# Controversies In STEMI Management

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Assembly of International Governors of ACC in Middle east & Africa

Stent Save a life Regional Africa Board

Chairman of ICC Hospital, Alexandria

### Disclosure Statement of Financial Interest

I, Mohamed Sobhy DO NOT have a financial interest/arrangement or affiliation with one or more organizations that could be perceived as a real or apparent conflict of interest in the context of the subject of this presentation.

### Controversies In STEMI

- Culprit versus complete revascularization in MVD
- Culprit versus complete revascularization in cardiogenic shock

## Controversies In STEMI

- Culprit versus complete revascularization in MVD
- Culprit versus complete revascularization in cardiogenic shock

# Multivessel Stenting in STEMI

# Senario 1

# Recent posterior STEMI





# Management

**Culprit only** 

# PCI LCX (DES)

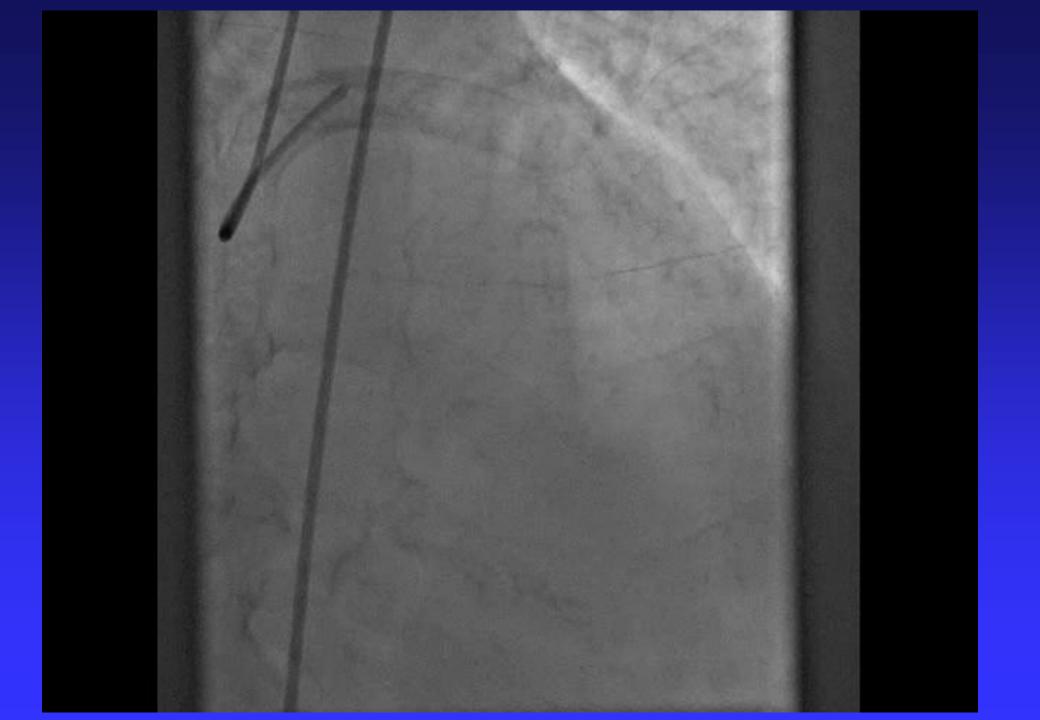


# **Senario 2**

**Anterior STEMI & double infarction** 

### **Cath Lab**





## Management

**Complete Revascularization** 

# PCI LAD (DES)



# PCI RCA (DES)



# Senario 3

Inferior & RV STEMI MVD NO Cardiogenic shock





## Management

What to do???

# PCI LAD (2DES)



# PCI RCA (DES) When to do it?!



### Background

- 30-50% of STEMI patients have additional stenoses other than the infarct related artery<sup>1,2</sup>
- Current guidelines support culprit vessel PCI only
- Contemporary studies have, however, suggested preventive revascularization <sup>3,4</sup>

<sup>&</sup>lt;sup>1</sup> Jong JA al. Coronary Artery disease 2006

<sup>&</sup>lt;sup>2</sup> Muller DW et al. Am Heart J 1991

<sup>&</sup>lt;sup>3</sup> Wald et al. NEJM 2013

<sup>&</sup>lt;sup>4</sup> Gershlick et al. ESC 2014

### **PRAMI**

The NEW ENGLAND JOURNAL of MEDICINE

#### ORIGINAL ARTICLE

# Randomized Trial of Preventive Angioplasty in Myocardial Infarction

David S. Wald, M.D., Joan K. Morris, Ph.D., Nicholas J. Wald, F.R.S., Alexander J. Chase, M.B., B.S., Ph.D., Richard J. Edwards, M.D., Liam O. Hughes, M.D., Colin Berry, M.B., Ch.B., Ph.D., and Keith G. Oldroyd, M.D., for the PRAMI Investigators\*

#### ABSTRACT

#### **BACKGROUND**

In acute ST-segment elevation myocardial infarction (STEMI), the use of percutaneous coronary intervention (PCI) to treat the artery responsible for the infarct (infarct, or culprit, artery) improves prognosis. The value of PCI in noninfarct coronary arteries with major stenoses (preventive PCI) is unknown.

