

STEMI with MVD; How can we treat?

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Evolving Guidelines of STEMI with MVD



2010 Revascularization

With the exception of Cardiogenic Shock(CS), PCI should be **limited to the culprit stenosis**.

IIa

B

2014 Revascularization

Immediate revascularization of significant non-culprit lesions during the same procedure as primary PCI of the culprit vessel may be **considered in selected patients**.

IIb

B

2020 ACS

Complete Revascularization(CR) should be considered in **NSTE-ACS patients** without CS and with MVD

IIa

C

Culprit Only

Complete Revascularization

2012 STEMI

The best strategy for STEMI patients with MVD, who underwent primary PCI of the IRA in the acute phase with remaining MVO, is **still not well established**.

2017 STEMI & 2018 Revascularization

Routine revascularization of non-IRA lesions should be considered in STEMI patients with MVD **before hospital discharge**.

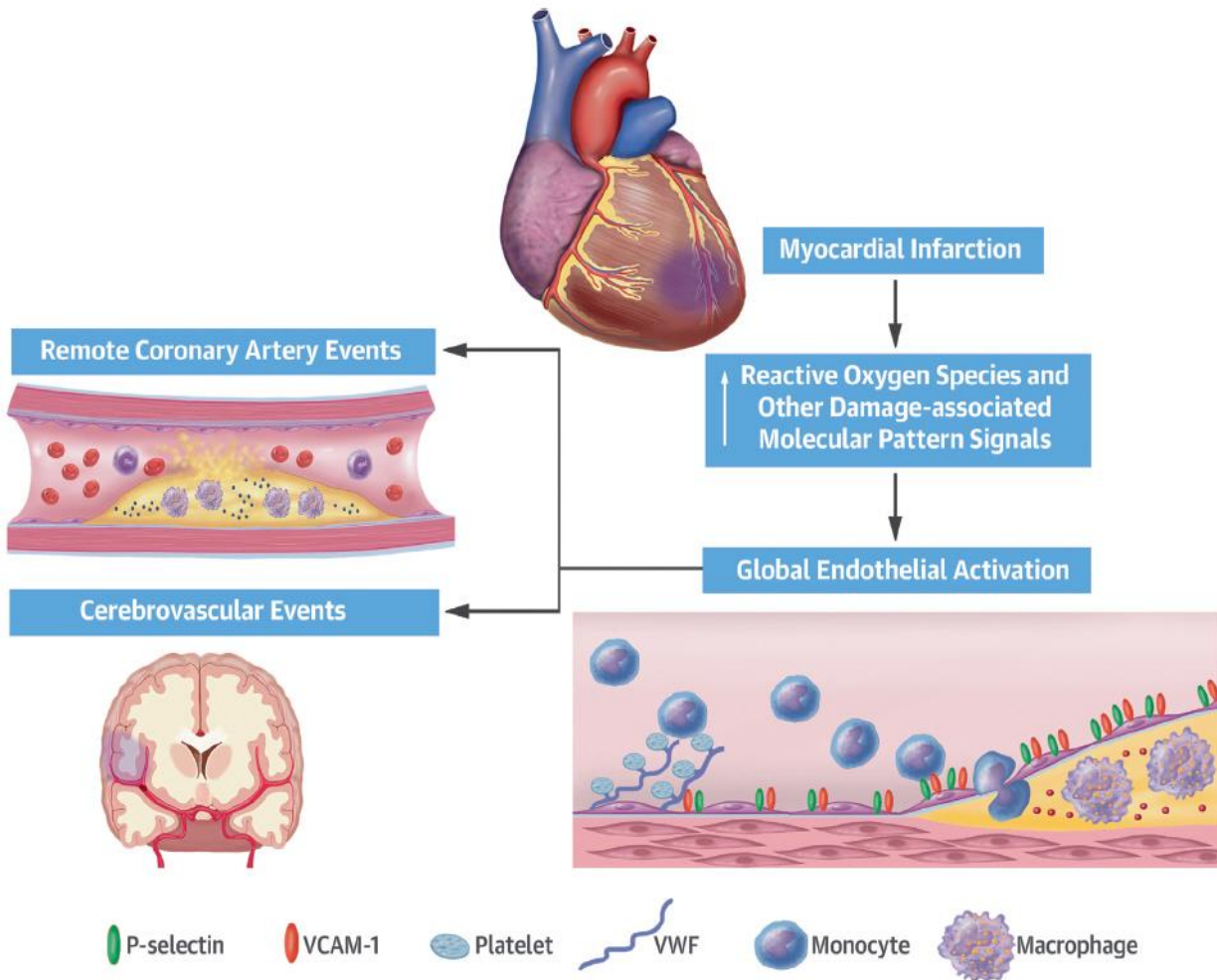
IIa

A

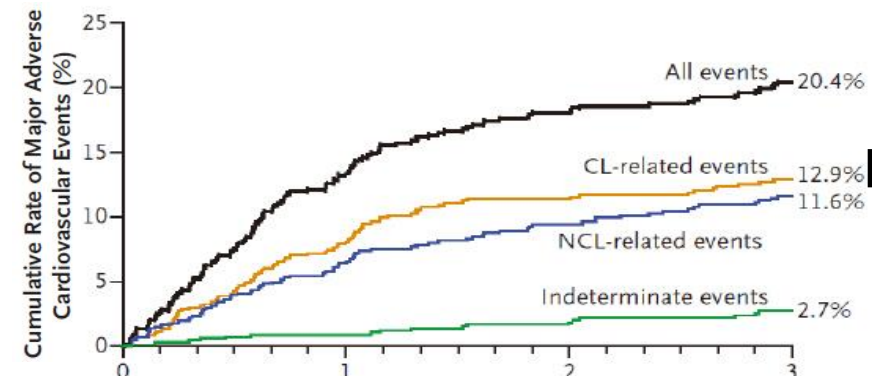
Routine revascularization of non-IRA lesions in Myocardial Infarction with **Cardiogenic shock**. III,B

What happen to NCLs in MI

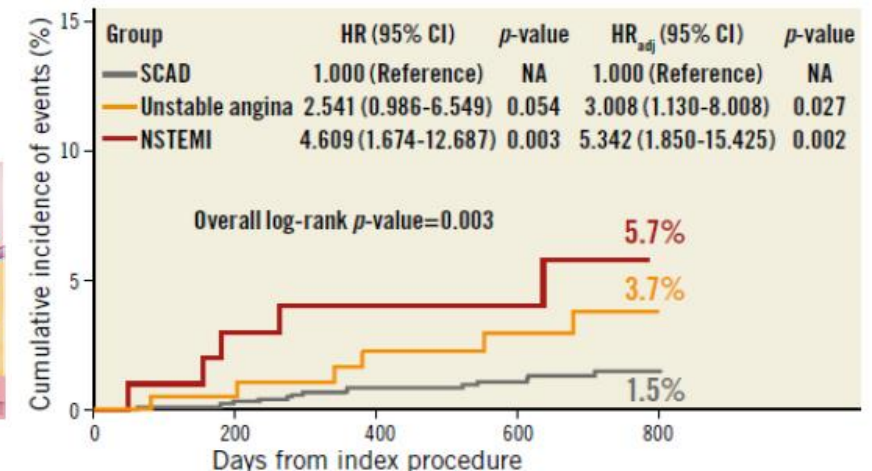
MI triggers systemic effects,
Remote Endothelial Activation after MI



Non- Culprit Leions(NCLs) were associated with
More endothelial dysfunction
More inflammation



Plaque progression



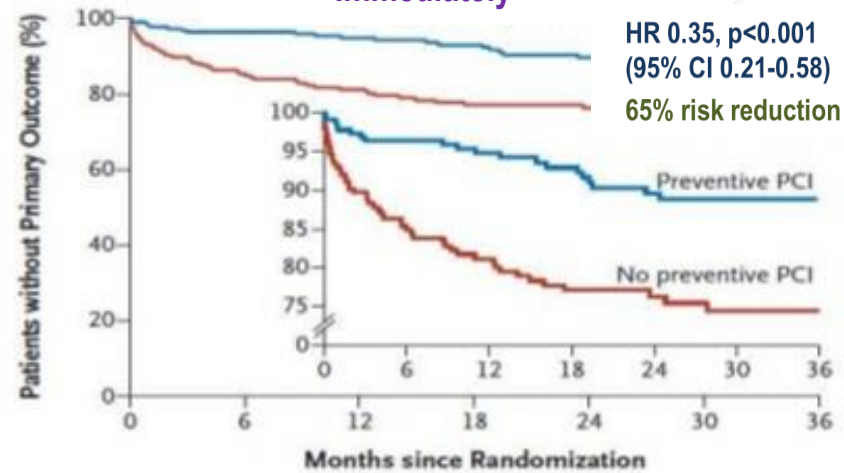
Worse outcomes

Evidence of CR in STEMI

Clinical Outcomes of Angiography-Guided Non-Culprit PCI

PRAMI – cardiac death, non-fatal MI, refractory angina

Immediately

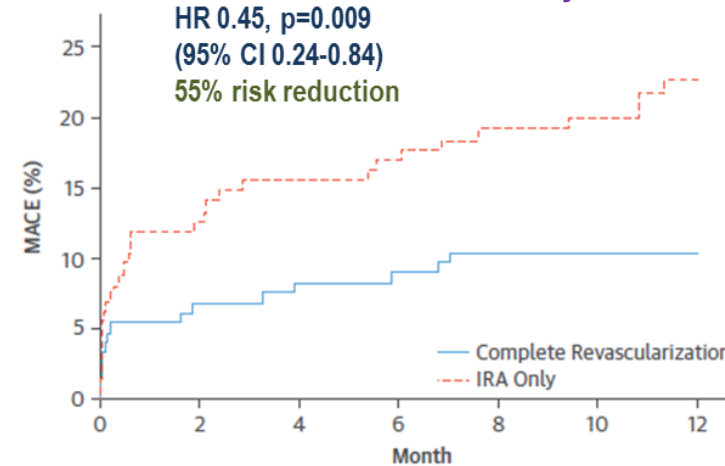


No. at Risk		0	6	12	18	24	30	36
Preventive PCI	234	196	166	146	118	89	67	
No preventive PCI	231	168	144	122	96	74	50	

