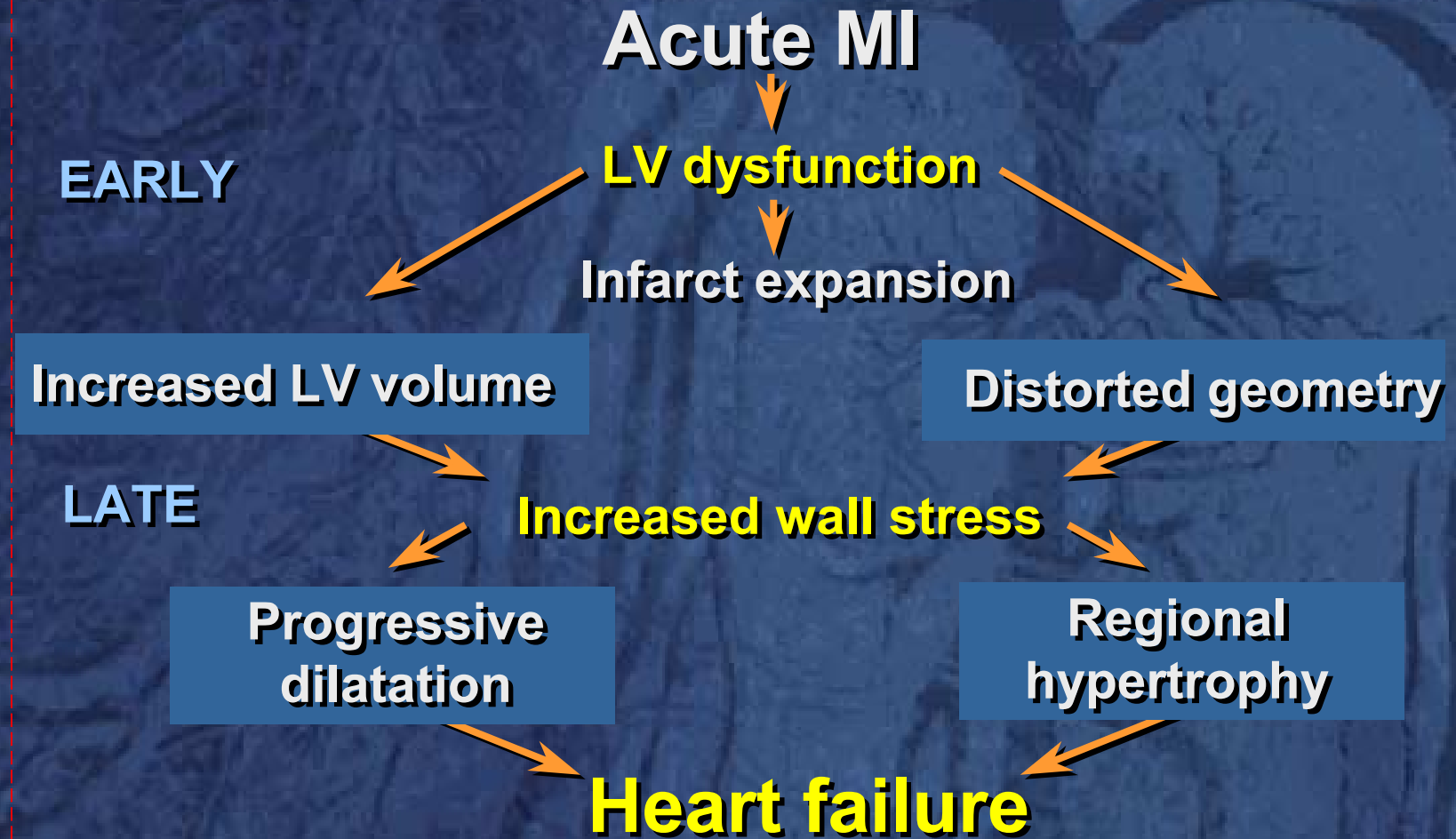


Element 24.00

Agati

The X's are ones that will be in workbook only. Keep the file intact now.

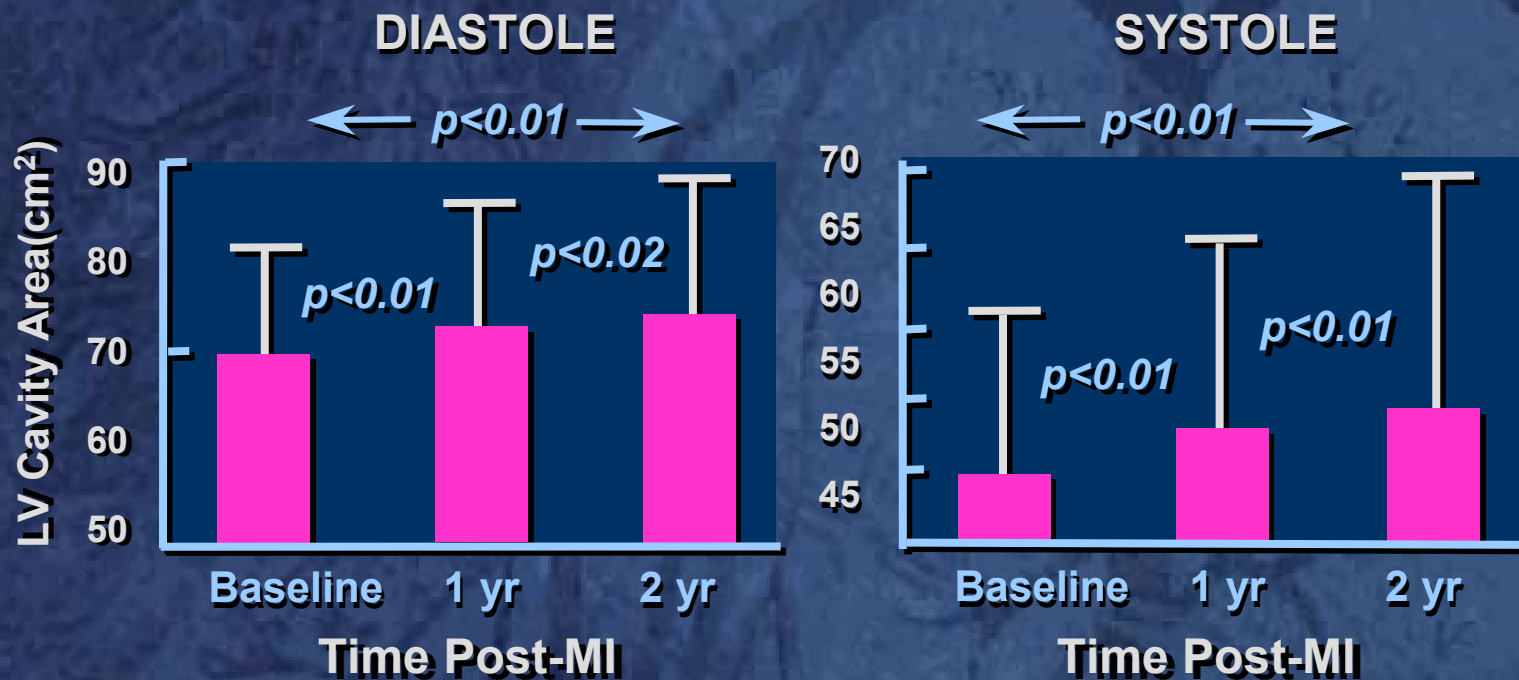
LV Remodeling after AMI



Timing of LV Dilatation after AMI

Kostuk	<i>Circ 1973</i>	96 pts	4 d - 1 mo	chest x ray
Erlebacher	<i>AJC 1982</i>	13 pts	10 d - 3-30 mo	echo
Mc Kay	<i>Circ 1986</i>	30 pts	10 h - 2 wk	angiography
Warren	<i>JACC 1988</i>	36 pts	7 h- 10 mo	radionuclide
Sharpe	<i>Lancet 1988</i>	70 pts	1 d- 1 yr	echo
Gadsball	<i>AJC 1989</i>	57 pts	2 wk - 1 yr	radionuclide
Picard	<i>Circ 1990</i>	57 pts	hrs - 3 mo	echo
Ertl	<i>AJC 1991</i>	78 pts	3 d - 6 ms	radionuclide
Mitchell	<i>JACC 1992</i>	52 pts	3 wk - 1 yr	angiography
Gaudron	<i>Circ 1993</i>	70 pts	4 d - 3 yr	radionuclide

LV Remodeling in SAVE Study



St. John Sutton et al., Circ 1997

Timing of LV Remodeling

- Early remodeling (in-hospital):
from 24-48 hrs to pre-discharge
- Late remodeling:
from pre-discharge to 6 mos



