

## ACC Middle East Conference 2018

## Non-STEMI Conservative Management vs Early Invasive Management

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# **Disclosures: None**









## **ACS-NSTEMI : Risk Assessment**

#### Focused history physical examination

#### 12-Lead EKG

#### Troponins

#### Use risk scores to assess prognosis in patients with NSTE-ACS

1. Find Points for Each Predictive Factor:

Killip Point: Class	5		SBP, mm Hg		Points		Heart Beats		Points	E)		Age, y	Poi	nts		Creatini Level, r		Point	s	
I 0 Ⅱ 20 Ⅲ 39 Ⅳ 59			≤80 80-99 100-11 120-13 140-15 160-19 ≥200	9	58 53 43 34 24 10 0		≤50 50-6 70-8 90-1 110- 150- ≥200	9 9 09 -149 -199	0 9 15 24 38 46			≤30 30-39 40-49 50-59 60-69 70-79 80-89 ≥90		1 8 5 1		0.80	-0.79 -1.19 -1.59 -1.99 -3.99	1 4 7 10 13 21 28		
Other Risk Factors			Points																	
Cardiac Arrest at Admis ST-Segment Deviation Elevated Cardiac Enzyn		3	39 28 14																	
m Points for All Predictiv	e Factors	i.																		
Killip <sub>+</sub> SBP <sub>+</sub> Class	Heart Rate	÷	Age	٠	Creatini Level	ne +	Arres			T-Segme eviation		Elevate Enzym	ed Card e Level	ac = s	Total Points					
ook Up Risk Correspondin	g to Tota	Points	5.																	
Total Points	≤60	70	80	90	100	110	120	130	140	150	160	170	180	190	200	210	220	230	240	2
Probability of In-Hospital Death, %	≤0.2	0.3	0.4	0.6	0.8	1.1	1.6	2.1	2.9	3.9	5.4	7.3	9.8	13	18	23	29	36	44	25





#### **GRACE Score For Risk Of Death In Non-ST Elevation Acute Coronary Syndrome**

Risk Factor	Finding		Points
Event Type	Non-ST Elevation ACS		
Killip Class	III (pulmonary edema)	•	39
Systolic Blood Pressure	100-119	•	43
Heart Rate	70-89	•	9
Age	60-69	۲	58
- Creatinine Level (mg%)	0.80-1.19	•	7
Cardiac Arrest At Admission	Absent	•	0
ST-Segment Deviation	Present	¥	39
Elevated Cardiac Enzyme Levels*	Present	•	14
,	Point Total		209
	In-Hospital Mortality Ris Category	k	High

https://qxmd.com/calculate/calculator\_262/grace

https://www.mdcalc.com/grace-acs-risk-mortality-calculator





# **Conservative Management**

#### Aspirin

- Non-enteric-coated aspirin to *all* patients promptly after presentation
- Aspirin maintenance dose continued indefinitely

#### P2Y<sub>12</sub> inhibitors

- Clopidogrel loading dose followed by daily maintenance dose in patients unable to take aspirin
- P2Y<sub>12</sub> inhibitor, in addition to aspirin, for up to 12 mo for patients treated initially with either an early invasive or initial ischemia-guided strategy:
- Clopidogrel
- Ticagrelor\*

## Low-molecular weight or unfractionated heparin

- SC enoxaparin for duration of hospitalization or until PCI is performed
- IV UFH for 48 h or until PCI is performed

#### Anti-anginal therapy

Initiate oral beta blockers within the first 24 h in the absence of HF, low-output state, risk for cardiogenic shock, or other contraindications to beta blockade

Administer sublingual NTG every 5 min  $\times$  3 for continuing ischemic pain and then assess need for IV NTG Administer IV NTG for persistent ischemia, HF, or hypertension

#### **Cholesterol management**

Initiate or continue high-intensity statin therapy in patients with no contraindications









revascularization

**ESC/EACTS GUIDELINES** 

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#### FOCUS ISSUE: CARDIAC INTERVENTION

