

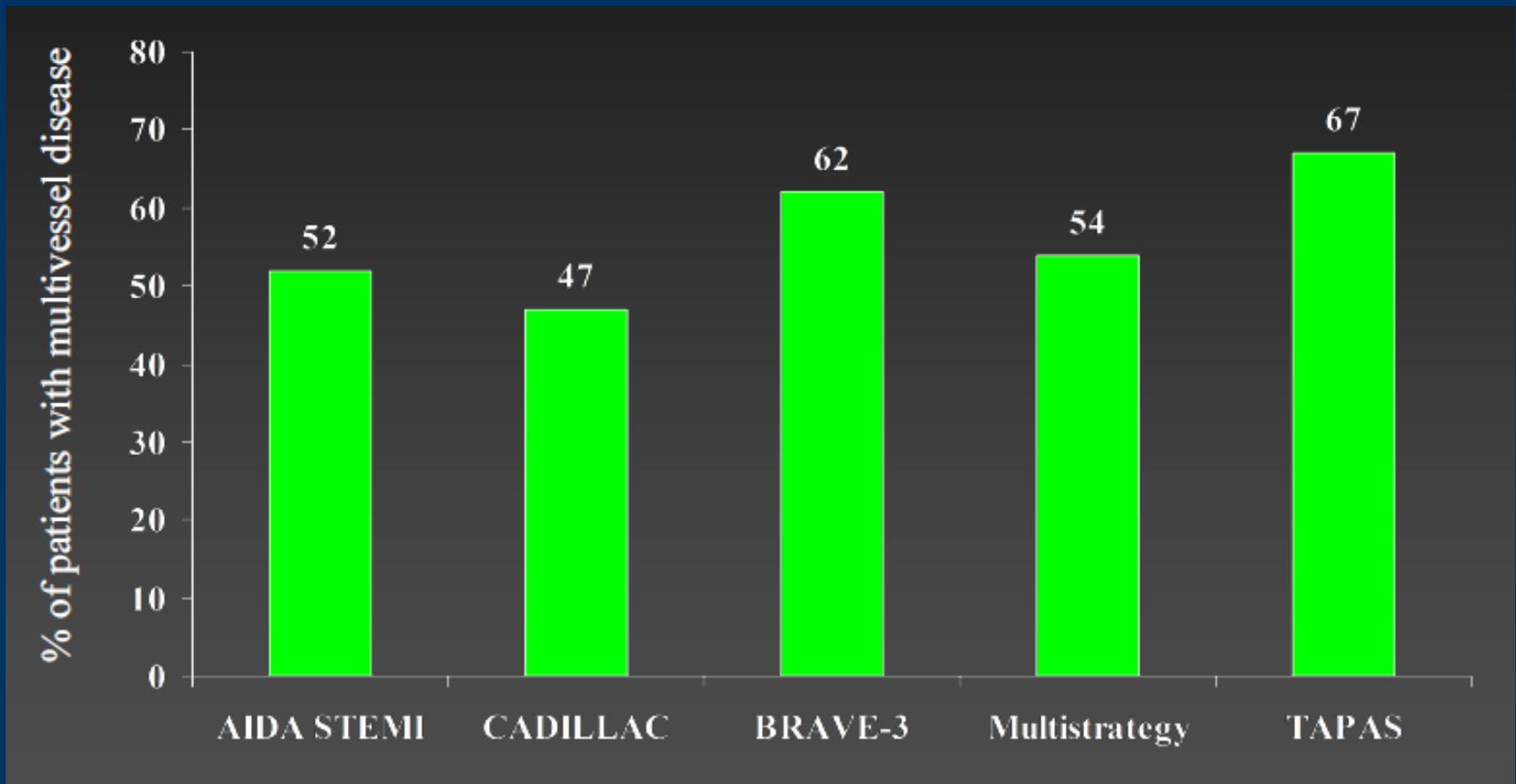


# Impact of Multivessel Revascularization on Health Status Outcomes in ST-segment Elevation Myocardial Infarction Patients

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# Incidence of Multivessel CAD

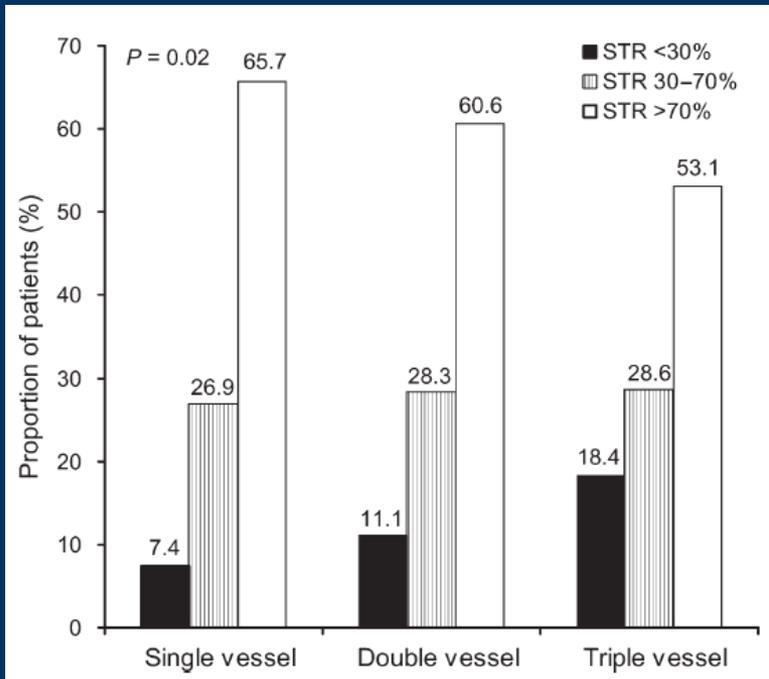
## STEMI without Shock



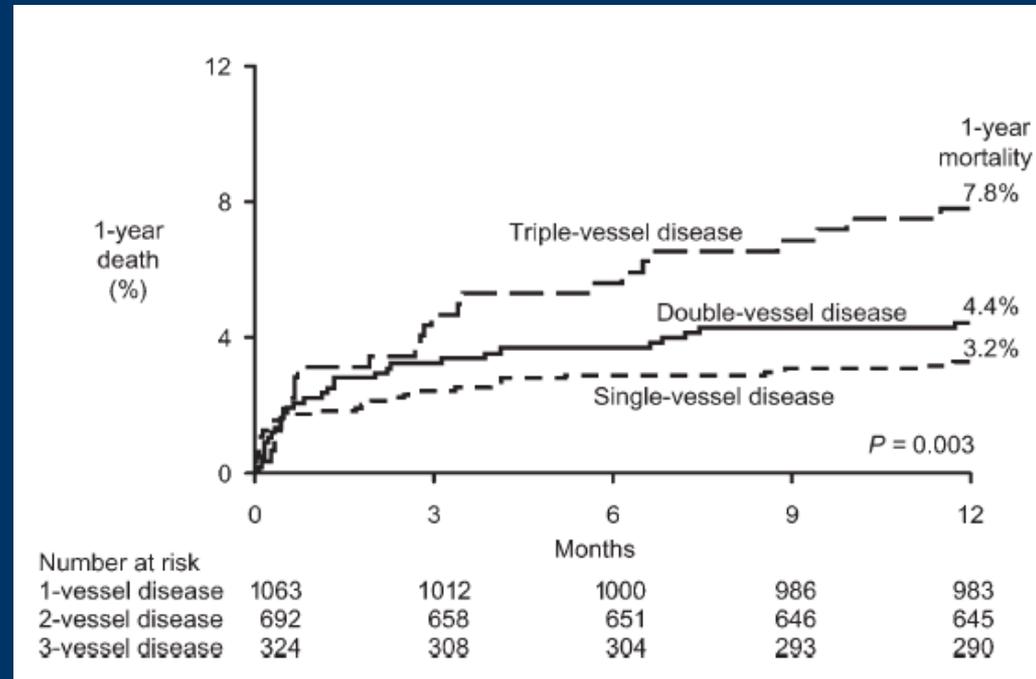
# CADILAC trial

2,082 non-shock STEMI <12 h

## ST-Resolution



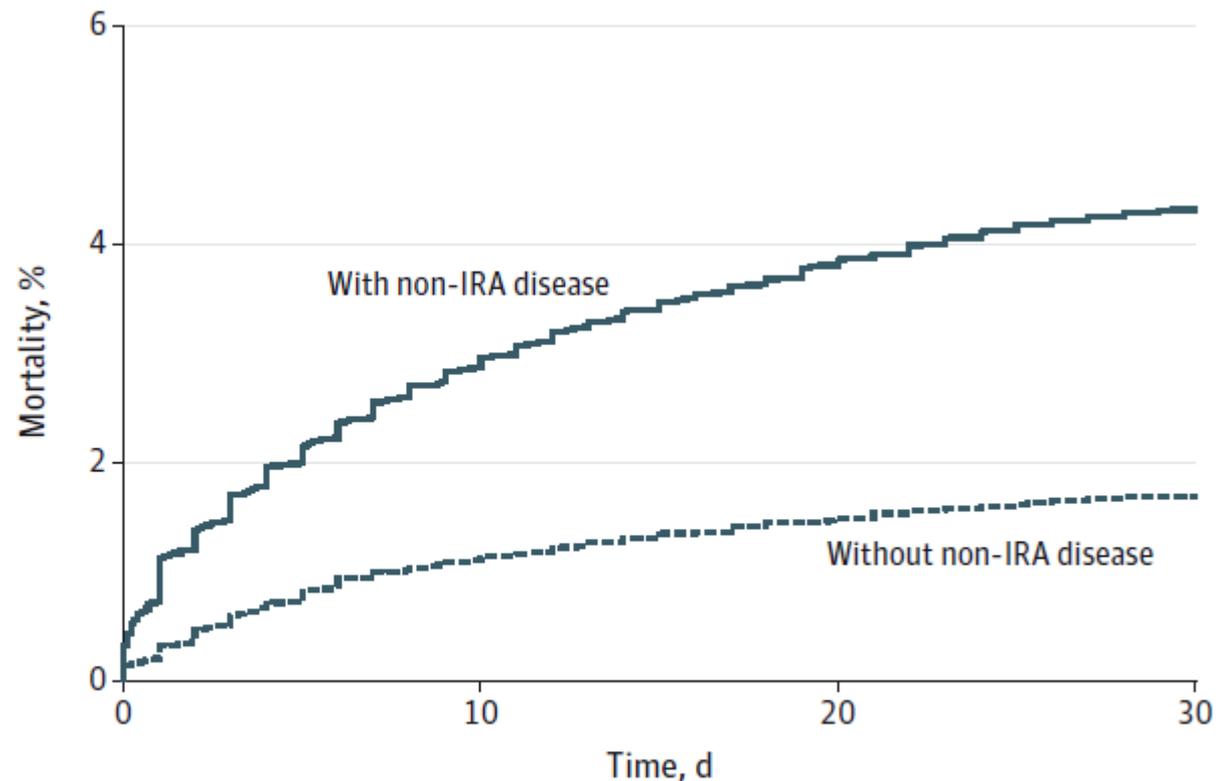
## 1-Year Mortality



# Non-Infarct-Related CAD in STEMI Pts

Pooled Analysis of 28,282 Pts from 8 RCTs

30-day Mortality: With vs. Without *non-IRA disease*



No. at risk	0	10	20	30
With non-IRA disease	14 916	14 479	14 335	14 115
Without non-IRA disease	13 351	13 201	13 141	13 001

