



Hospital Universitario
de La Princesa

Preliminary Program

**Madrid Microcirculation Meeting
-4th Edition-**

Treatment of microvascular



Madrid Microcirculation
Meeting -4th Edition-



Hospital Universitario
de La Princesa

obstruction in STEMI

Salvatore Brugaletta, MD, PhD, FESC



**Clínic
Barcelona**



UNIVERSITAT DE
BARCELONA

Disclosures/Conflicts of interest

Within the past 12 months, I or my spouse/partner have had a financial interest/arrangement or affiliation with the organization(s) listed below.

Affiliation/Financial Relationship

- Consulting Fees/Honoraria
- Speaker
- Research grant to my institution



Madrid Microcirculation Meeting - 4th Edition -

Company

- Boston Scientific, Insight Lifetech, iVascular, Novonordisk, Zoll,
- Abbott Vascular, General electric, Medis, Siemens
- Miracor



A need for treatment beyond reperfusion

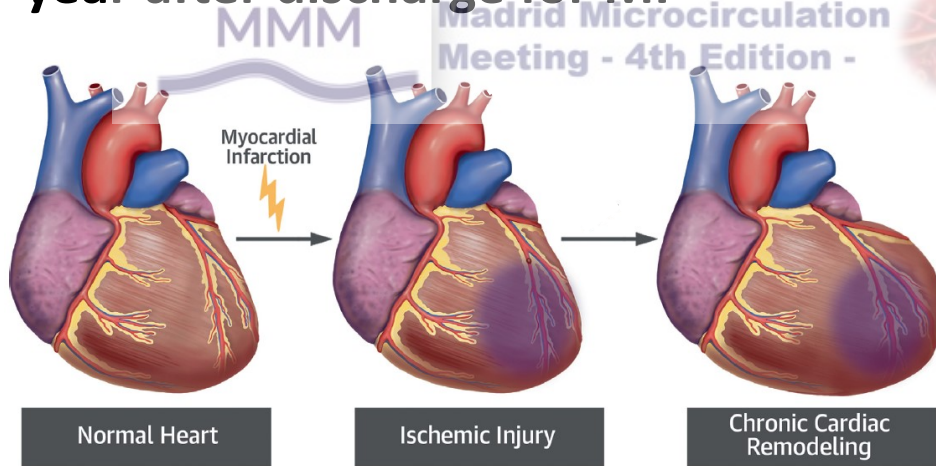
Treatment success of Acute Myocardial Infarction (AMI) patients is at a
10-year plateau



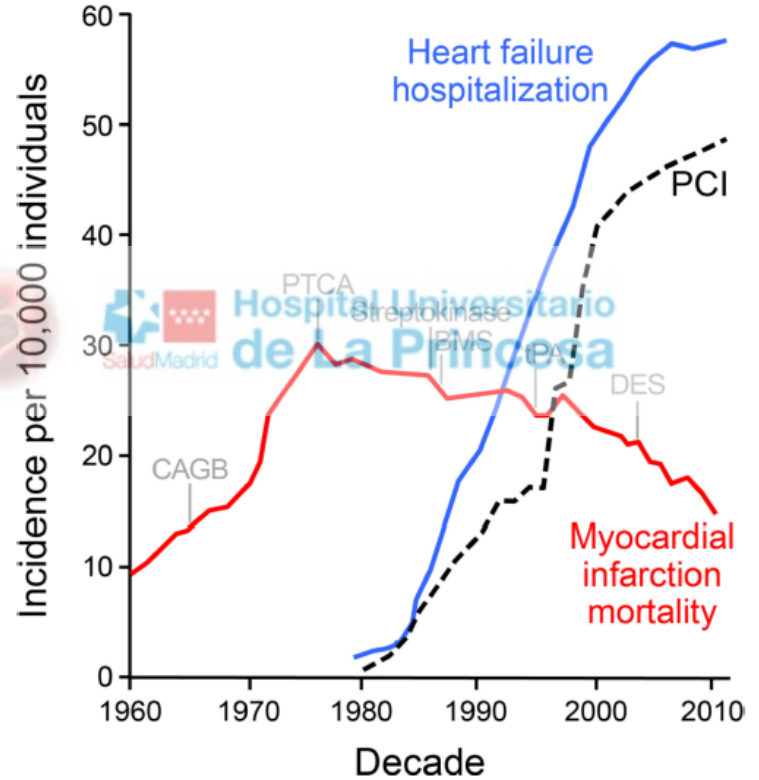
Up to 30% of patients develop HF within 1 year post-AMI, despite best practice PCI, due to **suboptimal myocardium salvage**

A paradox of medical success

Heart failure development is diagnosed in approximately 13% of patients at 30 days and 20–30% at 1 year after discharge for MI

