

Le Sindromi Coronariche Acute

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Sindromi Coronariche Acute

Classificazione

SINDROME CORONARICA ACUTA

Senza soprasl. ST

Con soprasl. ST

IMA senza ST soprasl.
(NSTEMI)

Angina instabile

Infarto miocardico

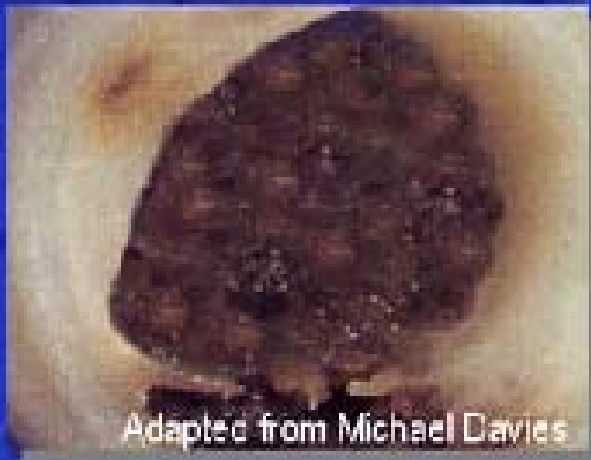
Non-Q

Con onda Q

Sindromi Coronariche Acute

Fisiopatologia

**SCA con persistente
ST sopraslivellato**



CK- MB or Troponin ↑

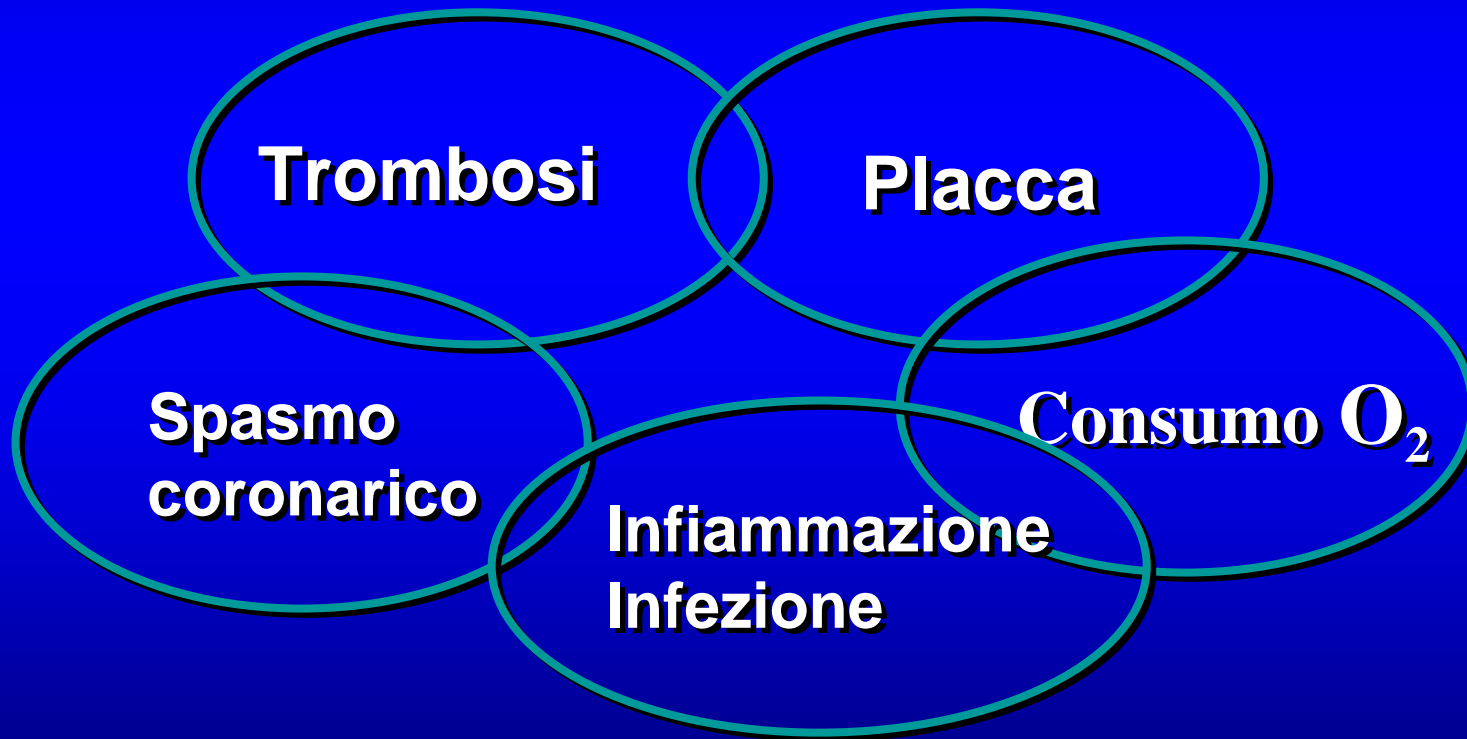
**SCA senza persistente
ST sopraslivellato**