Intervention in Acute Coronary Syndromes

Benefit of Early Invasive Therapy in Acute Coronary Syndromes A Meta-Analysis of Contemporary Randomized Clinical Trials Anthony A. Bavry, MD, MPH,\* Dharam J. Kumbhani, MD, SM,† Andrew N. Rassi, MD,‡ Deepak L. Bhatt, MD,\* Arman T. Askari, MD\* *Cleveland, Obio; and Philadelphia, Pennsylvania* 

### **AHA/ACC Guideline**

2018 ESC/EACTS Guidelines on myocardial

2014 AHA/ACC Guideline for the Management of Patients With Non–ST-Elevation Acute Coronary Syndromes A Report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines

> Developed in Collaboration With the Society for Cardiovascular Angiography and Interventions and Society of Thoracic Surgeons

Optimal timing of an invasive strategy in patients with non-ST-elevation acute coronary syndrome: a meta-analysis of randomised trials

Alexander Jobs, Shamir R Mehta, Gilles Montalescot, Eric Vicaut, Arnoud WJ van't Hof, Erik A Badings, Franz-Josef Neumann, Adnan Kastrati, Alessandro Sciahbasi, Paul-Georges Reuter, Frédéric Lapostolle, Aleksandra Milosevic, Goran Stankovic, Dejan Milasinovic, Reinhard Vonthein, Steffen Desch, Holger Thiele







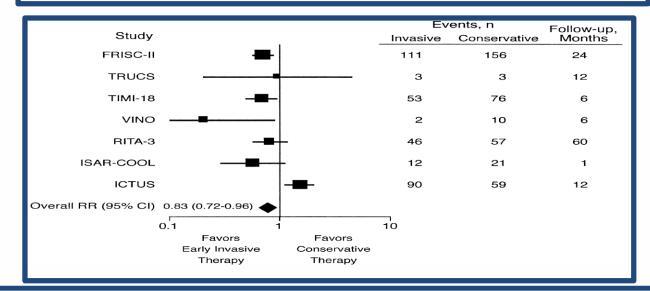
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Characteristic	FRISC-II	TRUCS	TIMI-18	VINO	RITA-3	ISAR-COOL	ICTUS
Enrollment period	1996-1998	1997-1998	1997-1999	1998-2000	1997-2001	2000-2002	2001-2003
Invasive/conservative patients, n	1222/1234	76/72	1114/1106	64/67	895/915	203/207	604/596
Age, yrs (mean)	66*	62	62	66	62	70*	62*
Women, %	30	27	34	39	38	33	27
Diabetes, %	12	29	28	25	13	29	14
Prior myocardial infarction, %	22	27†	39	26	28	23	23
Current smokers, %	30	31	NA	NA	32	21	41
Statin at randomization, %	10	21	52‡	43‡	45	NA	27
Statin at follow-up, %	55	NA	NA	NA	80	85	92
Thienopyridine with PCI, %	100§	NA	NA	100§	96	100¶	100
Elevated troponin at randomization, %	55	NA	54	100	75	67	100
Hours to angiography, median**	96/408	48/120++	22/79	6.2/1,464	48/1,020	2.4/86	23/283#
Glycoprotein IIb/IIIa inhibitor, type	NA	NA	Tirofiban	NA	NA	Tirofiban	Abciximab
Invasive undergoing PCI, %	10	95	94	0	25	100¶	93
Conservative undergoing PCI, %	NA	96	59	0	25	100¶	69
Conservative not undergoing PCI, %	NA	NA	99	0	0	100¶	0
Coronary stent use with PCI, %**	62/69	85/85	83/86	44/50	88/90	87/92	88/88
Overall PCI or CABG, %**	78/45	100/61	64/45	73/39	61/38	78/72	79/54
Relative difference in revascularization between treatment arms, %	73	64	42	87	61	8	46