#### **METHODS**

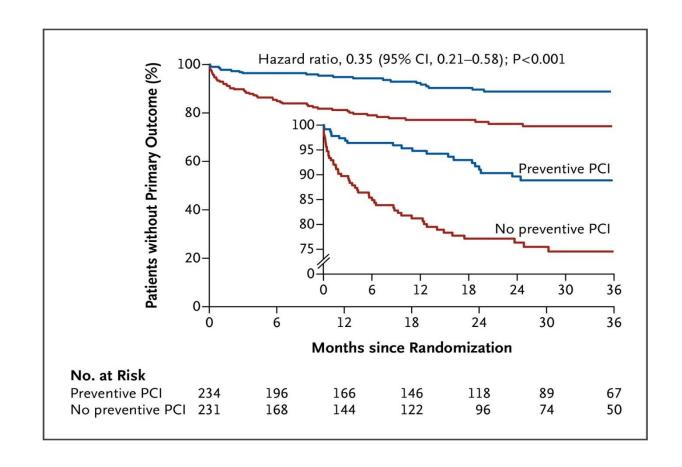
From 2008 through 2013, at five centers in the United Kingdom, we enrolled 465 patients with acute STEMI (including 3 patients with left bundle-branch block) who were undergoing infarct-artery PCI and randomly assigned them to either preven-

From the Wolfson Institute of Preventive Medicine, Barts and the London School of Medicine and Dentistry, Queen Mary University of London (D.S.W., J.K.M., N.J.W.), and London Chest Hospital (D.S.W.), London, Morriston Hospital, Swansea (A.J.C.), Freeman Hospital, Newcastle upon Tyne (R.J.E.), Norfolk and Norwich University Hospital, Norwich (L.O.H.), and Golden Jubilee National Hospital, Glasgow (C.B., K.G.O.) — all in the United Kingdom, Address really and the service of the control of the contr









# Criticism about the study:

- Sample size is small
- Larger number of patients were inferior infarcts
- •EF was not reported in the study
- •The study stopped prematurely???

#### ORIGINAL INVESTIGATIONS

# Randomized Trial of Complete Versus Lesion-Only Revascularization in Patients Undergoing Primary Percutaneous Coronary Intervention for STEMI and Multivessel Disease





#### The CvLPRIT Trial

Anthony H. Gershlick, MBBS,\* Jamal Nasir Khan, MB ChB,\* Damian J. Kelly, MB ChB, MD,†

John P. Greenwood, MB ChB, PhD,‡§ Thiagarajah Sasikaran, BSc, PhD,|| Nick Curzen, BM, PhD,¶

Daniel J. Blackman, MD,§ Miles Dalby, MBBS, MD,# Kathryn L. Fairbrother, BA,\*\* Winston Banya, MSc,††

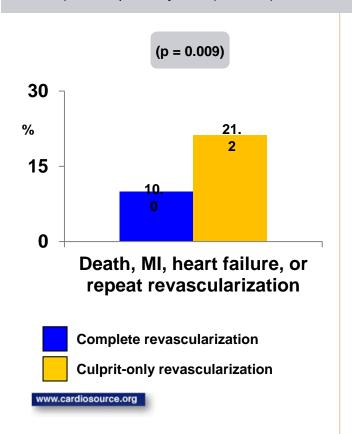
Duolao Wang, PhD,‡‡ Marcus Flather, MB BS,§§ Simon L. Hetherington, MB ChB, MD,|||

Andrew D. Kelion, BM BCh, DM,¶¶ Suneel Talwar, MB BS, MD,## Mark Gunning, MD,\*\*\* Roger Hall, MD,§§

Howard Swanton, MB BChir, MD,††† Gerry P. McCann, MB ChB, MD\*

#### **CVLPRIT**

**Trial design:** Participants with STEMI were randomized to complete revascularization (n = 150) vs. culprit-only PCI (n = 146).



#### Results

- Death, MI, heart failure, or ischemia-driven revascularization at 12 months: 10.0% of the complete revascularization group vs. 21.2% of the culprit-only group (p = 0.009)
- All-cause mortality: 1.3% vs. 4.1% (p = 0.14), respectively
- MI: 1.3% vs. 2.7% (p = 0.39), respectively
- Heart failure: 2.7% vs. 6.2% (p = 0.14), respectively
- Repeat revascularization: 4.7% vs. 8.2% (p = 0.2), respectively

#### **Conclusions**

- Among STEMI patients, complete revascularization appears beneficial at reducing major adverse cardiac events
- Benefit was primarily due to reduction in repeat revascularization procedures

Presented by Dr. Gershlick at ESC 2014

# Criticism about the study:

- Small study but significant outcome
- No FFR or IVUS of the N-IRA lesions
- Open study





The Third DANish Study of Optimal Acute Treatment of Patients with ST-segment Elevation Myocardial Infarction

PRImary PCI in MULTIvessel Disease - DANAMI3-PRIMULTI

Thomas Engstrøm, MD, DMSci, PhD Rigshospitalet, University of Copenhagen, Denmark







# Complete revascularisation versus treatment of the culprit lesion only in patients with ST-segment elevation myocardial infarction and multivessel disease (DANAMI-3—PRIMULTI): an open-label, randomised controlled trial



Thomas Engstrøm, Henning Kelbæk, Steffen Helqvist, Dan Eik Høfsten, Lene Kløvgaard, Lene Holmvang, Erik Jørgensen, Frants Pedersen, Kari Saunamäki, Peter Clemmensen, Ole De Backer, Jan Ravkilde, Hans-Henrik Tilsted, Anton Boel Villadsen, Jens Aarøe, Svend Eggert Jensen, Bent Raunqaard, Lars Køber, for the DANAMI-3—PRIMULTI Investigators\*

#### Summary

Background Patients with acute ST-segment elevation myocardial infarction (STEMI) and multivessel coronary disease have a worse prognosis compared with individuals with single-vessel disease. We aimed to study the clinical outcome of patients with STEMI treated with fractional flow reserve (FFR)-guided complete revascularisation versus treatment of the infarct-related artery only.

Methods We undertook an open-label, randomised controlled trial at two university hospitals in Denmark. Patients presenting with STEMI who had one or more clinically significant coronary stenosis in addition to the lesion in the infarct-related artery were included. After successful percutaneous coronary intervention (PCI) of the infarct-related artery, patients were randomly allocated (in a 1:1 ratio) either no further invasive treatment or complete FFR-guided revascularisation before discharge. Randomisation was done electronically via a web-based system in permuted blocks of varying size by the clinician who did the primary PCI. All patients received best medical treatment. The primary endpoint was a composite of all-cause mortality, non-fatal reinfarction, and ischaemia-driven revascularisation of lesions in non-infarct-related arteries and was assessed when the last enrolled patient had been followed up for 1 year. Analysis was on an intention-to-treat basis. This trial is registered with ClinicalTrials.gov, number NCT01960933.