# 2014 ESC/EACTS Guidelines on myocardial revascularization

The Task Force on Myocardial Revascularization of the European Society of Cardiology (ESC) and the European Association for Cardio-Thoracic Surgery (EACTS)

## Class IIa: (Benefit > Harm)

1. PCI should be limited to the **culprit vessel** with the exception of cardiogenic shock and persistent ischemia after PCI of the supposed culprit lesion. (*Level of Evidence: B*)
2. **Staged revascularization** of non-culprit lesions should be considered in STEMI patients with multivessel disease in case of symptom or ischemia within days to weeks after primary PCI (*Level of Evidence: B*)

## Class IIb: (Harm > Benefit)

1. **Immediate revascularization** of significant non-culprit lesions during the same procedure as primary PCI of the culprit vessel may be considered in selected patients. (*Level of Evidence: B*)

# 2013 ACCF/AHA Guidelines

**Circulation**  
JOURNAL OF THE AMERICAN HEART ASSOCIATION



**2013 ACCF/AHA Guideline for the Management of ST-Elevation Myocardial Infarction:  
A Report of the American College of Cardiology Foundation/American Heart Association  
Task Force on Practice Guidelines**

## **Class III: Harm**

- 1. PCI should not be performed in a noninfarct artery at the time of primary PCI in patients with STEMI who are hemodynamically stable. (*Level of Evidence: B*)**

*Circulation.* 2013;127:e362-e425

2015 ACC/AHA/SCAI Focused Update on Primary Percutaneous Coronary Intervention for Patients With ST-Elevation Myocardial Infarction: An Update of the 2011 ACCF/AHA/SCAI Guideline for Percutaneous Coronary Intervention and the 2013 ACCF/AHA Guideline for the Management of ST-Elevation Myocardial Infarction

## Class IIb

PCI of a noninfarct artery may be considered in selected patients with STEMI and multivessel disease who are hemodynamically stable, either at the time of primary PCI or as a planned staged procedure. *(Level of Evidence: B-R)*

- Not endorsing *routine* MV PCI in all patients with STEMI and MVD
- Integrate clinical data, lesion severity/complexity, and risk of CIN.

# Strategies for STEMI Pts with MV CAD

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## Culprit Only

- PCI to IRA only

## One-stage Multivessel PCI

- PCI to IRA and other significant non-IRA at primary PCI

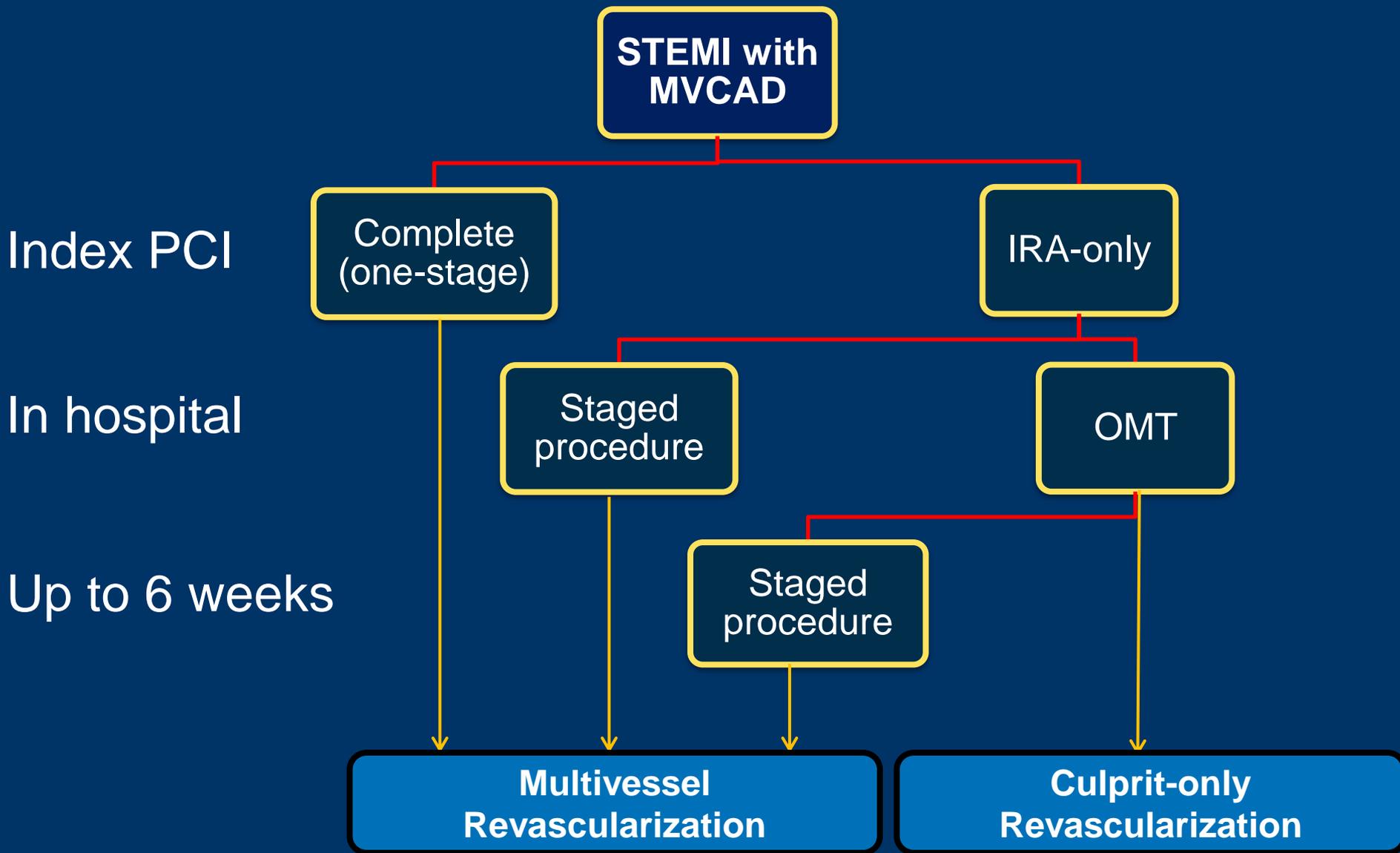
## Staged PCI within 1<sup>st</sup> stay

- PCI to IRA and other significant non-IRA in a later session during index hospital stay

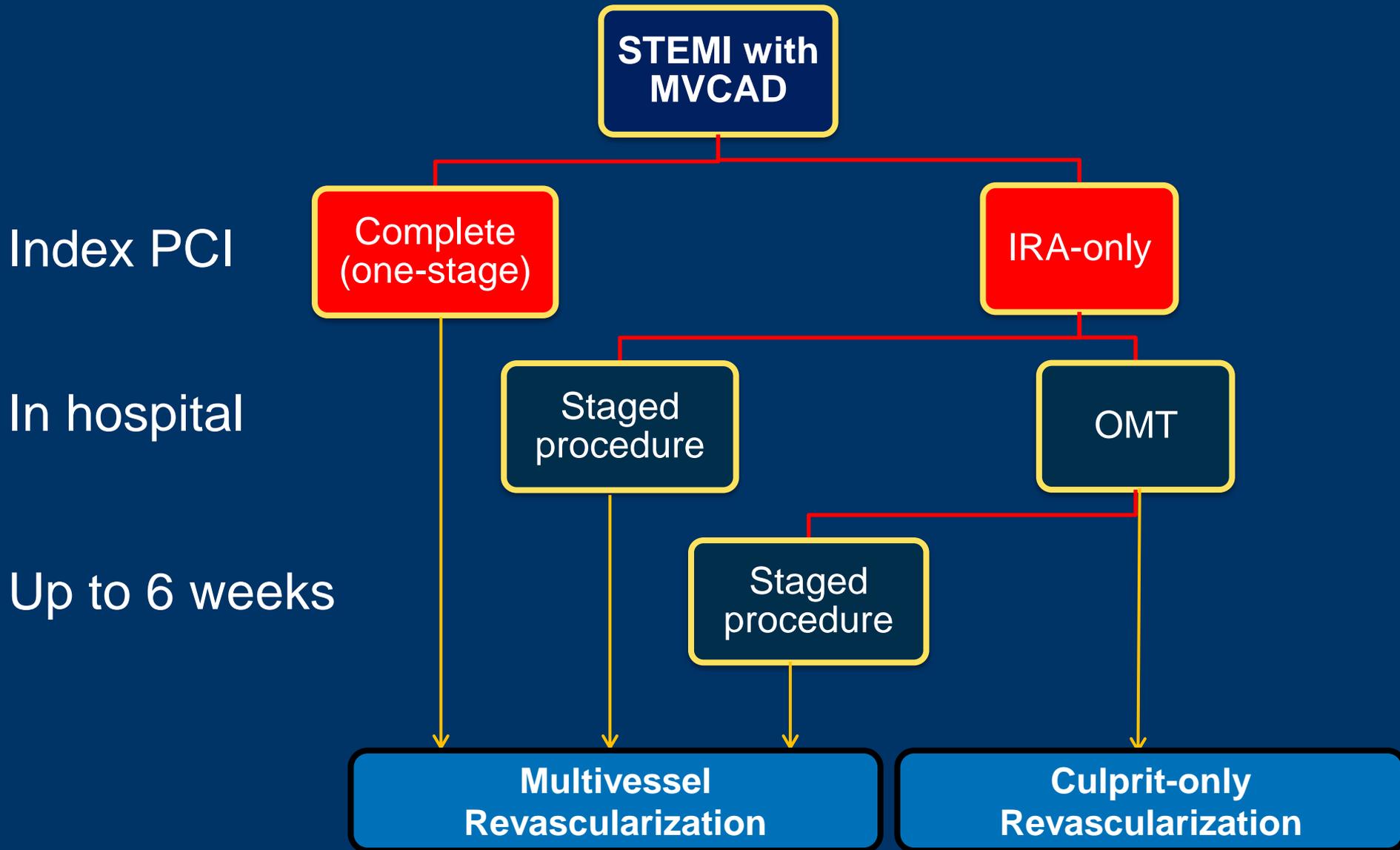
## Staged PCI within 2<sup>nd</sup> stay

