, MD, FHRSt, David O. Hodge, MS[§], Joshua P. Slusser, BS[§], Zhi-Yuan Song, MD^{†,‡},  , Douglas L. Packer, MD, FHRSt, Yong-Mei Cha, MD, FHRSt^{†,‡} 



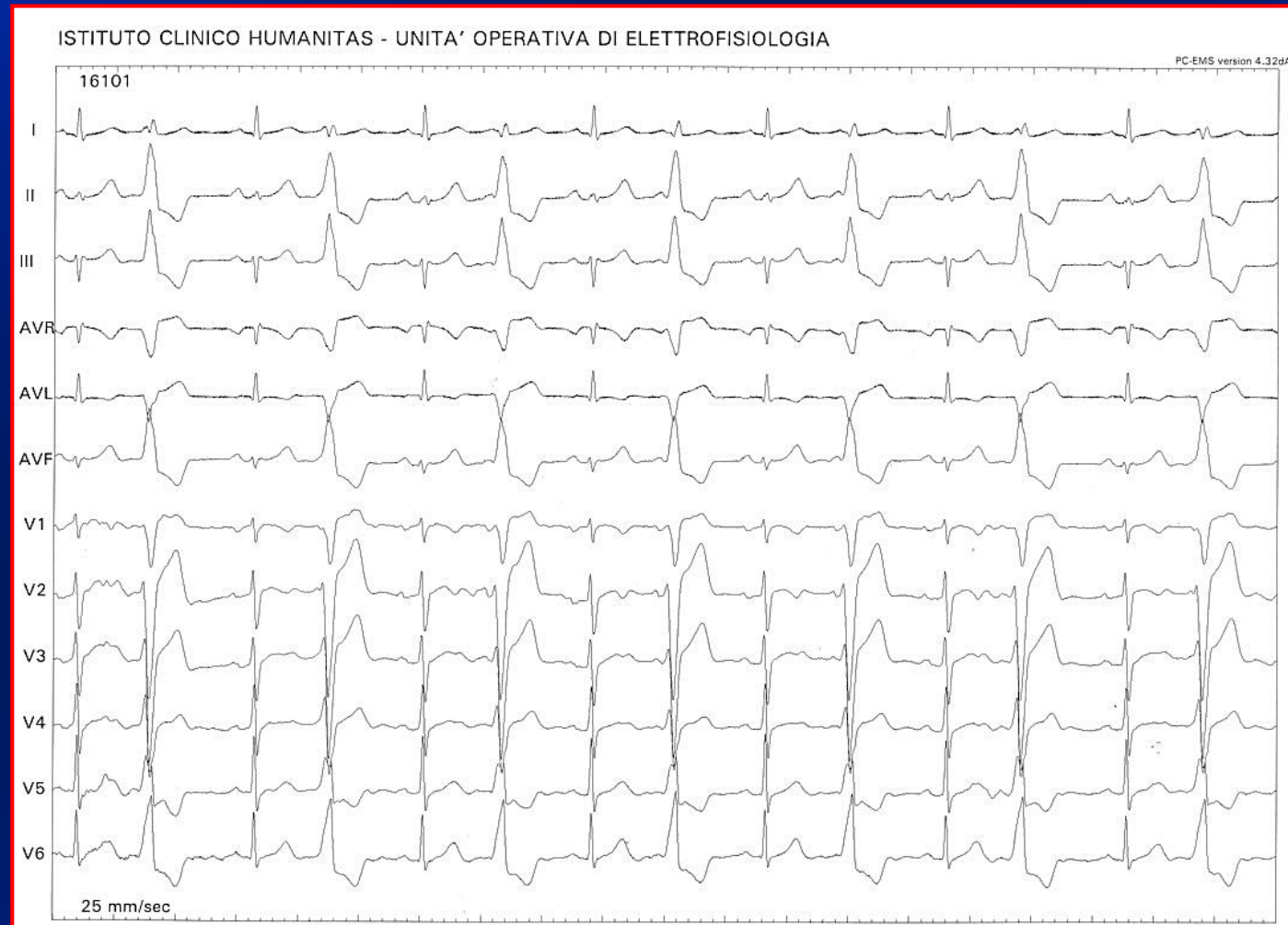
1) No structural heart disease

PVCs from RVOT

Male

47 yrs old