Findings From March, 2011, to February, 2014, we enrolled 627 patients to the trial; 313 were allocated no further invasive treatment after primary PCI of the infarct-related artery only and 314 were assigned complete revascularisation guided by FFR values. Median follow-up was 27 months (range 12–44 months). Events comprising the primary endpoint were recorded in 68 (22%) patients who had PCI of the infarct-related artery only and in 40 (13%) patients who had complete revascularisation (hazard ratio 0.56, 95% CI 0.38-0.83; p=0.004).

Interpretation In patients with STEMI and multivessel disease, complete revascularisation guided by FFR measurements significantly reduces the risk of future events compared with no further invasive intervention after primary PCI. This effect is driven by significantly fewer repeat revascularisations, because all-cause mortality and non-fatal reinfarction did not differ between groups. Thus, to avoid repeat revascularisation, patients can safely have all their lesions treated during the index admission. Future studies should clarify whether complete revascularisation should be done acutely during the index procedure or at later time and whether it has an effect on hard endpoints.

#### ublished Online

August 5, 2015 http://dx.doi.org/10.1016/ S0140-6736(15)60648-1

#### See Online/Comment

http://dx.doi.org/10.1016/ S0140-6736(15)60856-X

\*Listed at end of report and in the appendix (p 1)

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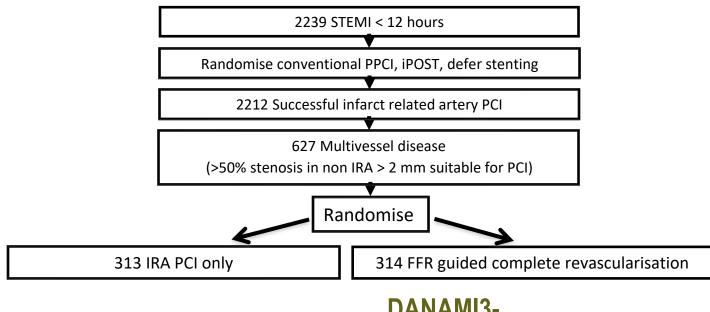
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### **DANAMI3-TRIAL PROGRAM**

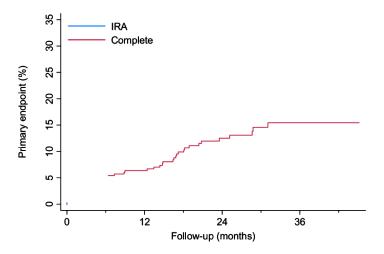


### DANAMI3-PRIMULTI





### **Primary endpoint**

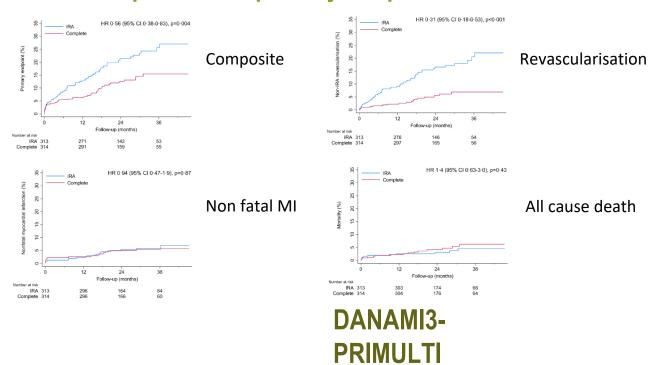


### DANAMI3-PRIMULTI



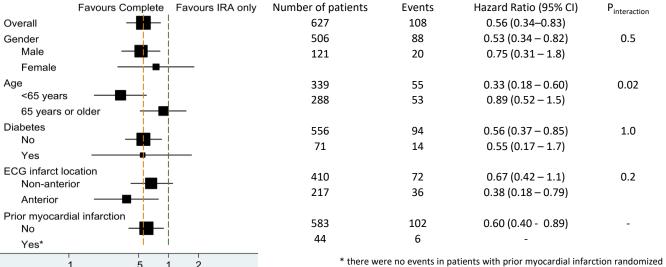


# **Individual components of primary endpoint**





# **Subgroup analysis**



to complete revascularization

# **DANAMI3-PRIMULTI**





### **Conclusions**

Complete FFR guided revascularisation of multivessel disease in STEMI patients, staged within the index admission, reduced the primary endpoint of all cause death, reinfarction and repeat revascularisation

40% of repeat revascularisations were urgent

However, the reduction in the primary endpoint was driven by repeat revascularisations and not by hard endpoints

Therefore, although complete revascularisation should be recommended, any condition that makes complex PCI unattractive may support a more conservative strategi of IRA PCI only

DANAMI3-PRIMULTI





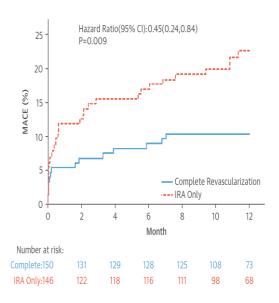
# The PRAMI, CvLPRIT, and DANAMI-3-PRIMULTI trials MACE

### The PRAMI trial

# Hazard ratio, 0.35 (95% CI, 0.21–0.58); P<0.001 Hazard ratio, 0.35 (95% CI, 0.21–0.58); P<0.001 Preventive PCI Solution No. at Risk Preventive PCI 234 196 168 146 118 89 67 No preventive PCI 231 168 144 122 96 74 50

