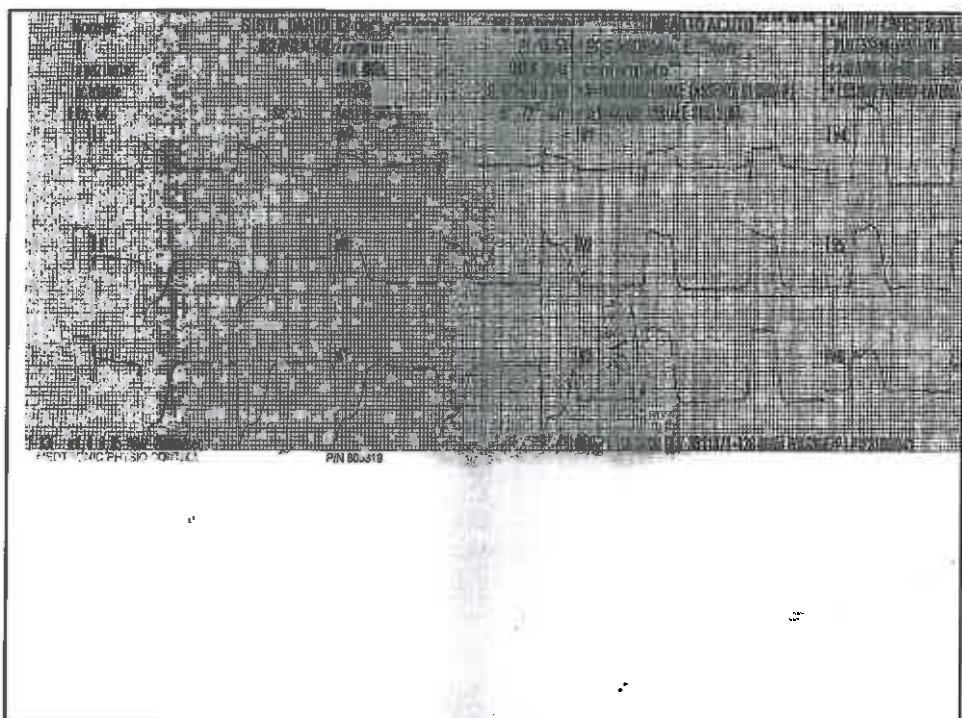
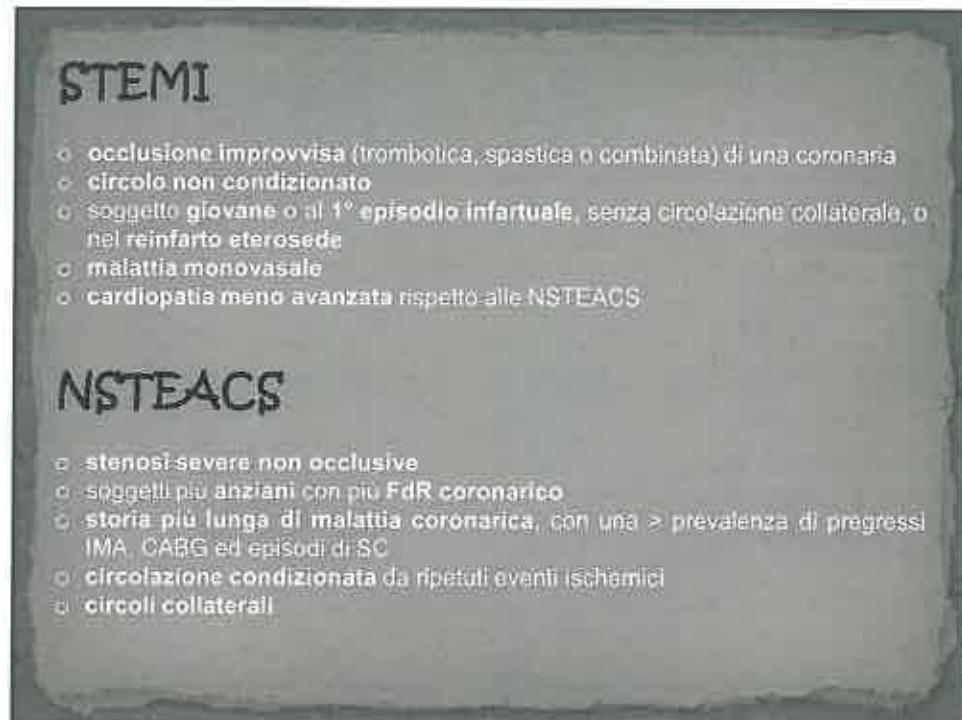


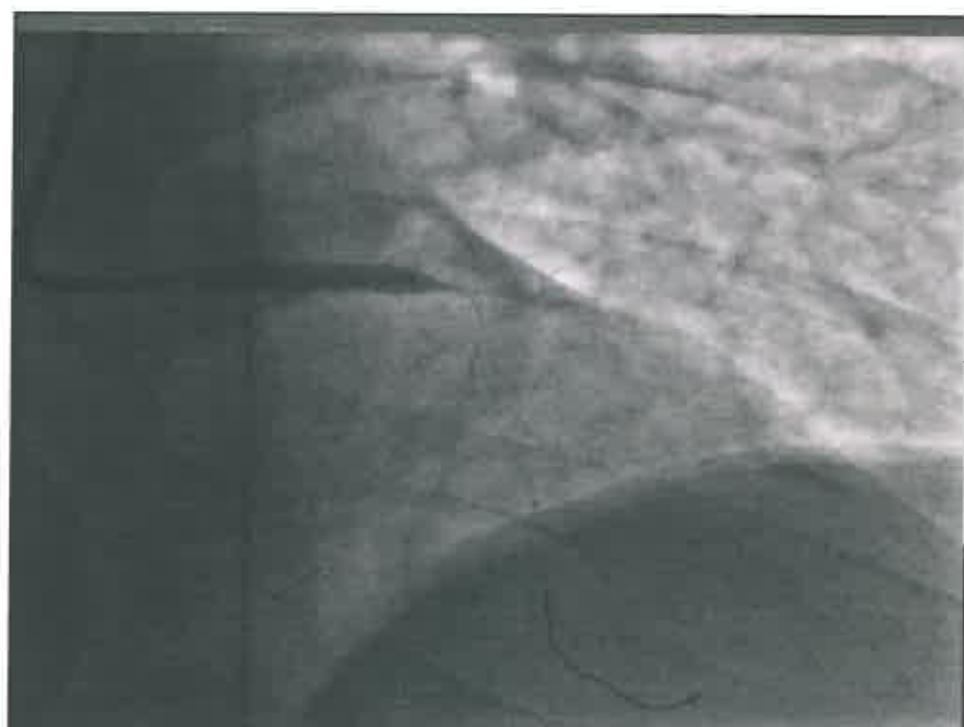
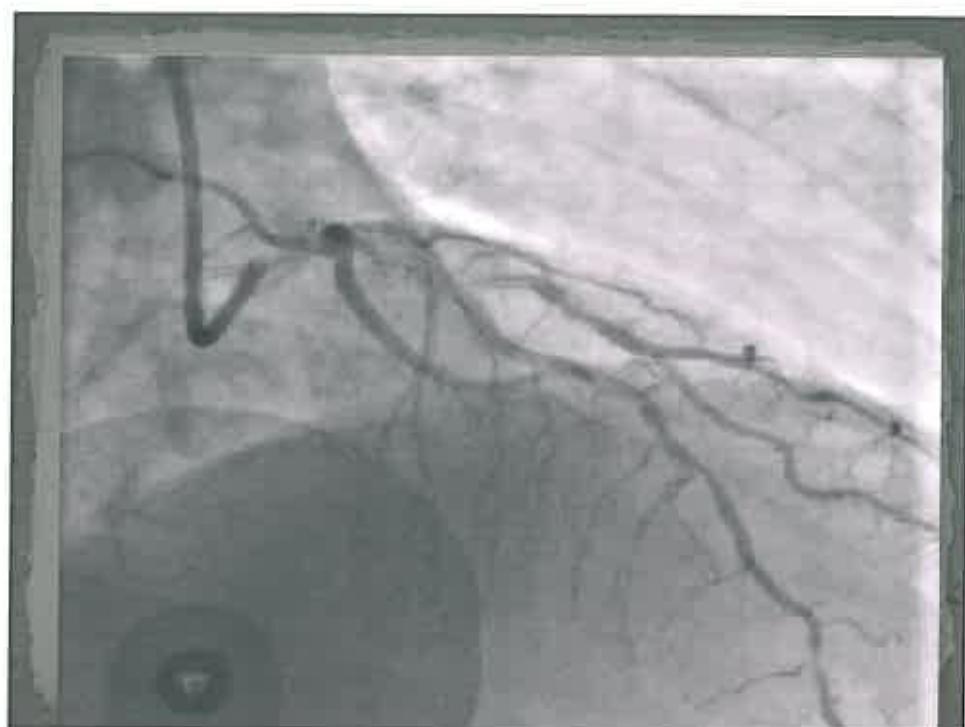
Criteria for Acute Myocardial Infarction

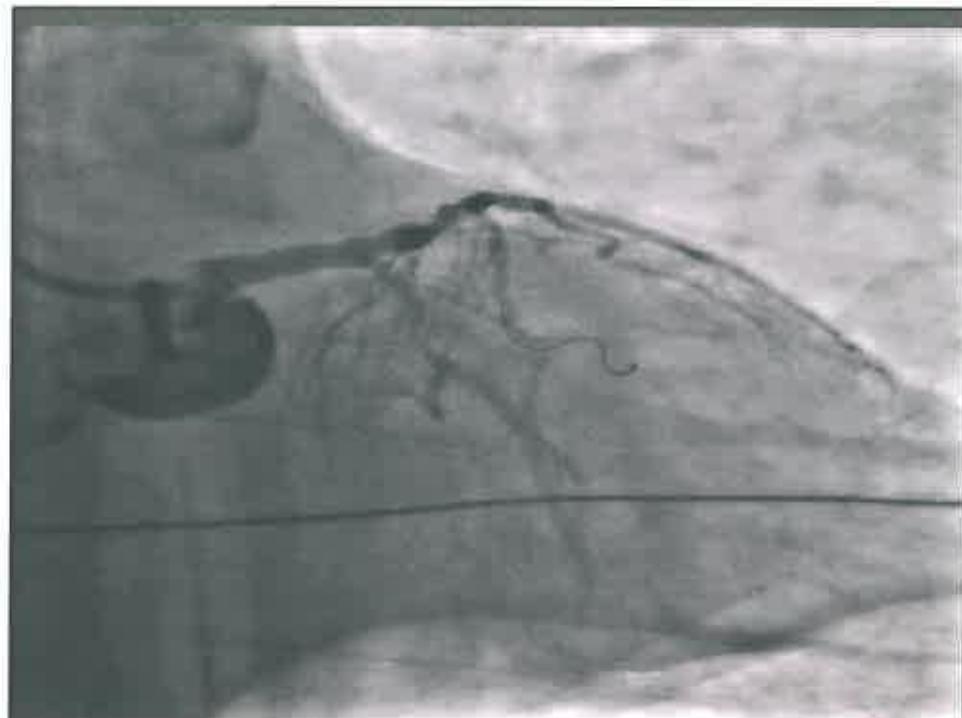
- Detection of a rise and/or fall of cardiac biomarker values (preferably cardiac troponin) with at least one value above the 99th percentile upper reference limit (URL) and with at least one of the following:
 - Ischaemic symptoms;
 - ECG changes of new ischaemia (new ST-T changes or new LBBB);
 - Development of pathologic Q waves in the ECG;
 - Imaging evidence of new loss of viable myocardium or new regional wall motion abnormality;
 - Identification of an intracoronary thrombus by angiography or autopsy.

Elevations of Cardiac Troponin Values because of Myocardial Injury

Injury related to primary myocardial ischaemia	Injury not related to myocardial ischaemia
<ul style="list-style-type: none"> • Plaque rupture. • Intraluminal coronary artery thrombus formation. 	<ul style="list-style-type: none"> • Cardiac contusion, surgery, ablation, pacing, or defibrillator shocks. • Rhabdomyolysis with cardiac involvement. • <u>Myocarditis</u>. • Cardiotoxic agents, e.g. anthracyclines, herceptin.
Injury related to supply/demand imbalance of myocardial ischaemia	
<ul style="list-style-type: none"> • <u>Tachy-/brady-arrhythmias</u>. • Aortic dissection or severe aortic valve disease. • Hypertrophic cardiomyopathy • Cardiogenic, hypovolaemic, or septic shock. • Severe respiratory failure. • Severe anaemia. • <u>Hypertension with or without LVH</u>. • Coronary spasm. • Coronary embolism or vasculitis. • Coronary endothelial dysfunction without significant CAD. 	Multifactorial or indeterminate myocardial injury <ul style="list-style-type: none"> • <u>Heart failure</u> • Stress (Takotsubo) cardiomyopathy • Severe <u>pulmonary embolism</u> or pulmonary hypertension. • Sepsis and critically ill patients. • <u>Renal failure</u>. • Severe acute neurological diseases, e.g. <u>stroke</u>, subarachnoid haemorrhage • Infiltrative diseases, e.g. amyloidosis, sarcoidosis. • Strenuous exercise.