Preventive PCI for non-culprit lesion >50% DS

CvLPRIT – all death, recurrent MI, HF, ischemia-revascularization

Immediately



Number at risk:		0	2	4	6	8	10	12
Complete Revascularization	150	131	129	128	125	108	73	
IRA Only	146	122	118	116	111	98	68	

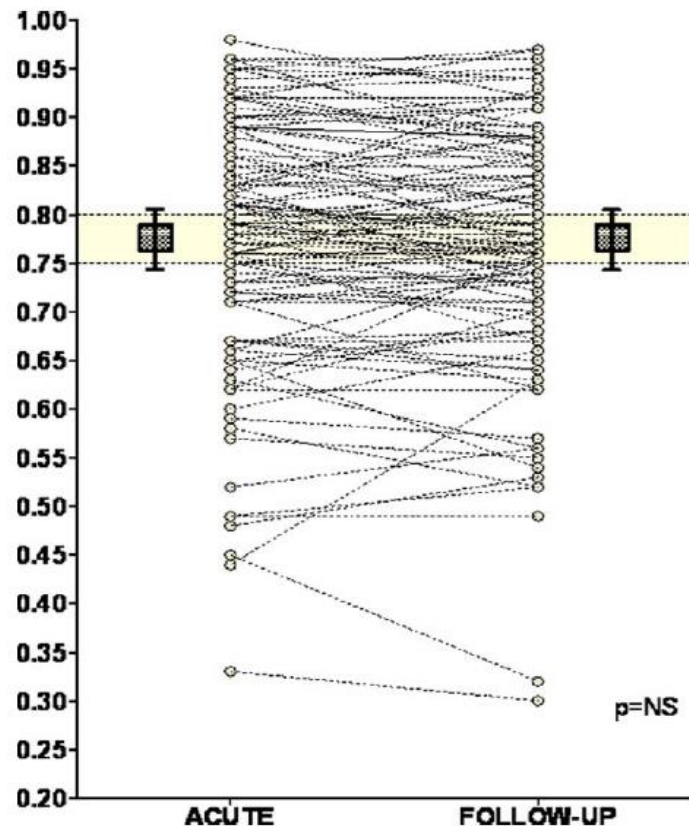
Preventive PCI for non-culprit lesion > 70% DS or > 50% DS in 2 views

Recent RCTs presented

“Angiography-guided” Complete Revascularization showed Significant benefit in Patient’s outcome than “Culprit-Only PCI”

Reliability of NCVs FFR in AMI

Non-culprit vessel of AMI Patient



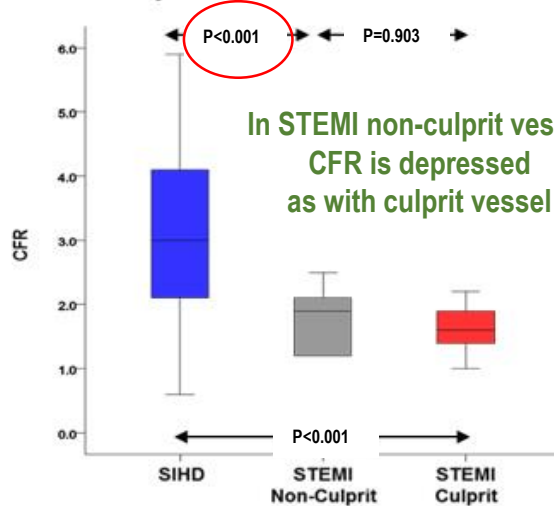
	Acute Phase (n=101)	1M Follow-Up (n=101)	P Value
LVEF (%)	59 ± 15	61 ± 14	NS
LVEDP (mmHg)	18 ± 7	17 ± 7	NS
FFR nonculprit	0.77 ± 0.13	0.77 ± 0.13	NS
IMR nonculprit (IU)	20 ± 3	24 ± 6	NS
DS nonculprit (%)	56 ± 14	55 ± 14	NS
TIMI flow nonculprit	2.93 ± 0.30	2.97 ± 0.20	NS
cTFC nonculprit	15 ± 6	15 ± 6	NS

In patients with acute MI (including STEMI and NSTEMI),
Non-culprit FFR did **not show significant change**.