7 Trials conducted between 1996-2003 BMS era. (n = 8375) JACC Vol. 48, No. 7, 2006 October 3, 2006:1319-25





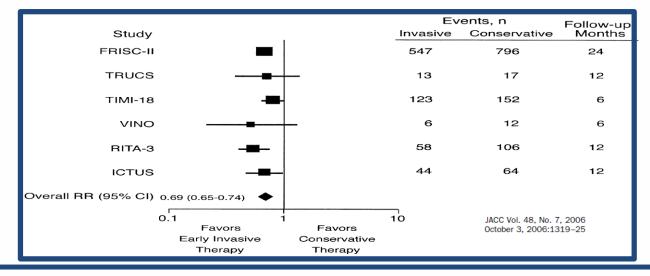


Relative risk of recurrent nonfatal <u>myocardial infarction</u> for early invasive therapy compared with conservative therapy at a mean follow-up of 2 years.









Relative risk of recurrent unstable angina resulting in rehospitalization for early invasive therapy compared with conservative therapy at a mean follow-up of 13 months

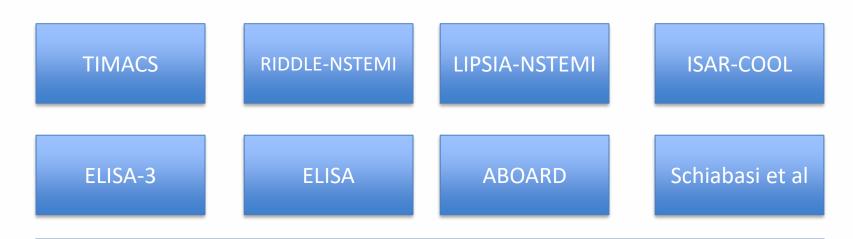




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Lancet 2017; 390: 737-46



8 Trials conducted between 2002 – 2016 DES Era (n = 5324) comparing an Early invasive versus a delayed invasive strategy

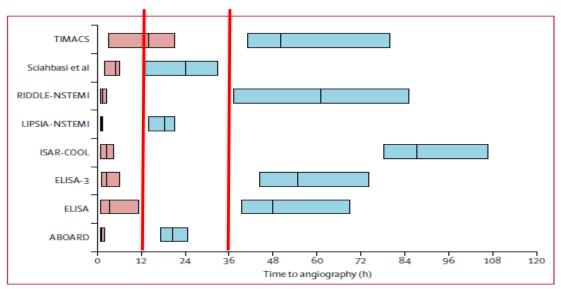




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	HR	95% CI	Weight
ABOARD	5-11	(0-60-43-72)	1.2%
ELISA	0.43	(0-11-1-66)	3.1%
ELISA-3	0.68	(0-36-1-29)	14.0%
ISAR-COOL	1.09	(0-46-2-56)	7-8%
LIPSIA-NSTEMI	0.74	(0-31-1-76)	7-6%
TIMACS	0.81	(0-60-1-11)	60.0%
RIDDLE-NSTEMI	0-88	(0-34-2-27)	6-3%
Random effects model	0-81	(0.64-1.03)	100-0%
Heterogeneity: I²=0%, τ²=0	), p=0.610	04	
Test for overall effect: p=0-	0879		

#### ALL-CAUSE MORTALITY

#### OUTCOMES AFTER AN EARLY INVASIVE V A DELAYED INVASIVE STRATEGY

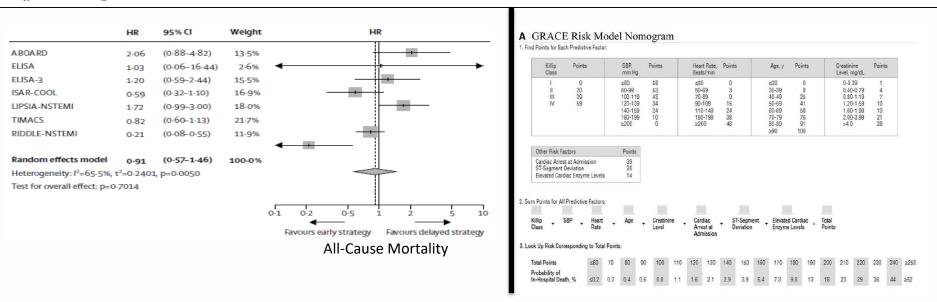




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Use risk scores to assess prognosis in patients with NSTE-ACS