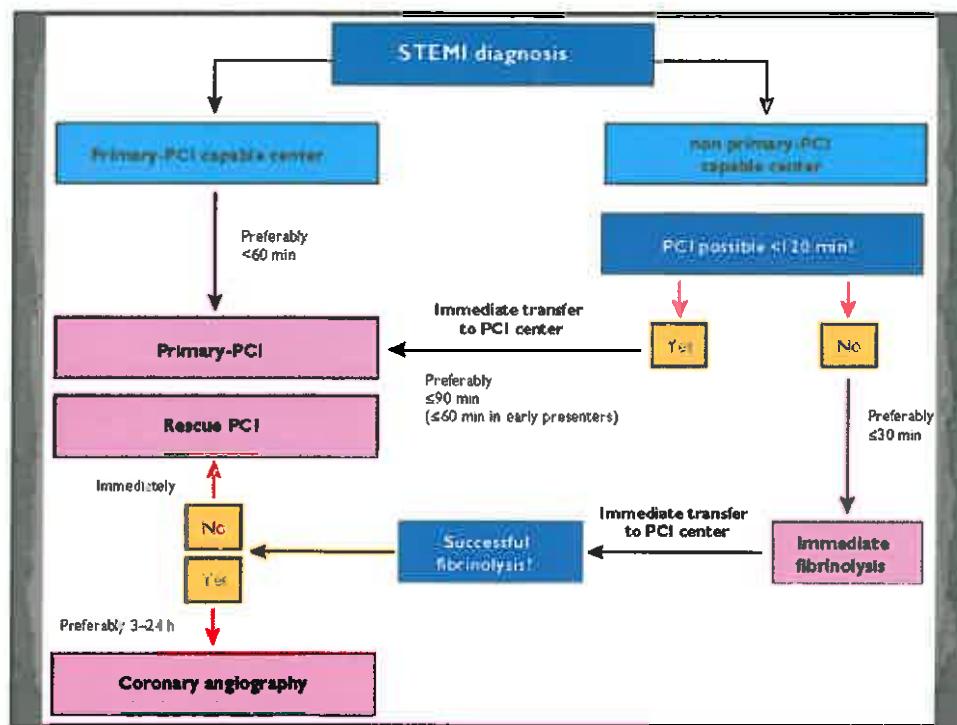




NSTEMI (MA NON Q) - INFARTO SUBENDOCARDICO

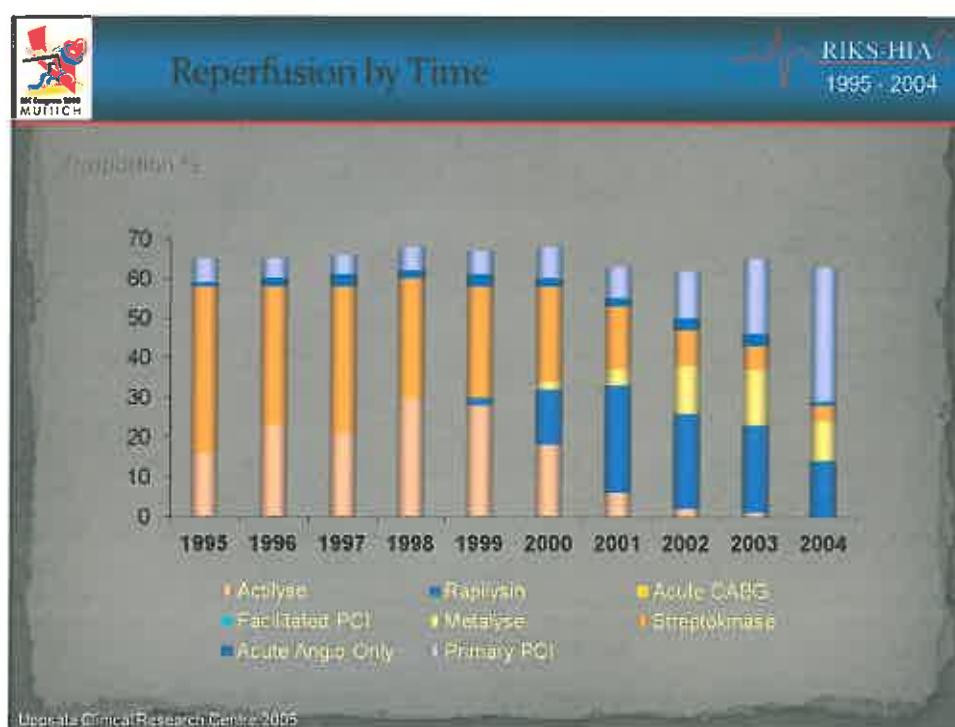
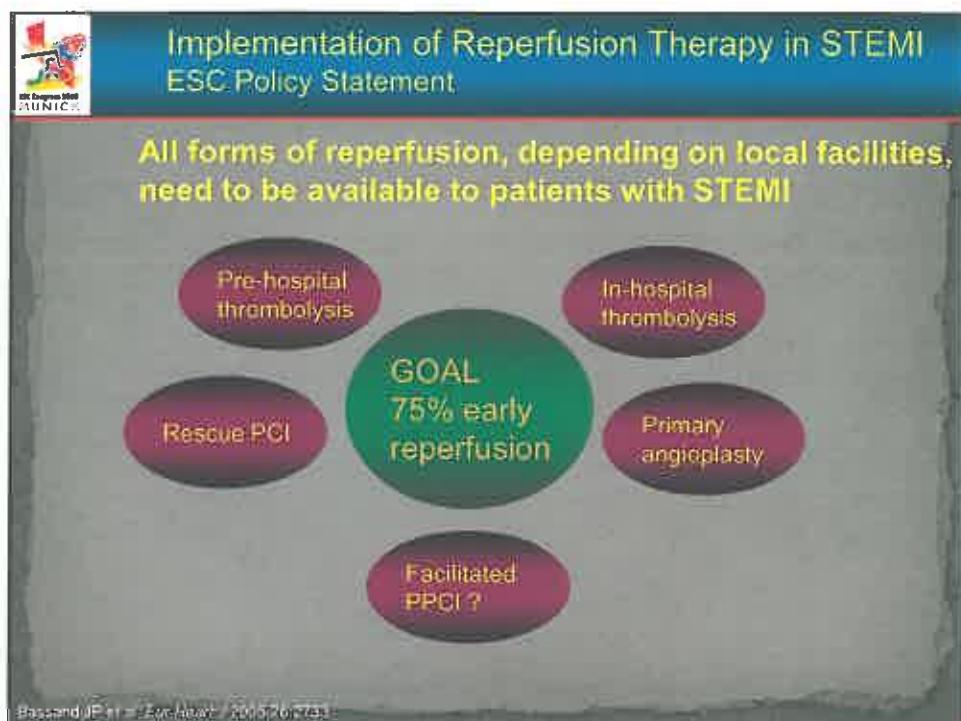
- ST **elevation**, anche se talvolta è possibile che ci sia solo **depressione acuta del livello** ?
- mancata comparsa di onde Q di necrosi, per cui l'NSTEMI è spesso denominato **IMA non Q**

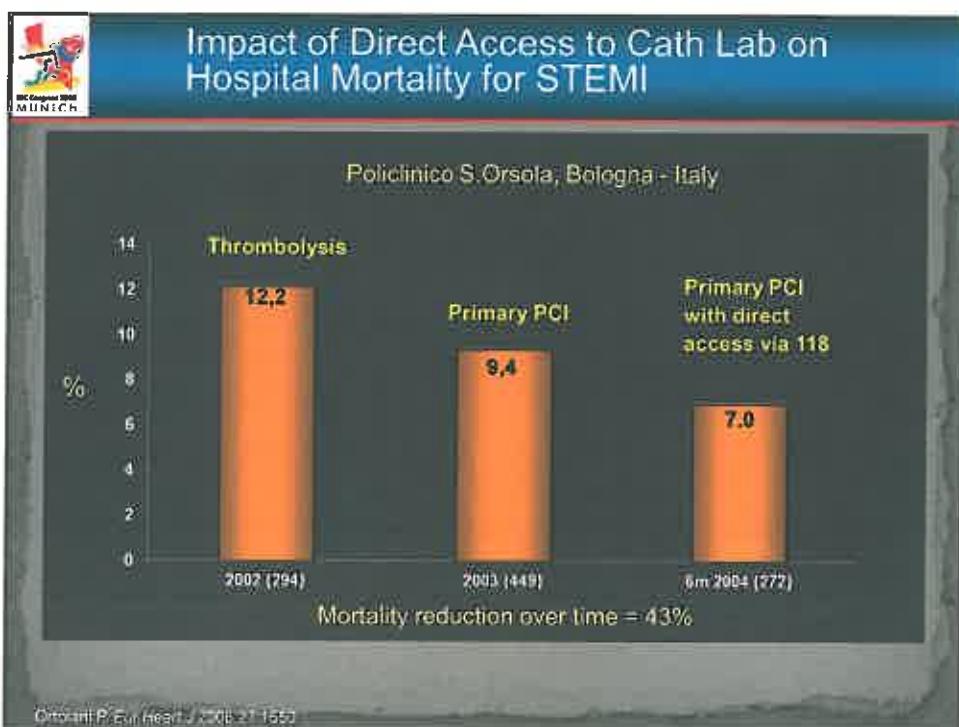
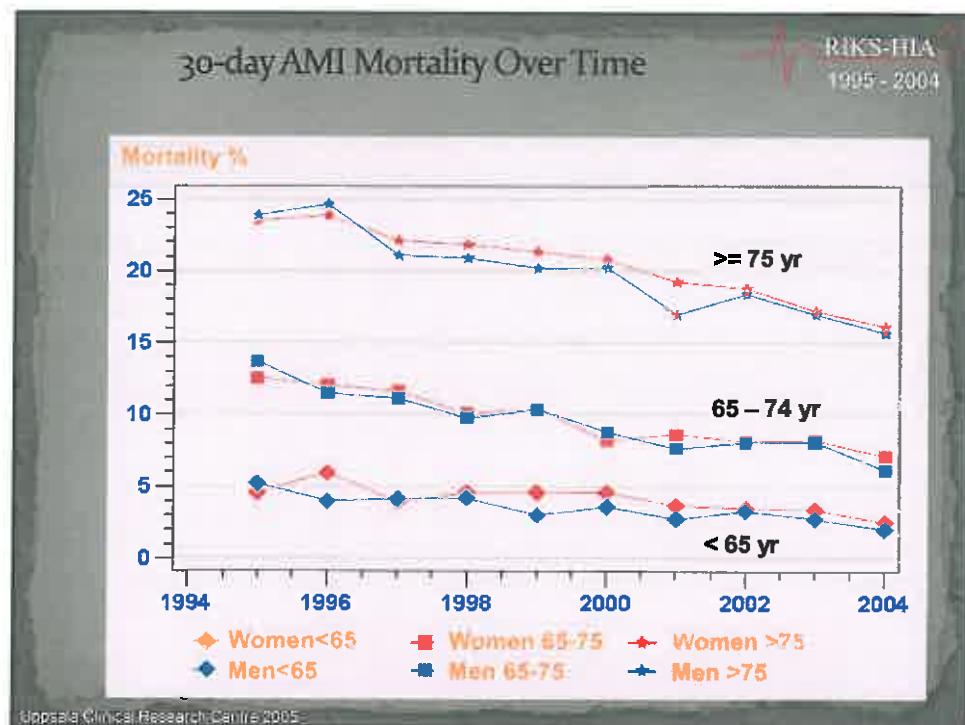
The figure displays three separate ECG leads. Each lead shows a normal P wave followed by a sharp Q wave, a tall R wave, and a prominent S wave. The ST segment and T wave are labeled. The first lead, labeled 'STEMI ST Elevation', shows a clearly elevated ST segment above the baseline. The second lead, labeled 'NSTEMI ST Depression', shows a ST segment that is slightly below the baseline. The third lead, also labeled 'NSTEMI ST Depression', shows a similar ST depression.