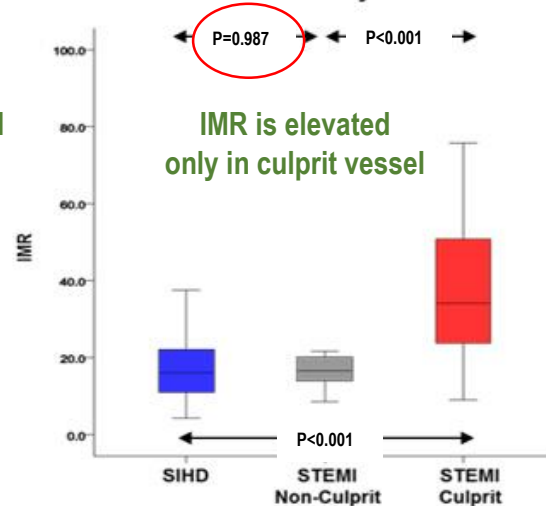
Reliability of NCVs FFR in STEMI



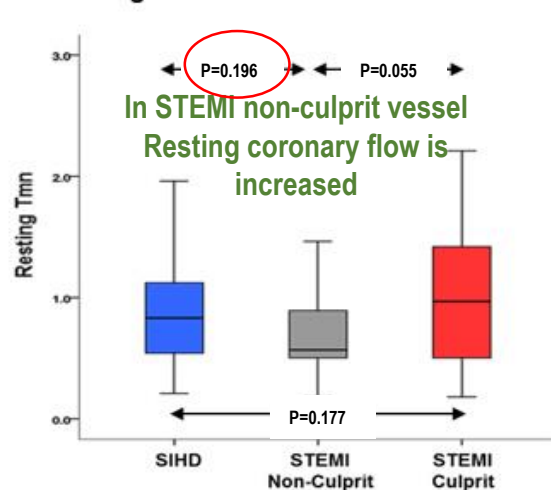
A. Coronary Flow Reserve



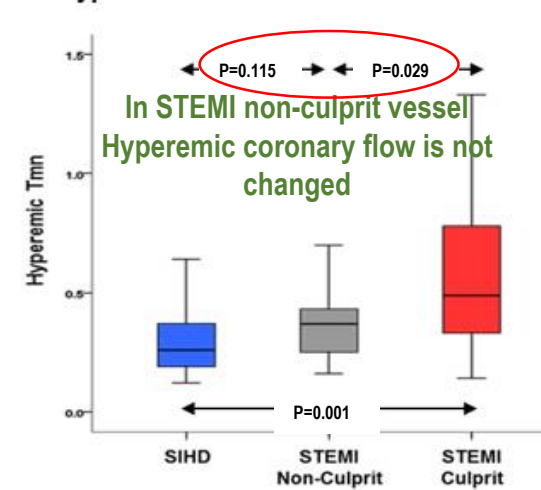
B. Index of Microcirculatory Resistance



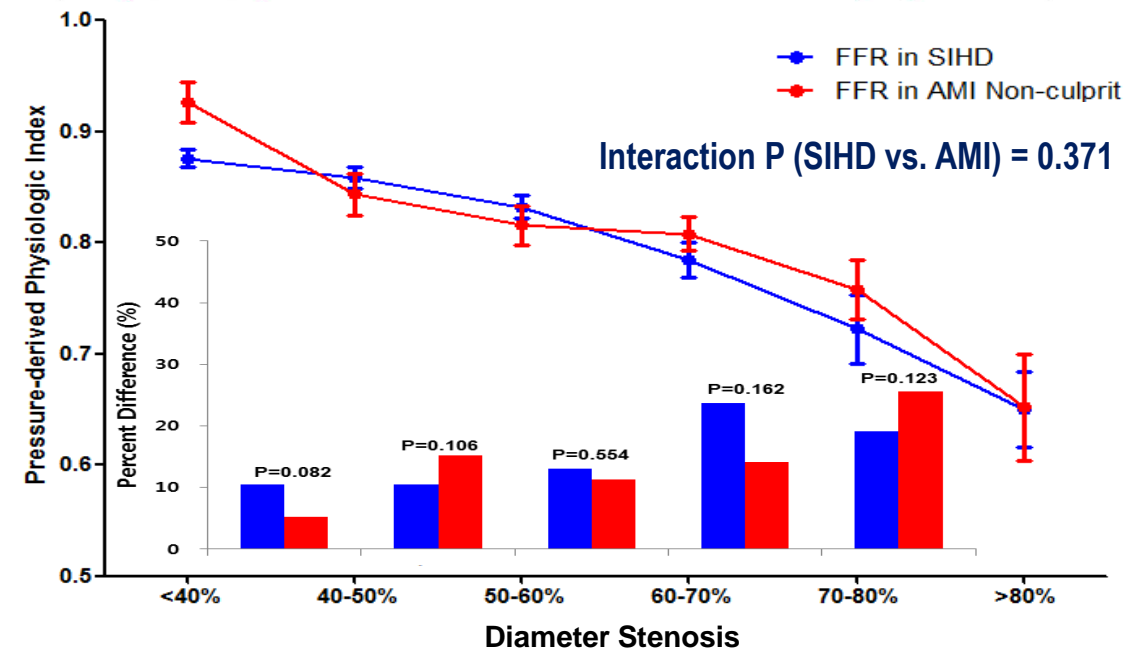
C. Resting Mean Transit Time



D. Hyperemic Mean Transit Time



Variables	Total (n=303)	SIHD (n=203)	AMI (n=100)	P value
Age (years)	60.8 ± 13.2	59.6 ± 13.7	63.3 ± 11.8	0.020
Male	230 (75.9%)	150 (73.9%)	80 (80.0%)	0.305
Clinical presentation				<0.001
Stable ischemic heart disease		203 (100%)		
Non ST-segment elevation myocardial infarction			66 (66.0%)	
ST-segment elevation myocardial infarction			34 (34.0%)	