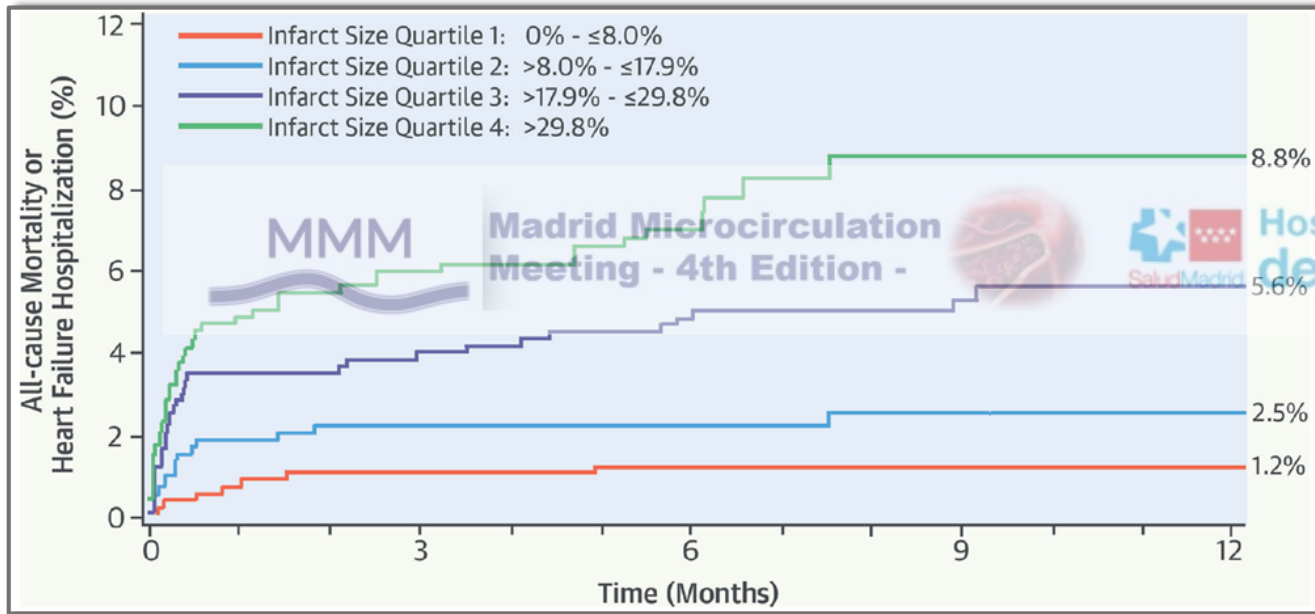


Sciarretta, S., et al. J Am Coll Cardiol, 2018. 71(18): p. 1999-2010.



Terzic, A. and A. Behfar, Trends in Cardiovascular Medicine, 2016. 26.

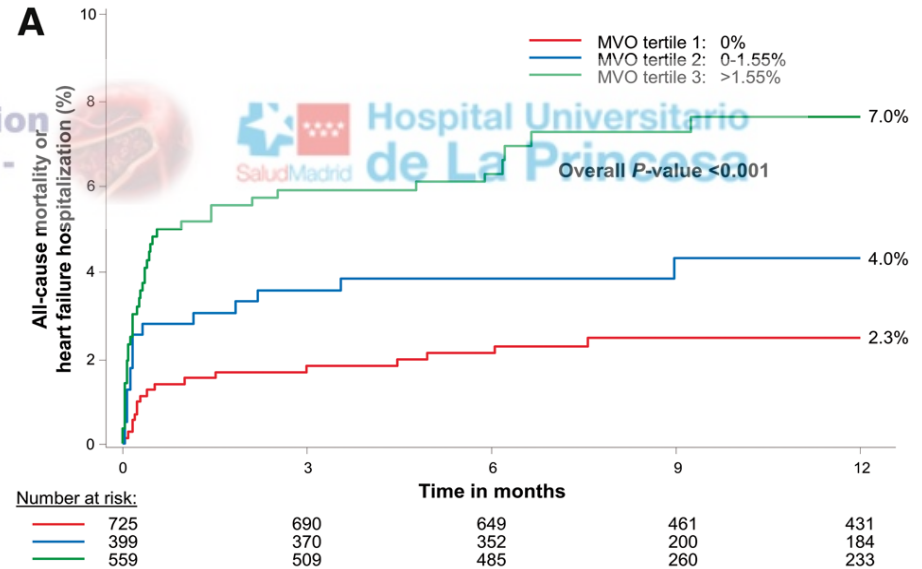
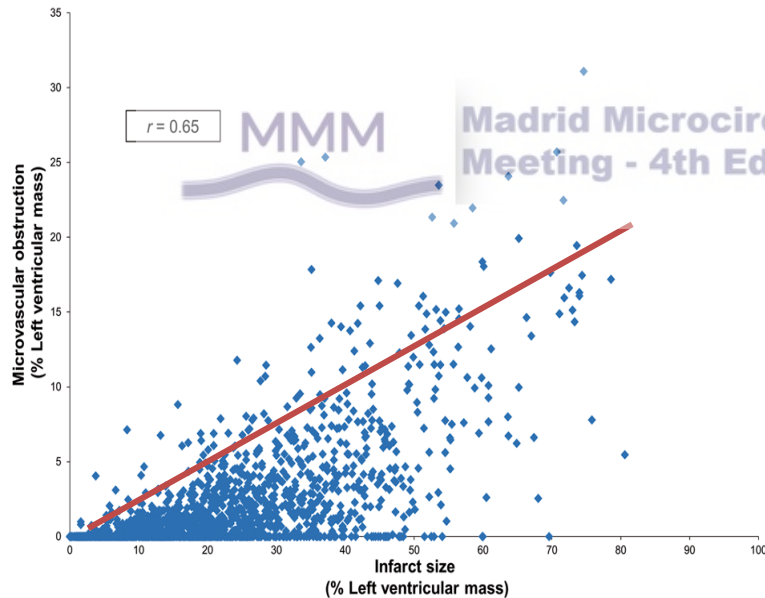
Infarct size is strongly associated with all-cause mortality & hospitalization for HF within 1 year