- PCI to IRA and other significant non-IRA during 2nd admission

# Timing of Revascularization



# IRA-only vs. One Stage MV PCI



# Intervention of the non-culprit vessels during primary PCI

## ADVANTAGES

- Reduction of total ischemic burden  
; **Better LVEF**
- Treatment of all unstable plaque by treating non-culprit vessel  
; **less future MACE**
- Less future hospitalizations and procedures  
; **cost saving**

## DISADVANTAGES

- Extension of infarcted m.  
; **acute complications (dissection, thrombosis)**
- Lesion severity in non-IRA  
**overestimated**  
(vasoconstriction and endothelial dysfunction)
- **Hemodynamic compromise**
- **Contrast load**

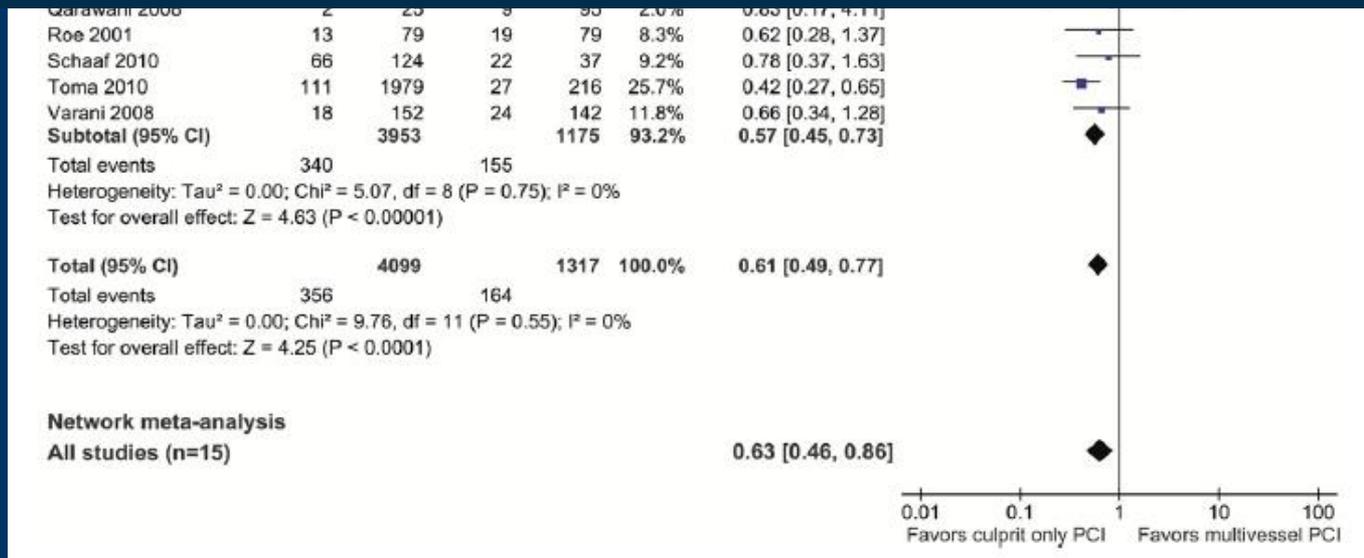
# Culprit only PCI vs. MV-PCI

## Pairwise and Network Meta-Analysis

**A**

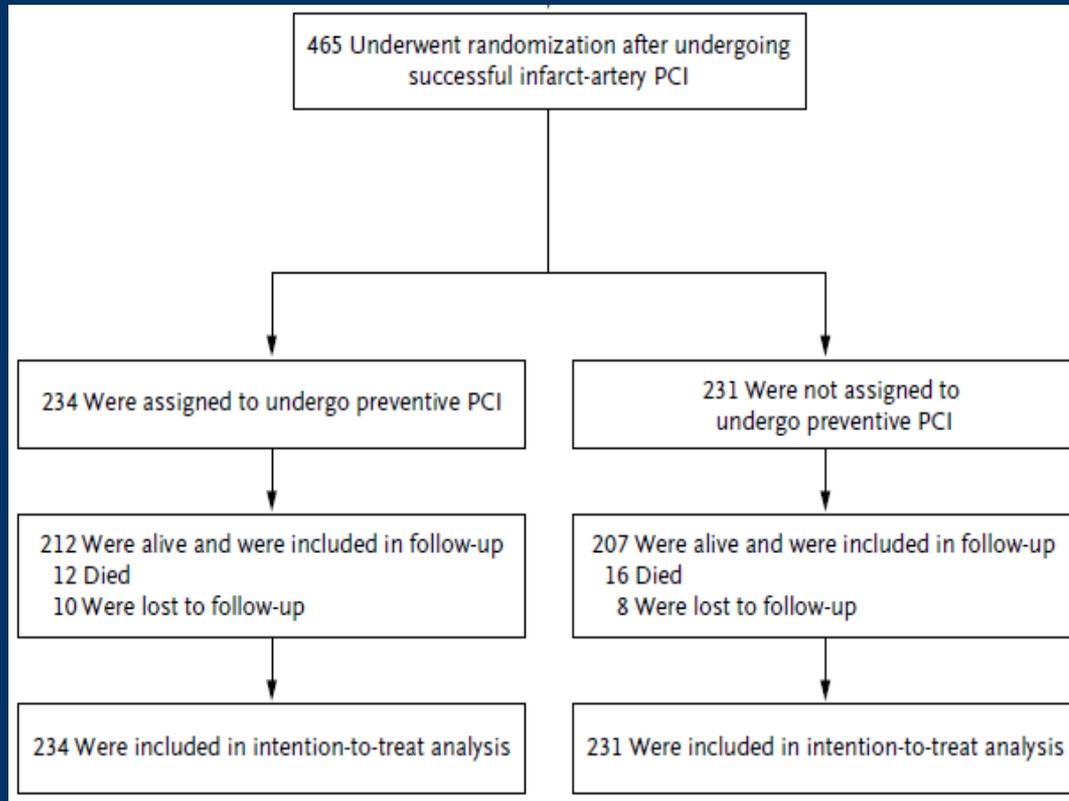
Study or Subgroup	Culprit only PCI		Multivessel PCI		Weight	Odds Ratio IV, Random, 95% CI	Odds Ratio IV, Random, 95% CI
	Events	Total	Events	Total			
<b>Prospective studies</b>							
Di Mario 2004	0	17	1	52	0.5%	0.98 [0.04, 25.20]	

**Long-term mortality favors  
Culprit Only PCI**



# PRAMI: “Preventative” PCI of Non-culprit Lesions after Culprit Lesion Primary PCI in STEMI

465 non-shock STEMI pts with MVD at 5 UK sites

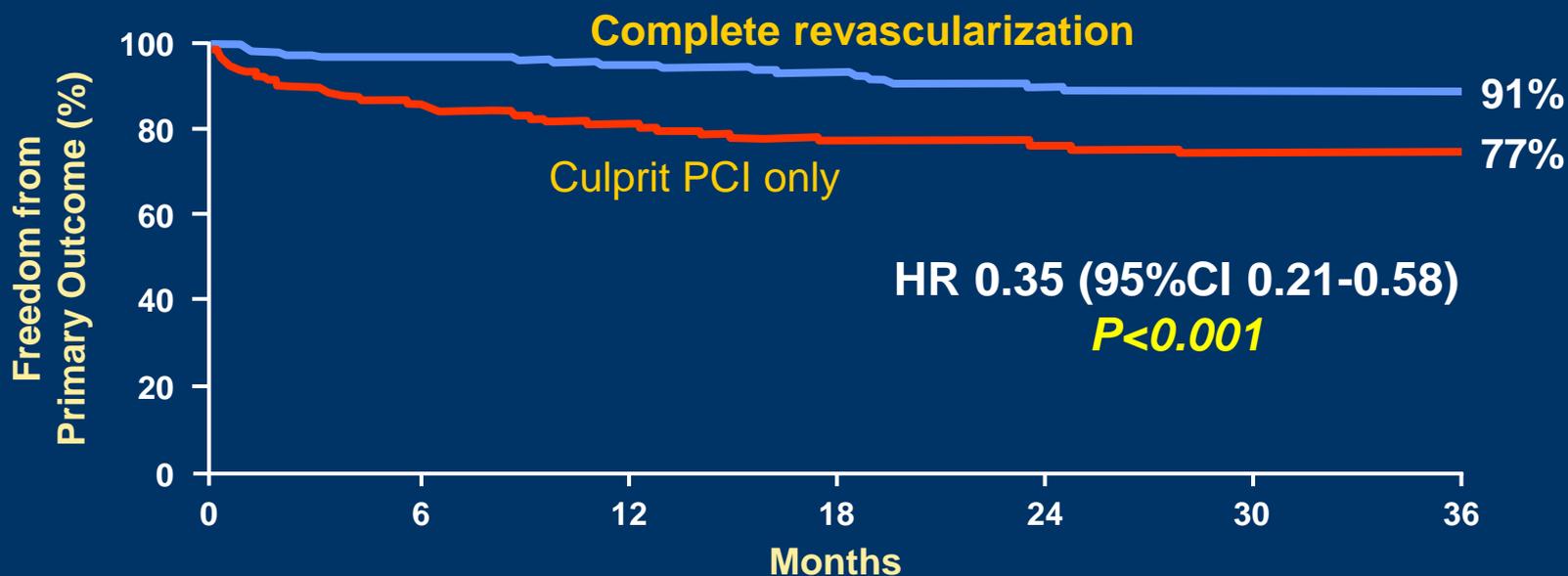


- **Staged PCI** in patients without angina was discouraged
- Further PCI only in cases of refractory angina