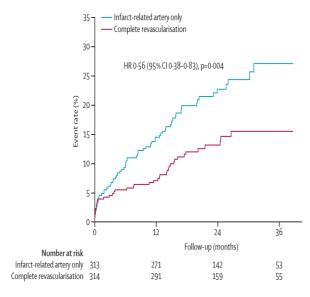
Wald DS et al. N Engl J Med 2013;369:1115-23

### The CvLPRIT trial



Gershlick AH et al. JACC 2015;65:963-72

### The DANAMI3-PRIMULTI trial



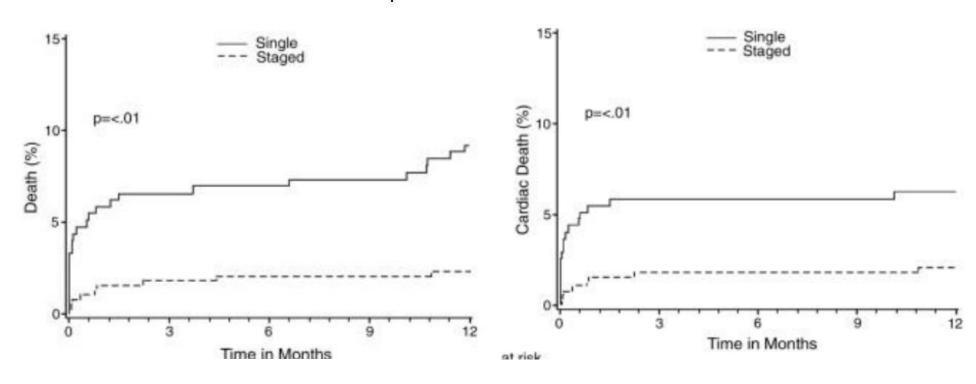
Engstrøm T et al. Lancet 2015;386:665-71