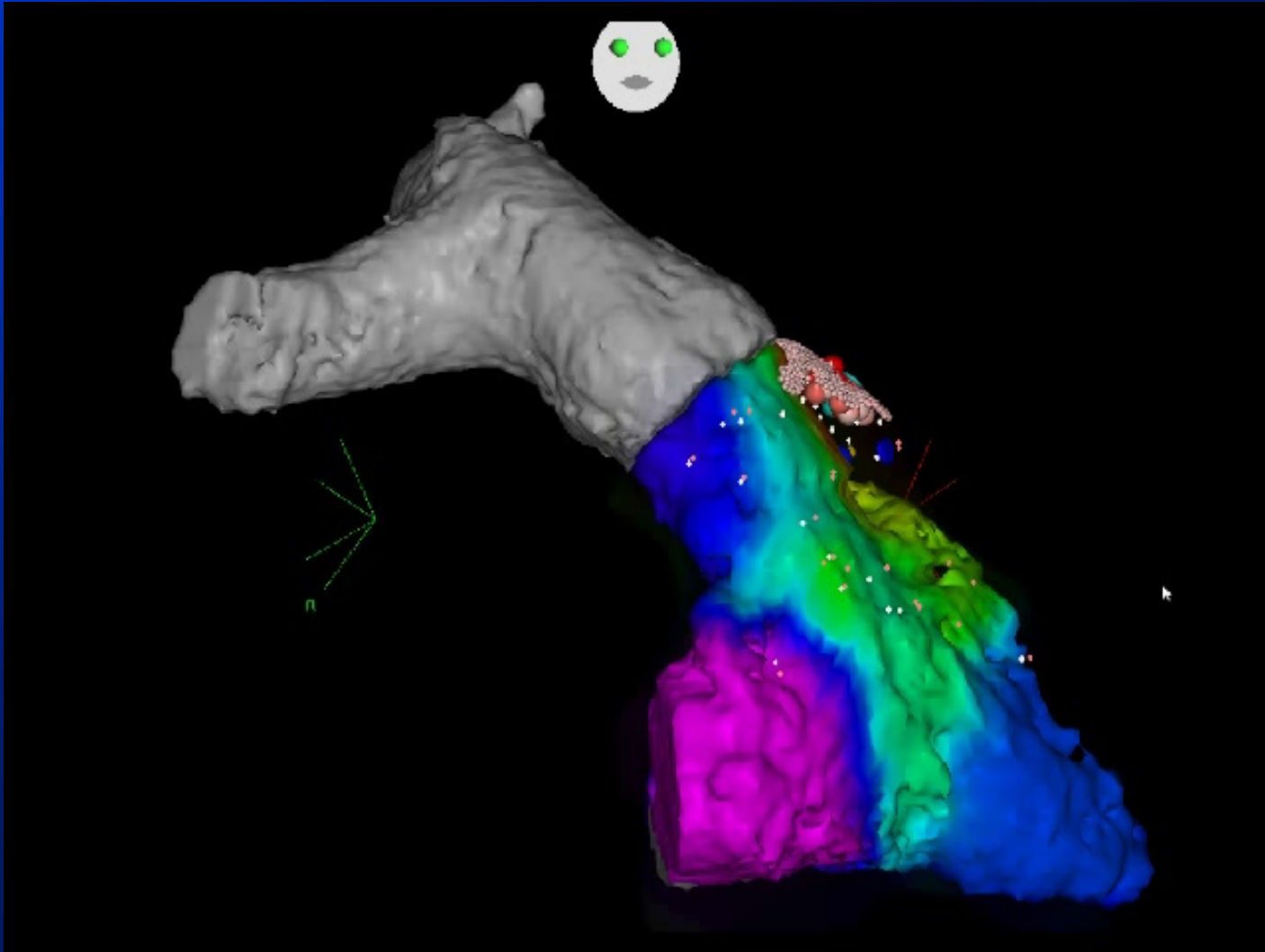
LVEF 45%

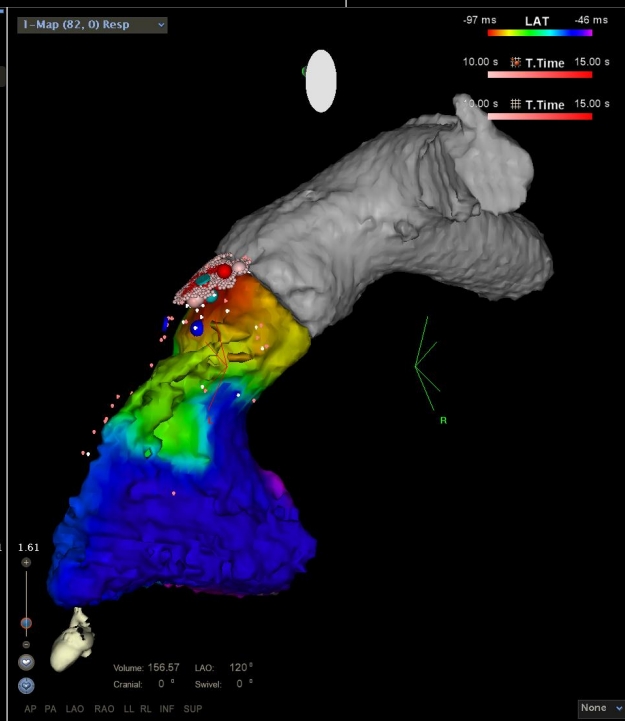
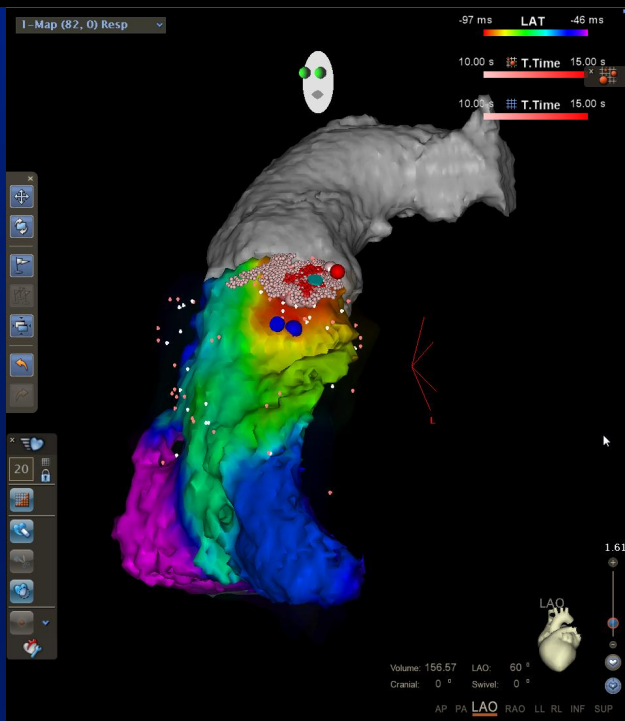
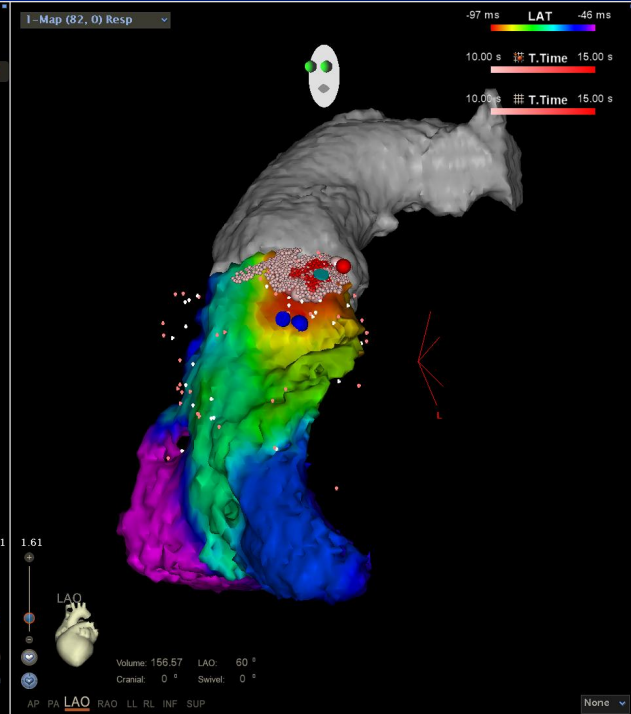
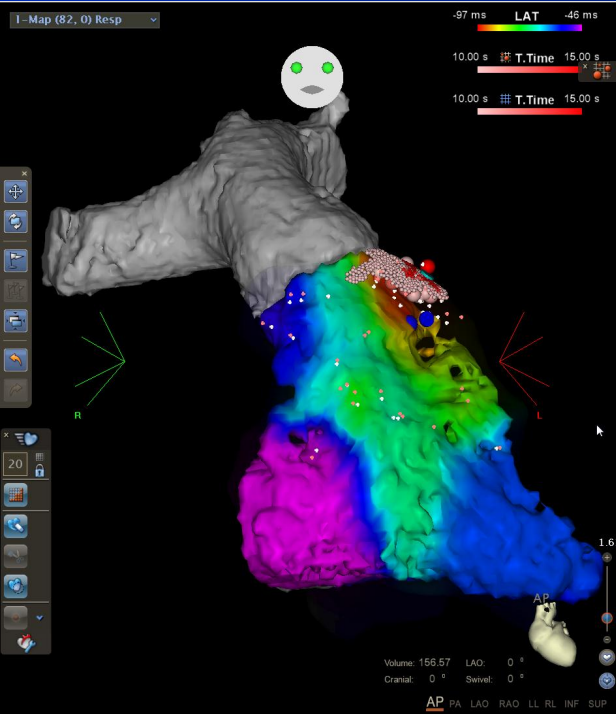
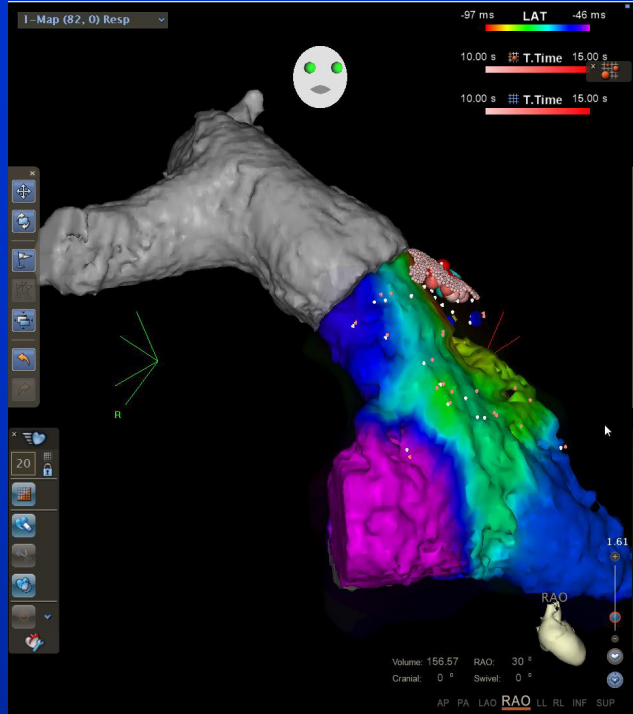