Primary endpoint: Cardiac death, MI or refractory angina

# PRAMI: “Preventative” PCI of Non-culprit Lesions after Culprit Lesion Primary PCI in STEMI

465 non-shock STEMI pts with MVD at 5 UK sites

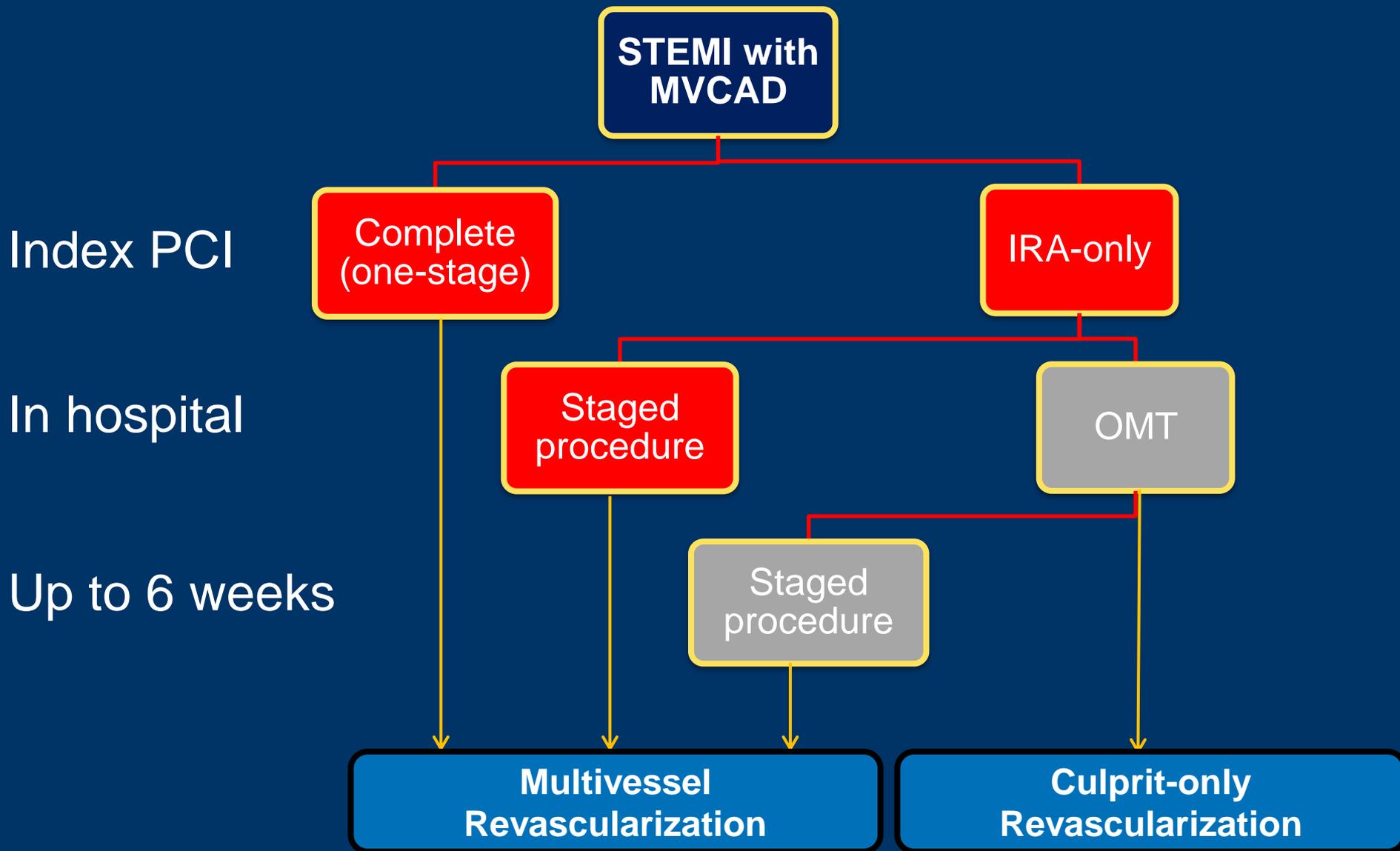


## No. at Risk

	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

600 pts planned; DSMB stopped trial early after 465 pts enrolled (2008-2013)

# IRA-only vs. MV PCI before discharge



# Should we intervene the non-culprit vessels as a Staged-PCI?

## ADVANTAGES

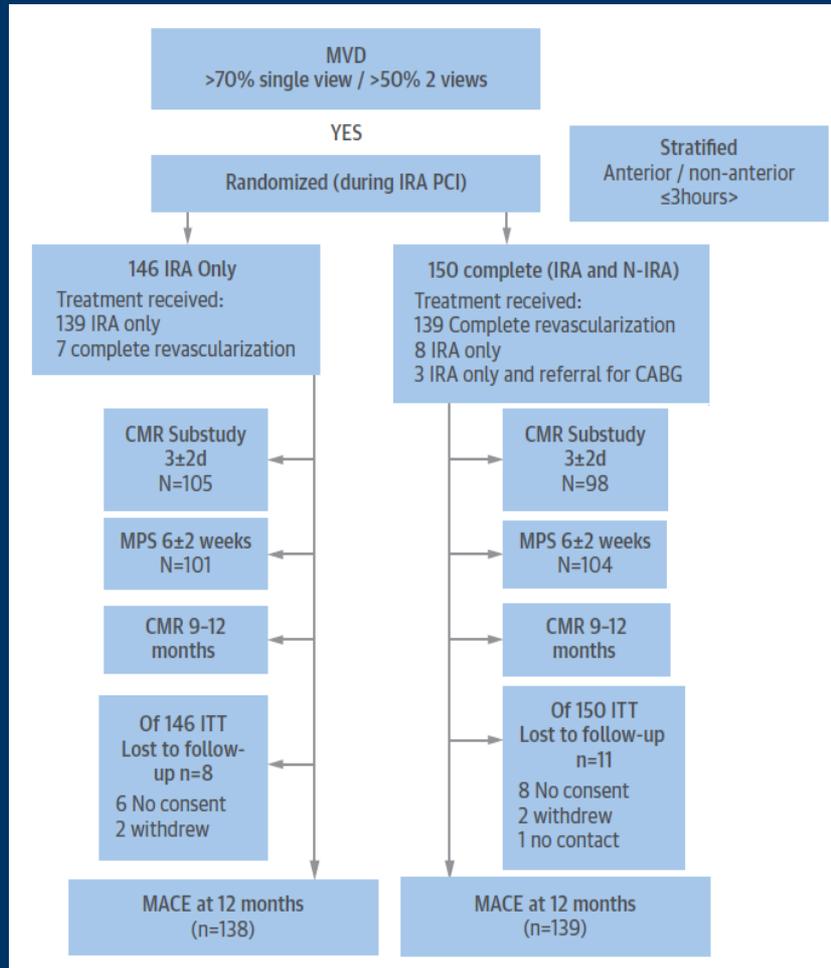
- Reduction of total ischemic burden  
; **Better LVEF**
- Treatment of all unstable plaque by treating non-culprit vessel  
; **less future MACE**
- **Increased safety** in stabilized patients

## DISADVANTAGES

- Increased **cost** by additional admission
- No proven benefit if patients are **asymptomatic**
- Expose patients to further **complication** of PCI
- **Uncertain timing** of 2<sup>nd</sup> PCI/admission

# CvLPRIT: Complete vs. Lesion-Only Primary PCI trial

## 296 STEMI pts at 7 UK Centers



- **CR Group**

- Treat IRA first
- CR recommended at same setting
- Staged procedure during the index admission

# CvLPRIT: Complete vs. Lesion-Only Primary PCI trial

## 296 STEMI pts at 7 UK Centers Clinical Outcomes at 12 Months

Variable	IRA only (N=146)	Complete Revascularisation (N=150)	HR (95% CI)	P value
<b>Time to First Event</b>				
MACE N= (%)	31 (21.2)	15 (10.0)	0.45 (0.24, 0.84)	0.009
Components N=(%)				
All-cause mortality	6 (4.1)	2 (1.3)	0.32 (0.06, 1.60)	0.14
Recurrent MI	4 (2.7)	2 (1.3)	0.48 (0.09, 2.62)	0.39
Heart failure	9 (6.2)	4 (2.7)	0.43 (0.13, 1.39)	0.14
Repeat Revascularisation	12 (8.2)	7 (4.7)	0.55 (0.22, 1.39)	0.2

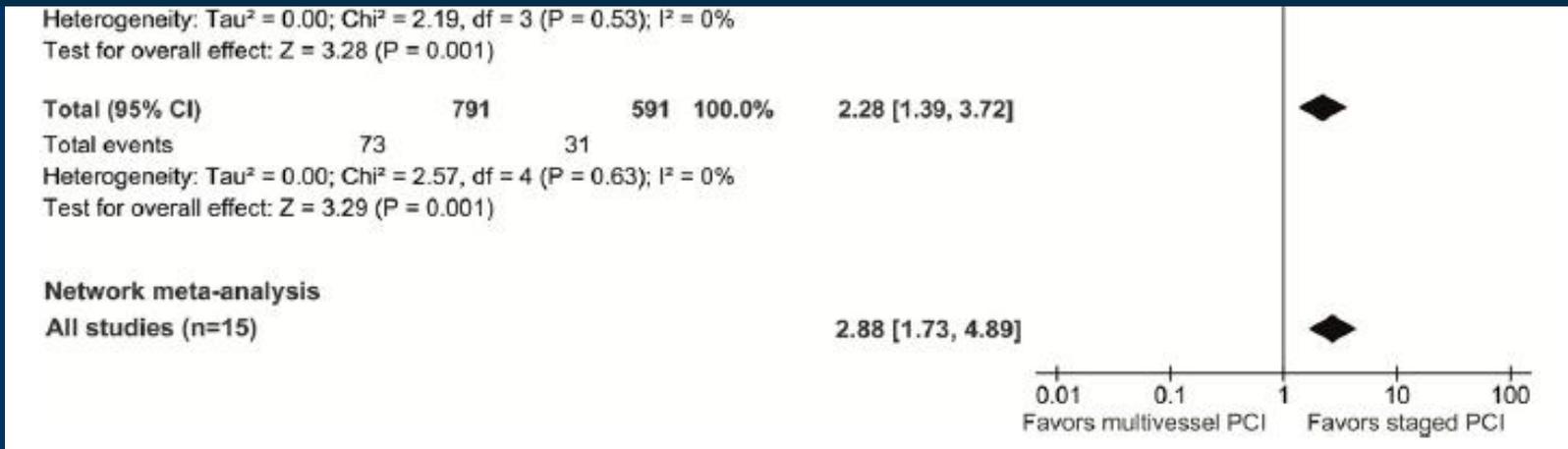
***55% reduced hazards of MACE by  
complete revascularization***