Reperfusion therapy

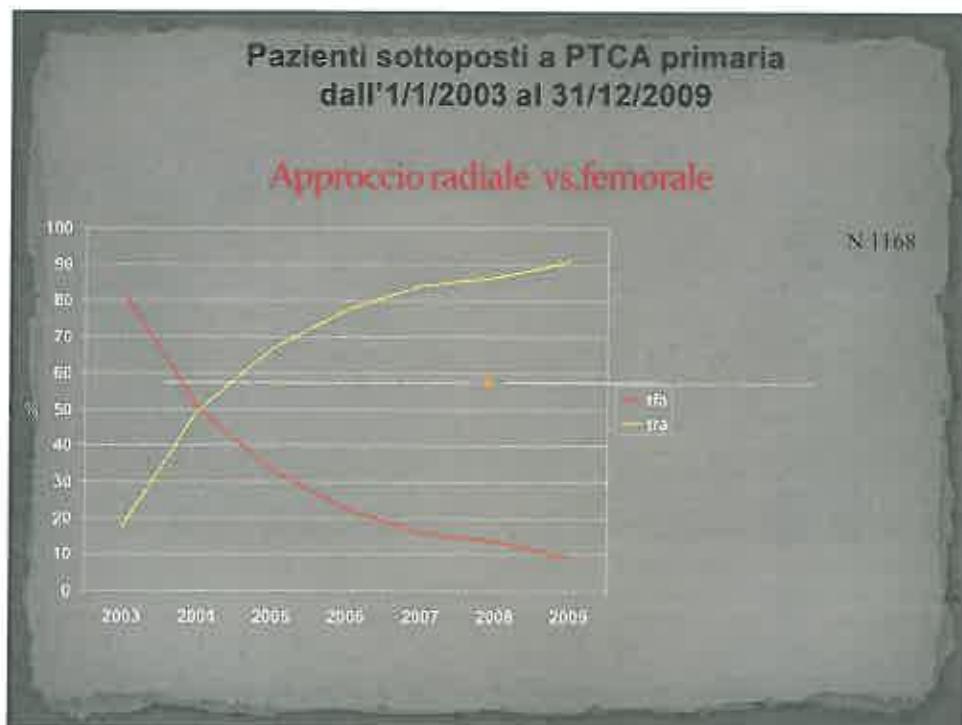
Recommendations	Class	Level
Reperfusion therapy is indicated in all patients with symptoms of <12 h duration and persistent ST-segment elevation or (presumed) new LBBB.	I	A
Reperfusion therapy (preferably primary PCI) is indicated if there is evidence of ongoing ischaemia, even if symptoms may have started > 12 h beforehand or if pain and ECG changes have been stuttering.	I	C
Reperfusion therapy with primary PCI may be considered in stable patients presenting 12-24 h after symptom onset.	IIb	B
Routine PCI of a totally occluded artery > 24 h after symptom onset in stable patients without signs of ischaemia (regardless of whether fibrinolysis was given or not) is not recommended.	III	A

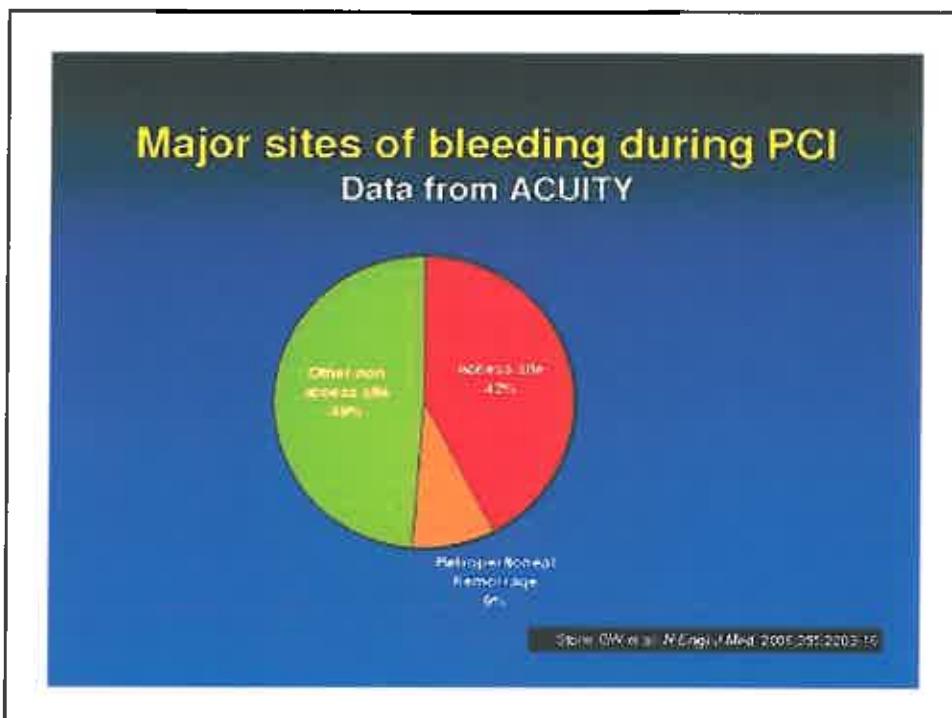
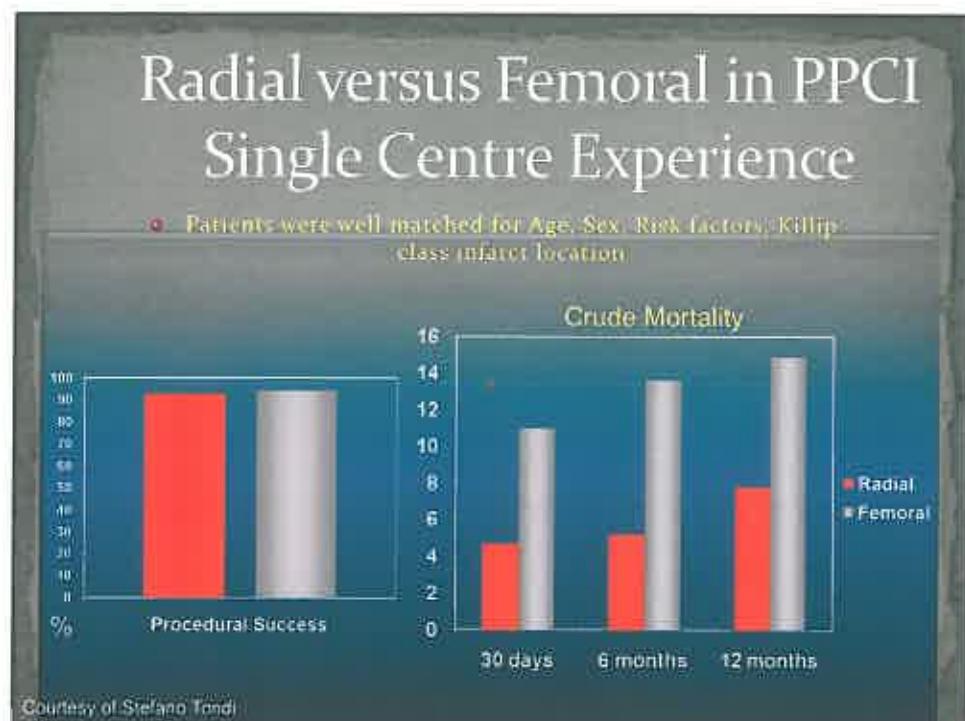


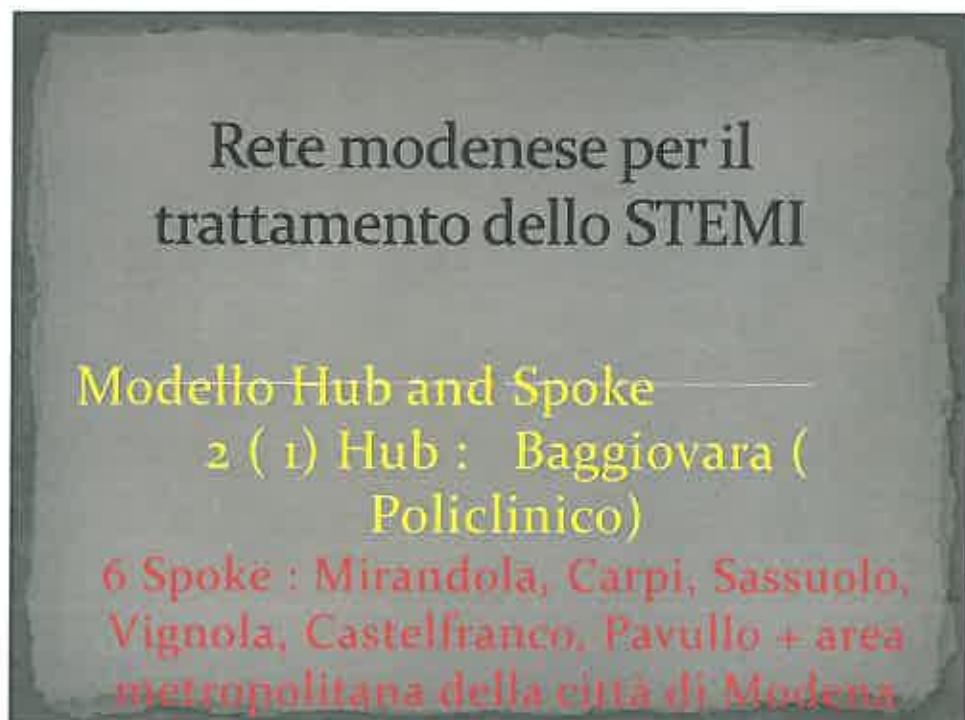
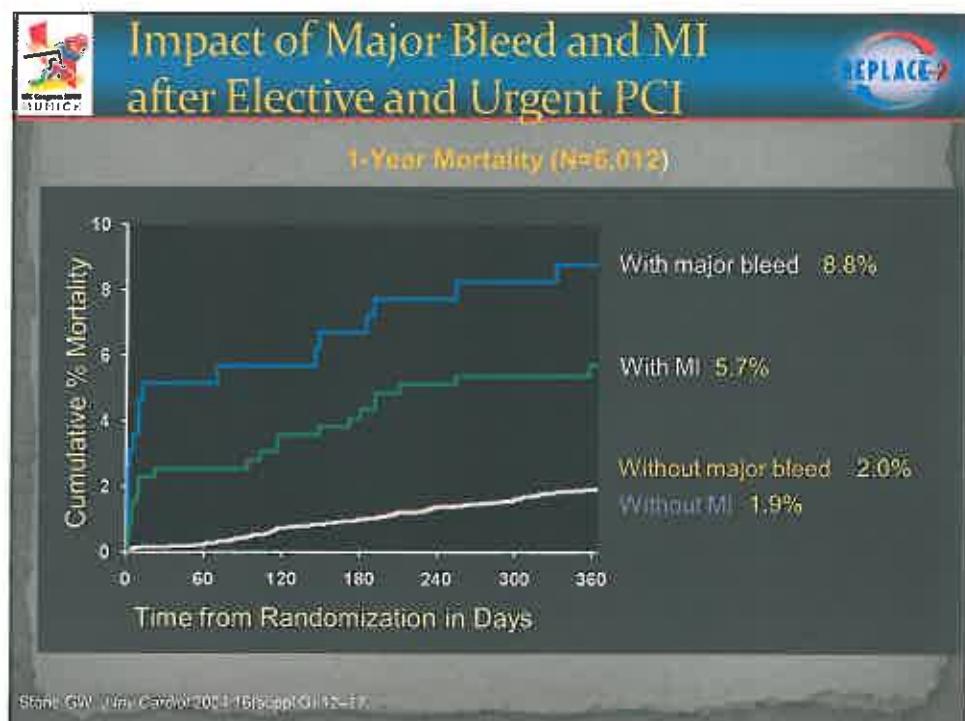


Procedural aspects of primary PCI

Recommendations	Class	Level
Procedural aspects of primary PCI		
Primary PCI should be limited to the culprit vessel with the exception of cardiogenic shock and persistent ischaemia after PCI of the supposed culprit lesion.	IIa	B
If performed by an experienced radial operator, radial access should be preferred over femoral access.	IIa	B
If the patient has no contraindications to prolonged DAPT (indication for oral anticoagulation, or estimated high long-term bleeding risk) and is likely to be compliant, DES should be preferred over BMS.	IIa	A
Routine thrombus aspiration should be considered.	IIa	B