2015 ESC Guidelines for the management of patients with ventricular arrhythmias and the prevention of sudden cardiac death

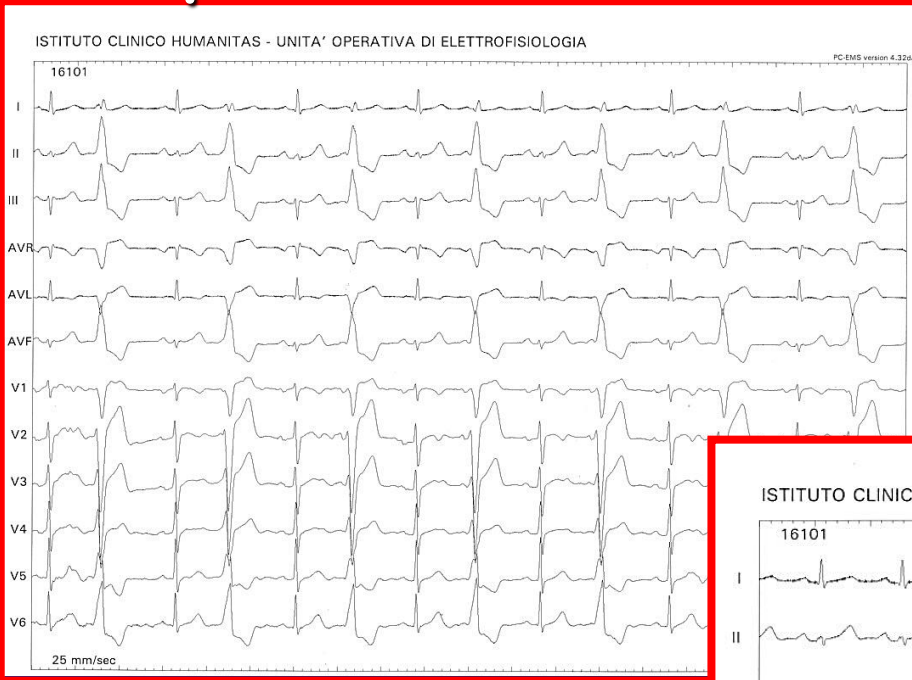
Treatment of outflow tract ventricular tachycardia

Recommendations	Class ^a	Level ^b	Ref
Catheter ablation of RVOT VT/PVC is recommended in symptomatic patients and/or in patients with a failure of anti-arrhythmic drug therapy (e.g. beta-blocker) or in patients with a decline in LV function due to RVOT-PVC burden.	I	B	525–528
Treatment with sodium channel blockers (class IC agents) is recommended in LVOT/aortic cusp/epicardial VT/PVC symptomatic patients.	I	C	529–531

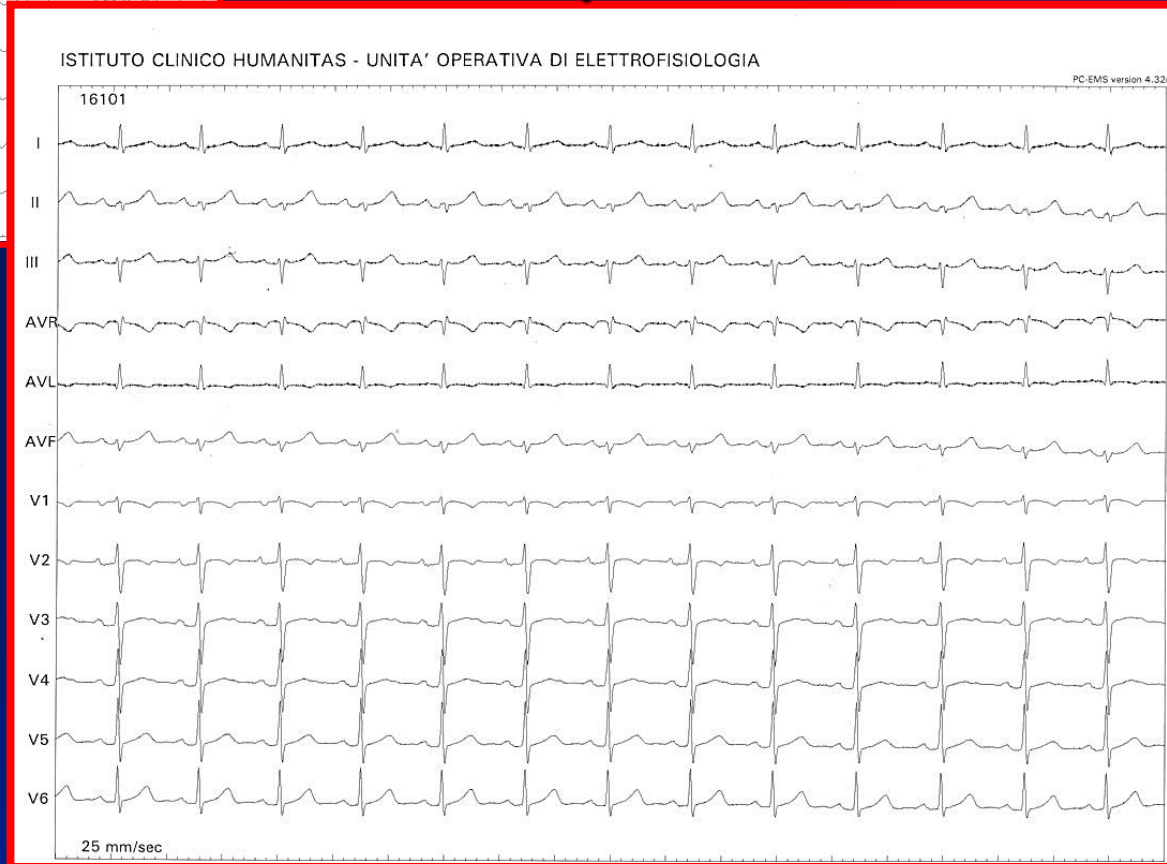




Pre procedure



Post procedure



3 months later

LVEF 55%

Relative efficacy of catheter ablation vs antiarrhythmic drugs in treating premature ventricular contractions: A single-center retrospective study*

Li Zhong, MD, PhD^{1,†}, Ying-Hsiang Lee, MD^{1,‡}, Xin-Miao Huang, MD, PhD^{1,§}, Samuel J. Asirvatham, MD, FHR^{1,¶}, Win-Kuang Shen, MD, FHR^{1,¶}, Paul A. Friedman, MD, FHR^{1,¶}, David O. Hodge, MS^{1,¶}, Joshua P. Slusser, BS^{1,¶}, Zhi-Yuan Song, MD^{1,¶}, Douglas L. Packer, MD, FHR^{1,¶}, Yong-Mei Cha, MD, FHR^{1,¶}