# MV-PCI vs. Staged PCI

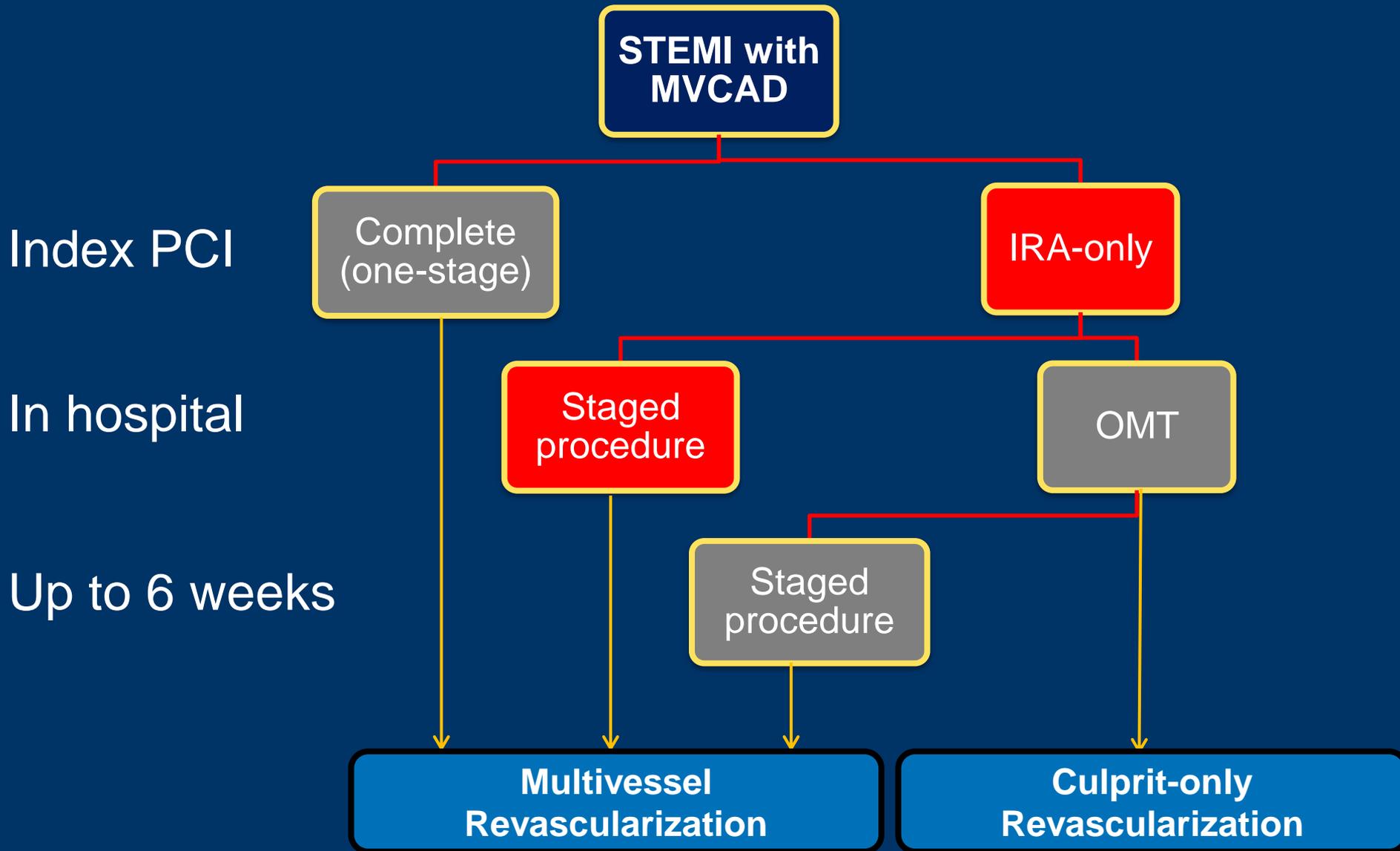
## Pairwise and Network Meta-Analysis



**Long-term mortality favors Staged PCI**



# IRA-only vs. In-hospital Staged PCI



# DANAMI3-PRIMULTI

627 STEMI Pts with MVD from 2011 to 2014

627 Multivessel disease  
(>50% stenosis in non IRA > 2 mm suitable for PCI)

Randomise

313 IRA PCI only

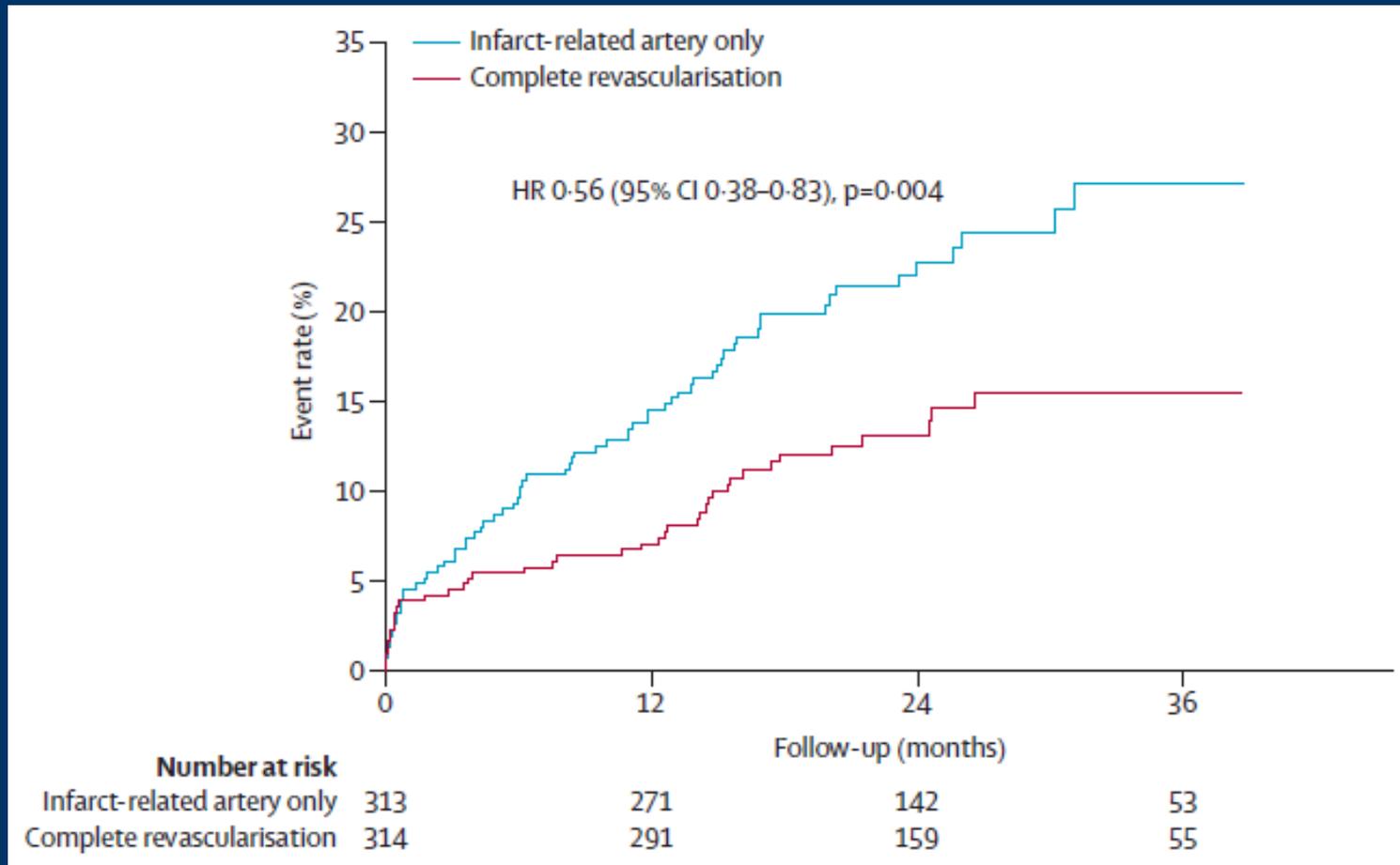
314 FFR guided complete  
revascularisation

\* Additional PCI procedures 2 days  
after the initial PCI before discharge

# DANAMI3-PRIMULTI

627 STEMI Pts with MVD from 2011 to 2014

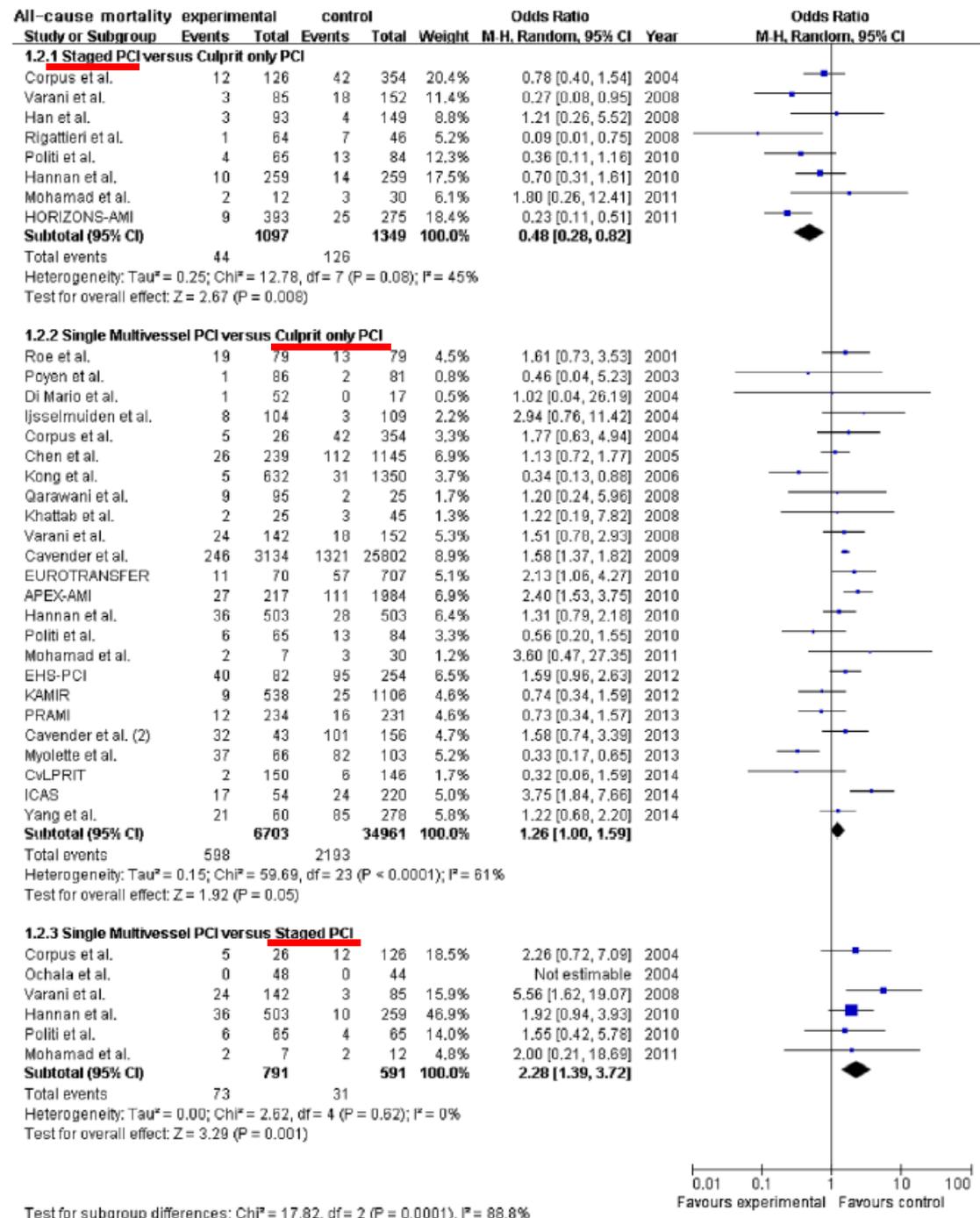
All-cause mortality, NFM, and Ischemia-driven revascularization



# Updated Meta-Analysis

## All-cause death

- ❖ Culprit only PCI < Staged
- ❖ One-time < Culprit only PCI
- ❖ One-time < Staged PCI



# Background

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- Previous studies have focused upon mortality and **no insights into the long-term health status** of STEMI patients managed with culprit-only or complete revascularization have been reported.