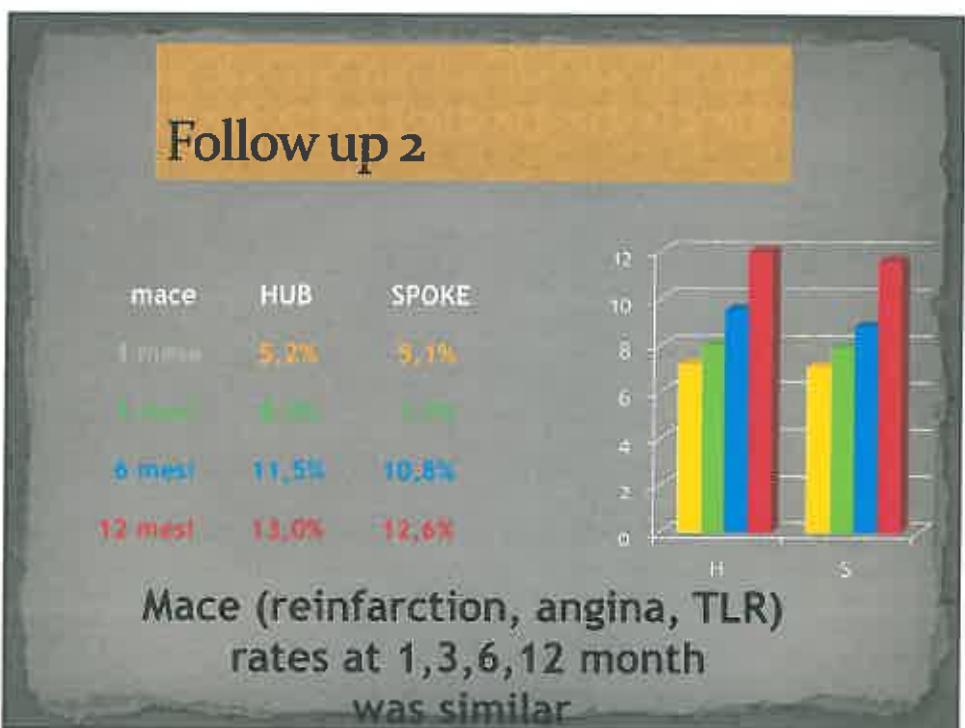
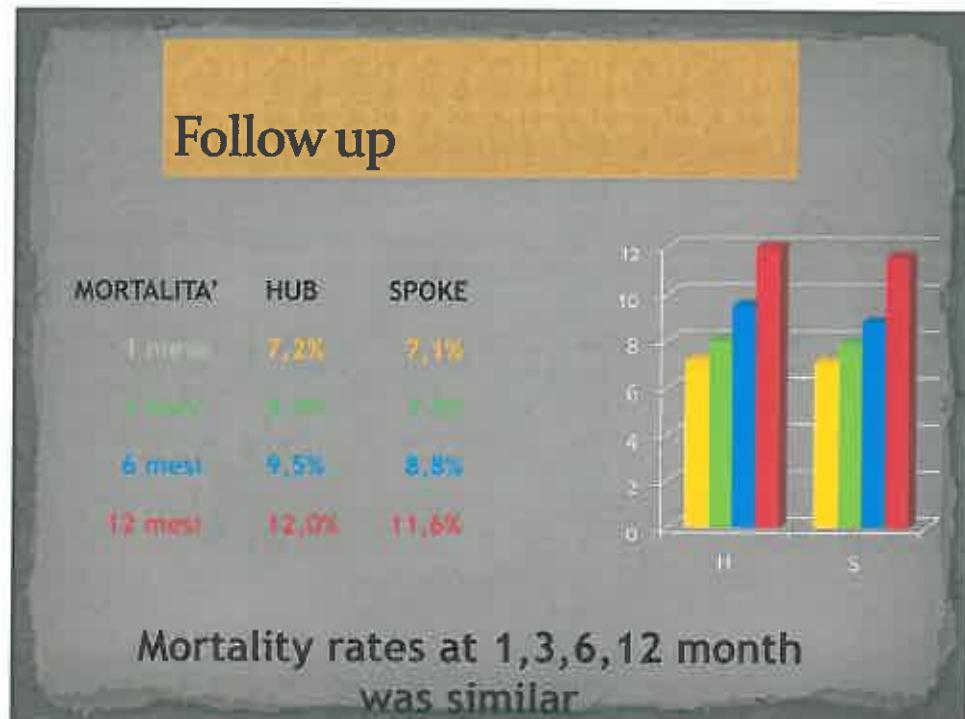


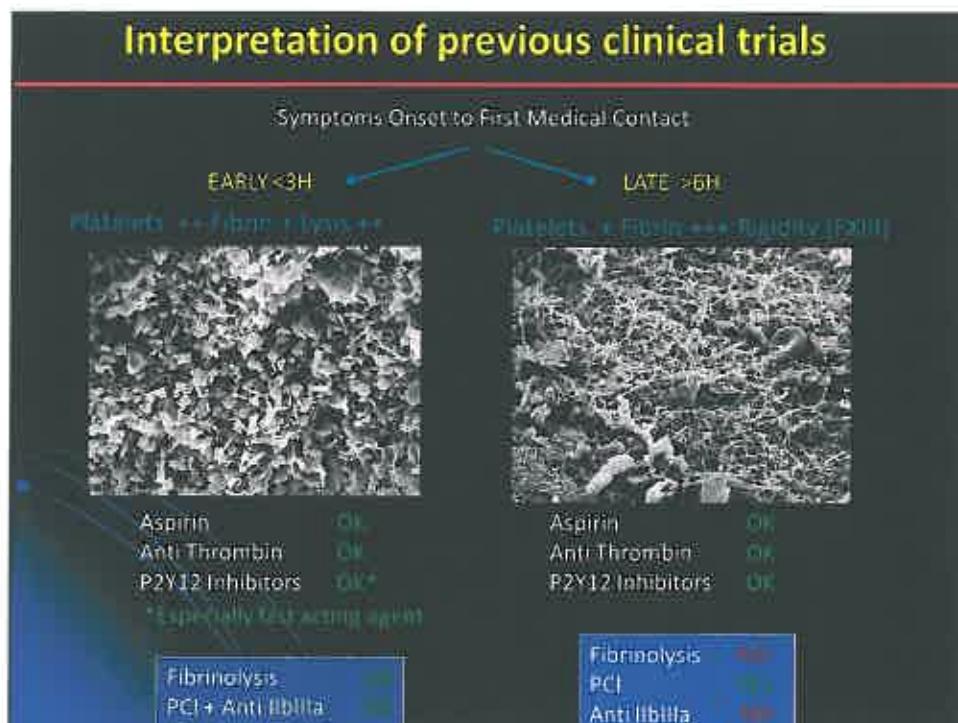
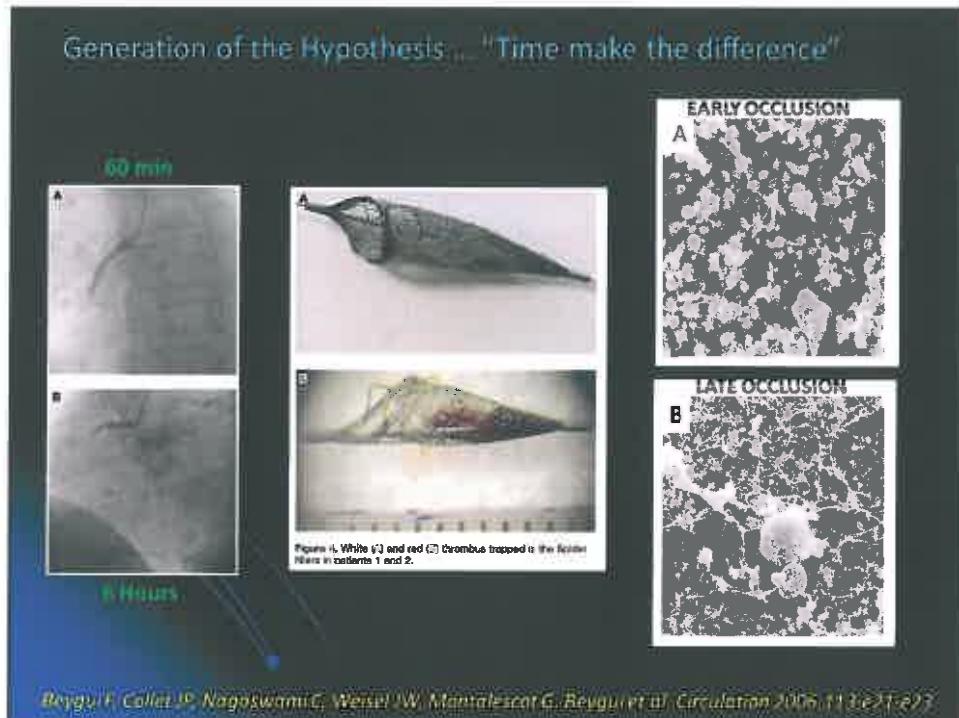
I pazienti che provengono dai centri periferici e quelli che giungono direttamente al NOCSAE sono risultati comparabili per età, sesso, principali fattori di rischio ecc.

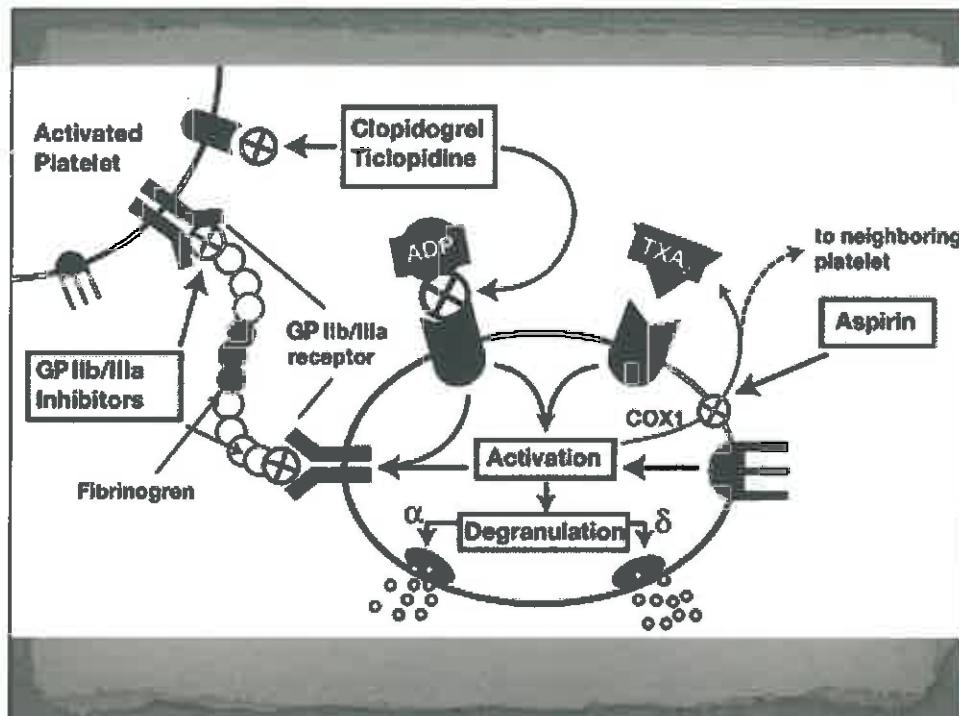
	GROUP A: HUB	GROUP B: SPOKE	p
Age (aa)	66,5	66,5	n.s.
sex (male %)	77,0	75,7	n.s.
diabetes (%)	13,1	14,7	n.s.
active smoker (%)	21,5	20,9	n.s.
hypertension (%)	28,5	25,1	n.s.
shock (%)	5,1	5,9	n.s.
Killip 1-2 (%)	96,3	95,8	n.s.