Magnitude of Remodeling

End-diastolic volume index changes

 $>20\%$ - severe dilation

$5.1 \leq \Delta \leq 20\%$ - mild/moderate dilation

$-5 \leq \Delta \leq 5\%$ - volume stability

$\leq -5\%$ - volume reduction



GISSI-3 Study design

Patients with AMI
within 24 hours



Randomization

Control

Nitrates

Lisinopril

Lisinopril
+ Nitrates



6 weeks evaluation
treatment withdrawal



6 months evaluation

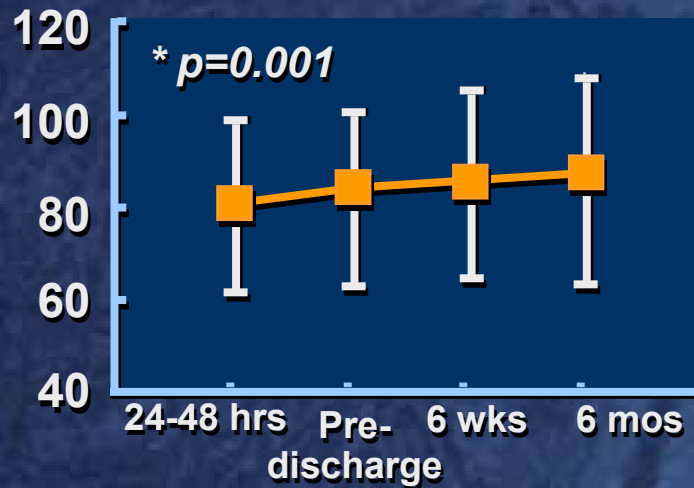


Study Population

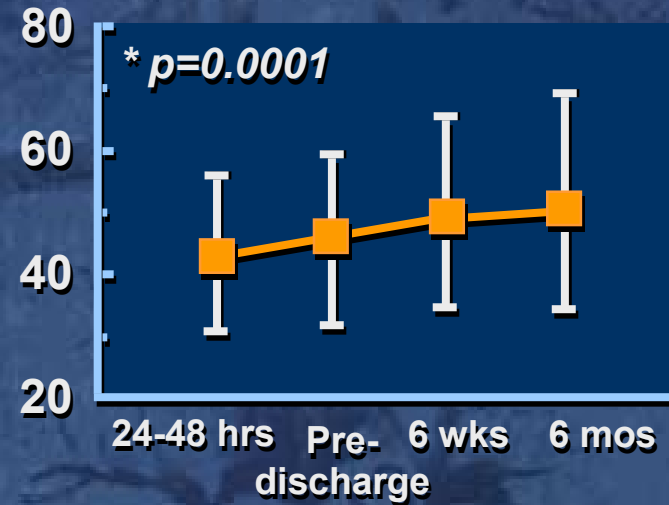
- **Subset of 614 patients enrolled in the GISSI 3-ECHO substudy with no cardiac events during 6-month follow-up**
- **Serial echo studies at 24-48 hrs, at hospital discharge, at 6 wks and at 6 mos after AMI**
- **Centralized analysis**



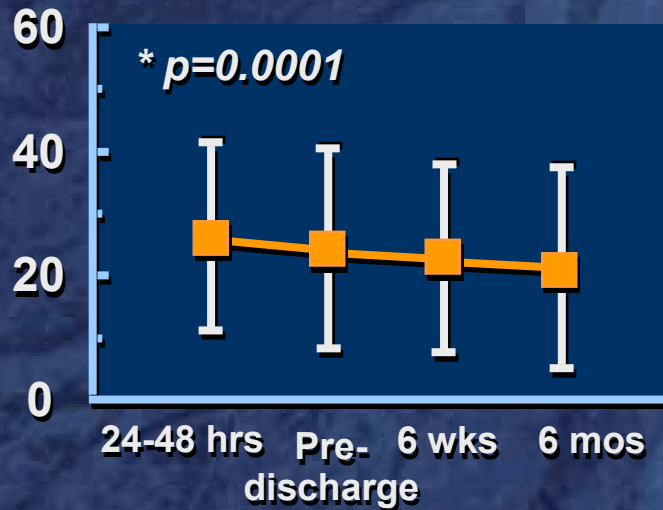
EDVi ml/m²



ESVi ml/m²

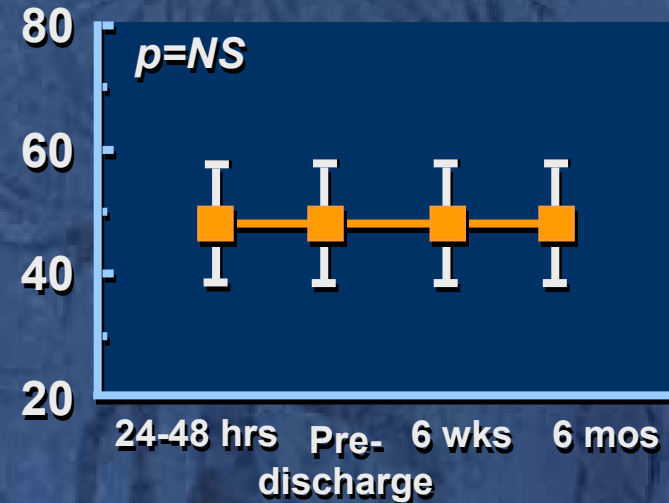


% WMA

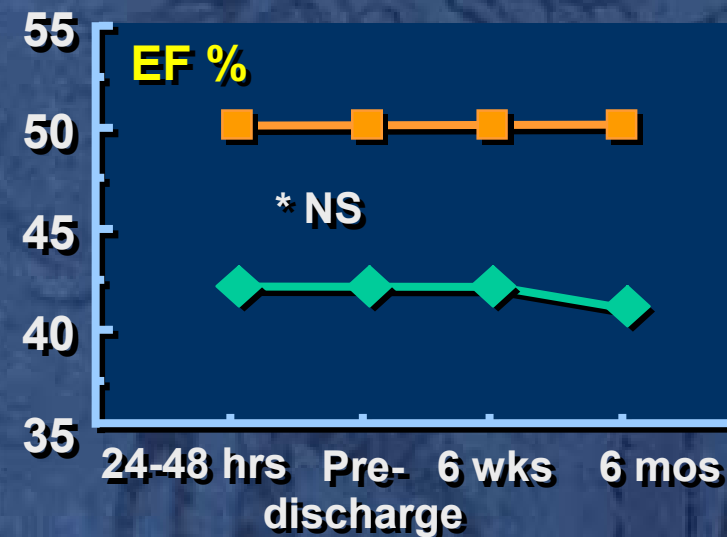
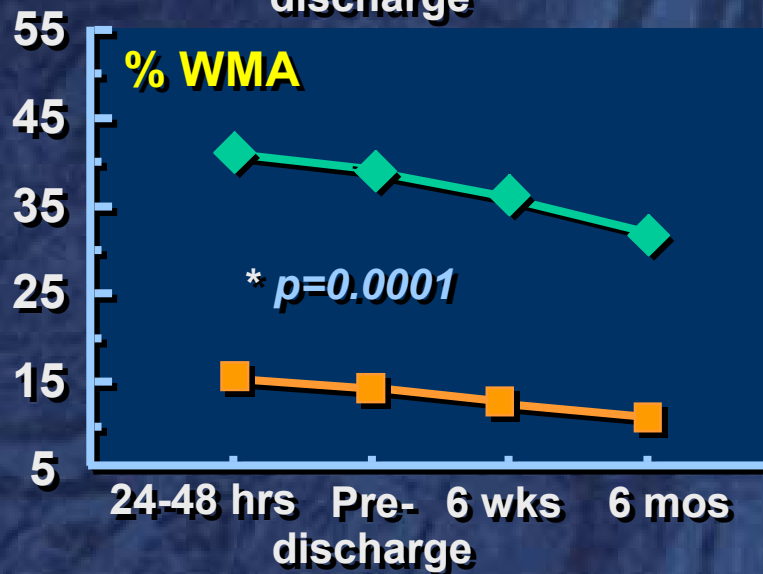
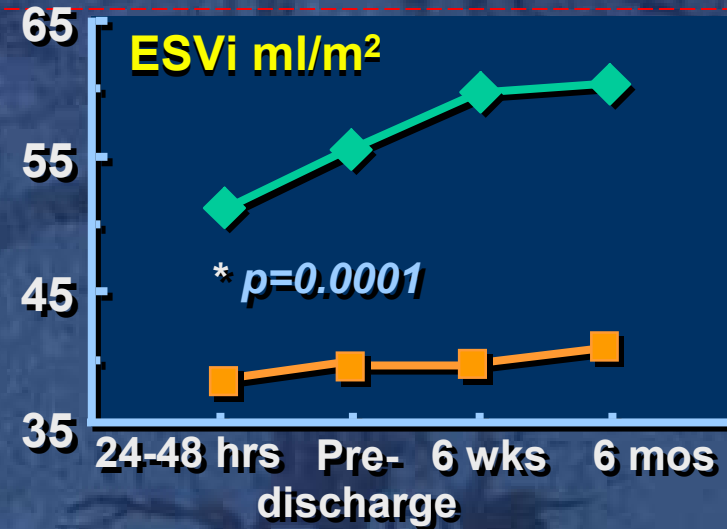
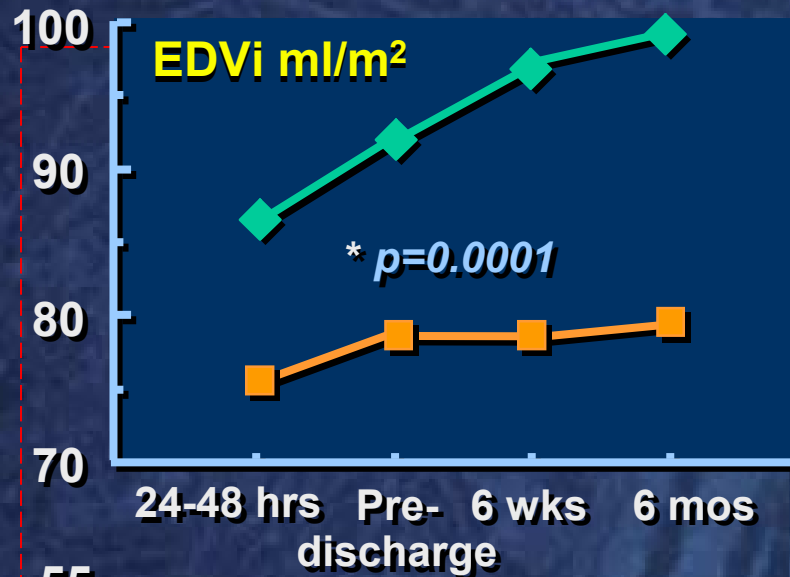