# Aims & Objectives

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- To define the potential patient-centered benefits of complete revascularization
  - the patterns of treating non-infarct vessels
  - patient characteristics associated with multivessel revascularization
  - variation in practice across hospitals
  - independent association of multivessel revascularization with 1-year health-related QoL and mortality

# Methods

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**TRIUMPH patients (All AMI patients,  
April 2005-December 2008), N = 4,340**



**Exclude patients with:**

- Prior CABG (n=537)
- No CAG, not MVCAD (n=2,218)
- No PCI (n=511)
- In-hospital death (n=4)
- NSTEMI (n=406)

**Final Study Population  
664, STEMI patients with multivessel CAD**

# QoL Outcomes

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- **Seattle Angina Questionnaire (baseline/1-year);**
  - 19-item patient-reported health status instrument
  - recall period of 4 weeks
- **Angina Frequency (SAQ AF)**
- **Quality of Life (SAQ QoL)**
- Physical Limitation
- Treatment Satisfaction

# Clinical Outcomes

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- **All-cause mortality:** phone follow-up and the Social Security death master index
- **Myocardial infarction**
- **Repeat revascularization procedures:** PCI or CABG
- **Severe angina:** having more than 3 episodes of angina per week as defined by a SAQ AF score of  $\leq 40$

# Statistical Analysis (1)

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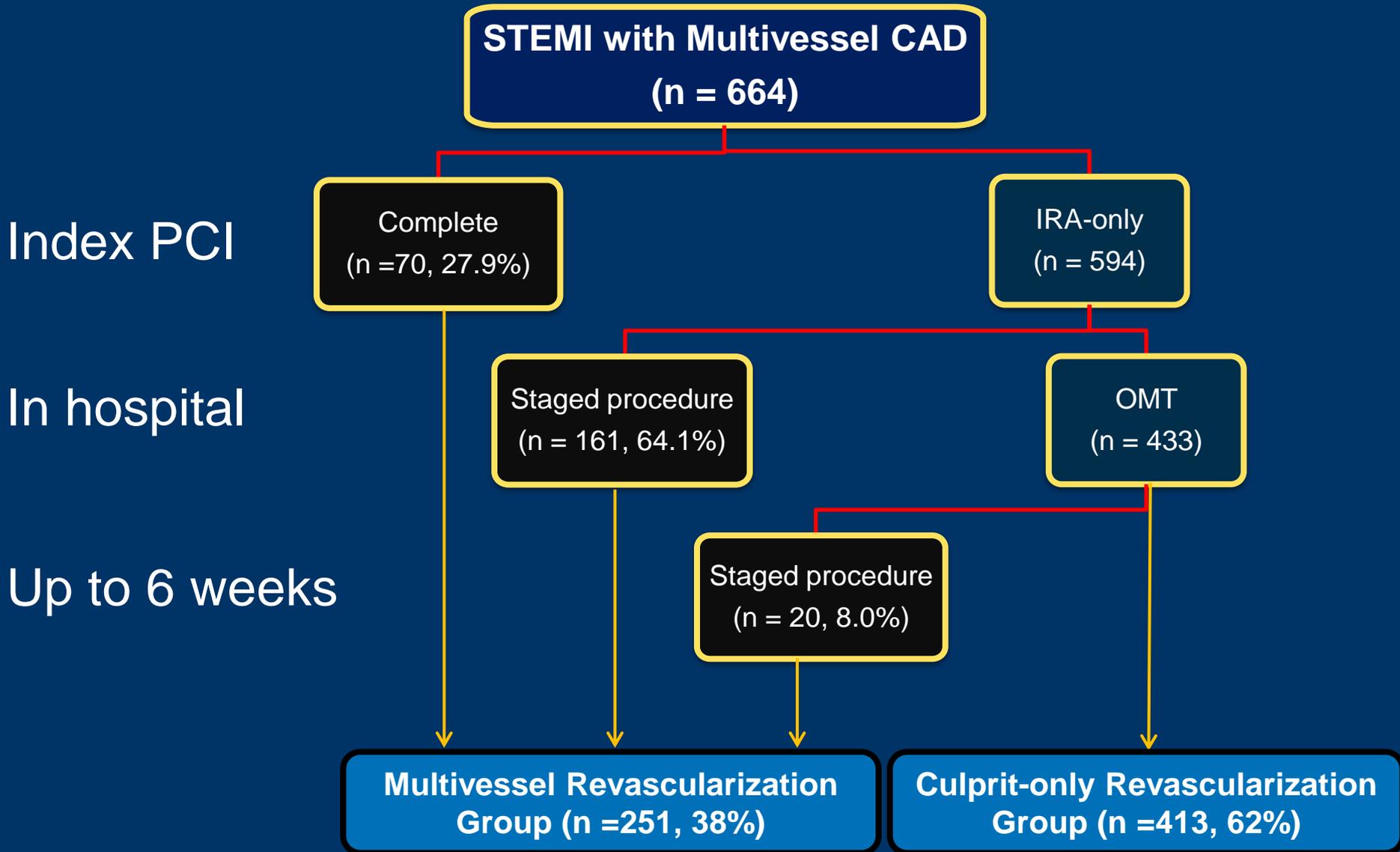
- Baseline clinical and demographic characteristics using **t-test or Mann-Whitney U tests** for continuous variables and **chi-square or Fisher's exact test** for categorical variables
- **Multivariable, hierarchical** (adjusting for site as a random effect) **modified Poisson regression model** to identify factors associated with multivessel revascularization
- **Median rate ratio (MRR)** to assess variation in the practice of multivessel revascularization across the study sites

# Statistical Analysis (2)

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- Hierarchical (adjusting for site as a random effect) **multivariable linear regression models** for each health status outcome (SAQ AF and SAQ QoL) to evaluate the association of multivessel revascularization with 1-year health status outcomes
- **Kaplan-Meier survival analysis** and the log-rank test to assess the associations of multivessel revascularization with 1-year mortality, myocardial infarction, and repeat revascularization
- **Sensitivity analysis** excluding patients undergoing CABG to determine whether our results were comparable in the PCI-only patients

# Timing of Revascularization



# Baseline Demographics

	Multivessel n = 251 (38%)	Culprit-only n = 413 (62%)	P-Value
Age	56.4 ± 10.0	58.7 ± 12.1	0.012
Caucasian	197 (79.4%)	299 (72.7%)	0.053
Female gender	62 (24.7%)	108 (26.2%)	0.678
Insurance: None/Self-Pay	58 (23.7%)	92 (22.9%)	0.830
Diabetes	59 (23.5%)	106 (25.7%)	0.532
Hypertension	144 (57.4%)	242 (58.6%)	0.756
Dyslipidemia	115 (45.8%)	190 (46.0%)	0.962
Prior PCI	29 (11.6%)	78 (18.9%)	0.012
Chronic heart failure	5 (2.0%)	8 (1.9%)	1.000
Peripheral vascular disease	5 (2.0%)	16 (3.9%)	0.179
Smoking	156 (62.2%)	252 (61.0%)	0.770
In-hospital heart failure	15 (6.0%)	15 (3.6%)	0.158
LV dysfunction (EF < 40%)	43 (18.4%)	79 (21.0%)	0.438
Peak troponin I/T (ng/dL): (Median)	23.8	13.7	< 0.001
Hemoglobin (g/dL): Initial (Median)	15.0	14.7	0.035
Systolic BP (mmHg): Initial (Median)	142.0	140.0	0.443

# Baseline Demographics

	Multivessel n = 251 (38%)	Culprit-only n = 413 (62%)	P-Value
Number of diseased vessels	2.5 ± 0.7	2.4 ± 0.6	< 0.001
Number of vessels treated	1.9 ± 0.6	1.0 ± 0.0	< 0.001
Distribution of culprit vessels			< 0.001
Left main coronary artery	1 (0.4%)	3 (0.7%)	
Proximal LAD artery	45 (17.9%)	31 (7.5%)	
Mid to distal LAD artery	42 (16.7%)	116 (28.1%)	
Left circumflex artery	23 (9.2%)	50 (12.1%)	
Right coronary artery	109 (43.4%)	181 (43.8%)	
LAD artery culprit	87 (34.7%)	147 (35.6%)	0.807
Distribution of non-culprit vessels			
Left main coronary artery	10 (4.0%)	18 (4.4%)	0.815
Proximal LAD artery	25 (10.0%)	12 (2.9%)	< 0.001
Mid to distal LAD artery	119 (47.4%)	169 (40.9%)	0.101
Left circumflex artery	112 (44.6%)	206 (49.9%)	0.188
Right coronary artery	74 (29.5%)	134 (32.4%)	0.424
Number of bare-metal stents	0.8 ± 1.2	0.7 ± 0.9	0.556
Number of drug-eluting stents	1.6 ± 1.6	0.7 ± 0.9	< 0.001

# Baseline and 1-year Health Status

	Multivessel n = 251 (38%)	Culprit-only n = 413 (62%)	P-Value
SAQ AF score (baseline)	89.6 ± 17.1	89.2 ± 16.8	0.77
SAQ AF score (1 year)	94.8 ± 14.2	92.8 ± 17.4	0.20
Mean changes in SAQ AF	5.2 ± 22.4	3.2 ± 20.8	0.34
SAQ QoL score (baseline)	62.3 ± 20.9	68.5 ± 22.9	< 0.001
SAQ QoL score (1 year)	85.0 ± 18.3	81.5 ± 20.7	0.07
Mean changes in SAQ QoL	22.3 ± 24.9	12.7 ± 26.5	< 0.001