Il tempo intercorso fra la diagnosi ECG di STEMI e l'arrivo nel CathLab era **significativamente maggiore** nel gruppo di pazienti giunti dalla periferia (tempo di trasporto)



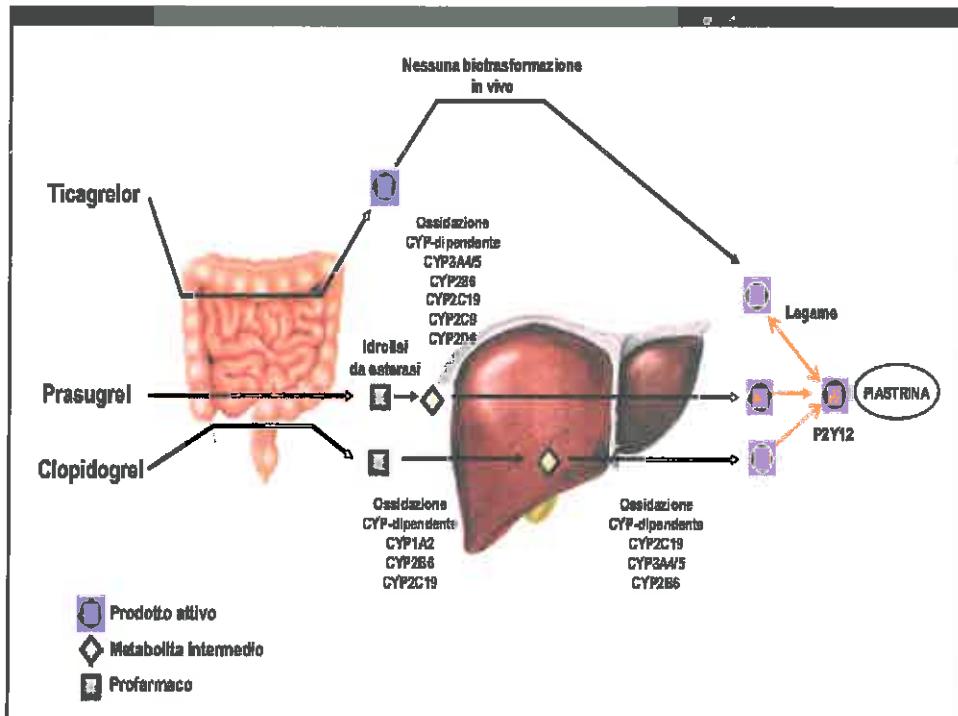






P2Y₁₂ Inhibitors

	Clopidogrel	Prasugrel	Ticagrelor
Class	Thienopyridine	Thienopyridine	Triazolopyrimidine
Reversibility	Irreversible	Irreversible	Reversible
Activation	Prodrug, limited by metabolism	Prodrug, not limited by metabolism	Active drug
Onset of effect ^a	2–4 h	30 min	30 min
Withdrawal before major surgery	5 days	7 days	5 days

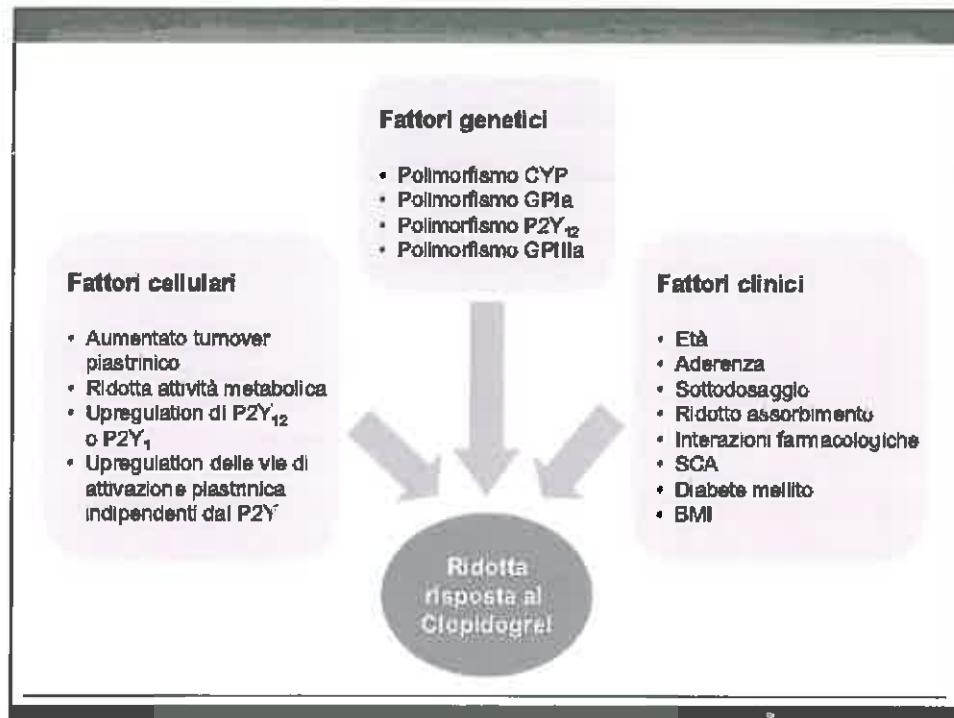


Clopidogrel (Plavix®) e Ticlopidina (Tiklid®)

- necessitano dell'attivazione epatica (P450)
- Ticlopidina può dare piastrinopenia e diarrea, è utilizzata spesso dai chirurghi vascolari per curare l'AOCP
- variabilità interindividuale della risposta terapeutica al Clopidogrel (spesso diabetici, anziani, pz con SCA sottoposti a PCI)

Prasugrel (Efient®) e Ticagrelor (Brilique®)

- aggiungono più rapidamente minore variabilità interindividuale dell'efficacia (p. es. meno interazioni farmacologiche (NO attivazione epatica))
- Prasugrel è più rapido ma ha > rischio di sanguinamento (Clopidogrel e Ticagrelor sono più sicuri)



Fibrinolytic therapy

Recommendations	Class	Level
Fibrinolytic therapy is recommended within 12 h of symptom onset in patients without contraindications if primary PCI cannot be performed by an experienced team within 120 min of FMC.	I	A
In patients presenting early (< 2 h after symptom onset) with a large infarct and low bleeding risk, fibrinolysis should be considered if time from FMC to balloon inflation is > 90 min.	IIa	B
If possible, fibrinolysis should start in the prehospital setting.	IIa	A
A fibrin-specific agent (tenecteplase, alteplase, reteplase) is recommended (over non-fibrin specific agents).	I	B
Interventions following fibrinolysis		
Rescue PCI is indicated immediately when fibrinolysis has failed (< 50 % ST-segment resolution at 60 min).	I	A
Emergency PCI is indicated in the case of recurrent ischaemia or evidence of reocclusion after initial successful fibrinolysis.	I	B

FIBRINOLISI

- * entro 2 h dall'inizio dei sintomi (comunque entro 12 h)
- * se sintomi insorti da non >2 h e PPCI non eseguibile entro 90 min.
- * senza controindicazioni al trattamento

CONTROINDICAZIONI

- 1 stroke emorragico o di crisi cognitiva
- 2 stroke ischemico nei precedenti 6 mesi
- 3 tumore cerebrale/metastasi SNC
- 4 trauma > 7 giorni cranico / chirurgia nelle 3 settimane precedenti
- 5 sanguinamento IC nel mese precedente
- 6 endocrinologia emorragica nota o in atto
- 7 grossopatia perniciosa
- 8 puntura non compattabile <24 h (biopsia epatica, puntura lombare)

In ogni caso, senza attendere il risultato della fibrinolisi, il pz deve essere trasportato presso un centro Hub per eseguire, in caso di mancata riperfusione, coronarografia.

Contraindications to fibrinolytic therapy

Relative

Transient ischaemic attack in the preceding 6 months.

Oral anticoagulant therapy

Pregnancy or within 1 week postpartum.

Refractory hypertension (systolic blood pressure > 180 mmHg and/or diastolic blood pressure > 110 mmHg).

Advanced liver disease.

Infective endocarditis.

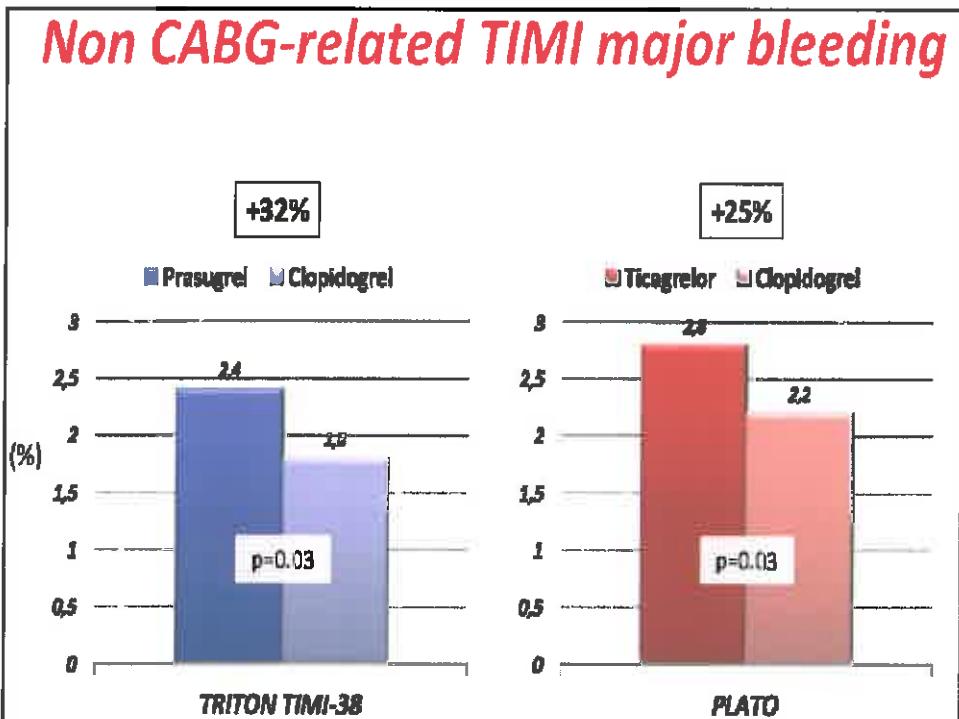
Active peptic ulcer.

Prolonged or traumatic resuscitation.

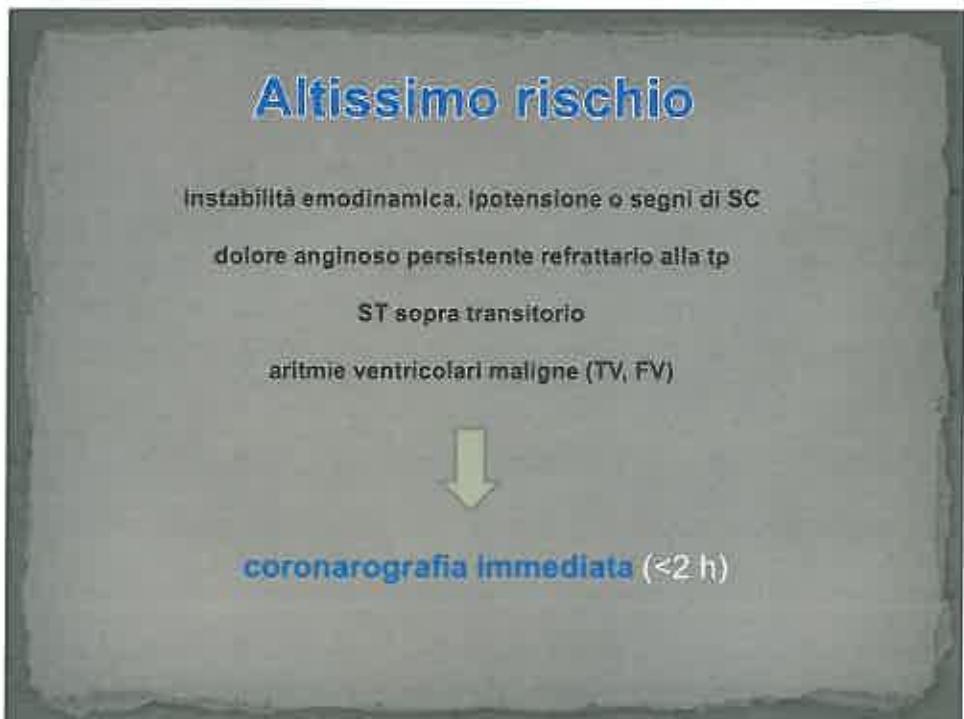
Routine therapies in the acute, subacute and long term phase of STEMI