* time effect

EF %



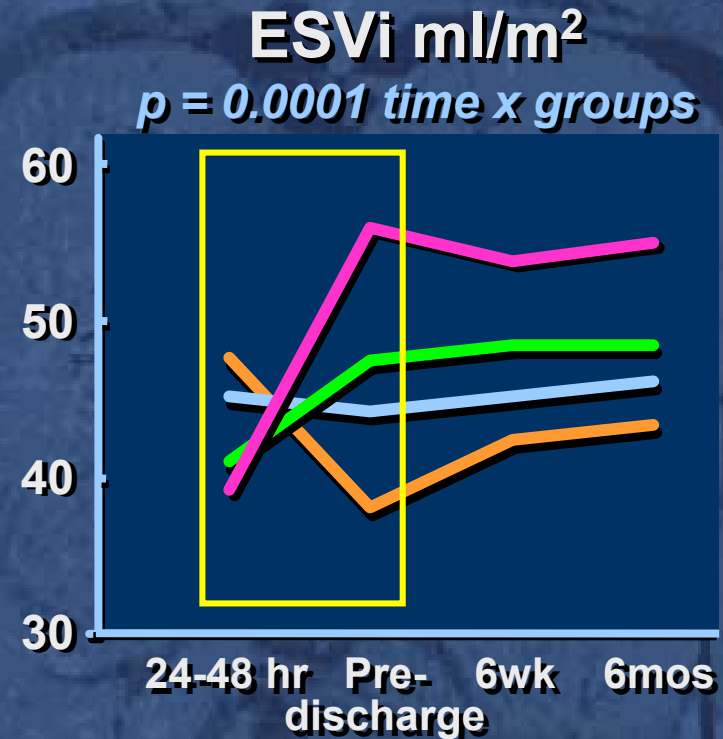
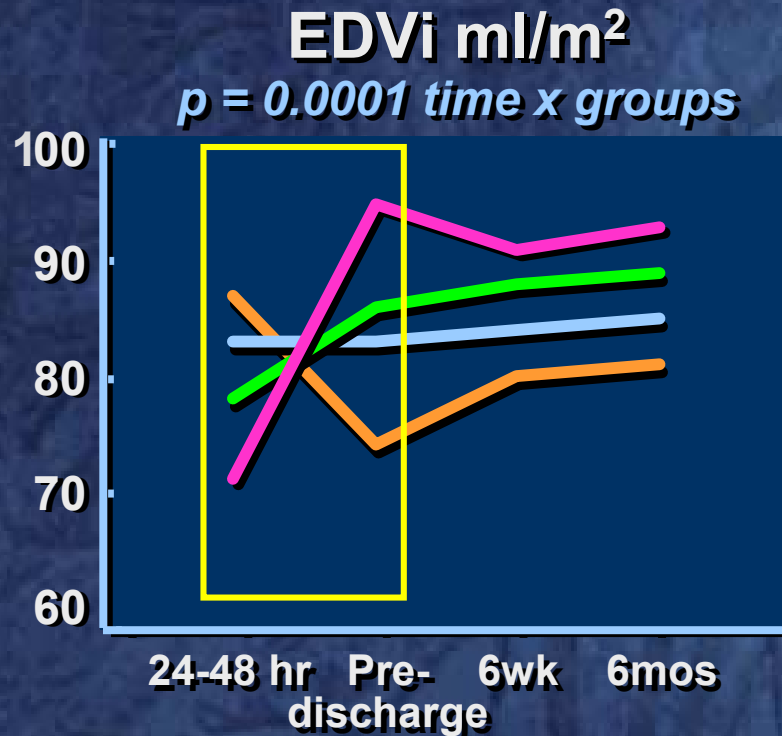
GISSI-3 Echo substudy. Am Heart J 2001; 141:131-8



■ %WMA <30% n. 407 ◆ %WMA ≥30% n. 280

GISSI-3 Echo substudy. Am Heart J 2001; 141:131-8

Early Remodeling

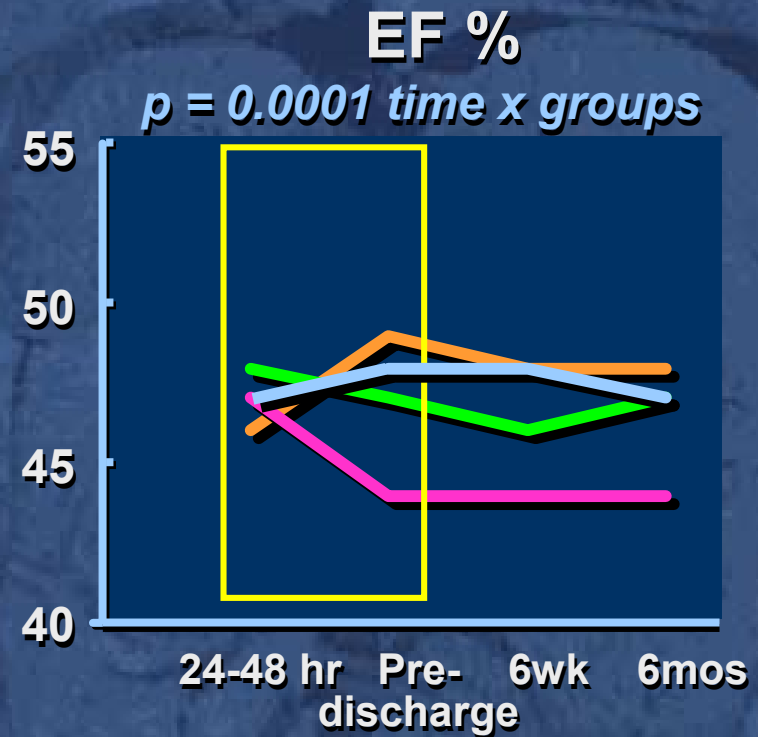
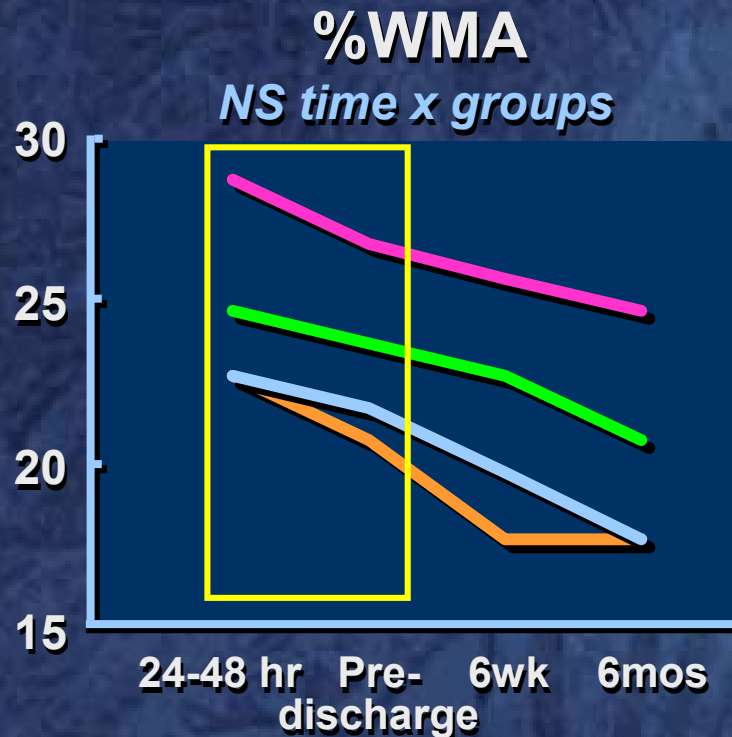


— volume reduction
— volume stability

— mild/moderate dilation
— severe dilation

GISSI-3 Echo substudy. Am Heart J 2001; 141:131-8

Early Remodeling



— volume reduction
— volume stability

— mild/moderate dilation
— severe dilation

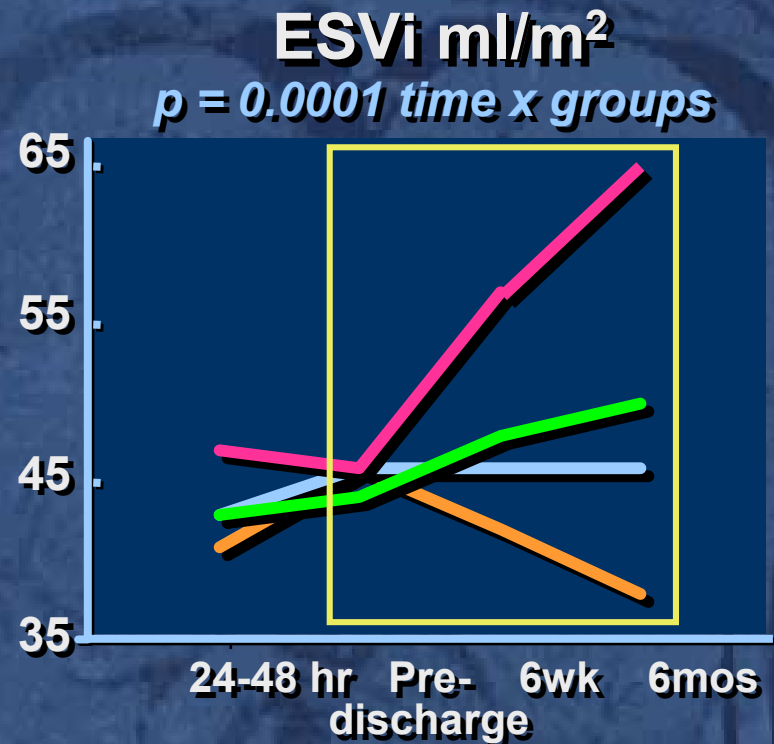
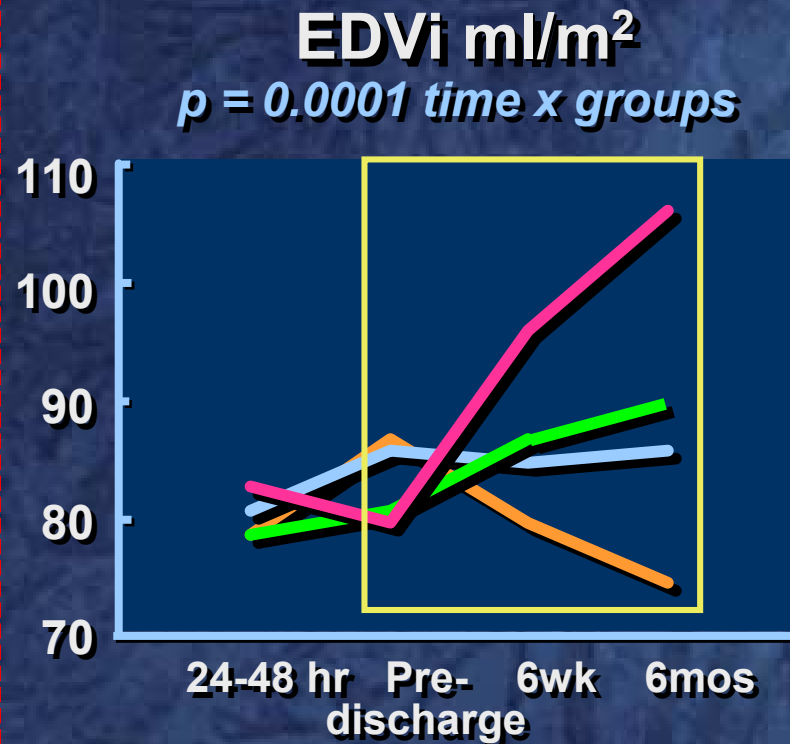
GISSI-3 Echo substudy. Am Heart J 2001; 141:131-8