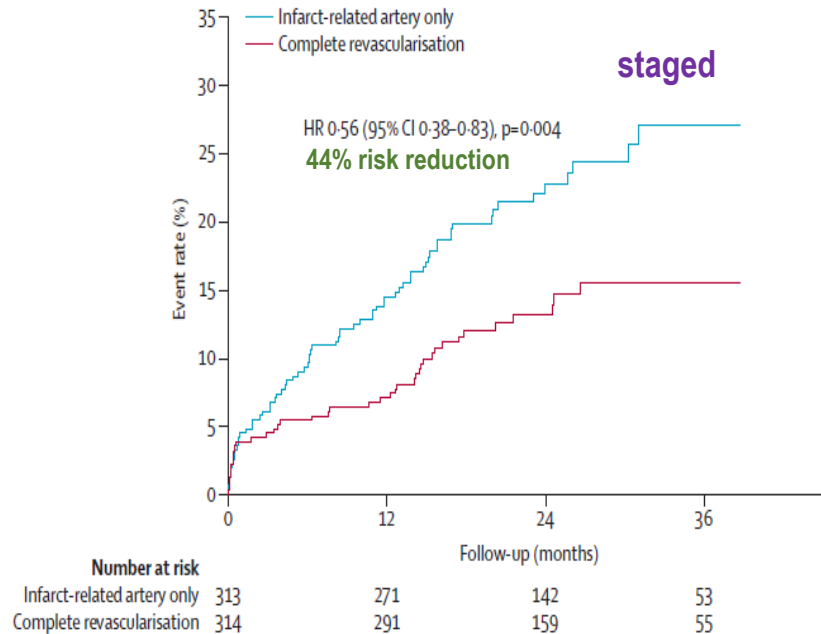


Local microvascular damage in culprit vessel was not extended to NCV territory, and **NCV FFR and IMR were not changed at all.**

Evidence of CR in STEMI

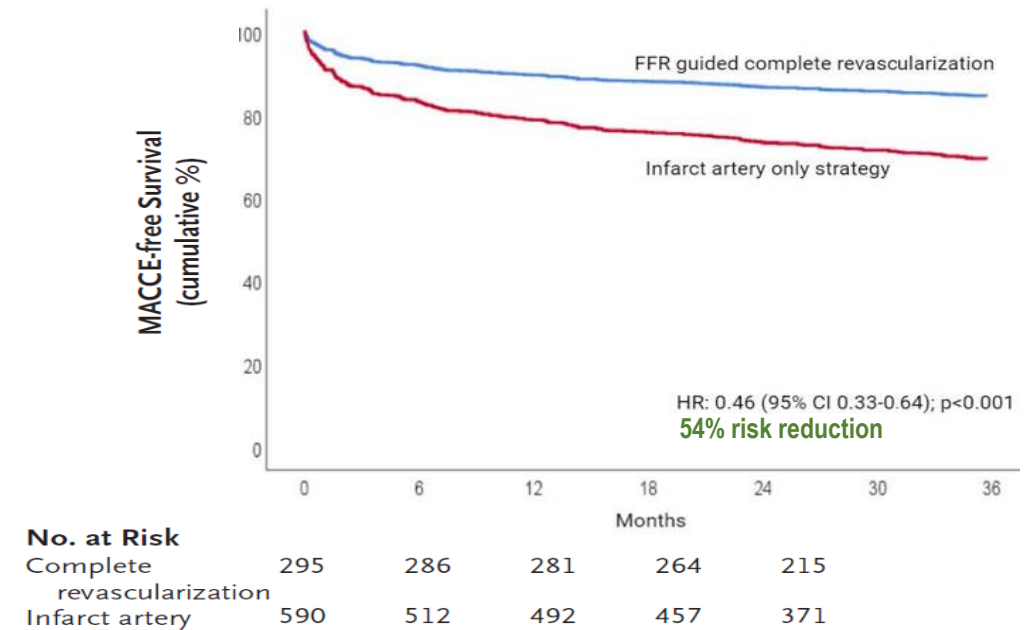
Clinical Outcomes of Physiology-Guided Non-Culprit PCI

DANAMI-3-PRIMULTI (Death, MI, IDR)



Mainly driven by ischemia-driven revascularization
(both urgent and planned)

COMPARE-ACUTE (Death, MI, Revascularization, Stroke)



Mainly driven by revascularization
(1/3 Unstable angina)

FFR-guided decision for non-culprit vessel in STEMI patients are strongly supported by 2 RCTs
(Both Staged Measurement and Acute phase Measurement)

Evidence of CR in STEMI



STEMI WITH MULTIVESSEL CAD AND SUCCESSFUL PCI TO THE CULPRIT LESION
 MVD defined as at least one additional non-culprit lesion ≥ 2.5 mm diameter and $\geq 70\%$ stenosis or 50-69% with FFR ≤ 0.80 **FFR Guided Enrollment <1%**