# Single vs Staged PCI in Ptes with MVD: Horizons AMI

### 668 ptes con IAM ST







# **COMPARE-ACUTE**



Randomised trial of

FFR-guided complete revascularization

versus

infarct artery only treatment

in

multivessel STEMI patients

On behalf of all COMPARE-ACUTE investigators

**Pieter Smits** 

Maasstad Hospital

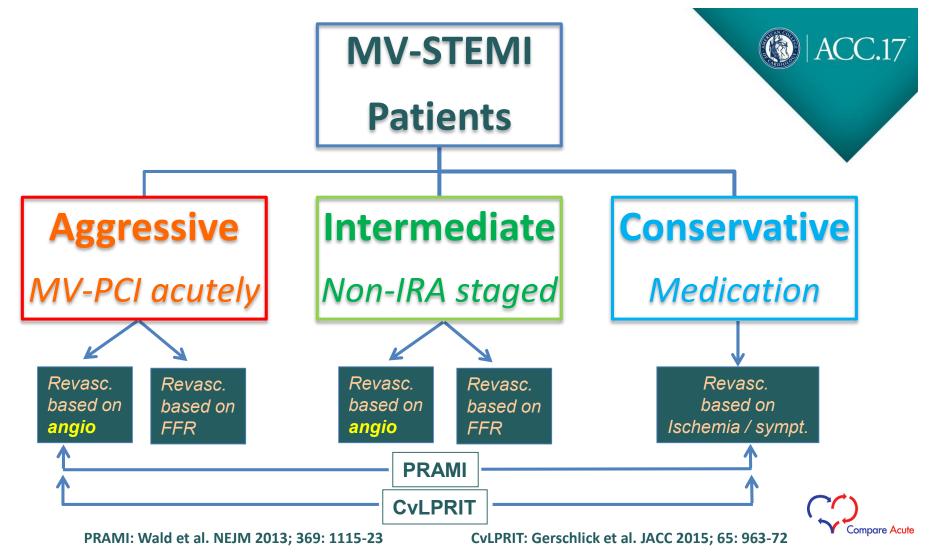
Rotterdam, The Netherlands







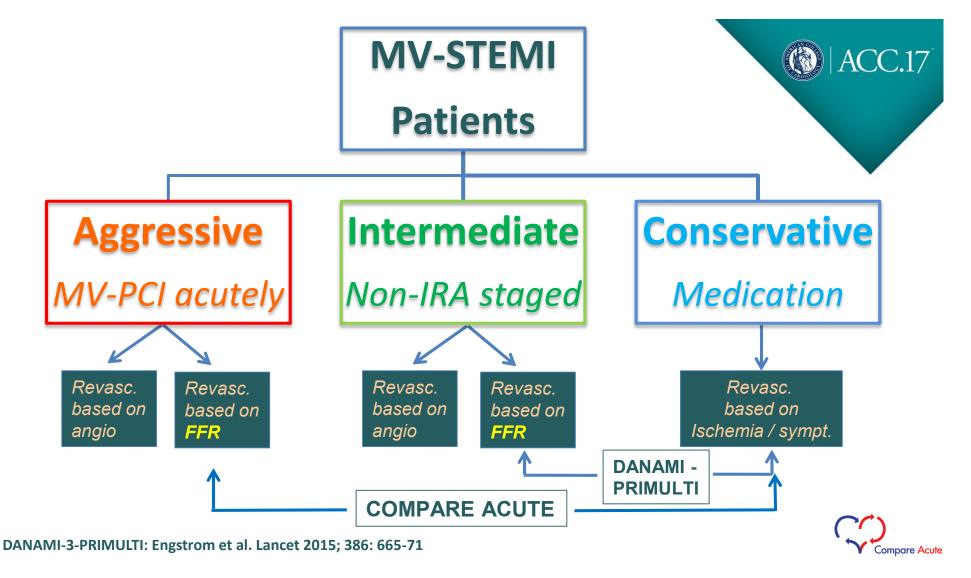




Pieter Smits, ACC 2017





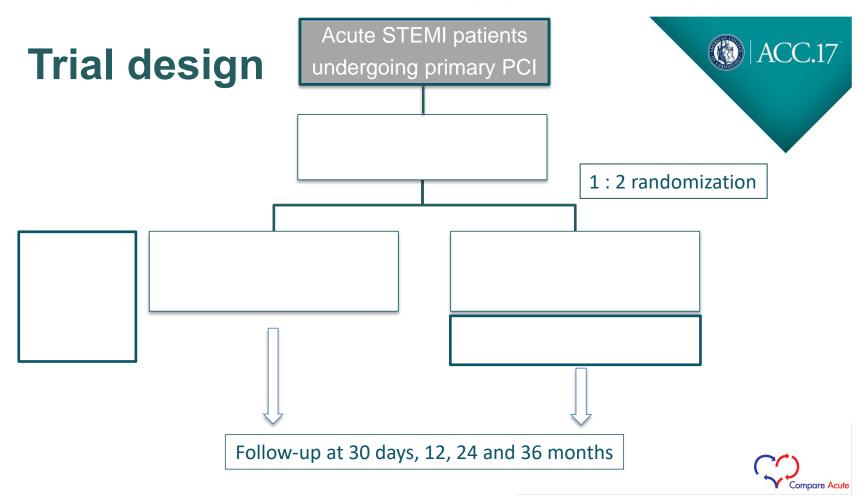


Pieter Smits, ACC 2017





# **COMPARE-ACUTE Trial**



P Smits. ACC 2017, March 18, 2017, at NEJM.org. DOI: 10.1056/NEJMoa1701067



# **COMPARE-ACUTE Trial**

- Approximately half (46.9%) of the lesions in non-related arteries considered significant on coronary angiography had an FFR value >0.8 and were therefore not physiologically significant.
- Deferring treatment of angiographically significant coronary lesions in non-related arteries with FFR>0.8 is safe and efficient.

P Smits. ACC 2017, March 18, 2017, at NEJM.org. DOI: 10.1056/NEJMoa1701067





# **Up-Coming Trials**

- COMPLETE Trial: N=3900
  - Culprit only vs staged procedures
- CROSS-AMI;
  - Culprit only vs. Stress echo-guided PCI





### CLINICAL DOCUMENT

# ACC/AATS/AHA/ASE/ASNC/SCAI/ SCCT/STS 2016 Appropriate Use



Indic	ation	Appropriate Use Score (1-9
Reva	scularization of the Presumed Culprit Artery by PCI (Primary PCI)	
1.	■ Less than or equal to 12 hours from onset of symptoms	A (9)
2.	<ul> <li>Onset of symptoms within the prior 12-24 hours AND</li> <li>Severe HF, persistent ischemic symptoms, or hemodynamic or electrical instability present</li> </ul>	A (8)
3.	■ Onset of symptoms within the prior 12–24 hours AND	M (6)
	■ Stable without severe HF, persistent ischemic symptoms, or hemodynamic or electrical instability	
	<ul> <li>Stable without severe HF, persistent ischemic symptoms, or hemodynamic or electrical instability</li> <li>essful Treatment of the Culprit Artery by Primary PCI Followed by Immediate Revascularization of 1 or More same Procedure</li> </ul>	Nonculprit Arteries During
the S	essful Treatment of the Culprit Artery by Primary PCI Followed by Immediate Revascularization of 1 or More	Nonculprit Arteries During  A (8)
	essful Treatment of the Culprit Artery by Primary PCI Followed by Immediate Revascularization of 1 or More same Procedure  Cardiogenic shock persisting after PCI of the presumed culprit artery	

The number in parenthesis next to the rating reflects the median score for that indication.

A = appropriate; CABG = coronary artery bypass graft; HF = heart failure; M = may be appropriate; PCI = percutaneous coronary intervention; R = rarely appropriate; STEMI = ST-segment elevation myocardial infarction.

### **CLINICAL DOCUMENT**

# ACC/AATS/AHA/ASE/ASNC/SCAI/ SCCT/STS 2016 Appropriate Use Criteria for Coronary Revascularization in Patients With Acute Coronary Syndromes



Indic	ation	Appropriate Use Score (1-9)		
	Successful Treatment of the Culprit Artery by Primary PCI or Fibrinolysis Revascularization of 1 or More Nonculprit Arteries During the Same Hospitalization			
Revascularization by PCI or CABG				
10.	<ul> <li>Spontaneous or easily provoked symptoms of myocardial ischemia</li> <li>One or more additional severe stenoses</li> </ul>	A (8)		
11.	<ul> <li>Asymptomatic</li> <li>Findings of ischemia on noninvasive testing</li> <li>One or more additional severe stenoses</li> </ul>	A (7)		
12.	<ul> <li>Asymptomatic (no additional testing performed)</li> <li>One or more additional severe stenoses</li> </ul>	M (6)		
13	<ul> <li>Asymptomatic (no additional testing performed)</li> <li>One or more additional intermediate stenoses</li> </ul>	R (3)		
14.	<ul> <li>Asymptomatic</li> <li>One or more additional intermediate (50%-70%) stenoses</li> <li>FFR performed and ≤0.80</li> </ul>	A (7)		

The number in parenthesis next to the rating reflects the median score for that indication.

A = appropriate; CABG = coronary artery bypass graft; FFR = fractional flow reserve; M = may be appropriate; PCI = percutaneous coronary intervention; R = rarely appropriate; STEMI = ST-segment elevation myocardial infarction.

# **Final Conclusions**

- Culprit-vessel PCI with in-hospital staged nonculprit PCI leads to lower risks of death and repeat revascularization at 2 years.
- The use od iFR/FFR may help to decide a better "complete functional Revascularisation" for STEMI PCI.
- Complete revascularisation should be done to Improve Prognosis and Reduce Costs!