Predictors of Early Remodeling

>20% LV dilation

	OR	CI 95%	p value
Baseline %WMA	1.030	1.013-1.048	0.0005
<i>Mitral reg. 3-4+ vs 1-2+</i>	1.739	0.694 -4.348	0.2365
<i>Anterior vs non-anterior MI</i>	1.399	0.861 -2.272	0.1752
<i>Thrombolysis Yes vs No</i>	1.275	0.793 -2.050	0.3165
<i>Nitrate Yes vs No</i>	1.233	0.814 -1.868	0.3234
<i>Lisinopril Yes vs No</i>	0.900	0.592 -1.368	0.6220
<i>Killip 2-3 vs Killip 1</i>	1.119	0.593 -2.110	0.7285
<i>Female vs Male</i>	0.921	0.520 -1.632	0.7773
<i>β-blockers i.v. no vs yes</i>	1.067	0.660 -1.726	0.7908

Late Remodeling

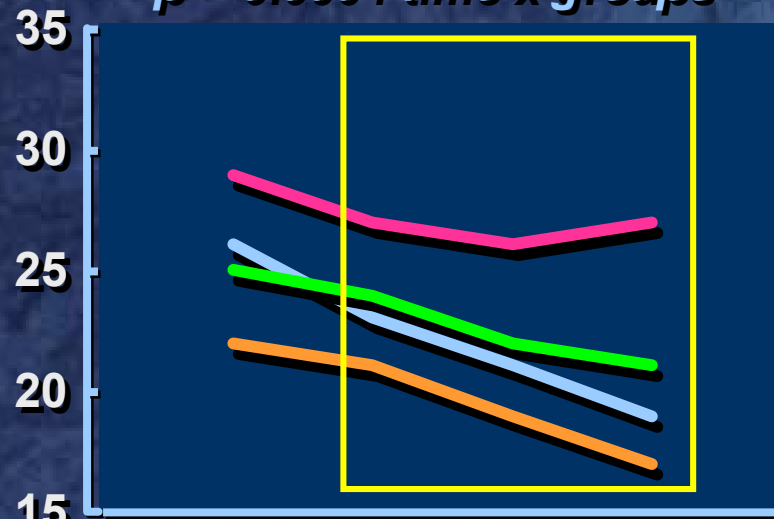


- volume reduction
- volume stability
- mild/moderate dilation
- severe dilation

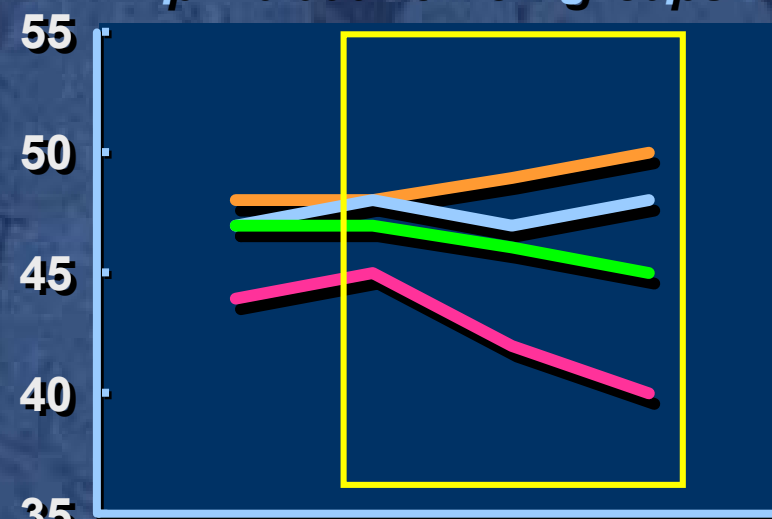
GISSI-3 Echo substudy. Am Heart J 2001; 141:131-8

Late Remodeling

%WMA
p = 0.0004 time x groups



EF %
p = 0.0001 time x groups



— volume reduction
— volume stability

— mild/moderate dilation
— severe dilation

GISSI-3 Echo substudy. Am Heart J 2001; 141:131-8

Predictors of Late Remodeling

>20% LV dilation

	OR	CI 95%	p value
Pre-discharge %WMA	1.026	1.008-1.045	0.0147
<i>Mitral reg. 3-4+ vs 1-2+</i>	2.261	1.031-4.958	0.0417
<i>Lisinopril no vs yes</i>	0.649	0.417-1.009	0.0546
<i>Age</i>	0.990	0.969-1.011	0.3322
<i>β-blockers i.v. no vs yes</i>	0.857	0.523-1.404	0.5396
<i>Thrombolysis no vs yes</i>	0.886	0.523-1.501	0.6515
<i>Nitrate no vs yes</i>	0.914	0.588-1.420	0.6887
<i>Undefined vs non anterior AMI</i>	1.146	0.537-2.445	0.7248

Remodeling

- Occurs after AMI *despite spontaneous recovery of regional function*
- With recovery of regional function, particularly evident in large infarcts

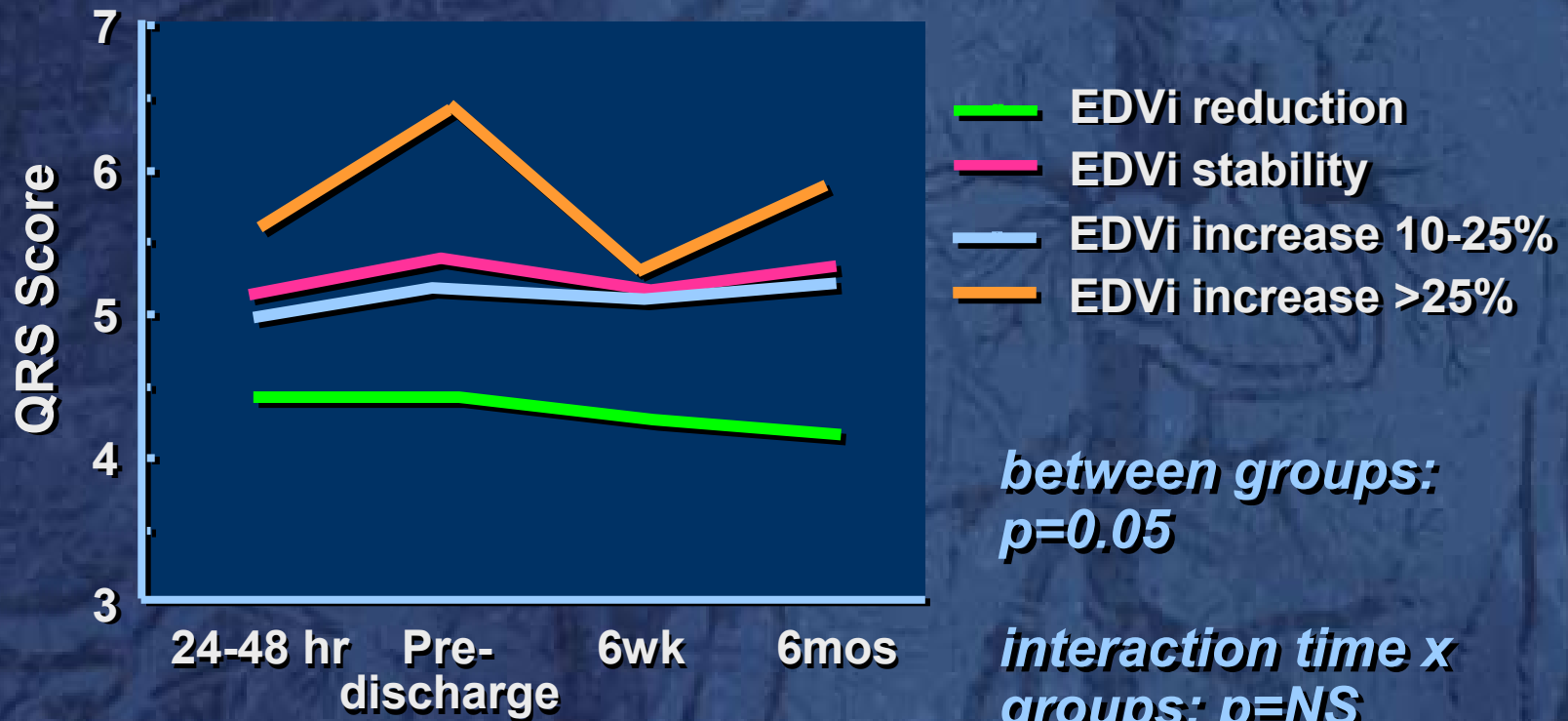
Remodeling Time

- **Early *is not predictive of subsequent dilation/dysfunction***
- **Late *associated with significant worsening of LV function***
- **Anytime *more extensive %WMA indicates higher risk for progressive LV dilation***