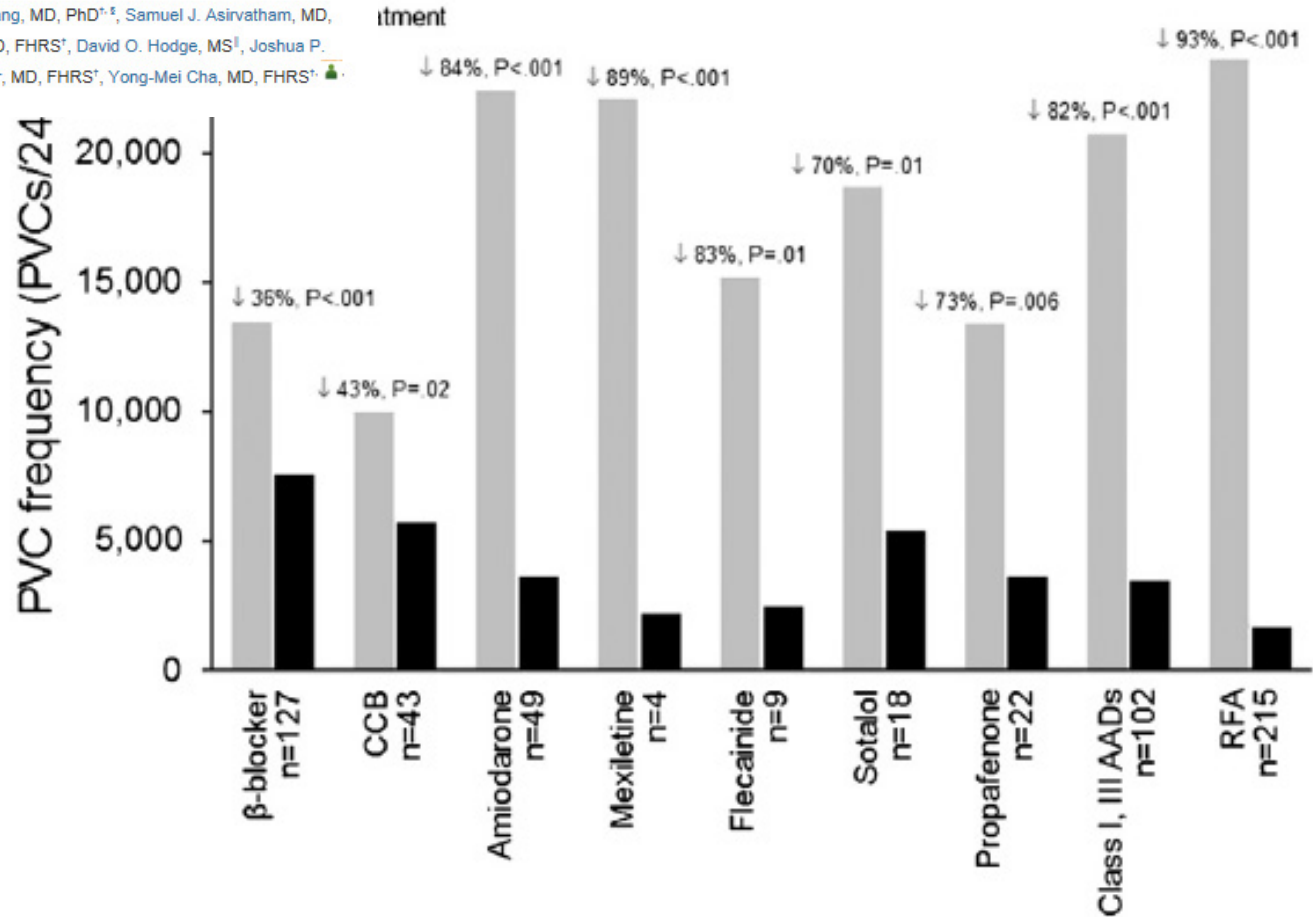


Figure 2 Reduction in the frequency of premature ventricular contractions (PVCs) after treatment with antiarrhythmic drugs (AADs) and radiofrequency ablation (RFA). CCB = calcium channel blocker.

The PVC frequency and burden significantly reduced by both RFA and AADs

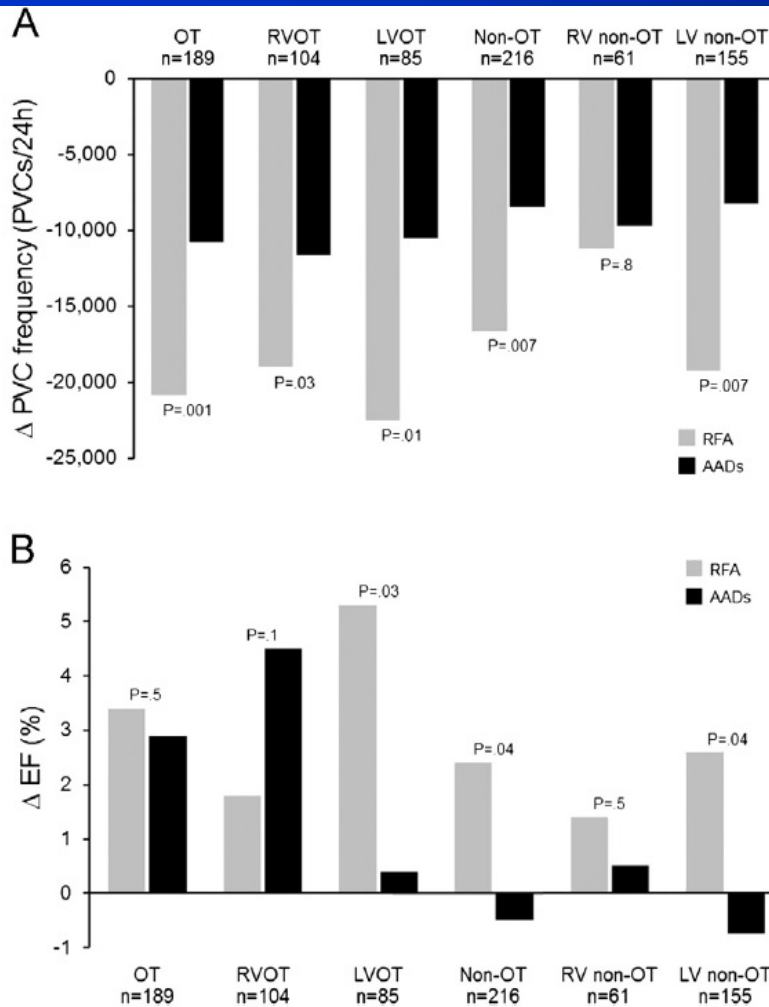


Figure 3 Changes in premature ventricular contraction (PVC) frequency and left ventricular ejection fraction (LVEF) after treatment by PVC origin groups. **A:** Comparison of PVC reduction after antiarrhythmic drug (AAD) and radiofrequency ablation (RFA) treatment. **B:** Comparison of changes in LVEF after RFA and AAD treatment. LV non-OT = left ventricular non-outflow tract; LVOT = left ventricular outflow tract; RV non-OT = right

In pts with lower PVC frequency, only RFA, not AADs, effectively reduced PVCs frequency

RFA more effective than AADs ($p < 0.001$)

Table 1 Comparison of baseline clinical characteristics between RFA and AAD treatment groups*

Characteristic	RFA (n = 215)	AADs (n = 295)	p†
Age (y)	47.2 ± 16.3	61.6 ± 17.8	<.001
No. of women/men	115/110	110/185	<.001
Comorbid conditions			
Hypertension	23 (11)	106 (36)	<.001
Diabetes	12 (6)	28 (9)	.121
CAD	13 (6)	38 (13)	.010
AF	12 (5)	45 (15)	.002
DCM	68 (32)	85 (29)	<.001
Sleep apnea	1 (0.4)	14 (5)	.074
History of syncope	15 (7)	13 (4)	.223
Symptoms			
Shortness of breath	32 (15)	43 (15)	.903
Dizziness	19 (9)	22 (7)	.645
Palpitations	58 (27)	59 (20)	.100
Chest discomfort	27 (13)	17 (6)	.010
PVC frequency (n/24 h)	23,554 ± 18,448	17,259 ± 14,512	<.001
PVC burden (%)	18.4 ± 15.2	12.1 ± 12.8	<.001
NSVT (runs/24 h)	275 ± 51	112 ± 43	.014
PVC origins			
RVOT	74 (34)	33 (11)	<.001
RV non-OT	22 (10)	39 (13)	.109
LVOT	47 (22)	39 (13)	.143
LV non-OT	51 (24)	106 (36)	<.001
Multiform PVC	9 (4)	18 (6)	.202
PVC QRS duration (ms)	148 ± 21	150 ± 19	.311
PVC coupling interval (ms)	488 ± 95	494 ± 104	.545
LVEF (%)	53.0 ± 11.9	52.1 ± 8.5	.210
LVESD (mm)	37.5 ± 8.7	37.2 ± 9.5	.723
LVEDD (mm)	53.9 ± 7.0	52.4 ± 6.7	.301
AADs			
β-Blocker	34 (16)	135 (46)	
CCB	6 (3)	21 (7)	
Mexiletine	2 (1)	13 (4)	
Flecainide	3 (1)	26 (9)	
Sotalol	2 (1)	12 (4)	
Amiodarone	5 (2)	41 (14)	
Propafenone	5 (2)	16 (5)	
Digoxin	3 (1)	25 (8)	<.001
ACEI/ARB	25 (12)	95 (32)	<.001

2) Structural heart disease

PVCs & structural heart disease (EF < 30%)



First step: OPT → CRT



If after CRT + OPT, PVC persist



Suboptimal BIV pacing →
absence or limited response to CRT

Biventricular Pacing < 90%

Cardiac resynchronization therapy and the relationship of percent biventricular pacing to symptoms and survival

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36000 pts
2 yrs f.u.