Almost Angiography-Guided Non-Culprit PCI

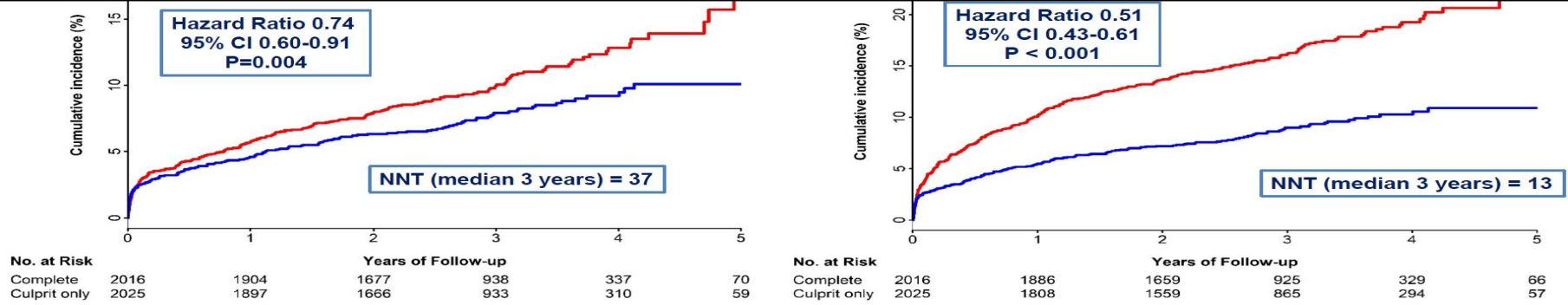
Exclusion Criteria: Intent to revascularize NCL, planned surgical revascularization, prior CABG

RANDOMIZATION

Stratified for intended timing of NCL PCI:
 During initial hospitalization or after discharge (max 45 d)

Actual Time to study NCL PCI in Complete Group (median)
 During initial hospitalization: 1 day (IQR 1-3)
 After hospital discharge: 23 days (IQR 12.5-33.5)

Benefits of CR in STEMI are 'No Doubt'



Reduced CV death or new MI by **26%** (P=0.004), NNT = 37
 Reduced CV death, new MI, or IDR by **49%** (P<0.001), NNT = 13

What is the Best Timing of CR?



Compare of 5 RCTs & Meta-Analysis(Subgroup)

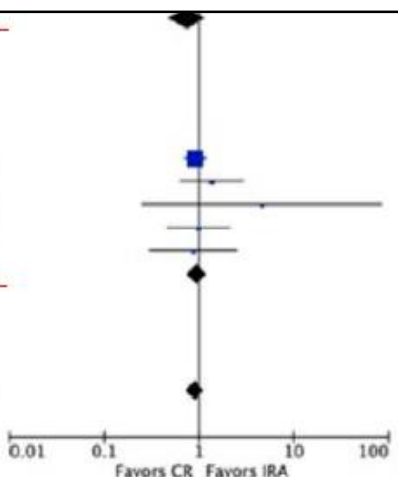
PRAMI	CvLPRIT	COMPARE-ACUTE	DANAMI-3-PRIMULTI	COMPLETE
Angiographic Guided	Angiographic Guided	FFR Guided	FFR Guided	Angiographic Guided

Immediately CR outcome seems to be **slightly better** than Staged CR. But, Comparative RCTs on this topic have **not yet been reported**. According to the experience of many experts, the Best Timing of CR depends on the **Patient & Lesion's condition**.

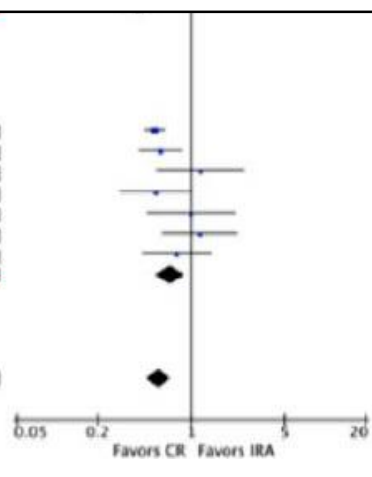
Im
CV De
HR 0.3
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Subtotal (95% CI)

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43-0.61)

Study	Total Events	CR	IRA	CR %	OR [95% CI]	
Staged Revascularization						
COMPLETE 2019	96	2016	106	2025	58.9%	0.91 [0.70, 1.19]
DANAMI-3-PRIMULTI 2015	15	314	11	313	7.3%	1.35 [0.63, 2.91]
Ghani 2012	4	79	0	40	0.5%	4.61 [0.25, 83.61]
Politi 2010	10	65	13	84	7.4%	0.99 [0.47, 2.12]
PRAGUE 2010	6	106	7	108	3.8%	0.87 [0.30, 2.51]
Subtotal (95% CI)	2580	2570	78.0%		0.96 [0.76, 1.21]	
Total events	131	137				
Heterogeneity: Tau ² = 0.00; Chi ² = 2.13, df = 4 (P = 0.71); I ² = 0%						
Test for overall effect: Z = 0.33 (P = 0.74)						
Total (95% CI)	3324	3621	100.0%		0.91 [0.74, 1.12]	
Total events	161	186				
Heterogeneity: Tau ² = 0.00; Chi ² = 4.88, df = 8 (P = 0.77); I ² = 0%						
Test for overall effect: Z = 0.89 (P = 0.37)						