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Patients with GRACE risk sco				_
ELISA-3	0.51	(0.25-1.07)	12-3%	
LIPSIA-NSTEMI	0-69	(0.27-1.79)	7-4%	
TIMACS	0.75	(0.51-1.09)	46-8%	
RIDDLE-NSTEMI	0.77	(0.28-2.18)	6.2%	
Random effects model	0.70	(0.52-0.95)	72.7%	\$
Patients with GRACE risk sco	ore ≤140			
ELISA-3	1.66	(0.40-6.96)	3.2%	
LIPSIA-NSTEMI	1.85	(0.17-20.35)	1.2%	<b>&gt;</b>
TIMACS	0.94	(0.55-1.61)	22-9%	·
RIDDLE-NSTEMI		/	0.0%	
Random effects model	1.04	(0.63-1.70)	27.3%	
Tast for subgroup differences	n 0.1930			
Test for subgroup differences:	p=0.1020			
				0.1 0.2 0.5 1 2 5 10
				$\leftarrow$
				Favours early strategy Favours delayed strategy

All-Cause Mortality

## GRACE Risk Score > 140









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Patients aged ≥75 years				
ABOARD	1.83	(0-17-20-20)	1-0%	<b>&gt;</b>
ELISA	0.36	(0-04-3-08)	1.3%	↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓
ELISA-3	0.41	(0-18-0-93)	8-8%	
ISAR-COOL	1.49	(0-47-4-71)	4.5%	
LIPSIA-NSTEMI	0.43	(0-11-1-63)	3-4%	
Sciahbasi et al			0-0%	
TIMACS	0.62	(0-38-1-01)	25.3%	
RIDDLE-NSTEMI	1-29	(0-39-4-24)	4.2%	
Random effects model	0.65	(0-46-0-93)	48-6%	
Designation and the second				
Patients aged <75 years ABOARD			0-0%	
ELISA	0.87	(0-12-6-16)	1.6%	
ELISA-3	1.97	(0-59-6-54)	4.2%	
ISAR-COOL	0.76	(0-20-2-84)	3.5%	
LIPSIA-INSTEMI	1.40	(0-40-4-97)	3.7%	
Sciahbasi et al	1.40	(0.40-4.97)	0-0%	
TIMACS	1-01	(0-67-1-52)	36-4%	
RIDDLE-NSTEMI	0.49	(0-09-2-66)	2-1%	
Random effects model	1.04	(0.74-1.46)	51.4%	
Random eneces model	1.04	(0) 4 1 40)	34.410	
Test for subgroup differences:	p=0.0647			
3 1				
				0.1 0.2 0.5 1 2 5 10
				$\leftarrow \rightarrow$
				Favours early strategy Favours delayed strategy

Age > 75

All-Cause Mortality









Alexander Jobs, Shamir R Mehta, Gilles Montalescot, Eric Vicaut, Arnoud W J van't Hof, Erik A Badings, Franz-Josef Neumann, Adnan Kastrati, Alessandro Sciahbasi, Paul-Georges Reuter, Frédéric Lapostolle, Aleksandra Milosevic, Goran Stankovic, Dejan Milasinovic, Reinhard Vonthein, Steffen Desch, Holger Thiele