### **COMPLETE Trial: Study Design** COMPLETE A randomized, comparative effectiveness study of complete versus culprit-only revascularization strategies to treat multi-vessel disease after 10 PCI for STEMI STEMI with successful PCI for STEMI (primary, rescue or pharmacoinvasive) + $\geq$ 70% stenosis or $\geq$ 50% with FFR < 0.80 RANDOMIZE Within 72 h of index STEMI PCI CULPRIT LESION-ONLY REVASC **COMPLETE REVASC Staged** PCI of all suitable No further revsac of non-culprit non-culprit lesions (< 45 d) lesions (OMT Alone) N=1950N=1950ALL patients receive OMT (ASA, Ticagrelor, Statin, Beta Blocker, RF Modification) Follow-up: Discharge, 30 Days, 6 mos, then Annually (avg. duration = 4 yrs)

Primary Outcome: CV Death / MI

Key Secondary Outcome: CV Death/MI/Ischemia driven revascularization

Randomization stratified for intended timing of PCI: within vs after initial hospitalization  $_{21}$ 







- The focused update does not, however, give a recommendation on when to perform PCI of the non-infract artery.
- The COMPLETE trial will evaluate the strategy of culprit lesion only revascularization versus complete revascularization using a staged PCI approach (within 45 days of the index procedure).



# Primary percutaneous coronary intervention for myocardial reperfusion in ST-elevation myocardial infarction: procedural aspects (strategy and technique)



Recommendations	Class	Level
Strategy		
Routinerevascularizationofnon-IRAlesionsshouldbeconsidered inpatientswithmultivesseldiseasebeforehospitaldischarge.	lla	Α
CABGshouldbeconsideredinpatientswithongoingischaemiaand largeareasofjeopardizedmyocardiumifPClofthelRAcannotbe performed.	lla	С
Technique		
Routineuse of thrombus as piration is not recommended.	Ш	A

# TCT Russia At TCT 2013 When To Acutely Perform Multivessel PCI in STEMI?

George D. Dangas, MD, PhD, FACC, FESC
Professor of Medicine
Mount Sinai Medical Center, New York, NY









# When to do MV PCI in STEMI?

- 1. When the culprit vessel has been treated successfully
- 2. And the non-culprit vessel has a proximal significant lesion with a lot of myocardium at risk
- 3. And the features of this lesion predict a good PCI result within 15-20min and <100cc additional contrast load
- 4. Then we may consider MV PCI in STEMI, particularly if the larger COMPLETE trial is also concordant

- Decision based on the individual patient remains the rule.
- Large contemporary meta-analyses, mostly from obsevational studies, before PRAMI/CVLPRIT are consistent in showing a benefit of staging non-IRA PCI versus either culprit only or immediate MV PCI strategies. The COMPLETE, DANAMI-3 and COMPARE-ACUTE trials will add other pieces to the puzzle.
- My practice for STEMI with MVD before & after PRAMI/CVLPRIT: Culprit only, then early planned, staged non-IRA PCI.

# Controversies In STEMI

- Culprit versus complete revascularization in MVD
- Culprit versus complete revascularization in cardiogenic shock

# Controversies In STEMI

Culprit versus complete revascularization in MVD

Culprit versus complete revascularization in cardiogenic shock





# CULPRIT-SHOCK: A Randomized Trial of Multivessel PCI in Cardiogenic Shock

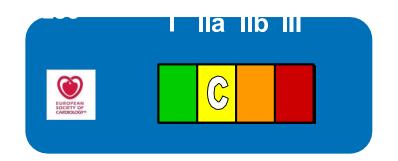
Holger Thiele, MD on behalf of the CULPRIT-SHOCK Investigators

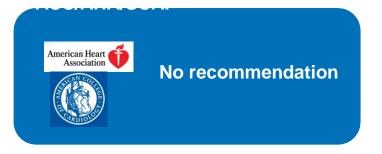




# Multivessel coronary artery disease present in up to 80% → higher mortality Guidelines

Multivessel PCI in Cardiogenic Shock European and American Recommendations 2017





# **Appropriate Use Criteria**



Ibanez et al. ESC STEMI Guidelines 2017. Eur Heart J 2017; epub



# Multivessel PCI in Cardiogenic Shock?



# **Metaanalysis Mortality – Registry-Data**

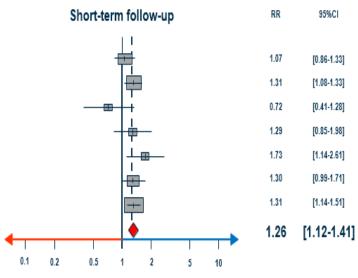
	MV-PCI		C-PCI	
	Events	Total	Events	Total
IABP-SHOCK II	75	167	119	284
ALKK	81	173	201	562
KAMIR	13	124	56	386
Yang et al.	19	60	68	278
Cavender et al.	20	43	42	156
EHS-PCI	40	82	95	254
NCDR	158	433	737	2654
Overall	406	1082	1318	4574
U-tit0 007 1:-24 00/	n=0.10			

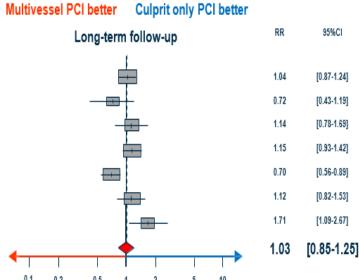
Heterogeneity:  $\tau^2=0.007$ ,  $t^2=31.0\%$ , p=0.19Test for overall effect: p=0.001

	MV-PCI		C-P	CI
	Events	Total	Events	Total
IABP-SHOCK II	91	167	149	284
KAMIR	16	124	69	386
Yang et al.	21	60	85	278
Cavender et al.	32	43	101	156
Mylotte et al.	37	66	82	103
van der Schaaf et al.	22	37	66	124
SHOCK	7	9	26	57
Overall	226	506	578	1387