Study	Total Events	CR	IRA	CR %	OR [95% CI]	
Staged Revascularization						
COMPLETE 2019	179	2016	339	2025	18.5%	0.53 [0.45, 0.63]
DANAMI-3-PRIMULTI 2015	40	314	68	313	12.7%	0.59 [0.41, 0.84]
Estevez Loureiro 2014	13	100	11	99	5.3%	1.17 [0.55, 2.49]
Ghani 2012	15	79	14	40	7.0%	0.54 [0.29, 1.01]
Politi 2010	10	65	13	84	5.2%	0.99 [0.47, 2.12]
PRAGUE 2010	17	106	15	108	6.7%	1.15 [0.61, 2.19]
Zhang 2015	18	215	23	213	7.5%	0.78 [0.43, 1.39]
Subtotal (95% CI)	2895	2882	63.0%		0.69 [0.54, 0.89]	
Total events	292	483				
Heterogeneity: Tau ² = 0.05; Chi ² = 11.60, df = 6 (P = 0.07); I ² = 48%						
Test for overall effect: Z = 2.83 (P = 0.005)						
Total (95% CI)	3639	3933	100.0%		0.57 [0.47, 0.70]	
Total events	364	730				
Heterogeneity: Tau ² = 0.05; Chi ² = 19.55, df = 10 (P = 0.03); I ² = 49%						
Test for overall effect: Z = 5.53 (P < 0.00001)						



How to define 'significant' NCLs?

Both Angio- and FFR- are supported by RCTs

Angiographic stenosis	FFR immediate phase	FFR staged phase
PRAMI	COMPARE-ACUTE	DANAMI-3-PRIMULTI
CvLPRIT		
COMPLETE		



FLOWER-MI

Flow Evaluation to Guide Revascularization in Multivessel ST-Elevation Myocardial Infarction

Randomized, open-label, multicenter trial

1,171 PATIENTS
PCI based on FFR ≤ 0.80 vs. >50% visual stenosis

concurrent

vs.

FFR-GUIDED REVASCULARIZATION (N=590)

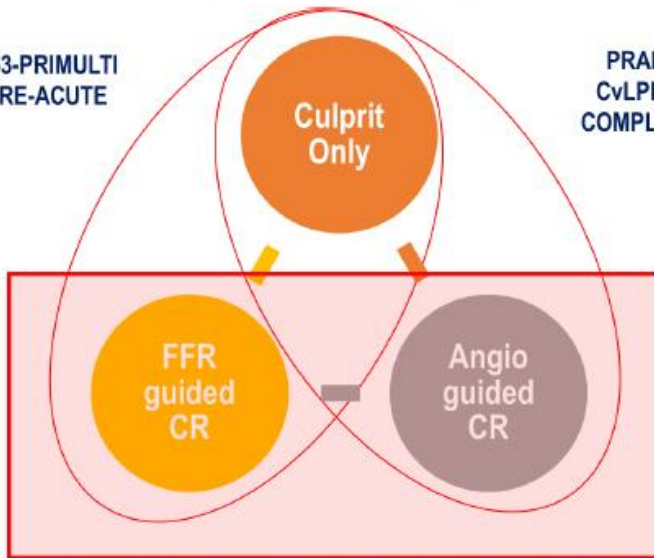
ANGIOGRAPHY-GUIDED REVASCULARIZATION (N=581)

INCLUSION CRITERIA: Patients with STEMI and multi-vessel non-culprit coronary disease

Profiles	FFR(n=586)	Angio(n=577)
Lesions with PCI (non-IRA)	546/980(55.7%)	806/891(90.5%)
Patients with ≥ 1PCI (non-IRA)	388/586(66.2%)	560/577(97.1%)
Mean Stent No. (non-IRA)	1.01 ± 0.99	1.50 ± 0.86
Non-IRA Post PCI TIMI3	686/980(70%)	827/891(92.8%)
Non-fatal MI	18(3.1%)	10(1.7%)
Periprocedural MI	7/18(38.9%)	2/10(20%)
Similar Clinical outcome, Given the 40% less PCI (Time, Cost, contrast, Procedure risk etc.) in FFR-guided group.		
Cardiac death	2/9(22.2%)	7/10(70%)
Non-cardiac death	7/9(77.8%)	3/10(30%)
Urgent revascularization	15(2.6%)	11(1.9%)
Non-IRA Treatment	8/15(53.5%)	3/11(27.3%)

DANAMI-3-PRIMULTI
COMPARE-ACUTE

PRAMI
CvLPRIT
COMPLETE



Question is How to Select Proper Non-IRA for PCI

FFR-Guided revascularization was not superior to angiography-guided revascularization

EDITORIALS



PCI for Nonculprit Lesions in Patients with STEMI — No Role for FFR

No. at Risk

Angiography-guided PCI	577	570	567	565	560	560	557	555	555	554	552	548	371
FFR-guided PCI	586	577	573	570	567	566	566	562	559	553	553	549	385

PCI strategy of MI with MVD in CS

CULPRIT-SHOCK(All cause Death)

Relative risk, 0.88 (95% CI, 0.76–1.01)