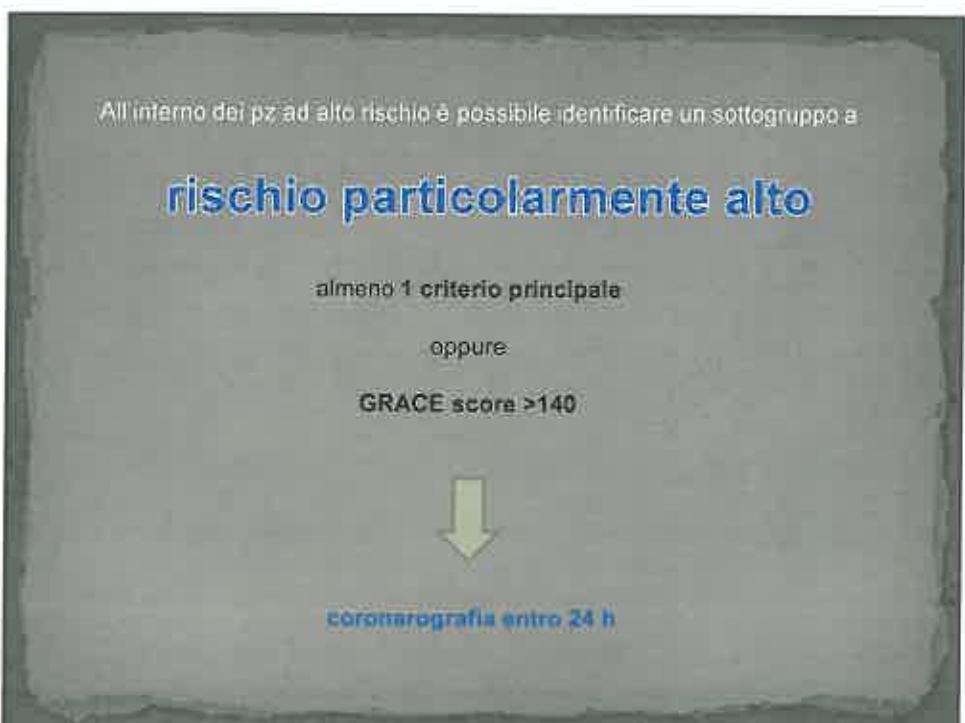
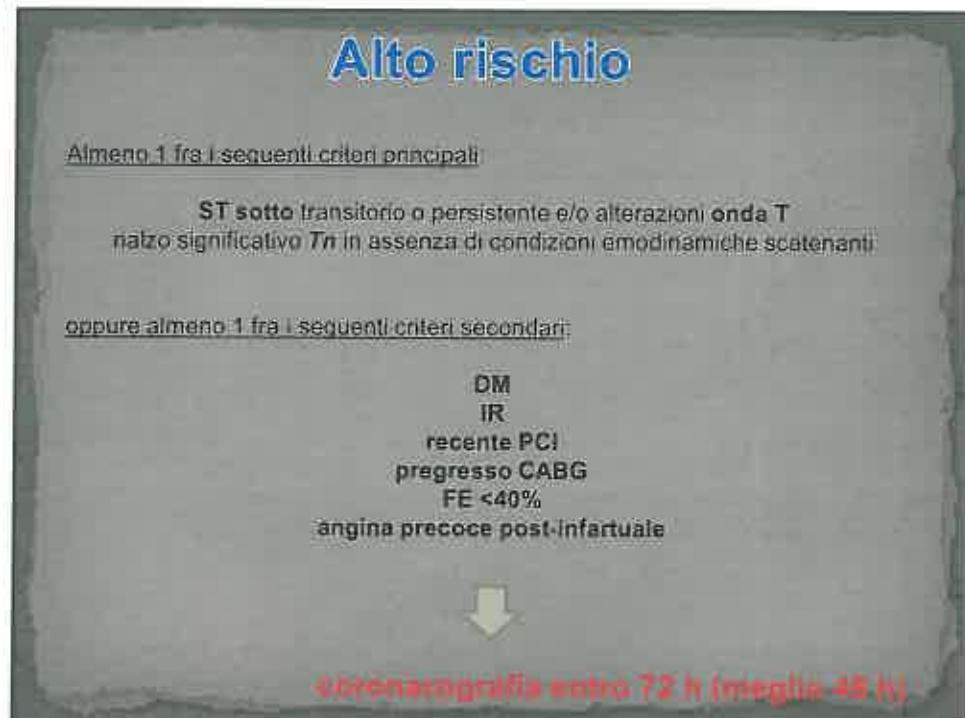
Recommendations	Class	Level
In patients with left ventricular thrombus, anticoagulation should be instituted for a minimum of 3 months.	Ia	B
In patients with a clear indication for oral anticoagulation (e.g. atrial fibrillation with CHA ₂ DS ₂ -VASc Score ≥ 2 or mechanical valve prosthesis), oral anticoagulation must be implemented in addition to antiplatelet therapy.	I	C
In patients require triple antithrombotic therapy, combining DAPT and OAC, e.g. because of stent placement and an obligatory indication for OAC, the duration of dual antiplatelet therapy should be minimized to reduce bleeding risk.	I	C
In selected patients who receive aspirin and clopidogrel, low-dose rivaroxaban (2.5 mg twice daily) may be considered if the patient is at low bleeding risk.	IIb	B

Non considerare univocamente a 12 mesi la durata della DAPT dopo STEMI, ma considerare durate minime di:
 - BMS: 3-6 mesi
 - DES: 1 anno



NSTEMI		
Recommendations	Class	Level
It is recommended to use established risk scores for prognosis and bleeding (e.g. GRACE, CRUSADE).	I	B
Additional ECG leads (V_3R , V_4R , V_7-V_9) are recommended when routine leads are inconclusive	I	C





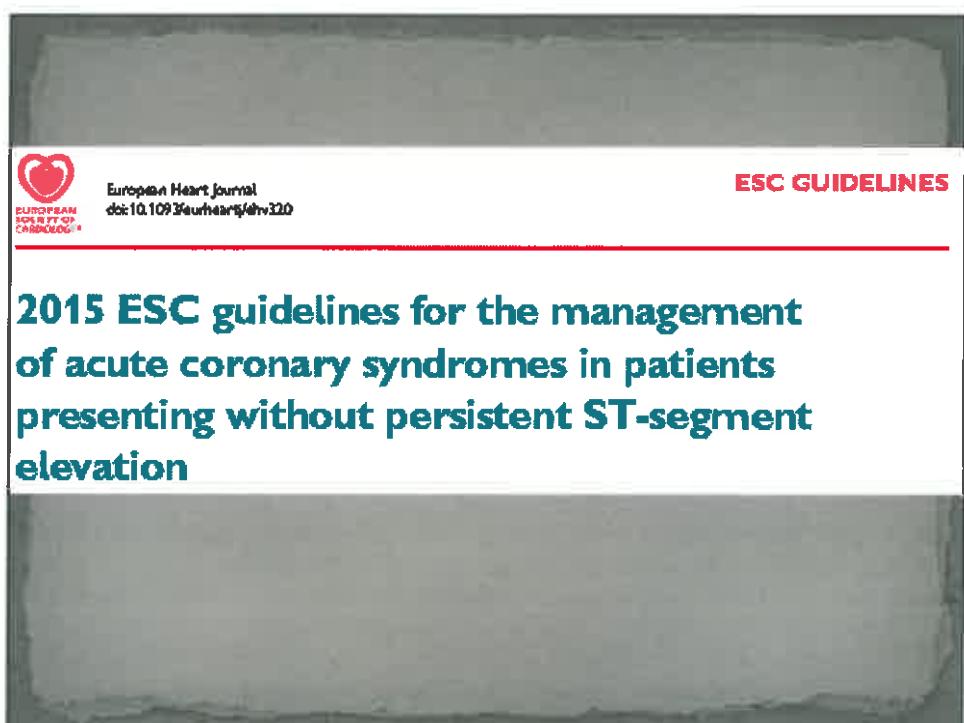
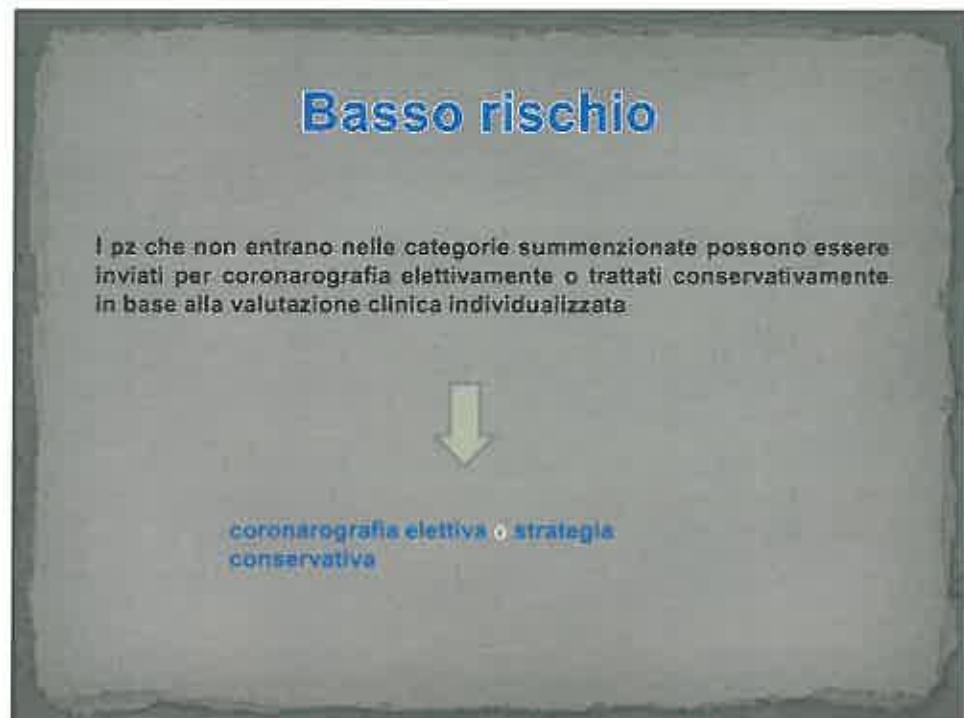


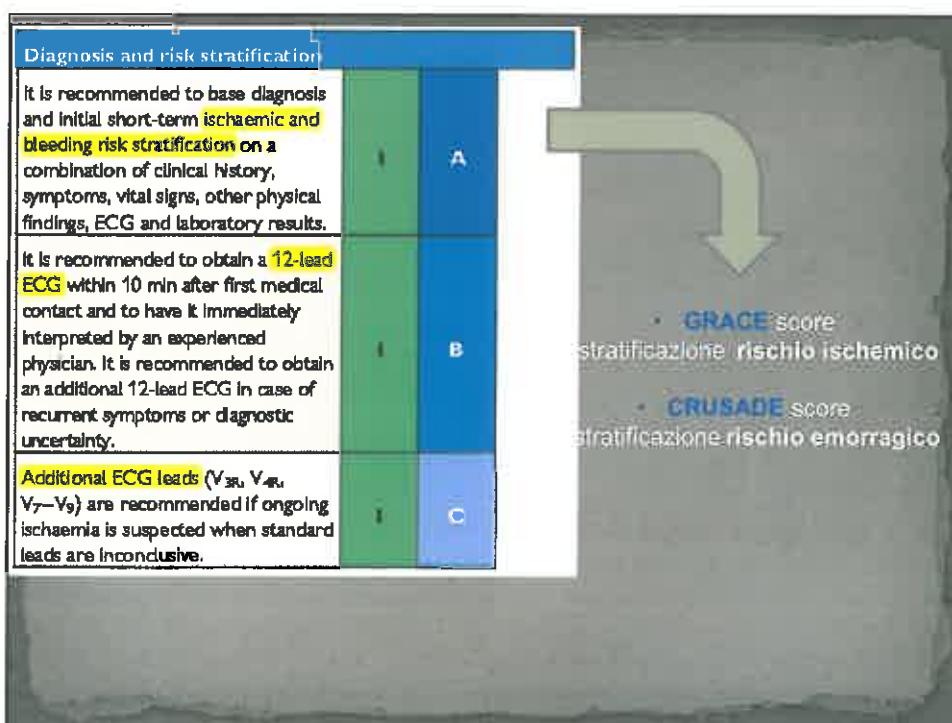
Table 7 Recommended unit and duration of monitoring according to clinical presentation after established NSTEMI-ACS diagnosis

Clinical Presentation	Unit	Rhythm monitoring
Unstable angina	Regular ward or discharge	None
NSTEMI at low risk for cardiac arrhythmias ^a	Intermediate care unit or coronary care unit	≤24 h
NSTEMI at intermediate to high risk for cardiac arrhythmias ^b	Intensive/coronary care unit; or Intermediate care unit	>24 h

NSTEMI — Non-ST-elevation myocardial infarction.

^aIf none of the following criteria: haemodynamically unstable, major arrhythmias, left ventricular ejection fraction <40%, failed reperfusion, additional critical coronary stenoses of major vessels or complications related to percutaneous revascularization.