Every 5% increase in infarct size increases the risk of hospitalization for HF by 20%

Relationship between microvascular obstruction & clinical outcomes following primary PCI

- Patient level meta-analysis 7 RCTs PPCI, N = 1688, MVO assessed within 7 days by CMR.
- Every 1% increase in MVO increases the risk of 1-year all-cause mortality by 14% and 1-year HF hospitalization by 8%





Pharmacological strategies to reduce MVO

B-blockers

Adenosine

Statins

Atrial natriuretic
peptide

Iib/IIIa
Inhibitors

↑ Coronary
microvascular
vasodilation

↑ Microvascular
dilation

Inhibition of neutrophil-
platelet coaggregation

↓ Neutrophil adherence
and neutrophil-
mediated
cellular damage

↓ Platelet aggregation

↓ Oxidative stress

↑ Endothelial function

↓ Platelet activation

↓ Inflammation

↓ Immune response

↓ Neutrophils-induced
endothelial cytotoxicity

Platelet inhibition

MMM Madrid Microcirculation Meeting - 4th Edition - Hospital Universitario de La Princesa



Pharmacological strategies to reduce MVO

B-blockers

Adenosine

Statins

Atrial natriuretic peptide

Iib/IIIa Inhibitors

↑ Coronary microvascular vasodilation

↑ Microvascular dilation

Inhibition of neutrophil-platelet coaggregation

↓ Neutrophil adherence and neutrophil-mediated cellular damage

↓ Platelet aggregation

↓ Oxidative stress

↑ Endothelial function

↓ Platelet activation

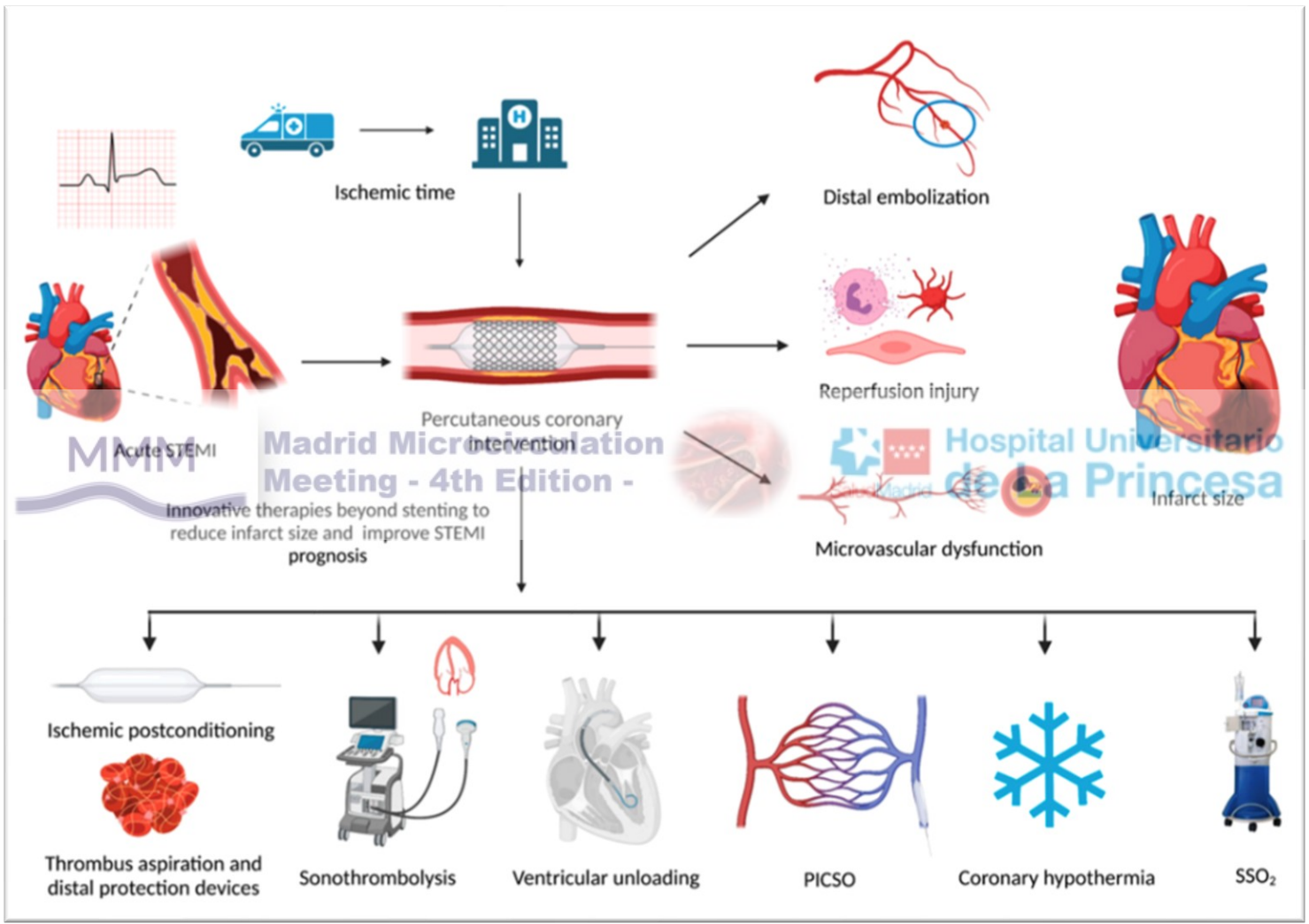
↓ Inflammation

↓ Immune response

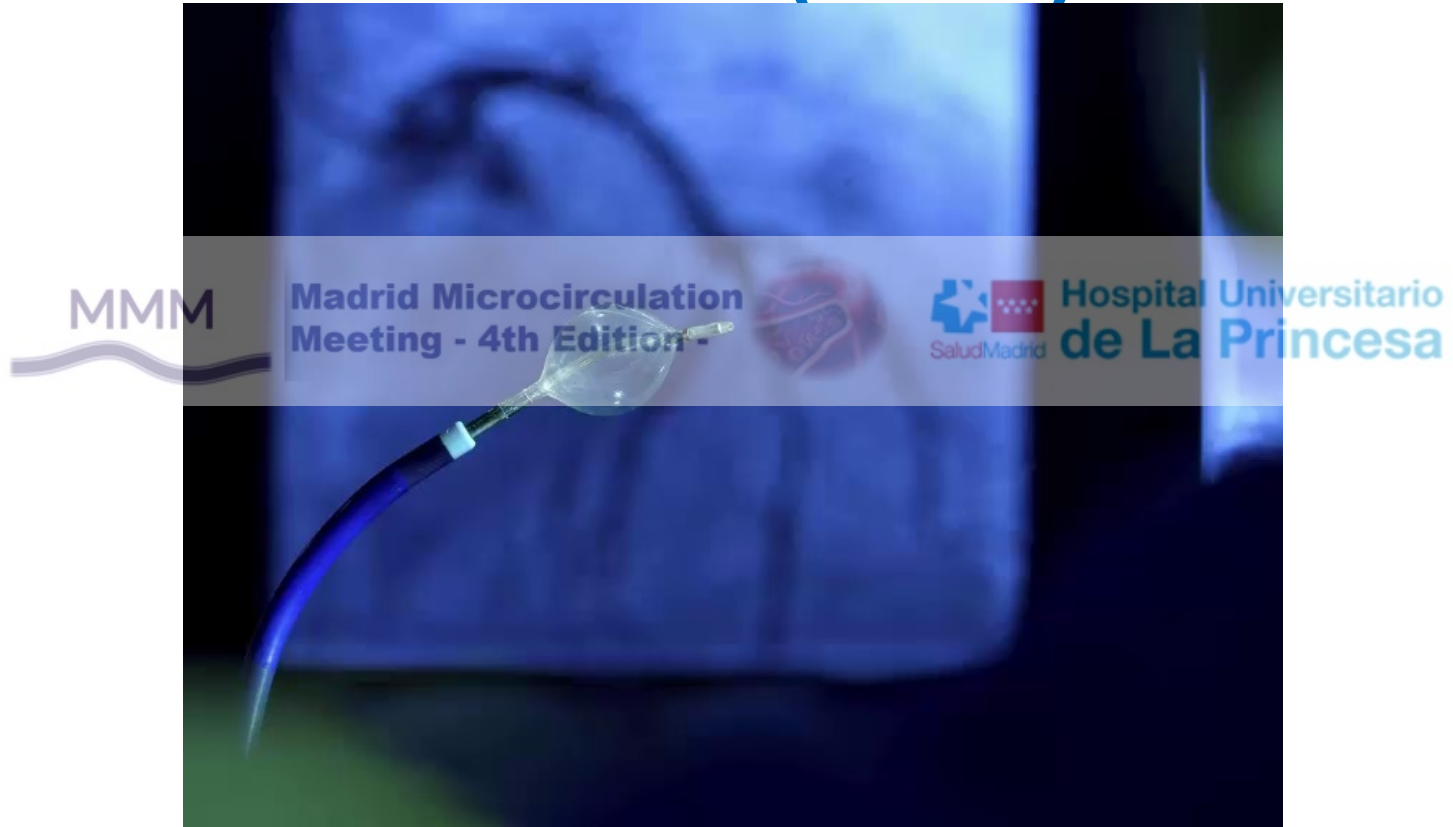
↓ Neutrophils-induced endothelial cytotoxicity

Platelet inhibition

MMM Madrid Microcirculation Meeting - 4th Edition - Hospital Universitario de La Princesa



Percutaneous intermittent coronary sinus occlusion (PiCSO)



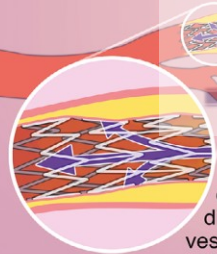
PiCSO

Diastole

Sistole



A 1. Plasma skimming & degranulation of platelets release miRNA into microcirculation