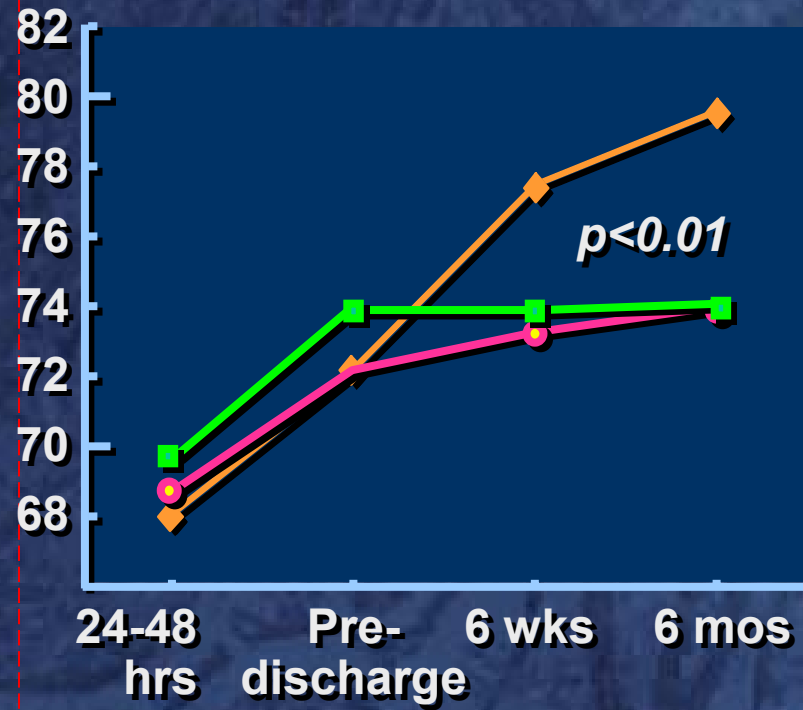
LV remodeling

QRS score changes

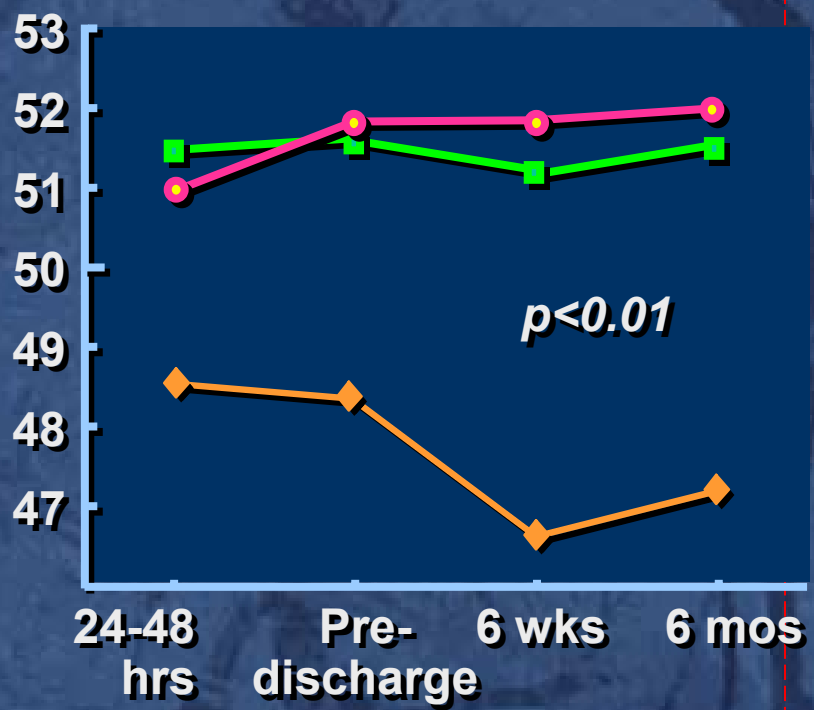


X LV remodeling according to the evolutionary changes of the negative T waves

EDVi (ml/m²)



EF (%)



● Negative T waves Reduction ■ Negative T waves No change ◆ Negative T waves Increase



Electrocardiographic Evolutionary Changes and Left Ventricular Remodeling After Acute Myocardial Infarction Results of the GISSI-3 Echo Substudy

Enzo Bosimini, MD,* Pantaleo Giannuzzi, MD,* Pier L. Temporelli, MD,* Francesco Gentile, MD,†
Donata Lucci, BS,‡ Aldo P. Maggioni, MD,‡ Luigi Tavazzi, MD,§ Luigi Badano, MD,||
Ioanna Stoian, MD,¶ Rita Piazza, MD,# Ioanna Heyman, MD,** Giacomo Levantesi, MD,††
Eugenio Cervesato, PHD,¶ Enrico Geraci, MD,‡‡ Gian L. Nicolosi, MD,¶
for the GISSI-3 Echo Substudy Investigators

JACC 2000;35:127-35

“The analysis of serial ECGs can predict postinfarct LV remodeling. Lack of resolution and late appearance of new negative T waves predict unfavorable remodeling with progressive deterioration of ventricular function”

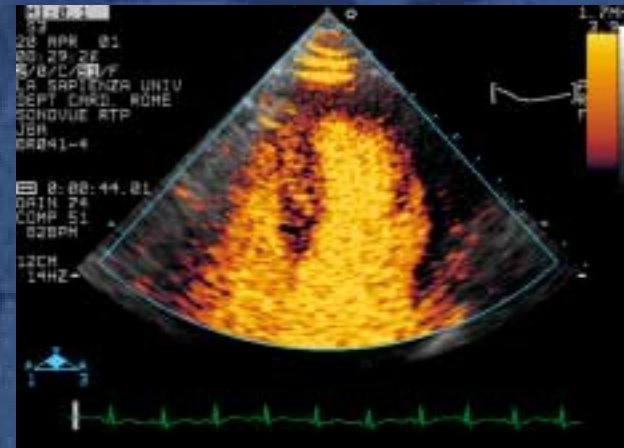
Determinants of LV Remodeling

Open Issues

- **Length of infarct segment**
- **IRA TIMI grade**
- **Role of diastolic filling pattern**
- **Role of microvascular damage**
(to be defined)

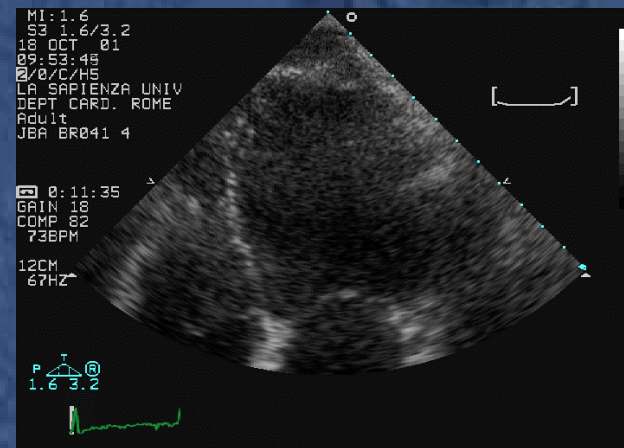
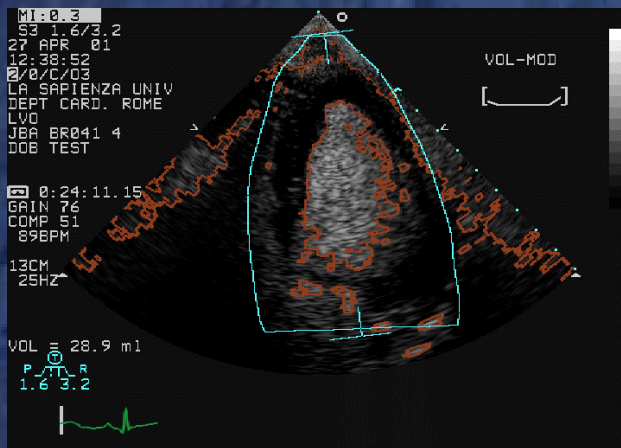
Influence of No-reflow on LV Remodeling

Day 1



Day 1

Day 7



F/U

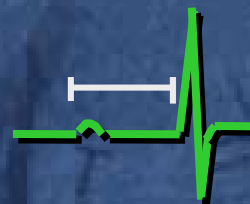


Mitral Doppler Flow Velocity Profile

Abnormal relaxation

Restrictive filling

Ecg

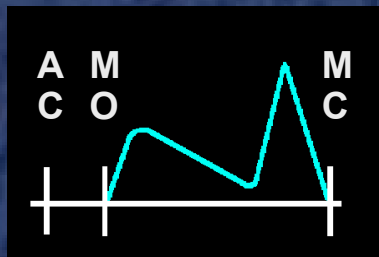


Phono

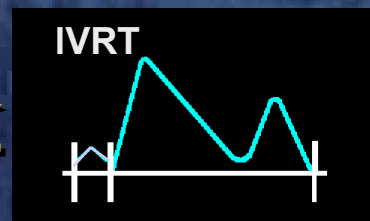


A1

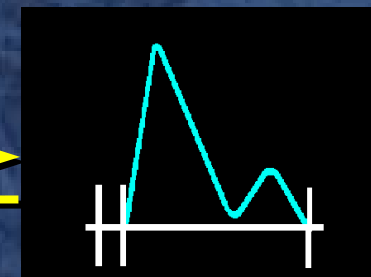
S1



Pattern I



Normal(ized)