^bIf one or more of the above criteria are present.



PCR At Admission Risk Model

At Admission (in-hospital/to 6 months)		At Discharge (to 6 months)	
Age	50-59	<input checked="" type="checkbox"/> Cardiac arrest at admission	
HR	70-89	<input checked="" type="checkbox"/> ST-segment deviation	
SBP	120-139	<input checked="" type="checkbox"/> Elevated cardiac enzymes/markers	
Creat.	1.6-1.99	Probability of	Death Death or MI
CHF	III (pulmonary edema)	In-hospital	27% 50%
		To 6 months	30% 70%
<input type="button" value="SI Units"/>		<input type="button" value="Reset"/>	
Calculator Instructions GRACE Info References Disclaimer			

Risk assessment by GRACE score:

We can assess risk by summation of score for all eight parameters.

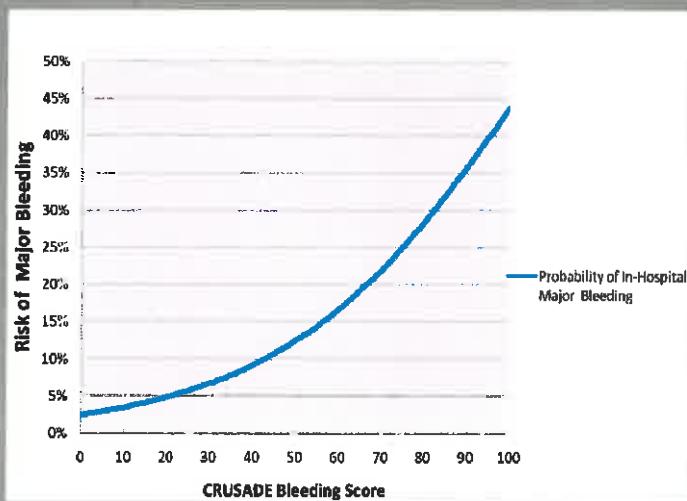
Total score	Risk assessment
≤ 100	Low risk patients – In-hospital death rate less than 1%
101-170	Medium risk patients – In-hospital death rate 1-9%
≥ 171	High risk patients – In-hospital death rate more than 9%

CRUSADE Bleeding Score Nomogram

Predictor	Range	Score
Baseline Hematocrit (%)	33-37 38-40 41-43 44-46 47-49 50-52	0 1 2 3 4 5
Creatinine Clearance (ml/min)	15-30 31-55 56-75 76-100 101-125 126-150	0 1 2 3 4 5
Heart rate (bpm)	60-80 81-90 91-100 101-110 111-120 121-130	0 1 2 3 4 5
Sex:	Male Female	0 1
Signs of CHF at presentation	No Yes	0 1
Prior Vascular Disease	No Yes	0 1
Diabetes Mellitus	No Yes	0 1
Systolic blood pressure (mm Hg)	120-130 131-140 141-150 151-160 161-170 171-180	0 1 2 3 4 5

Note: P=3 / 104 is number of 70 < 70 bpm.
 100% sensitivity defined as > 50 ml/min; Prior Vascular disease = defined as history of stroke, peripheral vascular disease, or coronary artery disease.

Risk of Major Bleeding Across the Spectrum of CRUSADE Bleeding Score



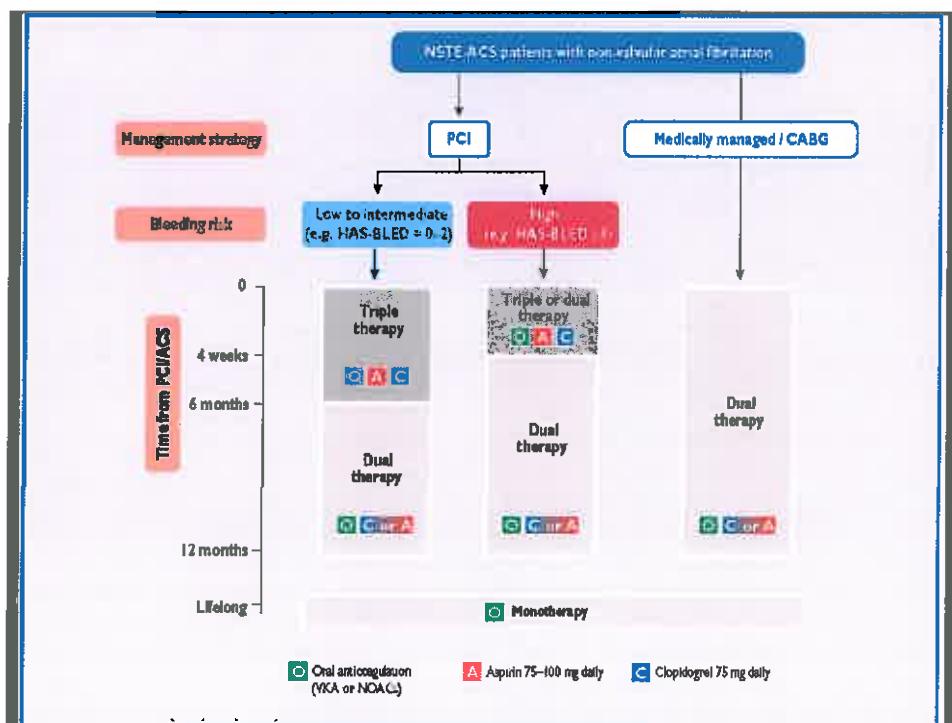
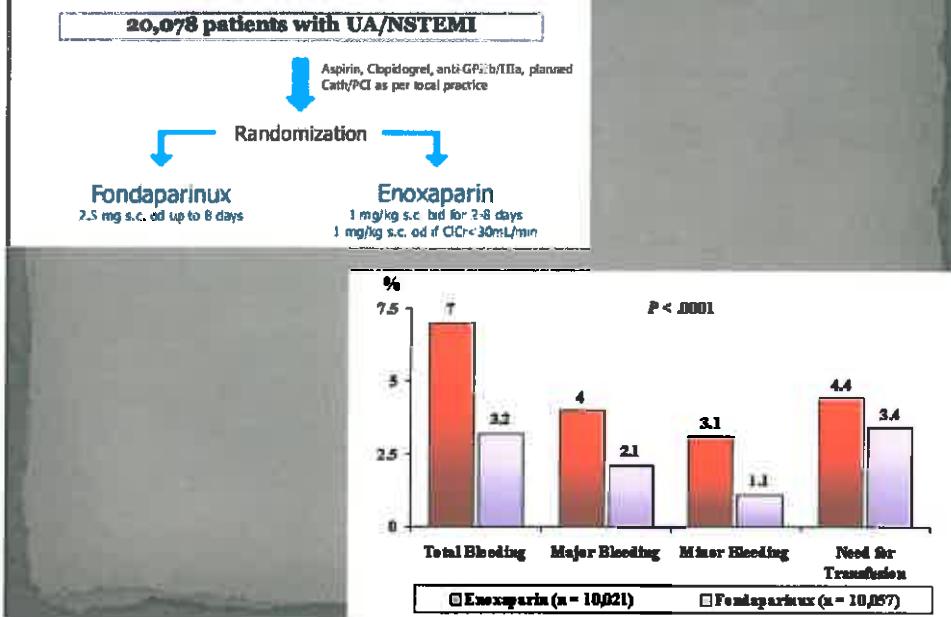
p<0.001 for trend; Derivation: C=0.71 Validation: C=0.70

Recommendations for platelet inhibition in non-ST-elevation acute coronary syndromes		
Recommendations	Class ^a	Level ^b
Oral antiplatelet therapy		
Aspirin is recommended for all patients without contraindications at an initial oral loading dose ^c of 150–300 mg (in aspirin-naïve patients) and a maintenance dose of 75–100 mg/day long-term regardless of treatment strategy.		
A P2Y ₁₂ inhibitor is recommended, in addition to aspirin, for 12 months unless there are contraindications such as excessive risk of bleed:		
• Ticagrelor (180 mg loading dose, 90 mg twice daily) is recommended, in the absence of contraindications, ^d for all patients at moderate-to-high risk of ischaemic events (e.g. elevated cardiac troponins), regardless of initial treatment strategy and including those pretreated with clopidogrel (which should be discontinued when ticagrelor is started).		
• Prasugrel (60 mg loading dose, 10 mg daily dose) is recommended in patients who are proceeding to PCI if no contraindication. ^e		
• Clopidogrel (300–600 mg loading dose, 75 mg daily dose) is recommended for patients who cannot receive ticagrelor or prasugrel or who require oral anticoagulation.		
P2Y ₁₂ inhibitor administration for a shorter duration of 3–6 months after DES implantation may be considered in patients deemed at high bleeding risk.	IIIb	
Intravenous antiplatelet therapy		
GP IIb/IIIa inhibitors during PCI should be considered for bailout situations or thrombotic complications.	IIIb	C
It is not recommended to administer GP IIb/IIIa inhibitors in patients in whom coronary anatomy is not known.	IIIb	A
Long-term P2Y₁₂ inhibition		
P2Y ₁₂ inhibitor administration (in addition to aspirin beyond 1 year) may be considered after careful assessment of the ischaemic and bleeding risks of the patient.	IIIb	A
General recommendations		
A proton pump inhibitor in combination with DAPT is recommended in patients at higher than average risk of gastrointestinal bleeds (i.e. history of gastrointestinal ulcer/haemorrhage, anticoagulant therapy, chronic NSAID/corticosteroid use or two or more of the following: age ≥65 years, dyspepsia, gastro-oesophageal reflux disease, Helicobacter pylori infection, chronic alcohol use).		

P2Y₁₂ Inhibitors

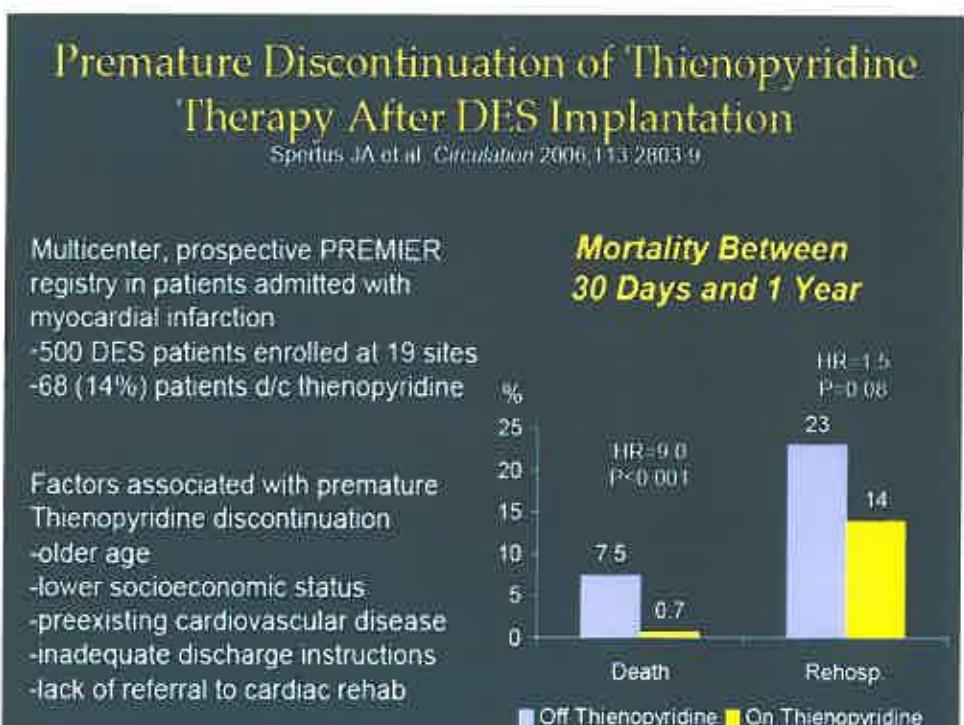
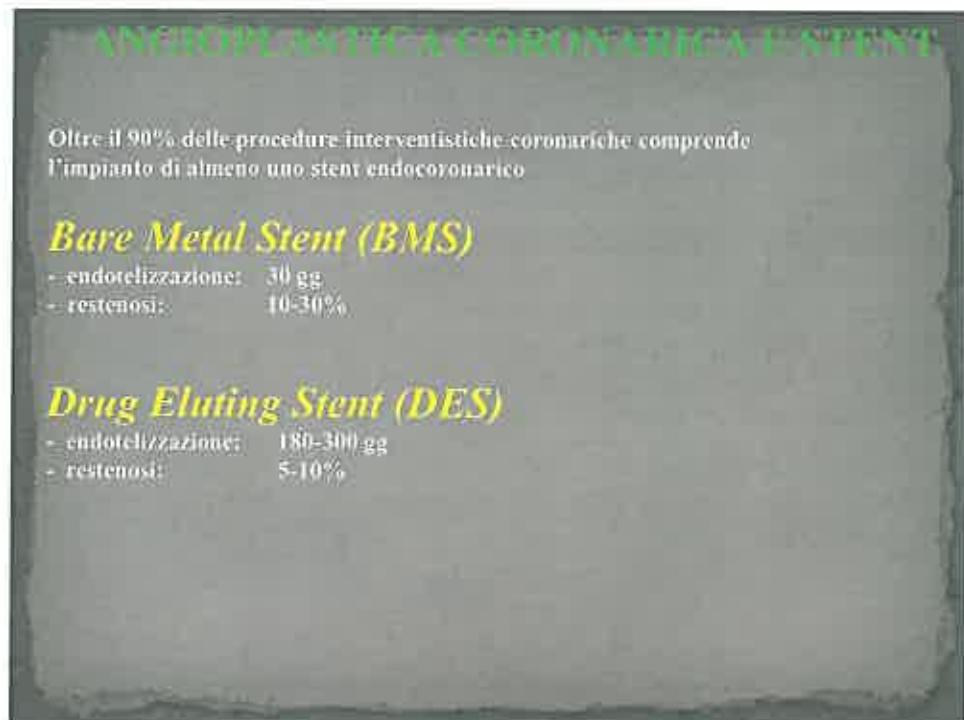
	Clopidogrel	Prasugrel	Ticagrelor
Class	Thienopyridine	Thienopyridine	Triazolopyrimidine
Reversibility	Irreversible	Irreversible	Reversible
Activation	Prodrug, limited by metabolism	Prodrug, not limited by metabolism	Active drug
Onset of effect ^a	2–4 h	30 min	30 min
Withdrawal before major surgery	5 days	7 days	5 days