5. Further opening of diseased vessel

4. Vasodilatation in border zones and increase in collateral blood flow

- 1) Flow redistribution through collateral
- 2) Plasma Skimming phenomenon
- 3) Release of growth and vasodilatation factors

1. Cilia bending produces pleiotropic molecular signalling



PiCSO Balloon release



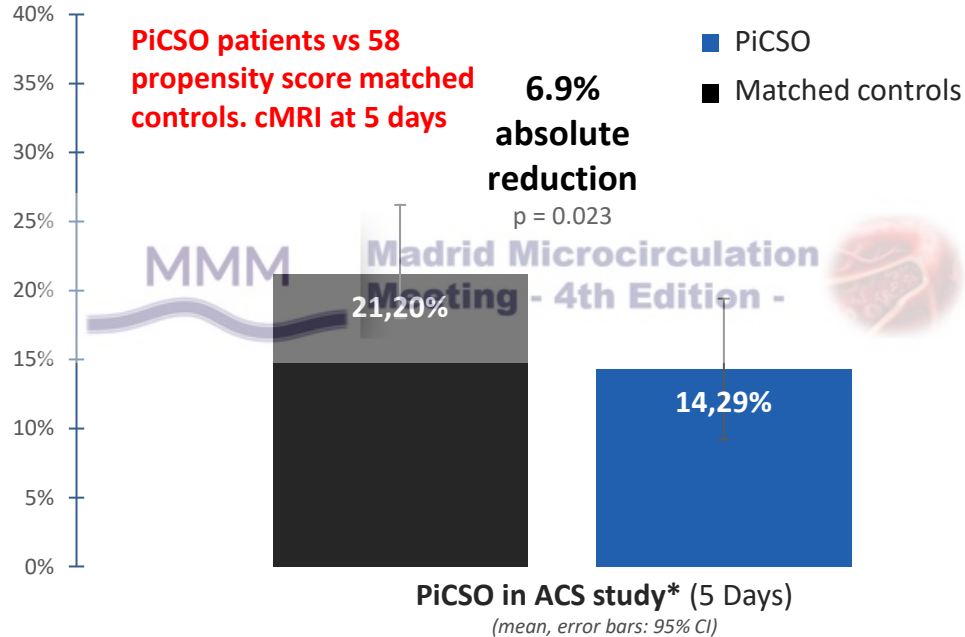
2. Pericyte stretching produces molecular signals

ema wash-out

- 2) Release of growth and vasodilators factors

PiCSO CE Mark study demonstrated clinical benefit

Infarct Size (% LV)



33% relative infarct size reduction in STEMI patients¹
(7% absolute)

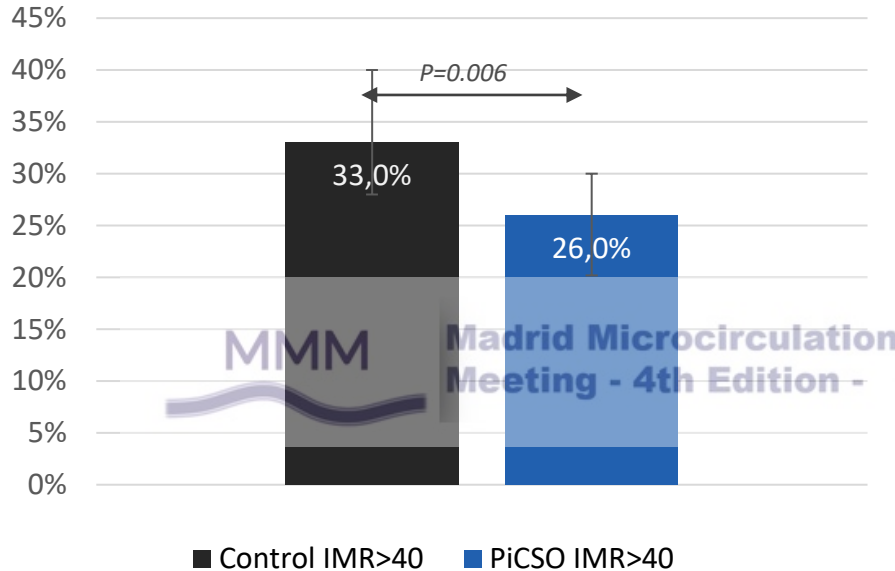
Hospital Universitario de La Princesa
Modeled on PiCSO results² at 1-year, could result in:

34% reduction in HF hospitalization &
25% reduction in mortality

1. Egred, M., et al., *Effect of Pressure-controlled intermittent Coronary Sinus Occlusion (PiCSO) on infarct size in anterior STEMI: PiCSO in ACS study*. IJC Heart & Vasculature, 2020. 28: p. 100526.
2. Stone et al: Polynomial regression analysis using PiCSO in ACS Study results based on Stone, et al. (2016). J Am Coll Cardiol 67(14): 1674-1683.