Pattern II

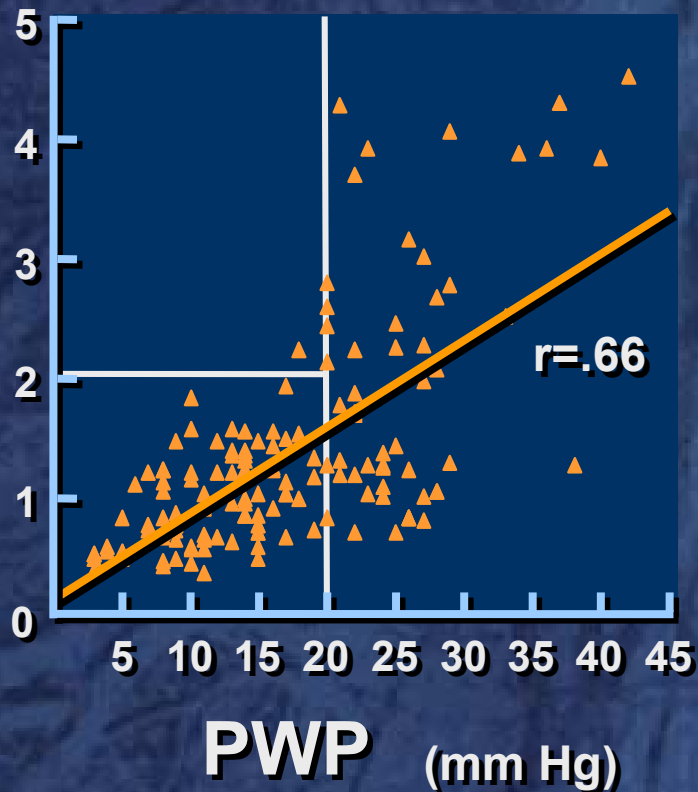
1 m/s

0

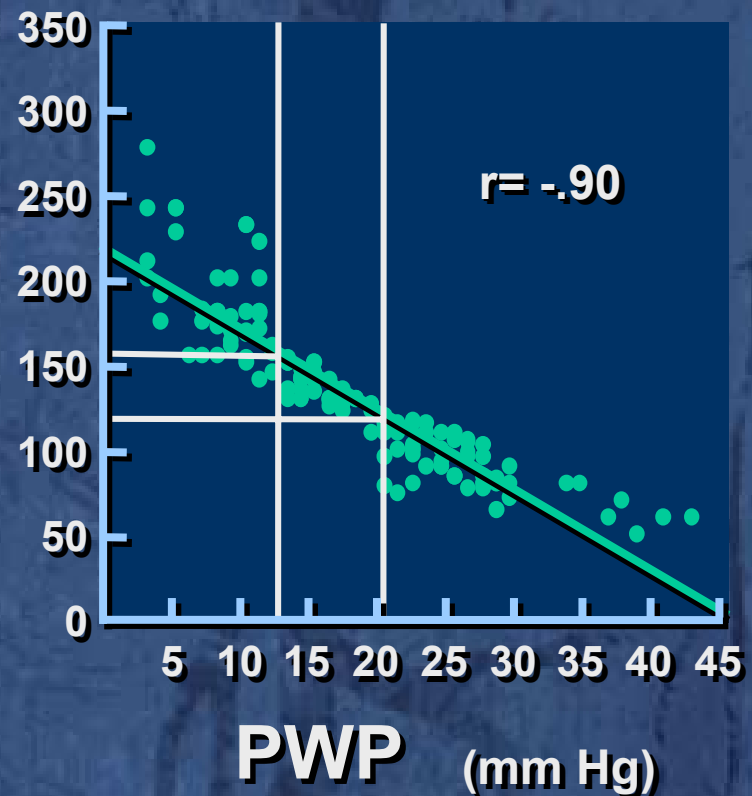


Doppler - hemodynamic correlations in 140 pts with AMI and severe LV dysfunction

E/A



DecT (ms)



Giannuzzi et al. J Am Coll Cardiol 1994



Deceleration time and LV Remodeling in the GISSI-3 echo substudy

- **571 patients with uncomplicated MI**
- **Serial Doppler echocardiographic study**

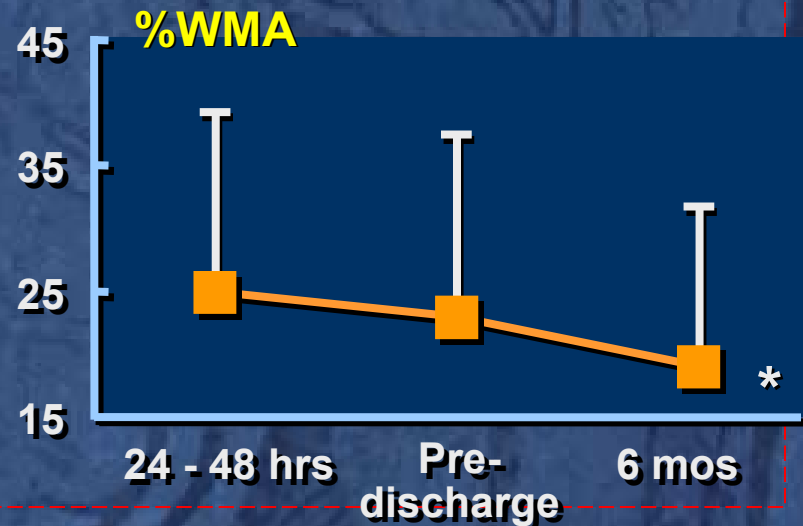
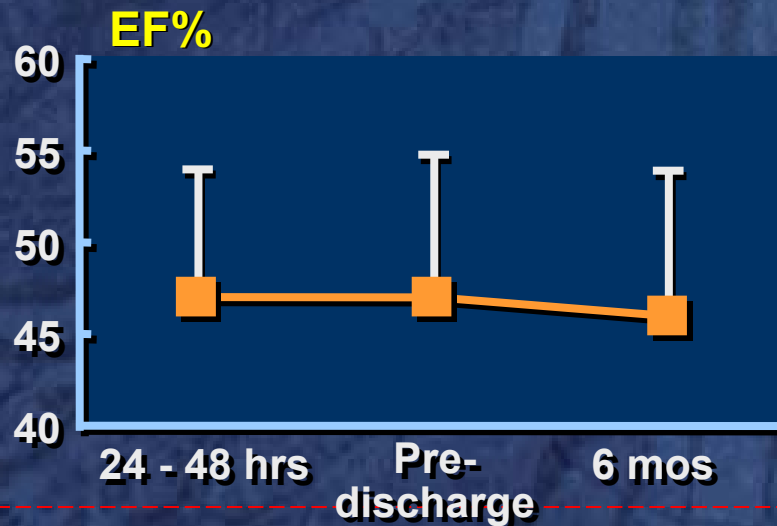
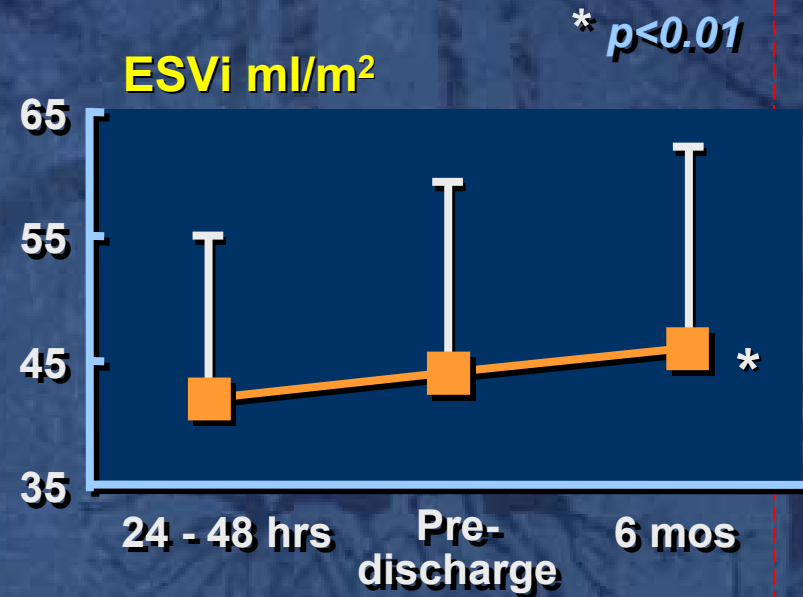
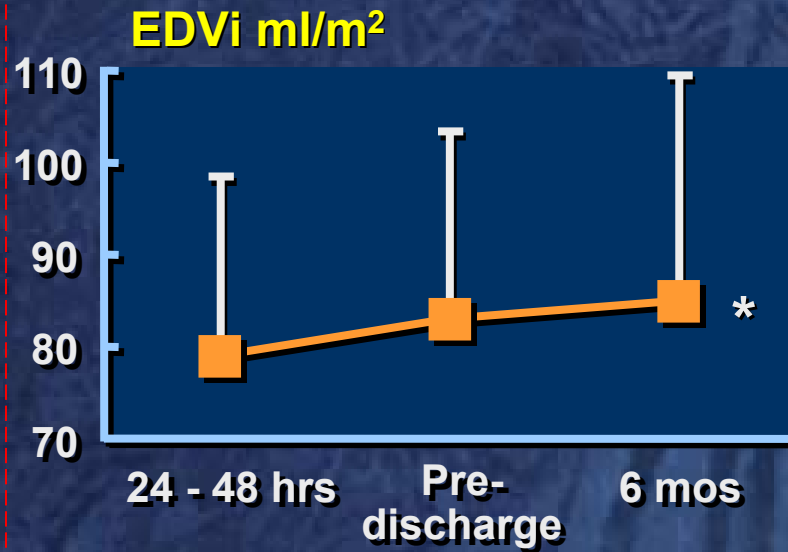
*at 24-48 hrs (mean 36 ± 8),
pre-discharge (mean 12 ± 5 days)
6 months (mean 194 ± 16 days) after*