Heterogeneity: T2=0.043, 12=67.8%, p=0.005

Test for overall effect: p=0.77



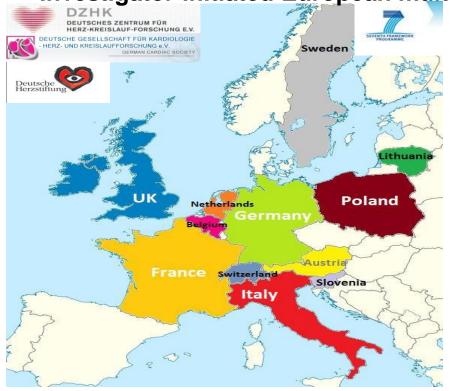


**Culprit only PCI better** 

Multivessel PCI better

# **CULPRIT-SHOCK Trial**

Investigator-initiated European multicenter trial; 1:1 randomization



**PI + Coordination:** 

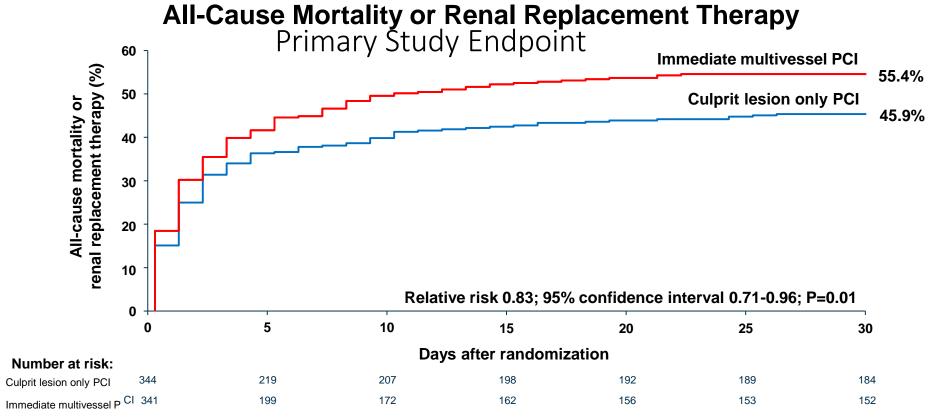
Holger Thiele

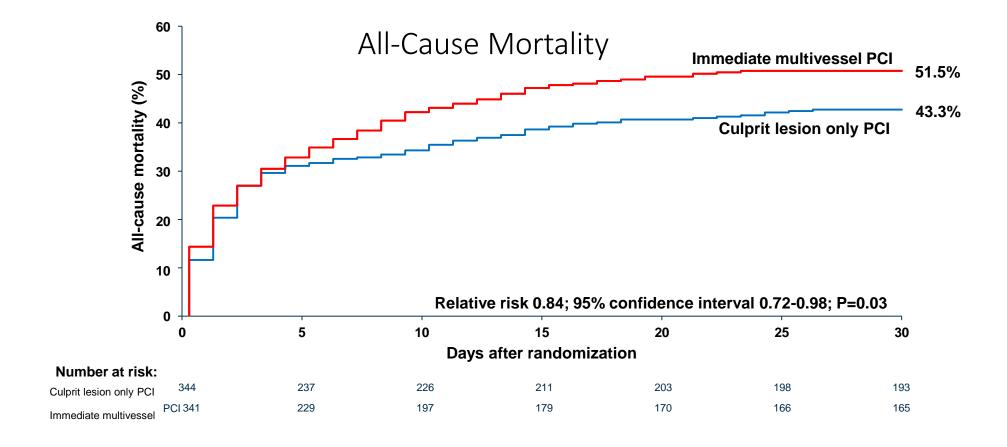
Co-PI:

Uwe Zeymer Steffen Desch

### **National Coordinators (83 centers):**

- Kurt Huber
- Gilles Montalescot
- → Jan Piek
- Holger Thiele
- Pranas Serpytis
- Janina Stepinska
- Christiaan Vrints
- Marko Noc
- Keith Oldroyd
- Stefan Windecker
- Stefano Savonitto





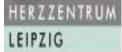
# **Conclusions**

- In patients with multivessel coronary artery disease and cardiogenic shock complicating acute myocardial infarction culprit lesion only PCI with possible staged revascularization reduced the composite of mortality or requirement for renal replacement therapy at 30 days.
- This effect in the primary outcome was mainly driven by a 30-day mortality reduction.
- This largest randomized European multicenter trial in cardiogenic shock complicating myocardial infarction challenges current guideline recommendations.

### ORIGINAL ARTICLE

# PCI Strategies in Patients with Acute Myocardial Infarction and Cardiogenic Shock

H. Thiele, I. Akin, M. Sandri, G. Fuernau, S. de Waha, R. Meyer-Saraei,
P. Nordbeck, T. Geisler, U. Landmesser, C. Skurk, A. Fach, H. Lapp, J.J. Piek,
M. Noc, T. Goslar, S.B. Felix, L.S. Maier, J. Stepinska, K. Oldroyd, P. Serpytis,
G. Montalescot, O. Barthelemy, K. Huber, S. Windecker, S. Savonitto,
P. Torremante, C. Vrints, S. Schneider, S. Desch, and U. Zeymer,
for the CULPRIT-SHOCK Investigators\*