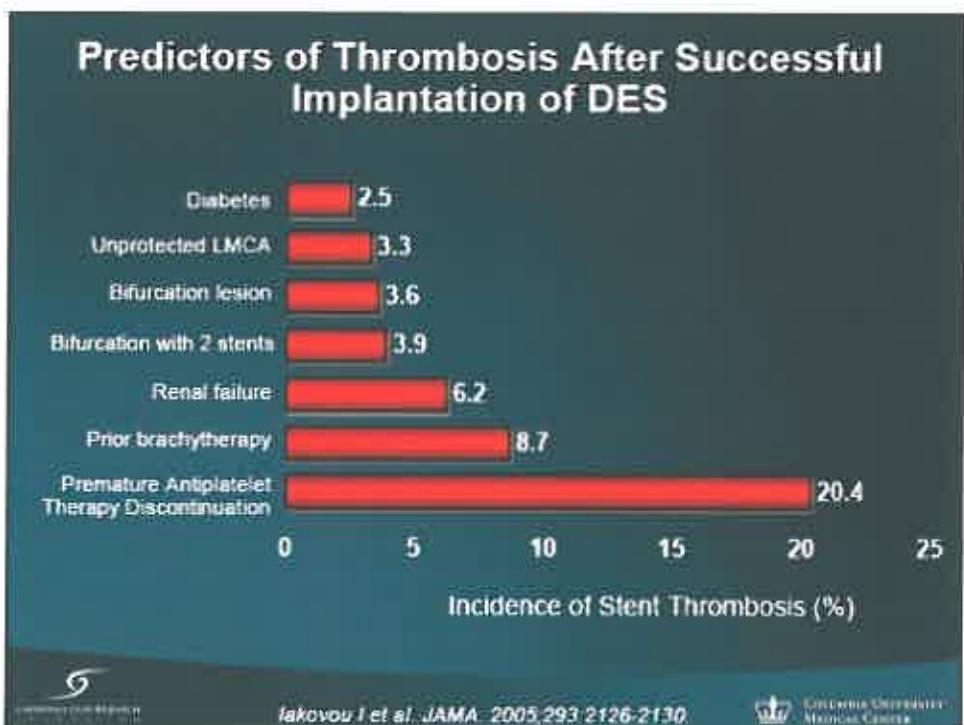
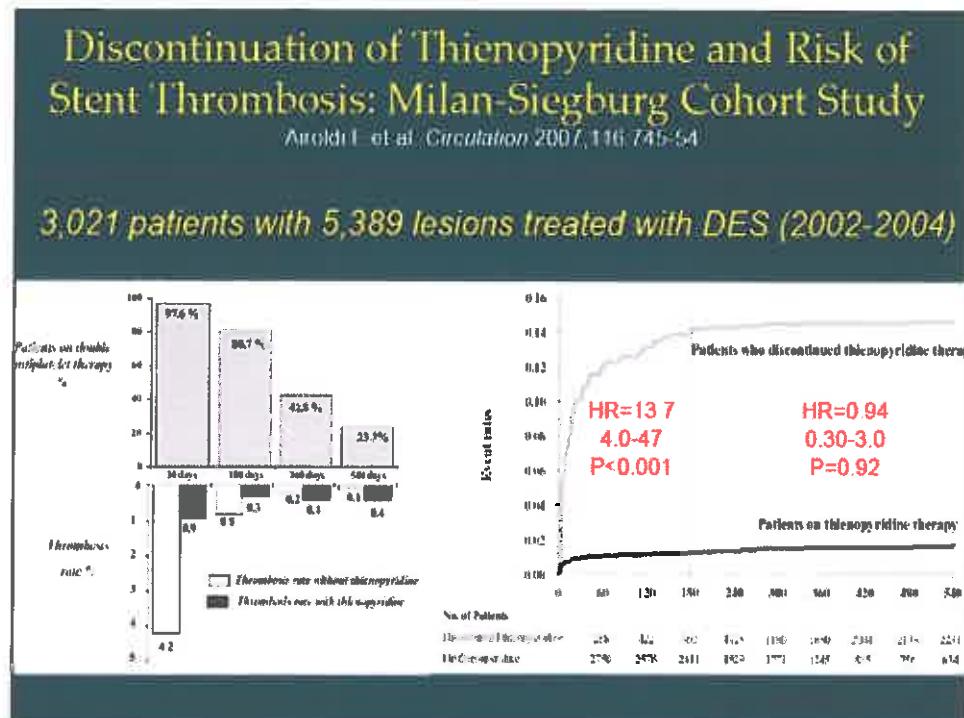
OASIS 5: An International, Multicenter, Randomized, Double-Blind, Double-Dummy Trial in 41 Countries



Risk criteria mandating invasive strategy in NSTE-ACS	
Very-high-risk criteria	<i>Immediate invasive strategy (<2 h)</i>
• Haemodynamic instability or cardiogenic shock	
• Recurrent or ongoing chest pain refractory to medical treatment	
• Life-threatening arrhythmias or cardiac arrest	
• Mechanical complications of MI	
• Acute heart failure	
• Recurrent dynamic ST-T wave changes, particularly with intermittent ST-elevation	
High-risk criteria	<i>Early invasive strategy (<24 h)</i>
• Rise or fall in cardiac troponin compatible with MI	
• Dynamic ST- or T-wave changes (symptomatic or silent)	
• GRACE score > 140	
Intermediate-risk criteria	<i>Invasive strategy (< 72 h)</i>
• Diabetes mellitus	
• Renal insufficiency (eGFR < 60 mL/min/1.73 m ²)	
• LVEF < 40% or congestive heart failure	
• Early post-infarction angina	
• Prior PCI	
• Prior CABG	
• GRACE Risk score > 109 and < 140	
Low-risk criteria	<i>Selective invasive strategy</i>
• Any characteristics not mentioned above	

GESTIONE DELLA TERAPIA ANTIAGGREGANTE	
<u>Condizioni ad alto rischio di trombosi</u>	
• nel primo 12 mesi dopo PCI/N + non medicato	
• prima di 6 settimane da PTCA + stent non medicato	
• prima di 2 settimane da PTCA senza stent	
• infarto miocardico acuto (< 7 gg) o recente (< 30 gg)	
• sindrome coronarica acuta	
• anatomia coronarica ad alto rischio	





Mortality rate associated with in-DES late thrombosis

45% Milan register
44.4% ERACI III register

Lakowicz JAMA

RISCHIO EMORRAGICO CHIRURGIA ORALE

	RISCHIO EMORRAGICO
Chirurgia Dentale (uso di tecniche di emostasi adeguate e acido tranexamico)	Estrazioni singole o multiple Protesi Endodonzia Igiene dentale Terapia periodontale
Chirurgia orale estesa	Basso Intermedio

Lakowicz JAMA 2014; 308(11):22
Juvek J et al. J Oral Maxillofac Surg 2013; 71(10):1357-1366

GESTIONE ANTIAGGREGANTI IN CHIRURGIA ORALE

La maggior parte delle pubblicazioni che ha considerato il rischio di sospendere vs continuare la terapia antiaggregante nelle procedure dentali ha concluso che la maggior parte dei pazienti può essere sottoposto a tali procedure senza interrompere la terapia antiaggregante, assicurandosi che metodi di tamponamento locale vengano usati per controllare eventuali sanguinamenti.

Dati su aspirina:

**Non dati su clopidogrel o
doppia antiaggregazione**

Per i dati sui clopidogrel e sulla doppia antiaggregazione si veda la presentazione di Dr. S. Cicali

Rischio emorragico basso (+++ frequente)

- Rischio trombotico basso : che fare?
- ASA: PROSEGUIRE
- INIBITORI RECETTORE P₂Y₁₂:
sospendere 5 giorni prima, riprendere
entro 24-72 ore con dose di carico

Rischio emorragico intermedio : implantologia e chirurgia orale

Rischio trombotico basso: che fare?

ASA :proseguire

Inibitori recettori P2Y₁₂: sospendere
5 giorni prima, riprendere entro 24-
72 ore con dose carico

Rischio trombotico intermedio: che
fare?

• **Chirurgia elettiva :** differire

• **Chirurgia non differibile;**

ASA: proseguire

Inibitori recettori P2Y₁₂: sospendere 5
giorni prima, riprendere entro 24-72 ore
con dose carico

Rischio trombotico basso: quando?

- > 6 mesi dopo impianto di BMS
- > 12 mesi dopo impianto di DES

Rischio trombotico intermedio: quando?

- > 1 mese < 6 mesi dopo PCI + BMS
- ≥ 6 mesi < 12 mesi dopo PCI + DES
- > 12 mesi dopo DES a rischio elevato (stent lunghi, multipli, in overlapping, piccoli vasi, biforcazioni, tronco comune, last remaining vessel)

Rischio trombotico alto

- < 1 mese dopo BMS
- < 6 mesi dopo DES
- < 12 mesi dopo DES a rischio elevato

- F
- Clinical benefit of a drug or intervention
 - reduces mortality



- Bleeding complications
 - increases mortality

- Careful patient selection and evaluation:
 - age, gender, past history of bleeding, low weight, renal insufficiency

Diapositiva 72

U4 Pertanto il trattamento del paziente con UA/NSTEMI passa attraverso una equilibrio molto delicato che deve prevedere una attenta selezione del paziente e delle sue caratteristiche cliniche al fine di valutare le caratteristiche potenzialmente positive dei farmaci e delle procedure invasive, tenendo conto di quella temibile variabile come il bleeding risk che può contribuire ad aumentare la mortalità

User; 20/09/2011

TAKE HOME MESSAGE

- **NSTEACS >> fondamentale la stratificazione del rischio**, riconoscendo l'opportunità di un intervento precoce solamente nei pz a > rischio. L'entità del vantaggio derivante dall'aggressività angiografica immediata è minore, anche perché i pz con NSTEACS sono più complessi e l'entità della coronaropatia può essere maggiore che nello STEMI, tant'è vero che alla coronarografia segue un'angioplastica solo in 50-70% casi. Anche la scelta della procedura di rivascolarizzazione (PCI vs CABG) pone maggiori dubbi rispetto allo STEMI, e le linee guida raccomandano una valutazione della scelta da parte dell'Heart Team (cardiologo clinico, cardiologo interventista e cardiochirurgo)
- **STEMI >> riparazione sistematica**, pressoché non basata sulla stratificazione del rischio, che viene raccomandata successivamente, più che altro al fine di valutare l'ischemia residua a carico dei territori non infarctuali e ai fini di prevenzione secondaria



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