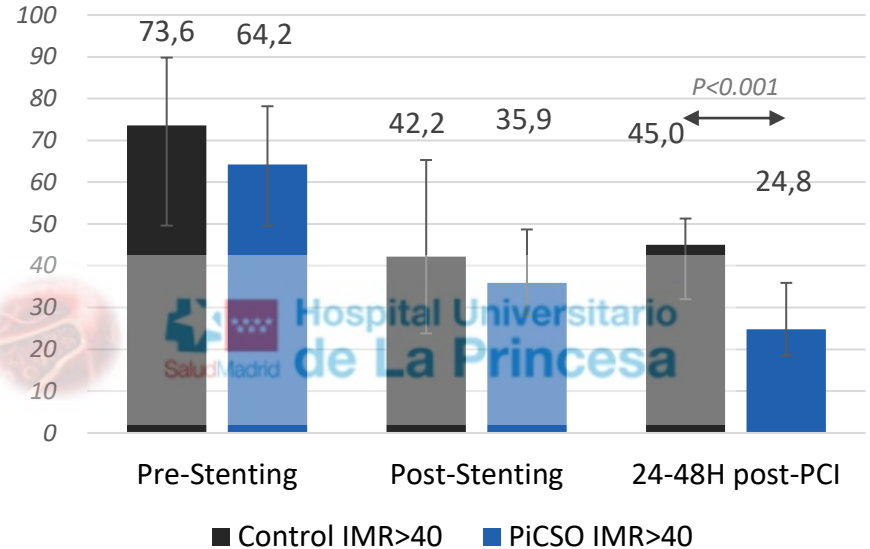
6-Month Infarct Size (% of LV)

(median, error bars: IQR)



IMR

(median, error bars: IQR)



- PiCSO-treated patients have better 6-months Infarct size (% of LV)
- PiCSO improved microcirculation - lower IMR @ 24-48h post-PCI

The PiCSO flow STEMI study

PROSPECTIVE, RANDOMIZED

46 anterior STEMI patients
within 12 hours



SoC



Microvascular
resistance (R_{μ})



endpoint
of PiCSO in
microvascular
immediately
its use

PiCSO-AMI-I trial

Design

International
Multicenter
Prospective
Randomized (1:1)
Controlled
Parallel-groups

PiCSO assisted pPCI

vs

Conventional pPCI

Primary Outcome

Infarct Size (%LV) at 5 ± 2 days CMR*

Secondary Outcome

MVO (%LV) at 5 ± 2 days CMR

IMH (%LV) at 5 ± 2 days CMR

Infarct Size (%LV) at 6 ± 1 months CMR

Myocardial Salvage 5 days CMR

Ejection Fraction 5 days /6 months CMR

ST segment resolution 60 – 90 min post flow restored

PiCSO Procedural Success rate

MACE at 6 months

*144 sample size
80% power, alpha 0.05
To detect 25% reduction in IS
Assuming IS of $26\% \pm 12$ in Control group and 20% drop-out rate

PiCSO-AMI-I trial

179 patients with anterior STEMI screened
between July 2019 and August 2022

30 patients with screening failure
(not fulfilling inclusion/exclusion criteria)

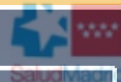
149 patients with anterior STEMI randomized

4 patients declined to remain in the study
post pPCI

145 patients fully enrolled and randomized



Madrid Microcirculation
Meeting - 4th Edition -



Hospital Universitario
de La Princesa

72
PiCSO-assisted pPCI

73
Conventional pPCI

11 patients (15.3%) – no 5 days CMR
6 patient (8.3%) – No infarct size measured

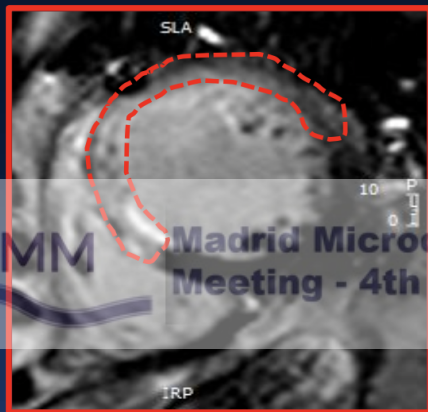
8 patients (11%) – no 5 days CMR
1 patient (1.4%) – No infarct size measured