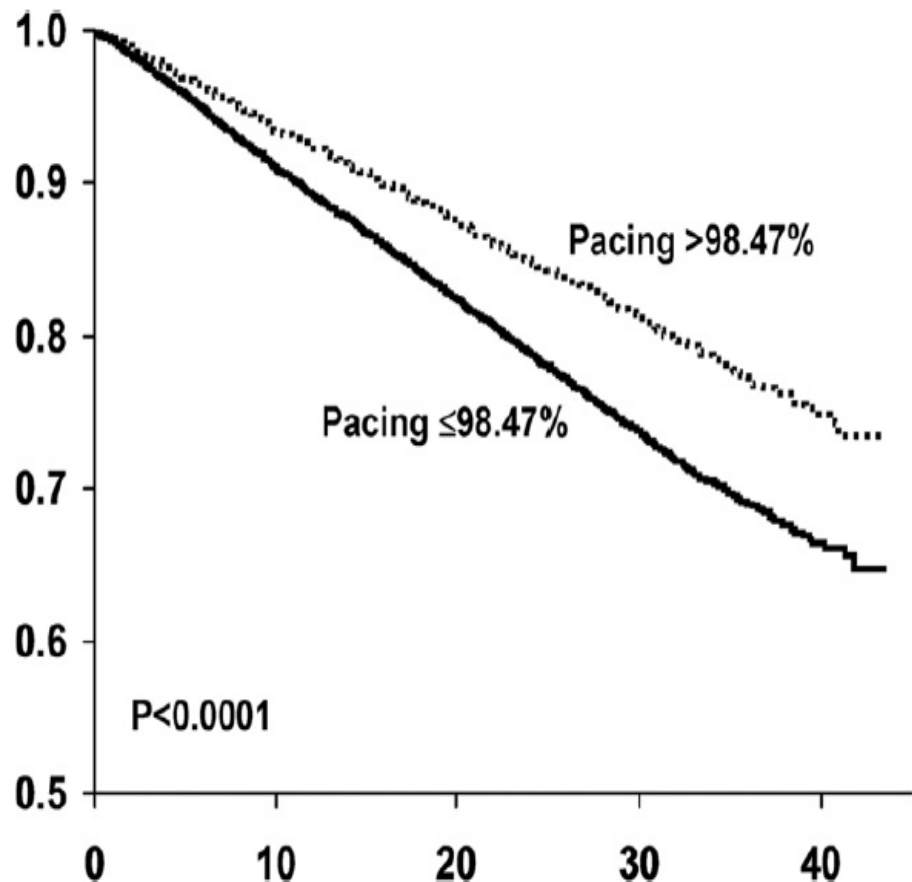
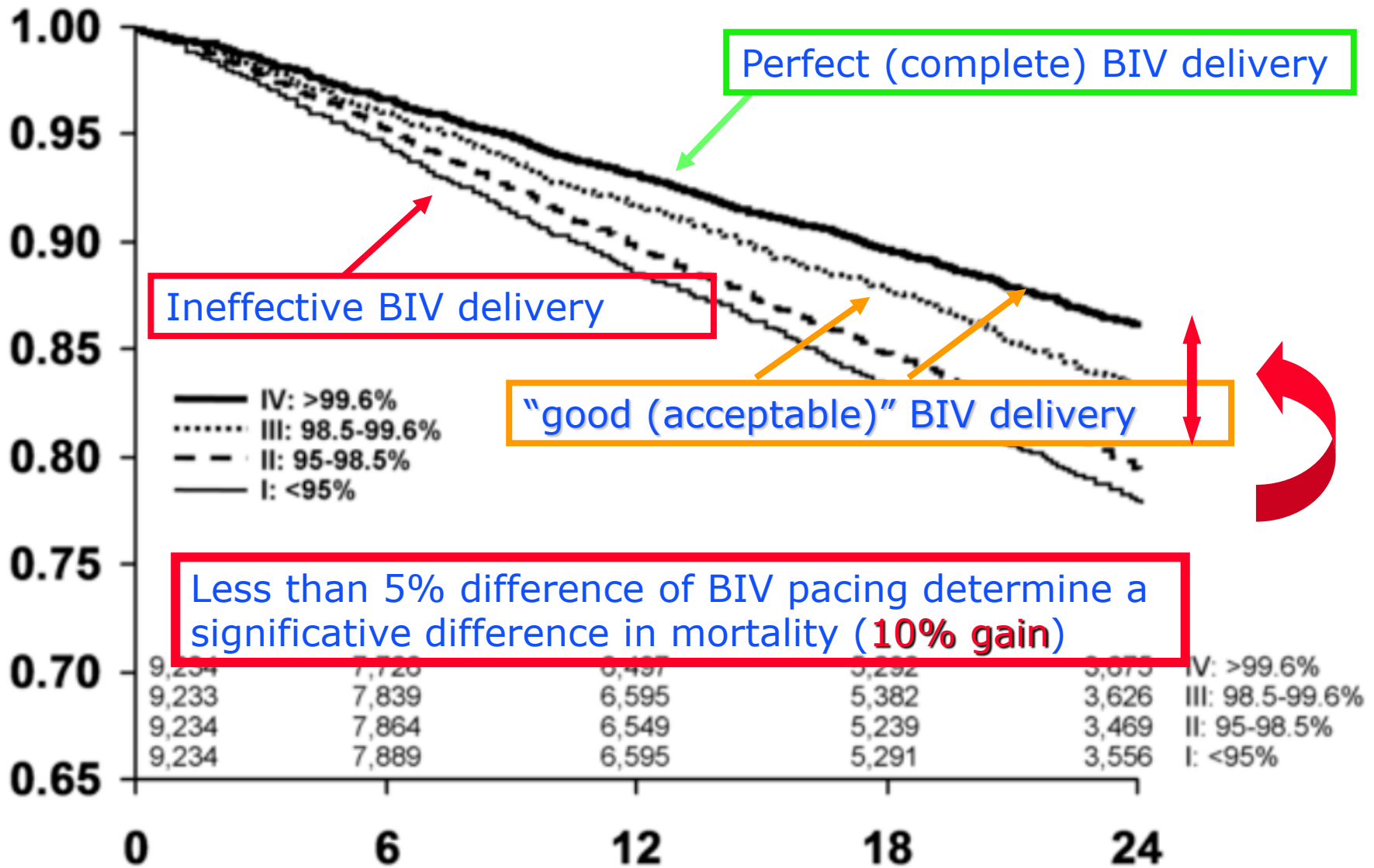
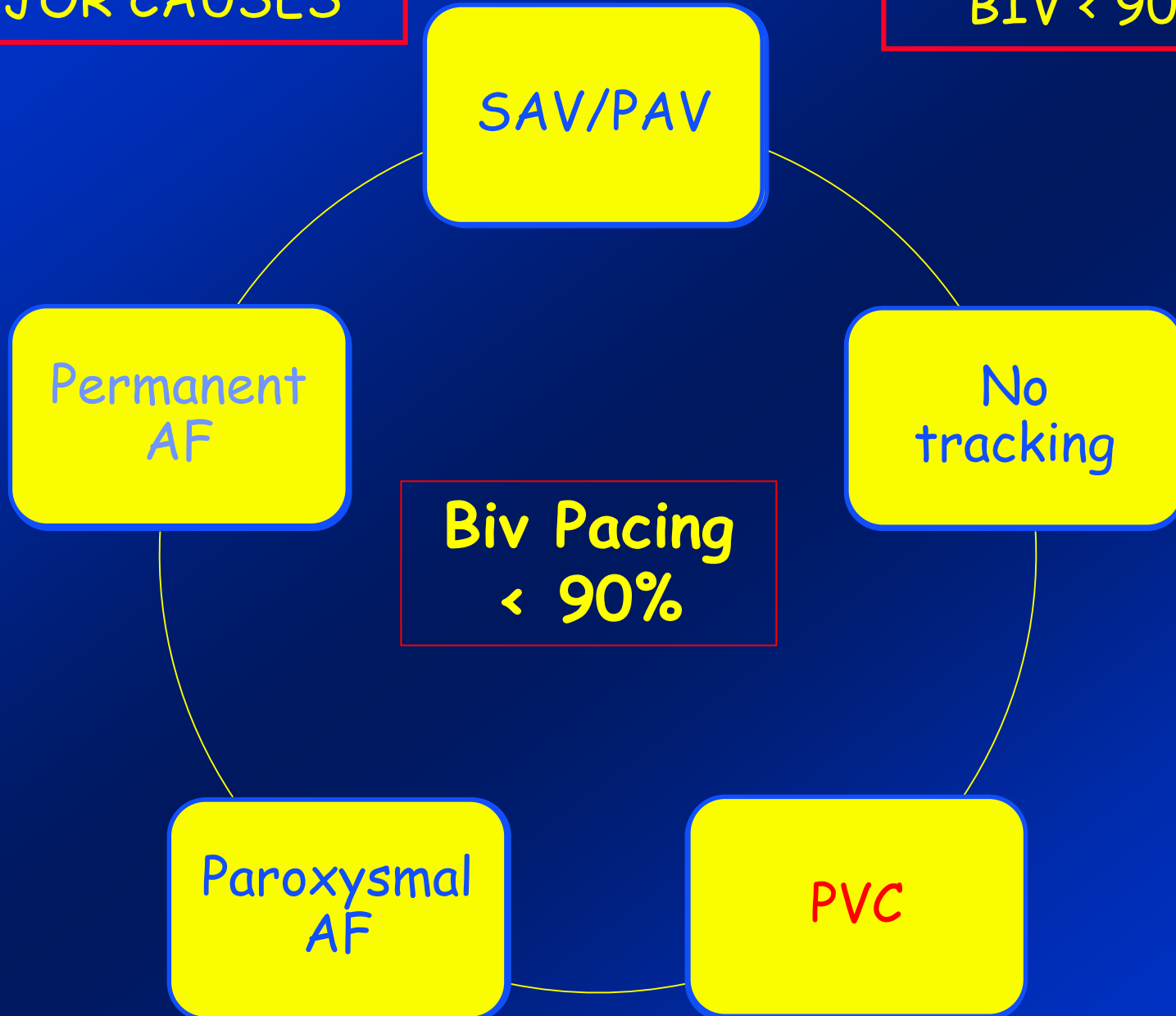