STEMI without Cardiogenic Shock STEMI with Cardiogenic Shock

Immediately >70%stenosis

Outcome **28.2% Mechanical support**

17.7% Staged Multivessel PCI & 12.5% Immediately no./total no. (%)

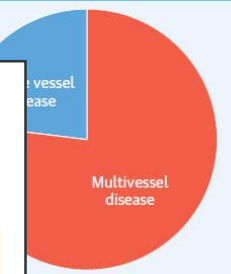
	Culprit-Lesion-Only PCI Group (N=344)	Multivessel PCI Group (N=341)	Relative Risk (95% CI)
Primary end point: death from any cause or renal-replacement therapy	158/344 (45.9)	189/341 (55.4)	0.83 (0.71–0.96)
Death from any cause*	149/344 (43.3)	176/341 (51.6)	0.84 (0.72–0.98)
Renal-replacement therapy	40/344 (11.6)	56/341 (16.4)	0.71 (0.49–1.03)
Death from any cause†	172 (50.0)	194 (56.9)	0.88 (0.76–1.01)
Renal-replacement therapy‡	40 (11.6)	56 (16.4)	0.71 (0.49–1.03)
Recurrent myocardial infarction	6 (1.7)	7 (2.1)	0.85 (0.29–2.50)
Death or recurrent infarction	175 (50.9)	199 (58.4)	0.87 (0.76–1.00)
Rehospitalization for congestive heart failure	18 (5.2)	4 (1.2)	4.46 (1.53–13.04)
Death, recurrent infarction, or rehospitalization for heart failure	190 (55.2)	203 (59.5)	0.87 (0.93–1.06)
Repeat revascularization			
Any	111 (32.3)	32 (9.4)	3.44 (2.39–4.95)
PCI	107 (31.1)	29 (8.5)	3.66 (2.50–5.36)

30d f/u

1year f/u

No. at Risk

	0	30	60	120	180	240	300	360	420
Multivessel PCI	165	161	160	156	152	149	131		
Culprit-lesion-only PCI	195	186	181	178	174	172	147		



KAMIR-NIH Nationwide Multicenter Registry (2011.11 - 2015.12)

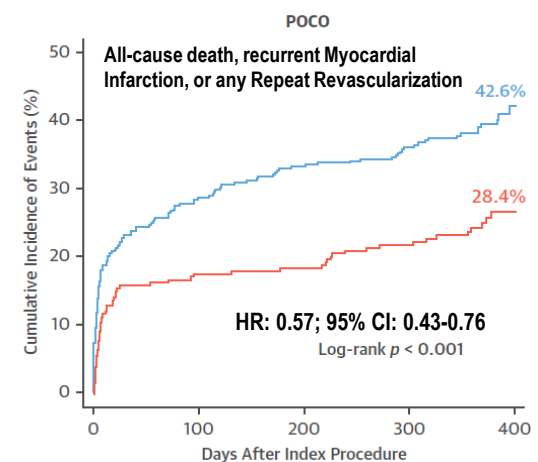
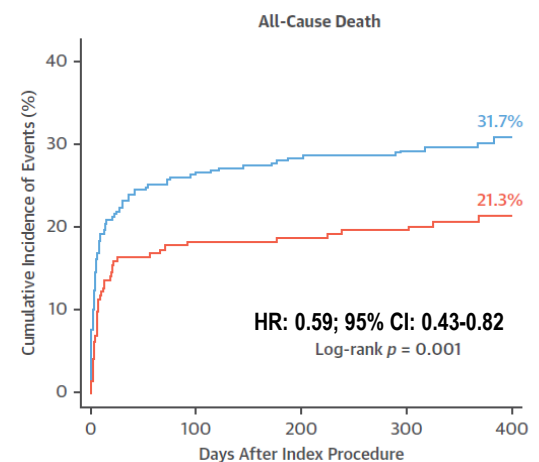
13,104 patients with Acute myocardial infarction

- Exclusion criteria
- NSTEMI, N = 6,804
 - STEMI onset >12h, N = 24
 - No cardiogenic shock, N = 4,923
 - Initial thrombolysis, N = 103
 - Single vessel disease, N = 549
 - Unsuccessful IRA PCI, N = 27
 - Lost to follow-up, N = 15

Study population N = 659
Median follow-up of 359.0 days

IRA-only PCI N = 399 Multivessel PCI N = 260

26.7% Mechanical support
Immediately PCI during Indexprocedure N=157(60.4%)
Staged PCI before discharge N=103(39.6%)



	0	100	200	300	400
IRA-only PCI	399	286	277	267	47
Multivessel PCI	260	213	207	203	37

	0	100	200	300	400
IRA-only PCI	399	275	256	239	42
Multivessel PCI	260	210	203	193	33

Conclusion

Previous abundant evidence exists regarding the **benefits of complete revascularization** for STEMI patients

Even in cardiogenic shock patients too. Of course, **hemodynamic stabilization is a priority.**

Further research is still needed on the best timing of CR.

Recent RCT demonstrated that **FFR-guided CR did not have a significant benefit over an Angiography-guided CR** with respect to the risk of death, MI, urgent revascularization. But, given the **40% less PCI (Time, Cost, contrast, Procedure risk etc.)** in FFR-guided CR.

Thank You
For your attention