55 (76.4%)
Infarct size available at 5 days

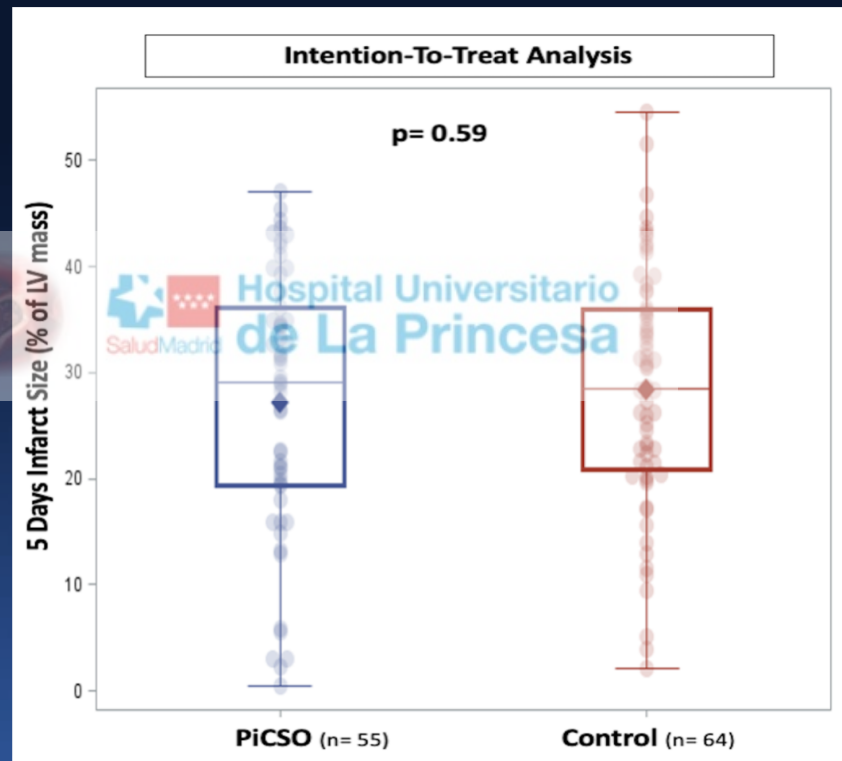
64 (87.4%)
Infarct size available at 5 days

PiCSO-AMI-I trial

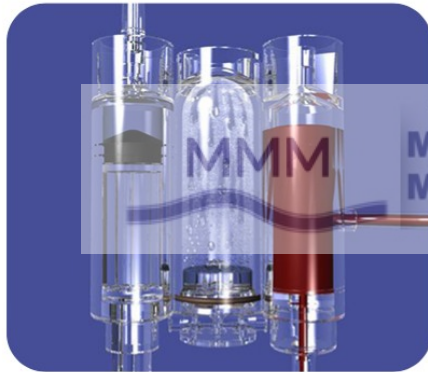
PRIMARY ENDPOINT: IS% @5days CMR – Intention to treat Analysis



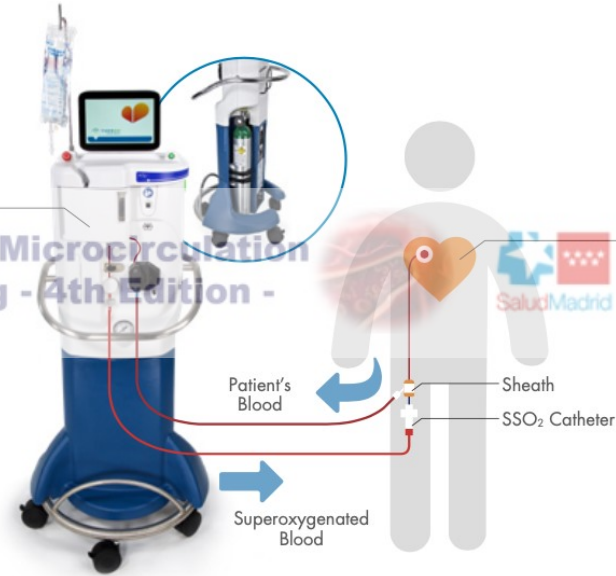
	PiCSO (n = 55)	Control (n = 64)	p
IS 5 days (LV%)	27.2% ± 12.4	28.3% ± 11.45	0.59