Changes in Doppler Variables From Baseline to 6 months

- Peak E from 64.8 - 65.1 cm/sec
- Peak A from 64.2 - 64.9 cm/sec
- E/A from 1.009 - 1.002
- DT from 158 - 177 msec* ($p < 0.01$)

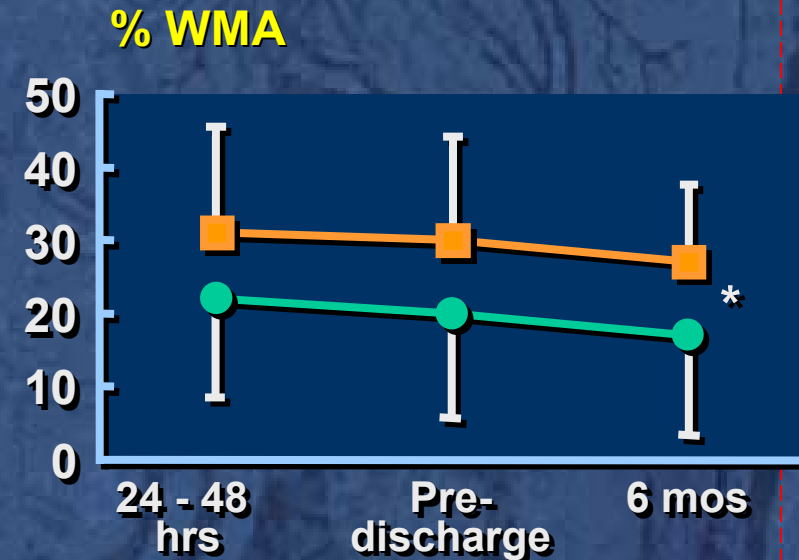
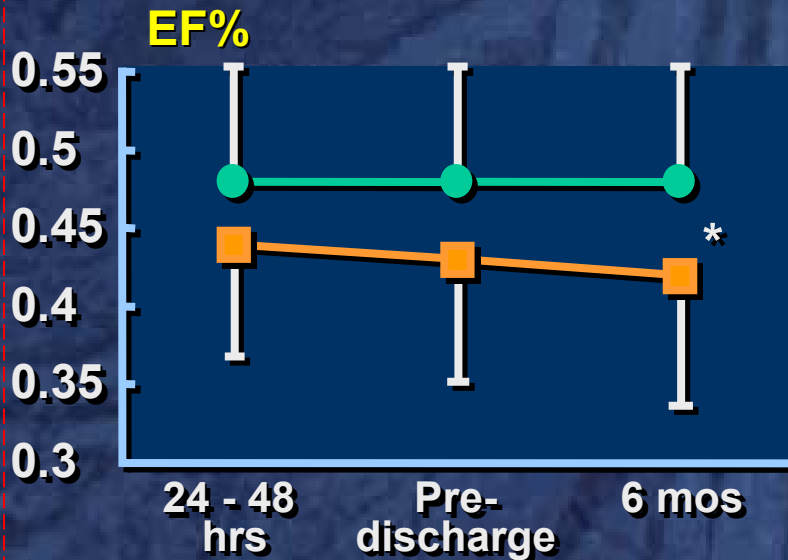
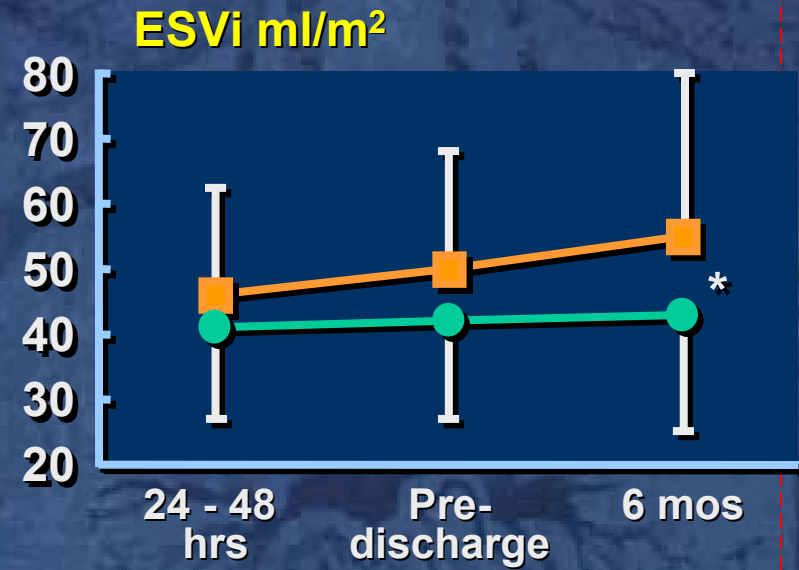
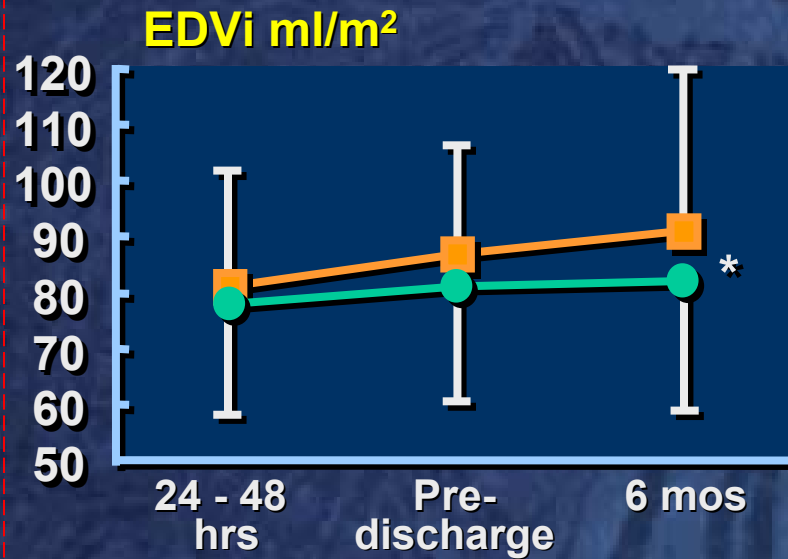


Total Population





Changes According to Baseline DT



* $p < 0.001$ interaction time x groups

■ Baseline DT <130 n= 147
● Baseline DT >130 n= 424



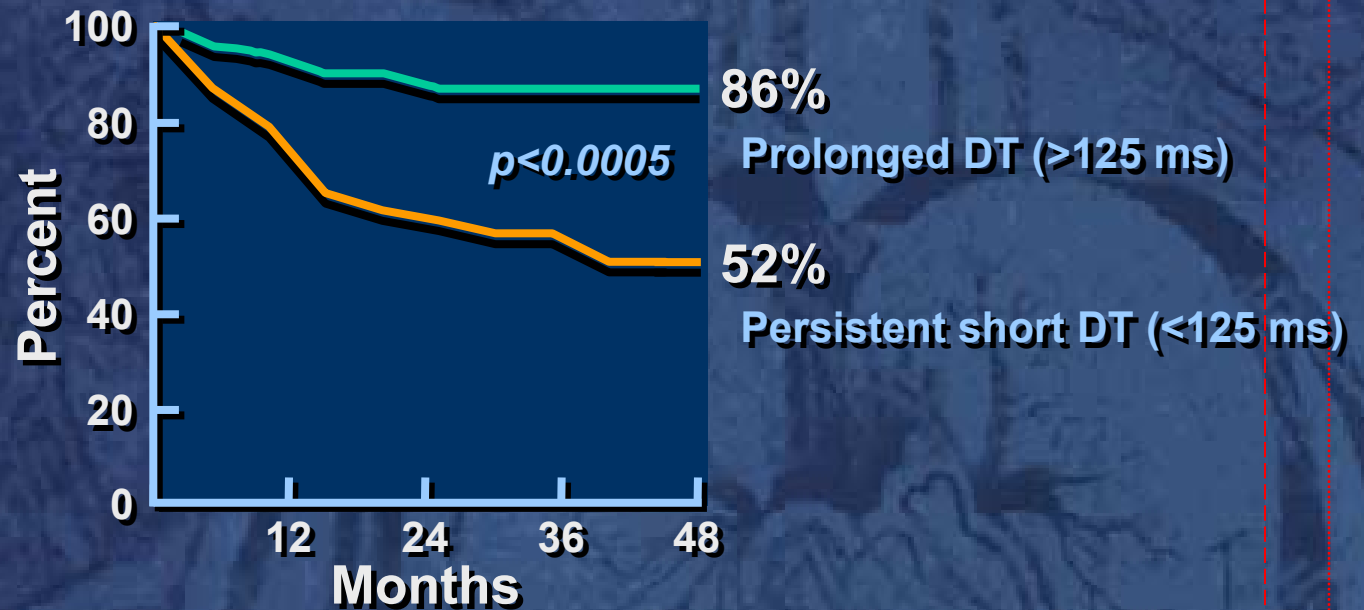
Independent Predictors of Dilation ($>5\%$)