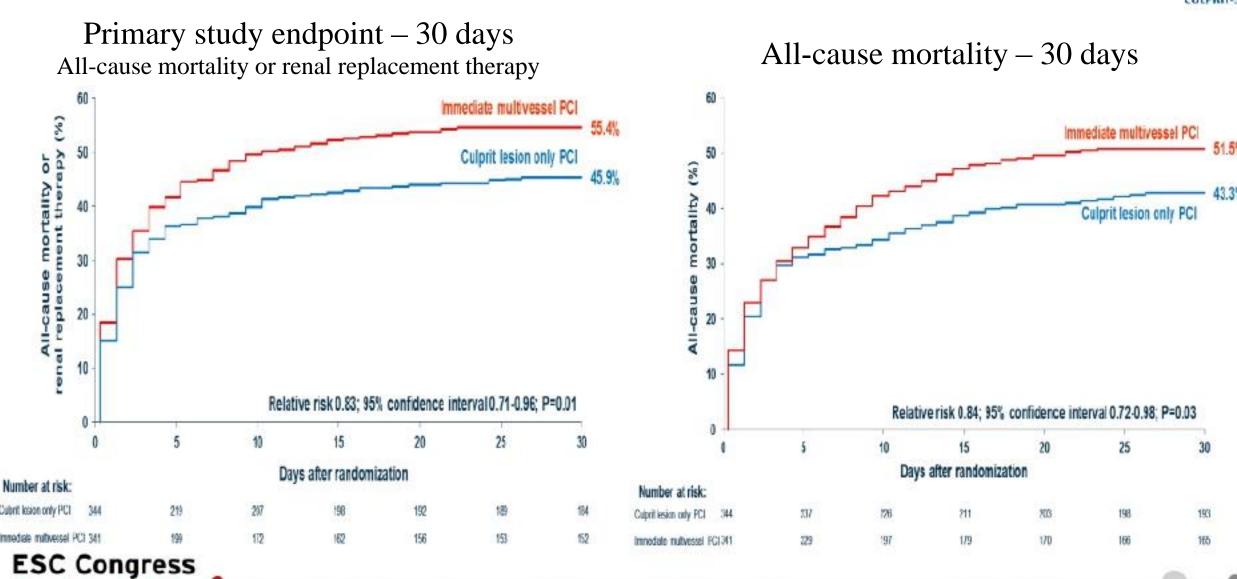


# CULPRIT-SHOCK: Culprit Lesion Only PCI versus Multivessel PCI in Cardiogenic Shock – 1-Year Results

Holger Thiele on behalf of the CULPRIT-SHOCK Investigators

# **CULPRIT-SHOCK Trial – 30-Day Results**



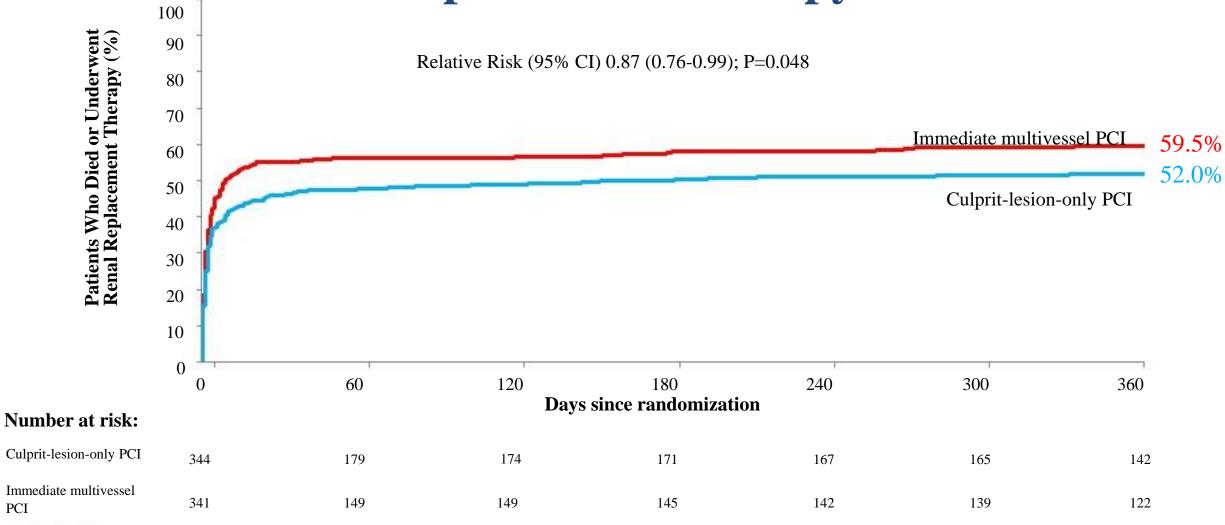


Munich 2018

Thiele et al. NEJM 2017; 377:2419-2432

# 1-Year All-Cause Mortality or Renal Replacement Therapy







**PCI** 

**N Engl J Med**; 2018 August 25, DOI: 10.1056/NEJMoa1808788

# www.nejm.org





### ORIGINAL ARTICLE

## One-Year Outcomes after PCI Strategies in Cardiogenic Shock

H. Thiele, I. Akin, M. Sandri, S. de Waha-Thiele, R. Meyer-Saraei, G. Fuernau, J. Eitel, P. Nordbeck, T. Geisler, U. Landmesser, C. Skurk, A. Fach, A. Jobs, H. Lapp, J.J. Piek, M. Noc, T. Goslar, S.B. Felix, L.S. Maier, J. Stepinska, K. Oldroyd, P. Serpytis, G. Montalescot, O. Barthelemy, K. Huber, S. Windecker, L. Hunziker, S. Savonitto, P. Torremante, C. Vrints, S. Schneider, U. Zeymer, and S. Desch, for the CULPRIT-SHOCK Investigators\*

ESC Congress Munich 2018





**#ESCCongress** 





# **ΘΕΔCTS** What is new in the 2018 Guidelines?

# **New recommendations**



## **UPGRADES**

Immediate coronary angiography and revascularization, if appropriate, in survivors of out-of-hospital cardiac arrest and an ECG consistent with STEMI

### **DOWNGRADES**

Bivalirudin for PCI in NSTE-ACS

Bivalirudin for PCI in STEMI

# **NEW RECOMMANDATIONS**

Routine revasularization of non-IRA lesions in myocardial infarction with cardiogenic shock

Changes compared with the 2014 version of the Myocardial Revascularization Guidelines that were due to updates for consistency with other ESC Guidelines published since 2014 are not shown.

Class I Class Ilb Class Ila Class III



# **EΔCTS** Percutaneous coronary intervention in ACS patients **ESC** with cardiogenic shock



Recommendations	Class	Level
Strategy		
Incardiogenicshock, routine revascularization of non-IRA lesions is not recommended during primary PCI.	III	В





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