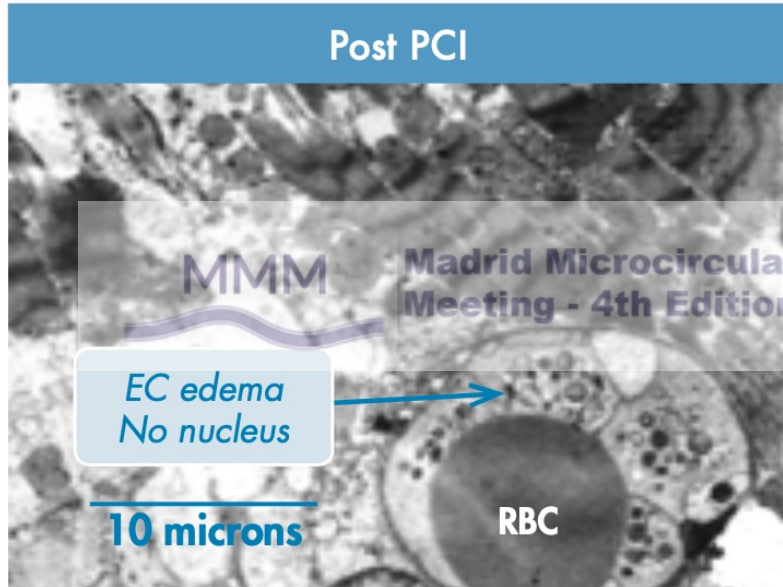
SSO₂ therapy



Madrid Microcirculation Meeting - 4th Edition -



SSO₂ therapy



SSO₂ therapy

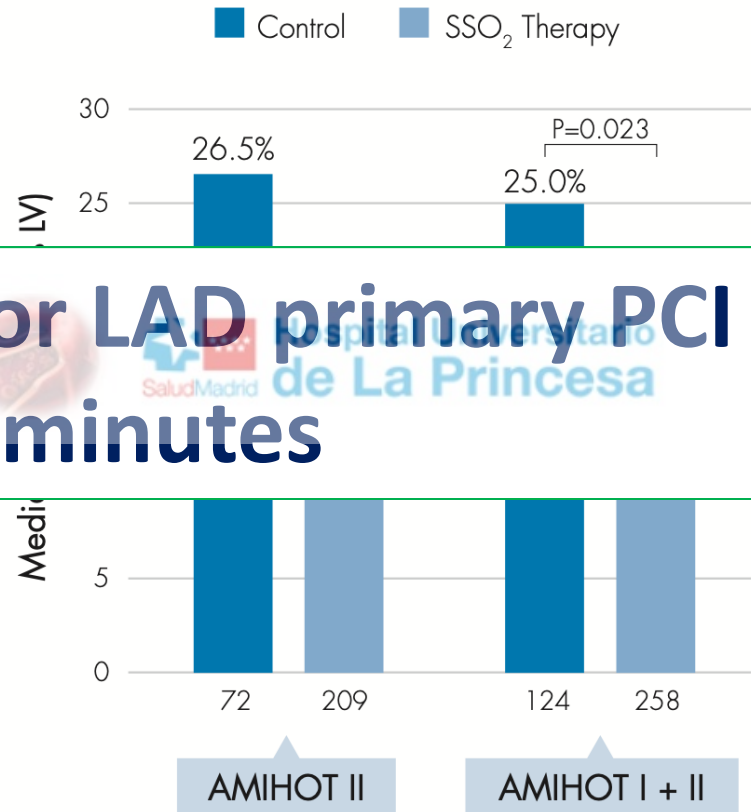
AMIHOT I

269 pts with acute or

**60 minutes infusion for LAD primary PCI
within 120 minutes**

AMIHOT II

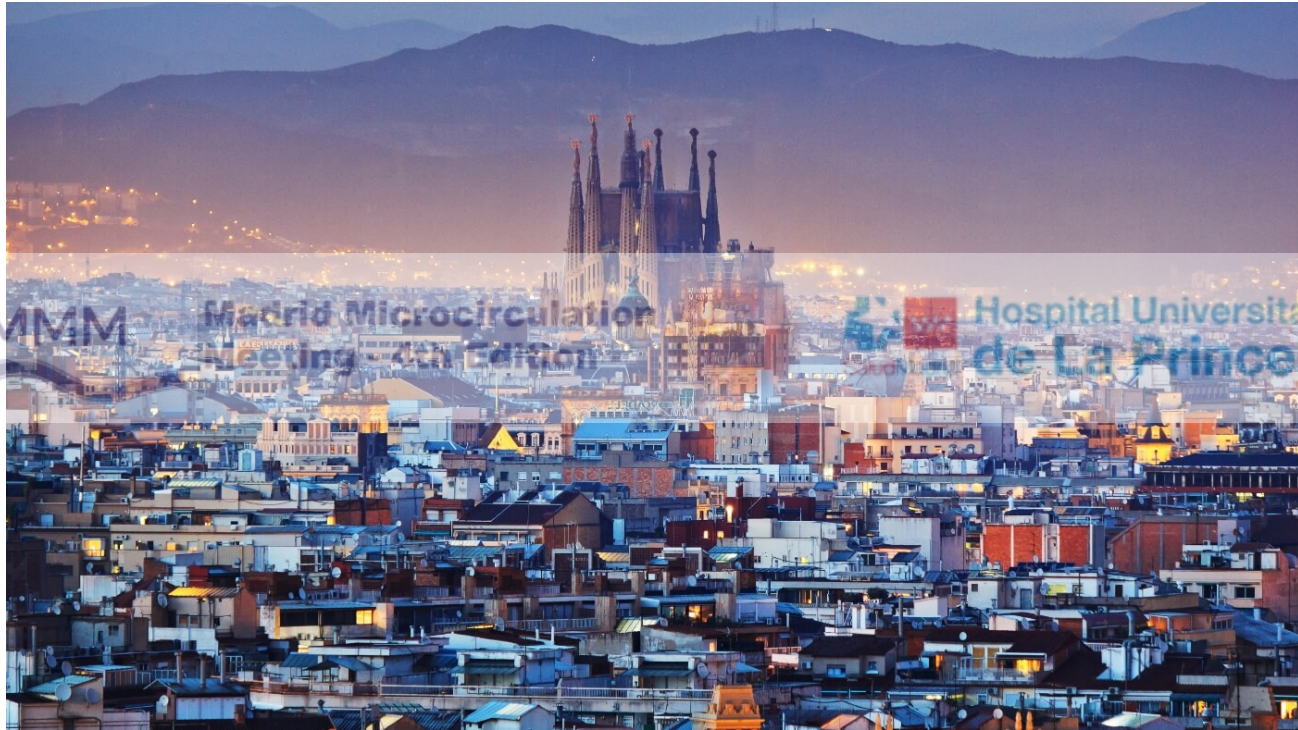
301 pts with anterior
STEMI



Take home-messages

- **MVO** as a primary therapeutic target in STEMI
- **Drugs & devices** may be useful tools
- There is a growing in the field of devices **specifically focused on treating MVO in STEMI patients**

Thank you!



@sbrugaletta

sbrugaletta@gmail.com