Figure 2 When assessed by quartiles, patients with biventricular pacing percentage above 99.6% experienced a 24% reduction in mortality compared with the other quartile groups (hazard ratio = 0.76, P < .001). Patients with <94.8% biventricular pacing had a 19% increase in mortality (hazard ratio = 1.19, P < .001).



5 MAJOR CAUSES

BIV < 90%



Biv Pacing
< 90%

Permanent
AF

No
tracking

PVC

Paroxysmal
AF

Vicious circle



Interaction of PVC & HF in CRT.



→ Loss of CRT

→ Heart failure worsening
→ More frequent hospitalizations

→ Inappropriate ICD shocks (?)

→ Increased sympathetic tone

→ Haemodynamic compromise

↑ PVCs
↑ Permanent AF



Association Between Frequency of Atrial and Ventricular Ectopic Beats and Biventricular Pacing Percentage and Outcomes in Patients With Cardiac Resynchronization Therapy

Martin H. Ruwald, MD, PhD,*† Suneet Mittal, MD,‡ Anne-Christine I James P. Daubert, MD,§ Scott McNitt, MS,* Amin Al-ahmad, MD,|| C Valentina Kutyifa, MD, PhD,* Jonathan S. Steinberg, MD,‡ Paul War Wojciech Zareba, MD, PhD*

Hot water
Discovery ...

From
MADIT CRT

TABLE 2 Association Between Number of Ectopic Beats (Ectopic Burden) and Likelihood of Low BIV Pacing (<97%) Versus High Percentage (≥97%)

	Odds Ratio: BIV Pacing <97% vs. BIV Pacing ≥97%	95% CI	p Value
Per percent increase in ectopic beats	1.18	1.14–1.23	<0.001
Per 0.1% increase in ectopic beats*	1.02	1.01–1.02	<0.001
<0.1% ectopic beats (<85 total APCs or PVCs)	1.00 = ref	NA	NA
0.1%–1.5% ectopic beats (85–1,347 total APCs or PVCs)	3.37	1.74–6.50	<0.001
>1.5% ectopic beats (>1,347 total APCs or PVCs)	13.42	7.02–25.66	<0.001
Separate analysis of APCs and PVCs			
Per percent increase in APCs/total beats	1.09	1.02–1.16	0.015
Per percent increase in PVCs/total beats	1.22	1.16–1.28	<0.001

Multivariate logistic regression model. Adjusted for age, left bundle branch block QRS configuration, prior ventricular arrhythmias, prior myocardial infarction, and creatinine level. *If estimated by use of an ordinal scale in the range from 0% to 1.5% by 0.1% increase and pooling patients with more than 1.5% ectopy: odds ratio: 1.16; 95% CI: 1.13 to 1.19; p < 0.001 per 0.1% increase in ectopic beats.

CI = confidence interval; NA = not applicable; ref = reference; other abbreviations as in Table 1.