Α	HR	95% CI	Weight	
Patients with elevated cardi	ac biomarker	s at baseline		
ABOARD	2.98	(0-31-28-67)	1.1%	
ELISA	0.33	(0.06-1.70)	2-1%	← · · · · · · · · · · · · · · · · · · ·
ELISA-3	0.57	(0-27-1-20)	10-4%	
SAR-COOL	1.04	(0.39-2.78)	6-0%	<del>``````</del>
JPSIA-NSTEMI	0.74	(0-31-1-76)	7.7%	
iciahbasi et al			0-0%	
TIMACS	0.77	(0.54-1.10)	45.7%	
RIDDLE-INSTEMI	0-88	(0.34-2.27)	6-3%	
Random effects model	0.76	(0.58-1.00)	79.3%	~
atients without elevated o	ardiac bioma	rkers at baseline		
ABOARD			0-0%	
ELISA	0-64	(0.06-7.08)	1.0%	·
LISA-3	1.19	(0-32-4-43)	3-3%	· · · · · · · · · · · · · · · · · · ·
SAR-COOL	1-31	(0.22-7.84)	1.8%	i
iciahbasi et al			0-0%	
TIMACS	0-97	(0.52-1.81)	14.6%	<del></del>
landom effects model	1.01	(0.59-1.70)	20.7%	
lest for subgroup differences:	D=0-3553			
<b>3</b> ,	1 - 2222			0.1 0.2 0.5 1 2 5 10
				Favours early strategy Favours delayed strategy
				ravous carly survey, ravous delayed survey
Elevated C	Cardia	c Marke	ers	All-Cause Mortality







#### **AHA/ACC Guideline**

#### 2014 AHA/ACC Guideline for the Management of Patients With Non–ST-Elevation Acute Coronary Syndromes A Report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines

Developed in Collaboration With the Society for Cardiovascular Angiography and Interventions and Society of Thoracic Surgeons

#### Table 8. Factors Associated With Appropriate Selection of Early Invasive Strategy or Ischemia-Guided Strategy in Patients With NSTE-ACS

Immediate invasive (within 2 h)	Refractory angina Signs or symptoms of HF or new or worsening mitral regurgitation
	Hemodynamic instability
	Recurrent angina or ischemia at rest or with low-level activities despite intensive medical therapy
	Sustained VT or VF
lschemia-guided strategy	Low-risk score (eg, TIMI [0 or 1], GRACE [<109]) Low-risk Tn-negative female patients
	Patient or clinician preference in the absence
	of high-risk features
Early invasive (within 24 h)	of high-risk features None of the above, but GRACE risk score >140 Temporal change in Tn (Section 3.4)
Early invasive (within 24 h)	None of the above, but GRACE risk score >140
Early invasive (within 24 h) Delayed invasive (within 25–72 h)	None of the above, but GRACE risk score >140 Temporal change in Tn (Section 3.4)
Delayed invasive (within	None of the above, but GRACE risk score >140 Temporal change in Tn (Section 3.4) New or presumably new ST depression None of the above but diabetes mellitus Renal
Delayed invasive (within	None of the above, but GRACE risk score >140 Temporal change in Tn (Section 3.4) New or presumably new ST depression None of the above but diabetes mellitus Renal insufficiency (GFR <60 mL/min/1.73 m <sup>2</sup> )
Delayed invasive (within	None of the above, but GRACE risk score >140 Temporal change in Tn (Section 3.4) New or presumably new ST depression None of the above but diabetes mellitus Renal insufficiency (GFR <60 mL/min/1.73 m <sup>2</sup> ) Reduced LV systolic function (EF <0.40)
Delayed invasive (within	None of the above, but GRACE risk score >140 Temporal change in Tn (Section 3.4) New or presumably new ST depression None of the above but diabetes mellitus Renal insufficiency (GFR <60 mL/min/1.73 m <sup>2</sup> ) Reduced LV systolic function (EF <0.40) Early postinfarction angina

## 2018 ESC/EACTS Guidelines on myocardial revascularization

Recommendations for invasive evaluation and revascularization in non-ST-elevation acute coronary syndrome