# Independent Correlates of MV Revascularization

## Age

- 40 vs. 50

- 60 vs. 50

- 70 vs. 50

- 80 vs. 50

Female

Caucasian

Avoid care due to costs

## Disease vessels (per 1 increment)

Non-LAD culprit vessel

History of CHF

In-hospital heart failure

History of atrial fibrillation

History of diabetes mellitus

LV systolic dysfunction (moderate or severe)

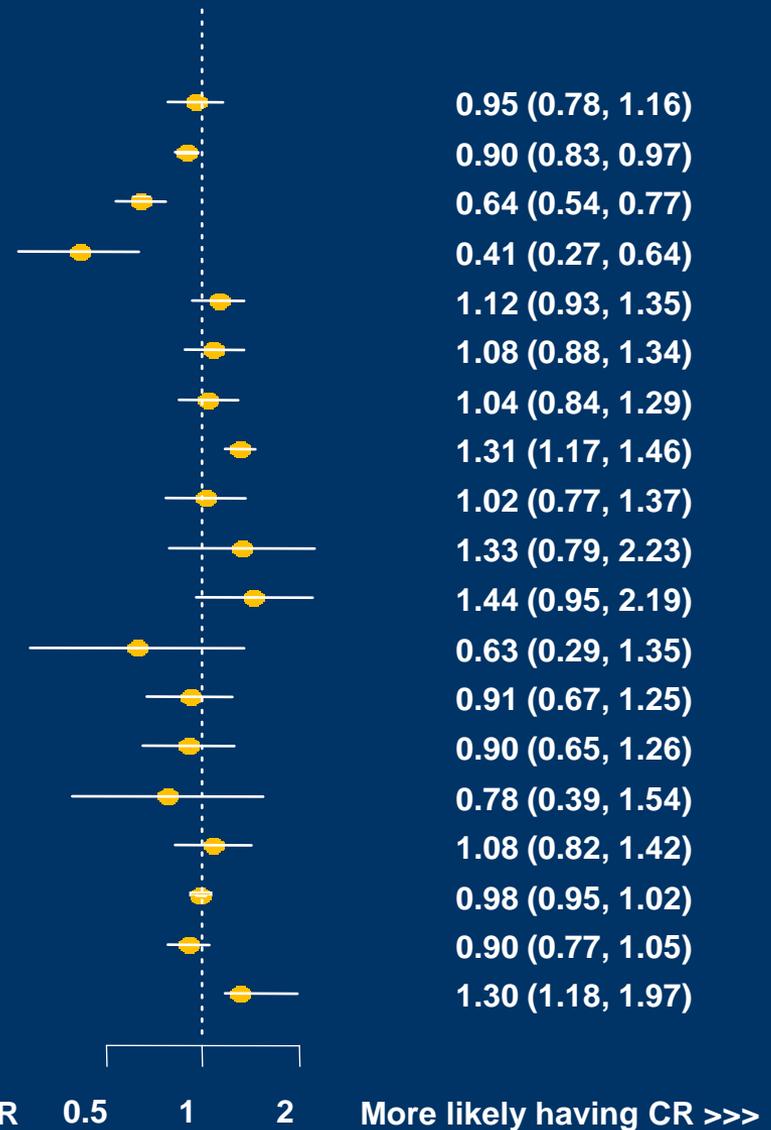
Initial creatinine (per 5 units increment)

Initial hemoglobin (per 5 units increment)

SF-12 PCS (per 5 units increment)

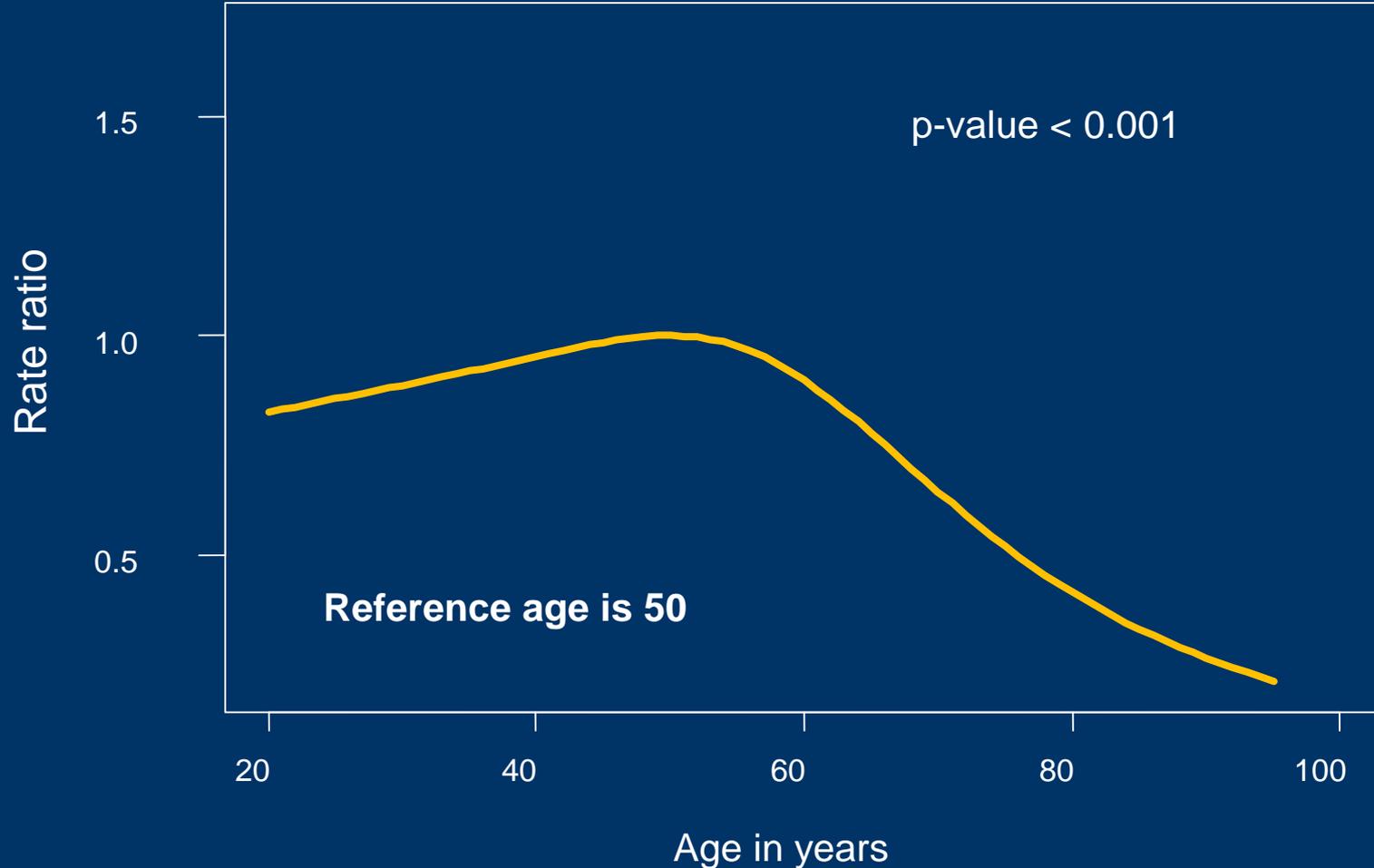
Angina at baseline

## Hospital site (median rate ratio)



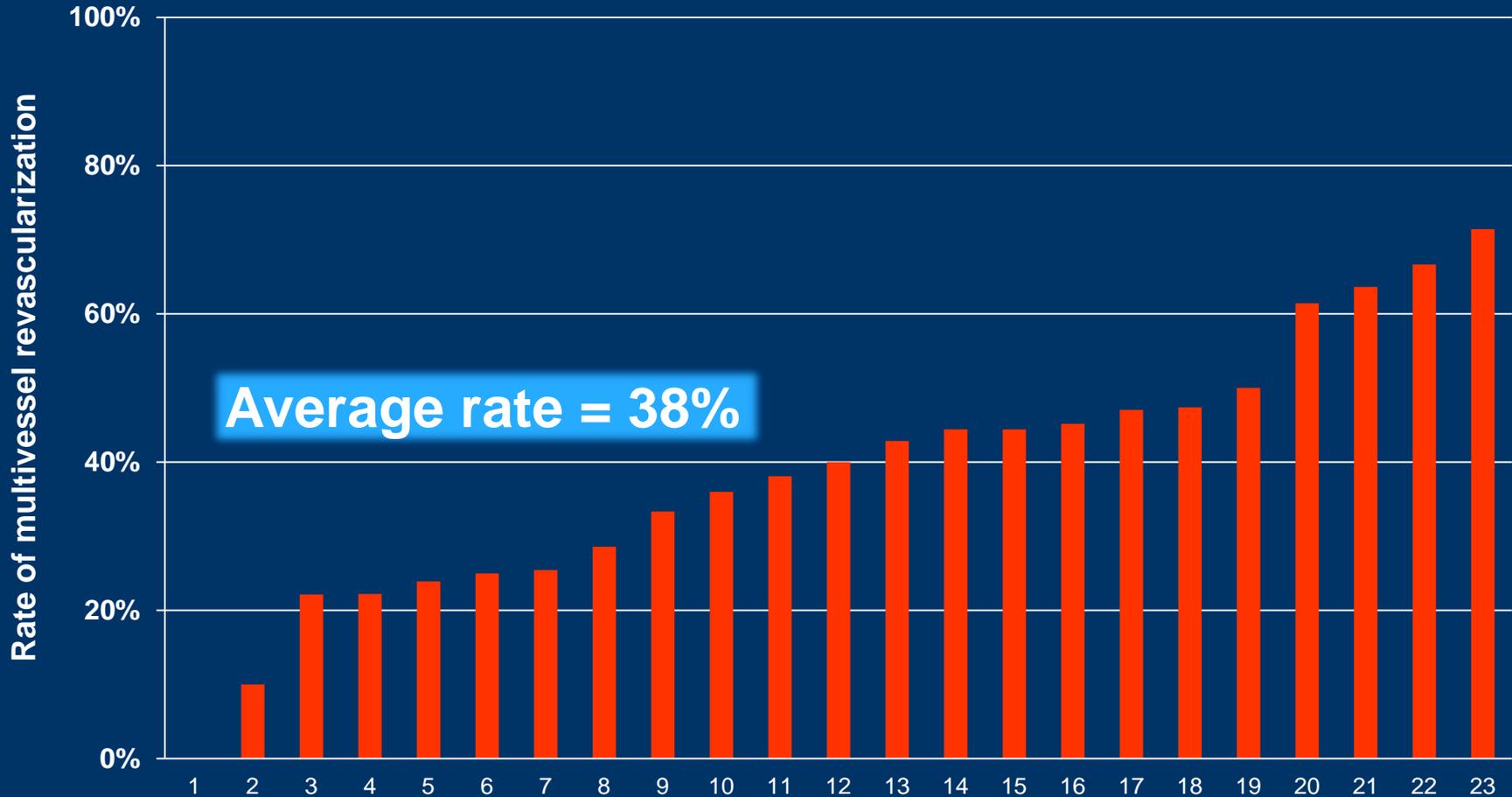
# Age and Likelihood of MV Revascularization