PVCs detection

Quick Look II

Dispositivo: Maximo™ II CRT-D D284TRK

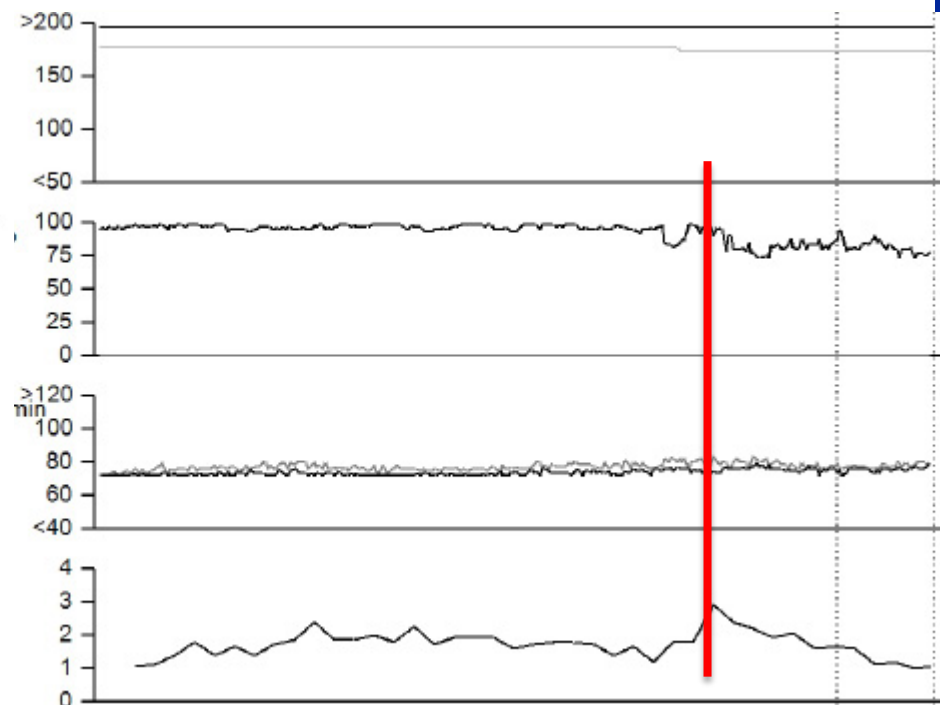
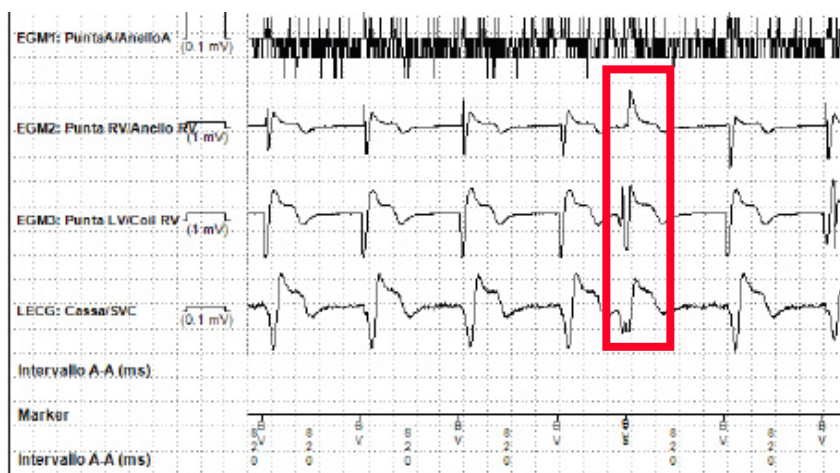
Numero di serie: PZP610776S