	Recommendatio	Class <sup>a</sup>	L ovol <sup>b</sup>			
	Urgent coronary an ommended in patie mic risk ( <i>Figure 4</i> ). <sup>19</sup>	1.1	с			
		trategy (<24 h) is recom- s with at least one high- re 4). <sup>164,174,176</sup>				
	An invasive strategy presentation) is ind with at least one int rion ( <i>Figure 4</i> ) or re symptoms. <sup>170,171</sup>	ı	<b>A</b>			
C	Invasive evalua	tion in Non-ST-Elevation Acute Corona	ary Syndromes			
	÷	↓ l		¥		
	Very High-Risk Haemodynamic instability or cardiogenic shock Recurrent/ongoing chest pain refractory to medical tut Life-threatening arrhythmias or cardiac arrest Mechanical complications of MI Acute heart failure Recurrent dynamic STF wave changes*	High-Risk • Established diagnosis of non-ST-elevation myocardial infarction based on cardiac troponins Dynamic ST/T-changes (symptomatic or silent) • GRACE score >140	Intermediate Risk  Diabetes mellitus or renal insufficiency  UVEF <40% or congestive heart failure confidention angina or prior ECU/00  GRACE risk sore >109 and <140 or recurrent symptoms/ischaemia on non-invasive testing.			
				÷	ç	
	Immediate Invasive (<2 hours) IC IA Invasive (<24 hours) IA Invasive (<72 hours) IA					

CABG = connary artery bypass grafting: GRACE = Global Registry of Acute Coronary Events; LVEF = left ventricular ejection fraction; MI =myocardial infarction PCI = percutaneous coronary intervention.

\*Particularly intermittent ST-elevation; \*Estimated glomerular filtration rate <60mL/min/1.73m2

According to ESC NSTE-ACS 2015 Guidelines



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# THANK YOU

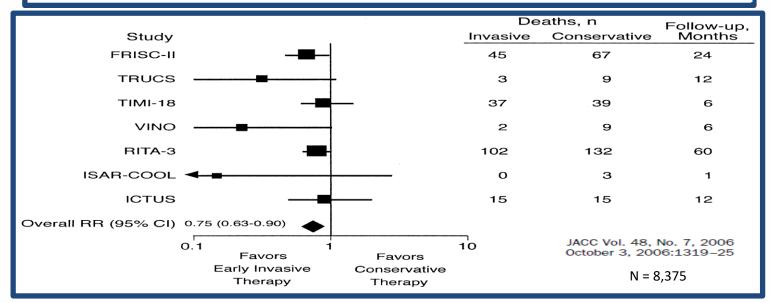






## Benefit of Early Invasive Therapy in Acute Coronary Syndromes

A Meta-Analysis of Contemporary Randomized Clinical Trials

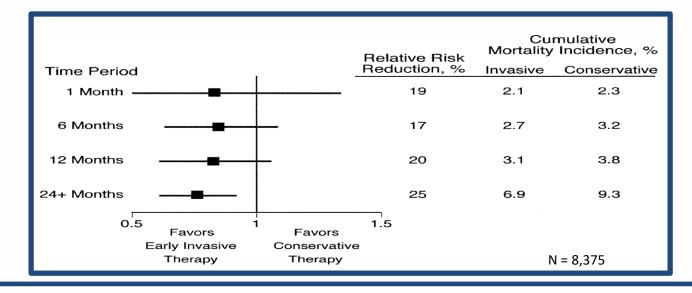


Relative risk of all-cause mortality for early invasive therapy compared with conservative therapy at a mean

follow-up of 2 years.



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Relative risk of all-cause mortality for early invasive therapy compared with conservative therapy as a function of

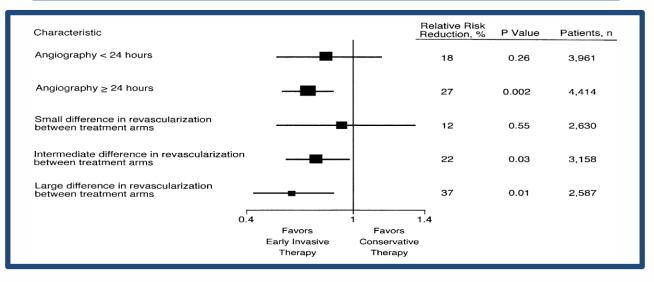
time

JACC Vol. 48, No. 7, 2006 October 3, 2006:1319-25









Relative risk of all-cause mortality based on time of angiography and the extent of revascularisation

JACC Vol. 48, No. 7, 2006 October 3, 2006:1319-25