Variable	Odds ratio	p value
Baseline EDVi	0.96	0.0001
Baseline DT <130 ms	2.31	0.0001
Baseline %WMA	1.02	0.0001

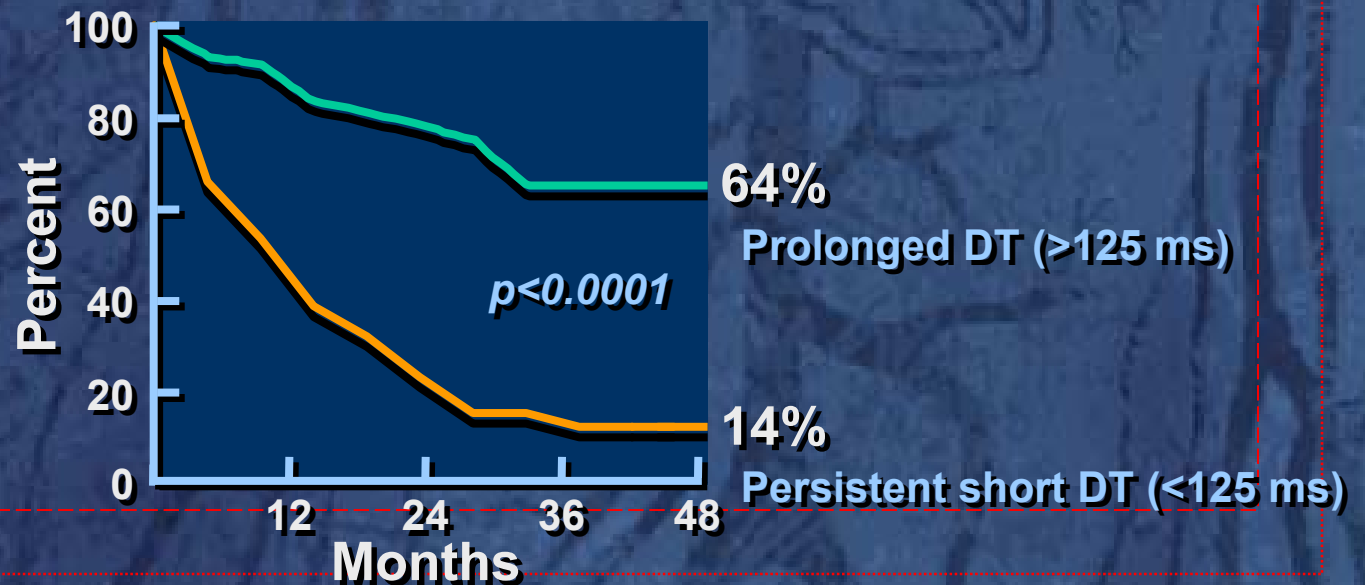
Temporelli et al. Submitted

X Restrictive Diastolic Filling in CHF

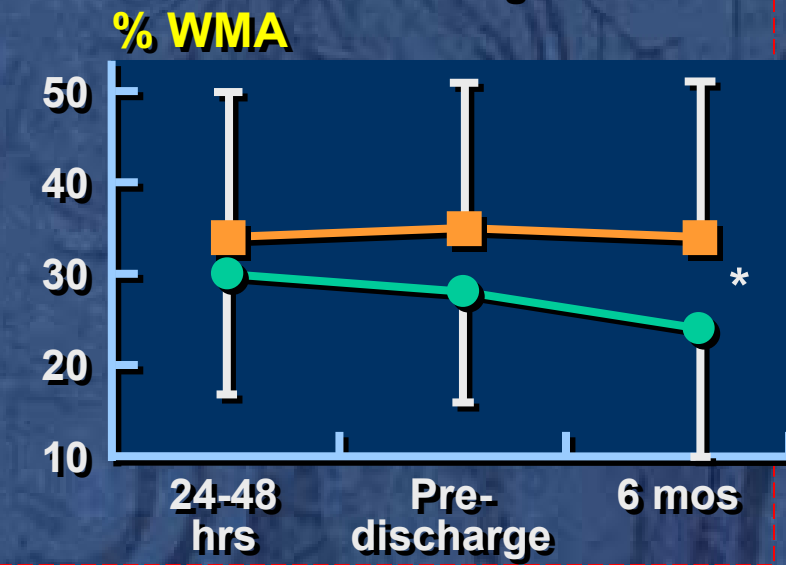
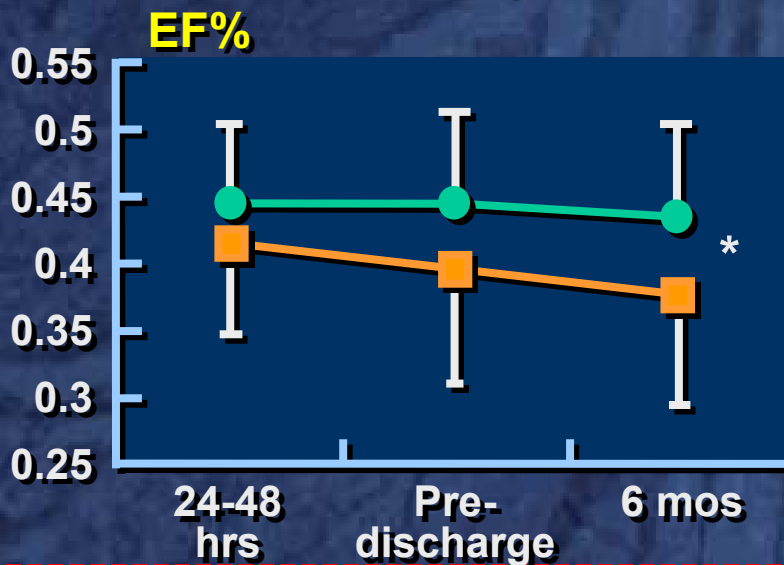
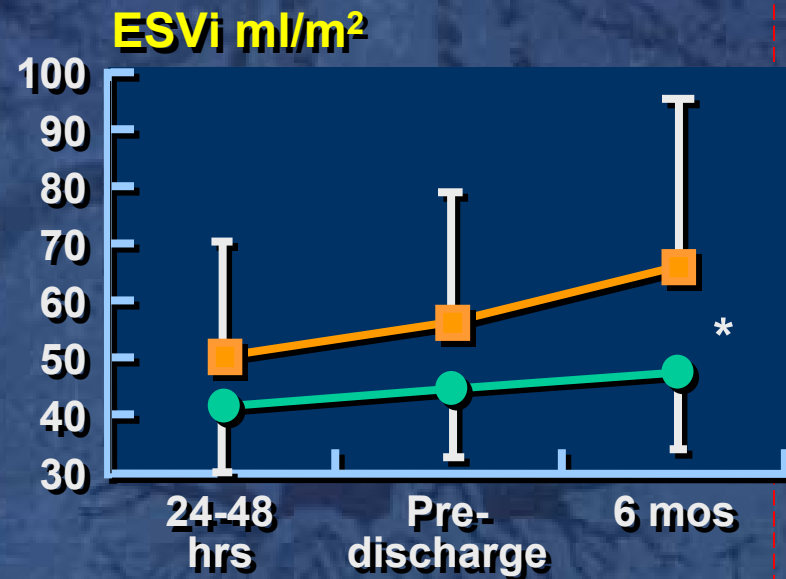
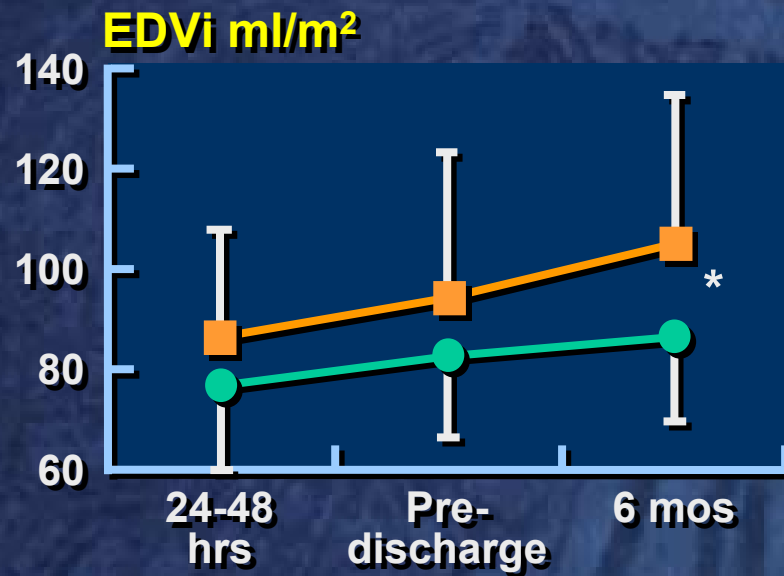
Cumulative Survival



Survival free of all Cardiac Events



X Changes According to Pre-Discharge DT



* $p < 0.001$ interaction time x groups

■ Pre-discharge DT <130 n= 56
● Pre-discharge DT >130 n= 91



Independent Predictors

Severe (>20%) dilation

Variable	Odds ratio	p value
Pre-discharge DT <130 ms	3.06	0.0001
Baseline EDVi	0.95	0.0001
Baseline %WMA	1.02	0.0001

Temporelli et al. Submitted



LV Changes

Post-AMI pts

- **LV dilation occurs**

even in uncomplicated MI, in spite of significant recovery of %WMA and improvement in LV filling

- **A short (<130 ms) DT**

occurs at baseline, which persists at pre-discharge, allows identification of pts more compromised and at high risk of progressive late dilation



LV Diastolic Filling

Post-AMI pts

- Adds important information to indices of global and regional function
- Allows identification of those at increased risk of progressive LV dilation and dysfunction