Data interrogazione: 18-Dec-2012 10:28:40

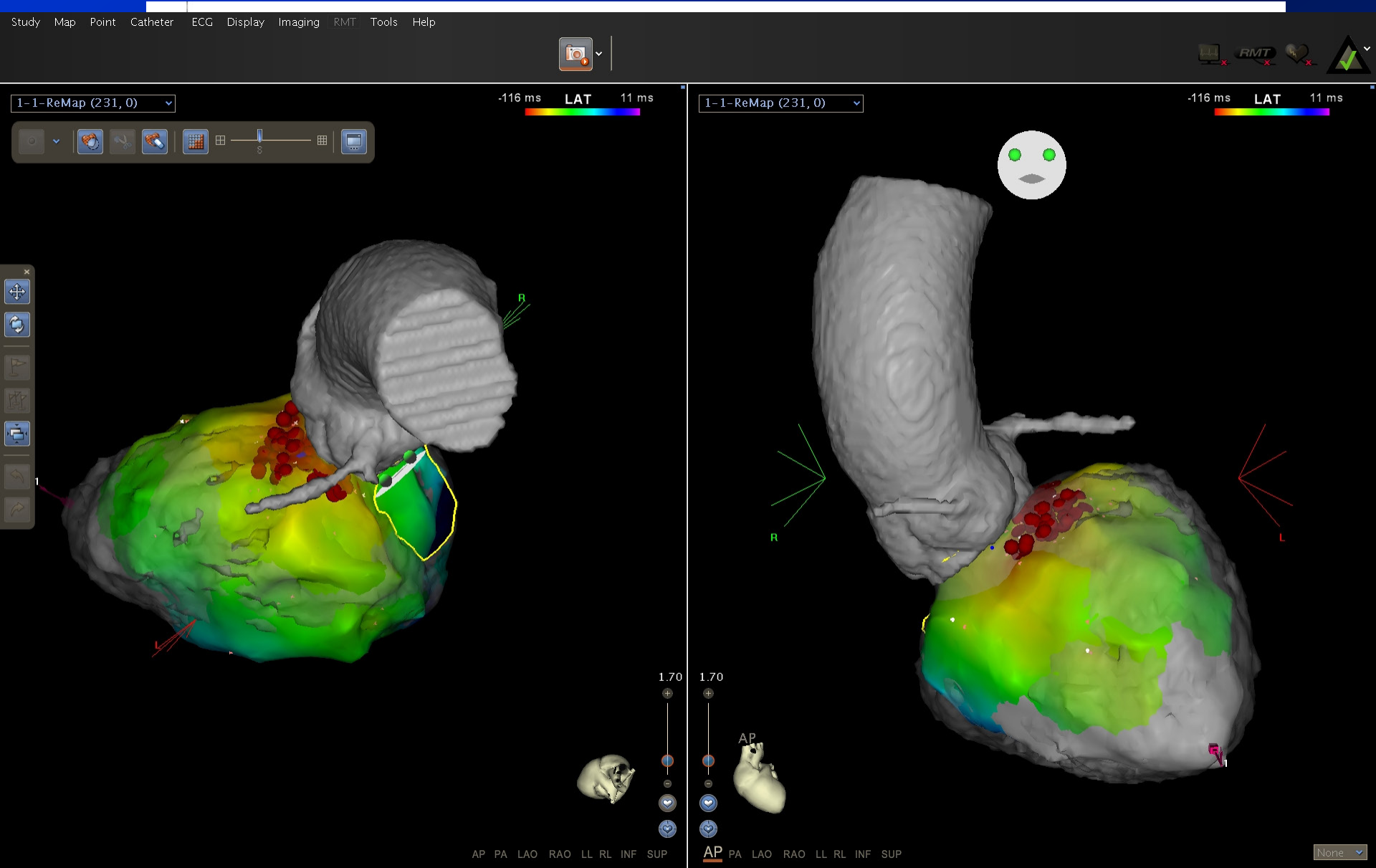
Medico: drGasparini Maurizio
0282244401

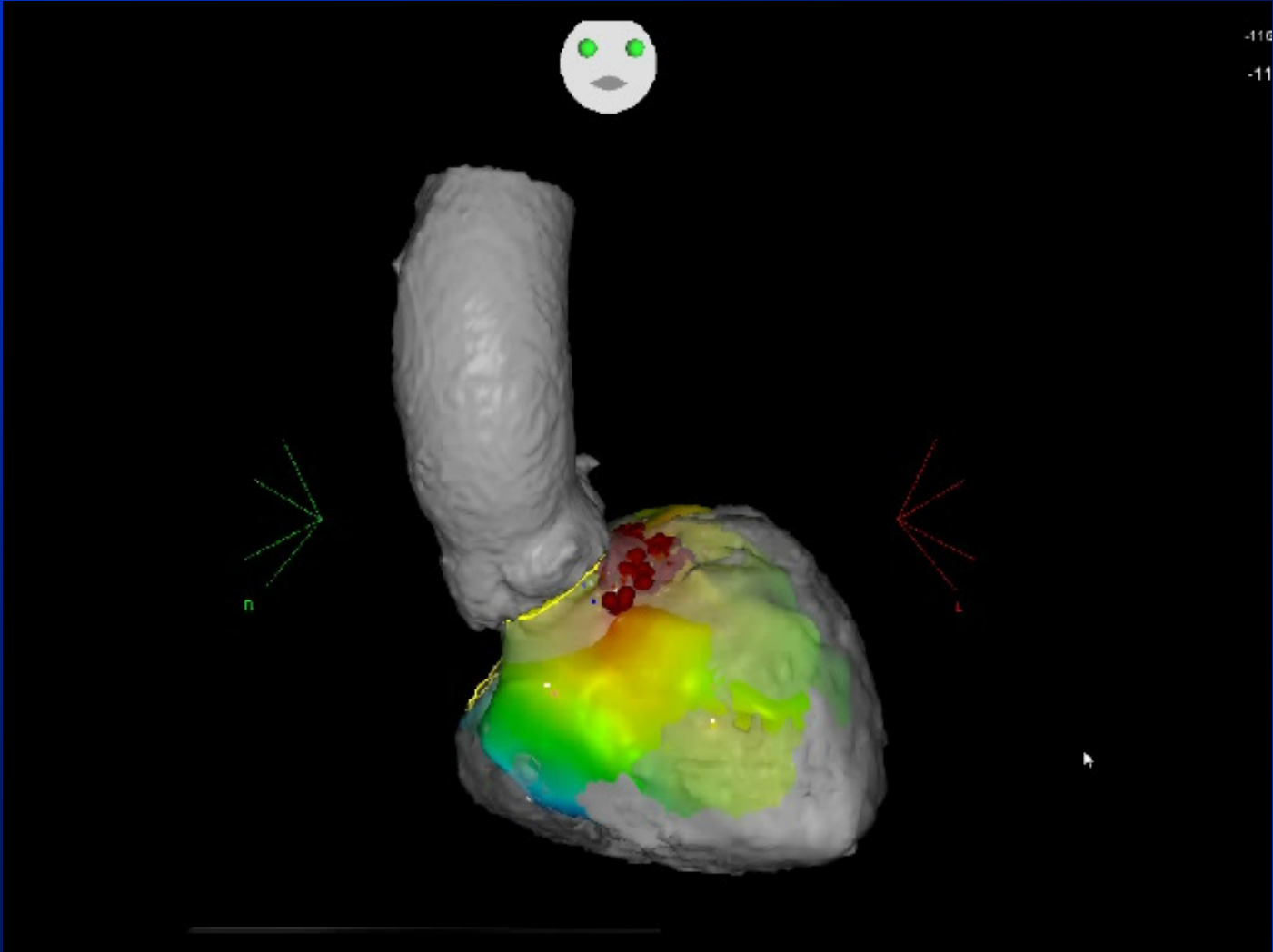
OSSERVAZIONI (1)

- Stimolazione V. (battiti V.) inferiore al 90%.



Ventricular extrasbeats





13210



After PVC ablation

Contatori Brady/CHF da ultimo azzeramento (27 Mar 2014)

- Atriale
- Ventricol. destra
- Ventricol. sinistra

1% stimolato
99% stimolato
100% stimolato

Clinical message



- CRT effective *only* when BIV pacing > 98%
- Consider arrhythmic substrate ablation (PVC ablation) if necessary.

Radiofrequency Ablation of Premature Ventricular Ectopy Improves the Efficacy of Cardiac Resynchronization Therapy in Nonresponders

Dhanunjaya Lakkireddy, MD,* Luigi Di Biase, MD, PhD,†‡§ Kay Ryschon, MS,* Mazda Biria, MD,* Vijay Swarup, MD,|| Yeruva Madhu Reddy, MD,* Atul Verma, MD,¶|| Sudharani Bommana, MPHIL,* David Burkhardt, MD,† Raghuvver Dendi, MD,* Antonio Dello Russo, MD, PhD,# Michela Casella, MD, PhD,# Corrado Carbucicchio, MD,# Claudio Tondo, MD, PhD,# Buddhadeb Dawn, MD,* Andrea Natale, MD†‡

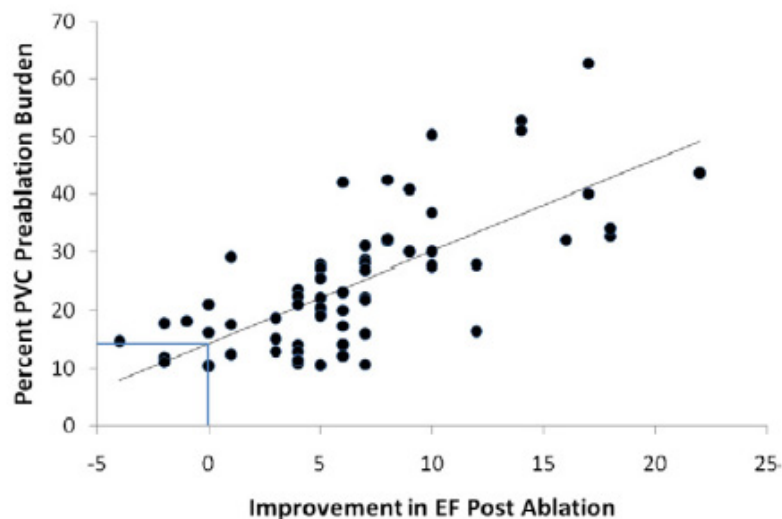


Figure 2

Correlation Between PVC Burden and EF Change Following Ablation

Percentage of pre-ablation premature ventricular contraction (PVC) burden and correlation with change in post-ablation ejection fraction (EF). The figure shows a Pearson correlation coefficient of 0.699, which is statistically significant at $p < 0.001$ (2-tailed). BIV = biventricular.

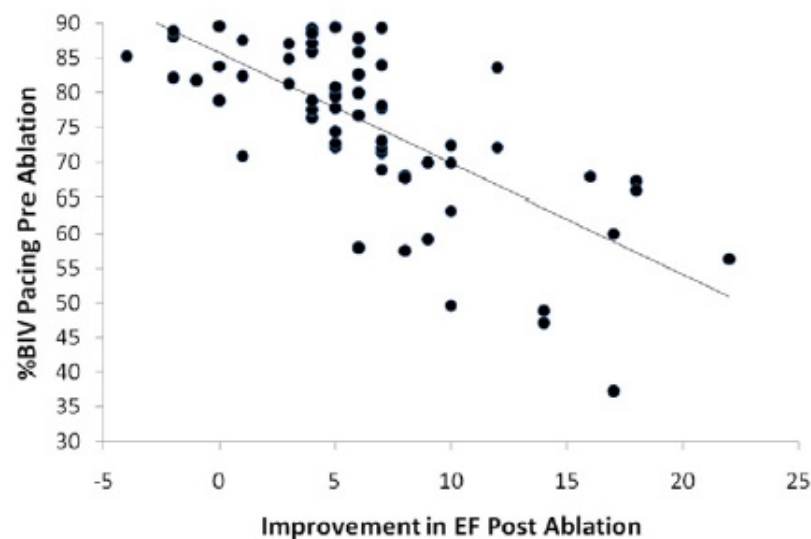


Figure 3

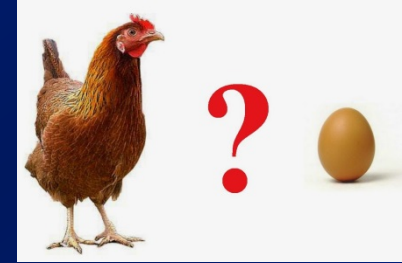
Correlation Between Pre-Ablation BIV Pacing % and Post-Ablation Improvement in EF

The figure shows a Pearson's correlation coefficient of 0.699 which is statistically significant at $p < 0.001$ (2-tailed). BIV = biventricular pacing; EF = ejection fraction.

Conclusions

Mandatory primum movens identification

PVC and HF: who comes first ?



PVC

HF Low EF

No heart disease

Structural heart disease

↓ First

↓ First

PVC RF ablation

CRT

(Class IA indication for RVOT) (no different indication in GL)