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# Hospital Variation of MV Revascularization

Median Rate Ratio = 1.30 (95% CI 1.18 to 1.97)

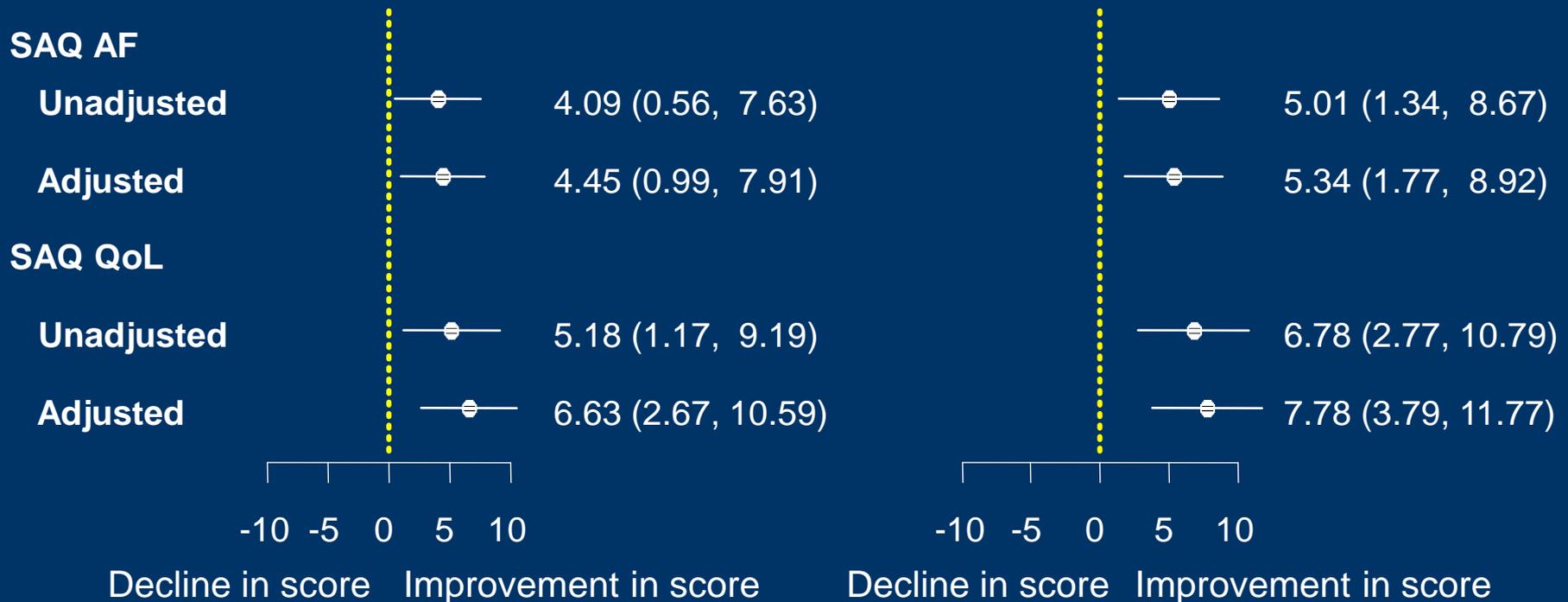


# Health status outcomes

## Multivessel vs. culprit-only revascularization

Entire cohort (PCI + CABG)

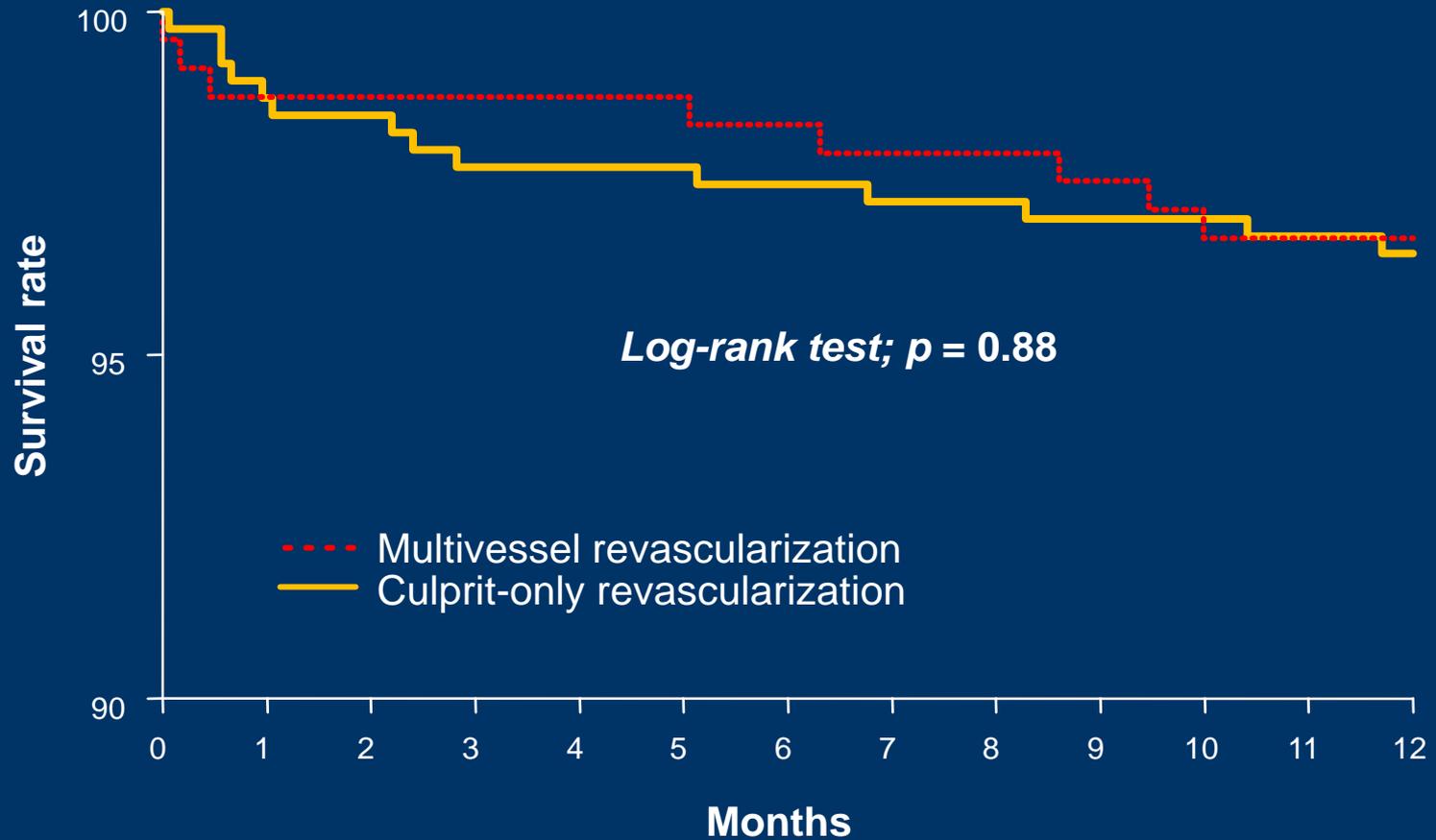
PCI only cohort



# Clinical Outcomes at 1 year

	Multivessel n = 251	Culprit-only n = 413	P-Value
<b>Mortality</b>	8 (3.6%)	14 (3.4%)	0.88
<b>Recurrent MI</b>	7 (3.5%)	4 (1.4%)	0.12
<b>Repeat revascularization</b>	17 (7.5%)	32 (9.1%)	0.50
<b>Severe angina</b>	10 (4.4%)	22 (6.3%)	0.34

# K-M Curves of 1-year Mortality



# Study Limitations

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- Potential for unmeasured confounding or selection bias
- No query to the clinicians as to why complete or culprit-only revascularization was performed
- Missing SAQ data in 1/3 of patients at 1 year
- No angiographic core laboratory assessing the severity of CAD
- Excluded in-hospital death. Cannot be extrapolated to extremely sick patients

# Conclusions

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- In a large, multicenter AMI registry,  
**Multivessel, complete revascularization in STEMI setting;**
  - common (n=251, 38% among 664 patients)
  - varied by patient characteristics and the treating hospital
  - improved both angina and QoL at 1 year
- **Future studies** of the potential benefits and harms of multivessel revascularization in STEMI patients should include **both symptoms and health-related QoL outcomes** so that more complete insights into the benefits of multivessel revascularization can be assessed

# Impact of Multivessel Revascularization on Health Status Outcomes in Patients With ST-Segment Elevation Myocardial Infarction and Multivessel Coronary Artery Disease



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## ABSTRACT

**BACKGROUND** Up to 65% of patients with ST-segment elevation myocardial infarction (STEMI) have multivessel coronary artery disease (MVCAD). Long-term health status of STEMI patients after multivessel revascularization is unknown.

**OBJECTIVES** This study investigated the relationship between multivessel revascularization and health status outcomes (symptoms and quality of life [QoL]) in STEMI patients with MVCAD.

**METHODS** Using a U.S. myocardial infarction registry and the Seattle Angina Questionnaire (SAQ), we determined the health status of patients with STEMI and MVCAD at the time of STEMI and 1 year later. We assessed the association of multivessel revascularization during index hospitalization with 1-year health status using multivariable linear regression analysis, and also examined demographic, clinical, and angiographic factors associated with multivessel revascularization.

**RESULTS** Among 664 STEMI patients with MVCAD, 251 (38%) underwent multivessel revascularization. Most revascularizations were staged during the index hospitalization (64.1%), and 8.0% were staged after discharge, with 27.9% performed during primary percutaneous coronary intervention. Multivessel revascularization was associated with age and more diseased vessels. At 1 year, multivessel revascularization was independently associated with improved symptoms (4.5 points higher SAQ angina frequency score; 95% confidence interval [CI]: 1.0 to 7.9) and QoL (6.6 points higher SAQ QoL score; 95% CI: 2.7 to 10.6). One-year mortality was not different between those who did and did not undergo multivessel revascularization (3.6% vs. 3.4%; log-rank test  $p = 0.88$ ).

**CONCLUSIONS** Multivessel revascularization improved angina and QoL in STEMI patients with MVCAD. Patient-centered outcomes should be considered in future trials of multivessel revascularization.

(J Am Coll Cardiol 2015;66:2104-13) © 2015 by the American College of Cardiology Foundation.

**Thank you for your attention!**