Troponin elevated or not

SINDROMI CORONARICHE ACUTE:

Meccanismi eziopatogenetici comuni per l'angina instabile e l'infarto miocardico acuto

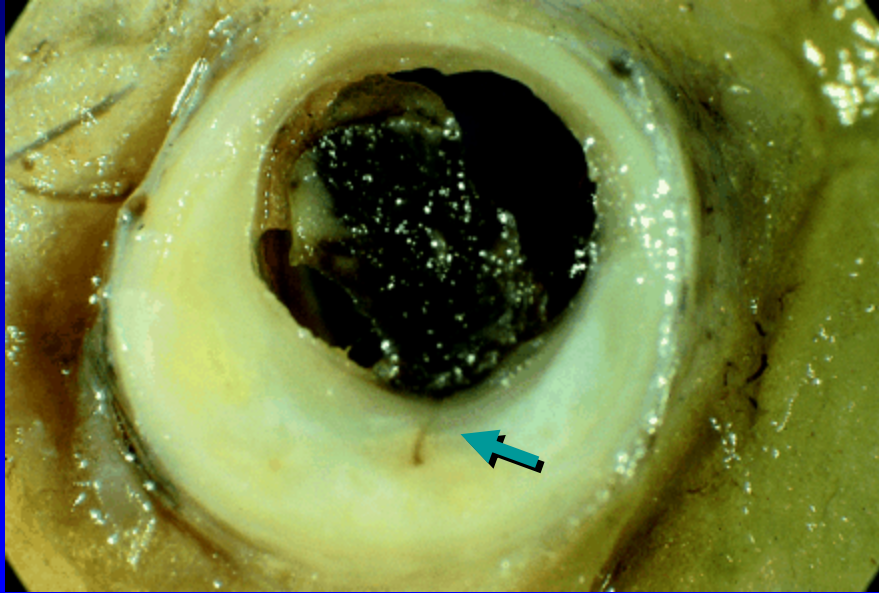


Eventi coronarici acuti

- sono innescati dalla trombosi della placca colpevole
- responsabili della trombosi sono 2 meccanismi:
 - erosione: ampia denudazione dell'endotelio che ricopre la placca
 - rottura: della capsula fibrosa che racchiude il nucleo lipidico

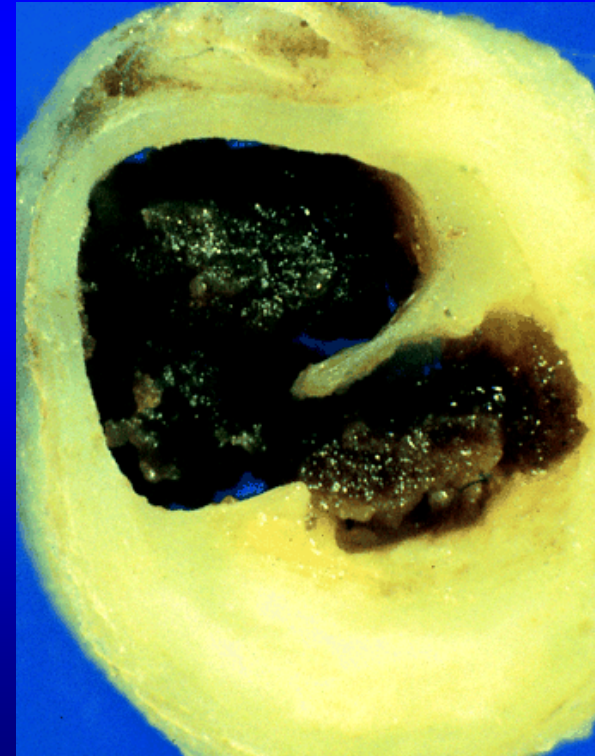
Eventi coronarici acuti

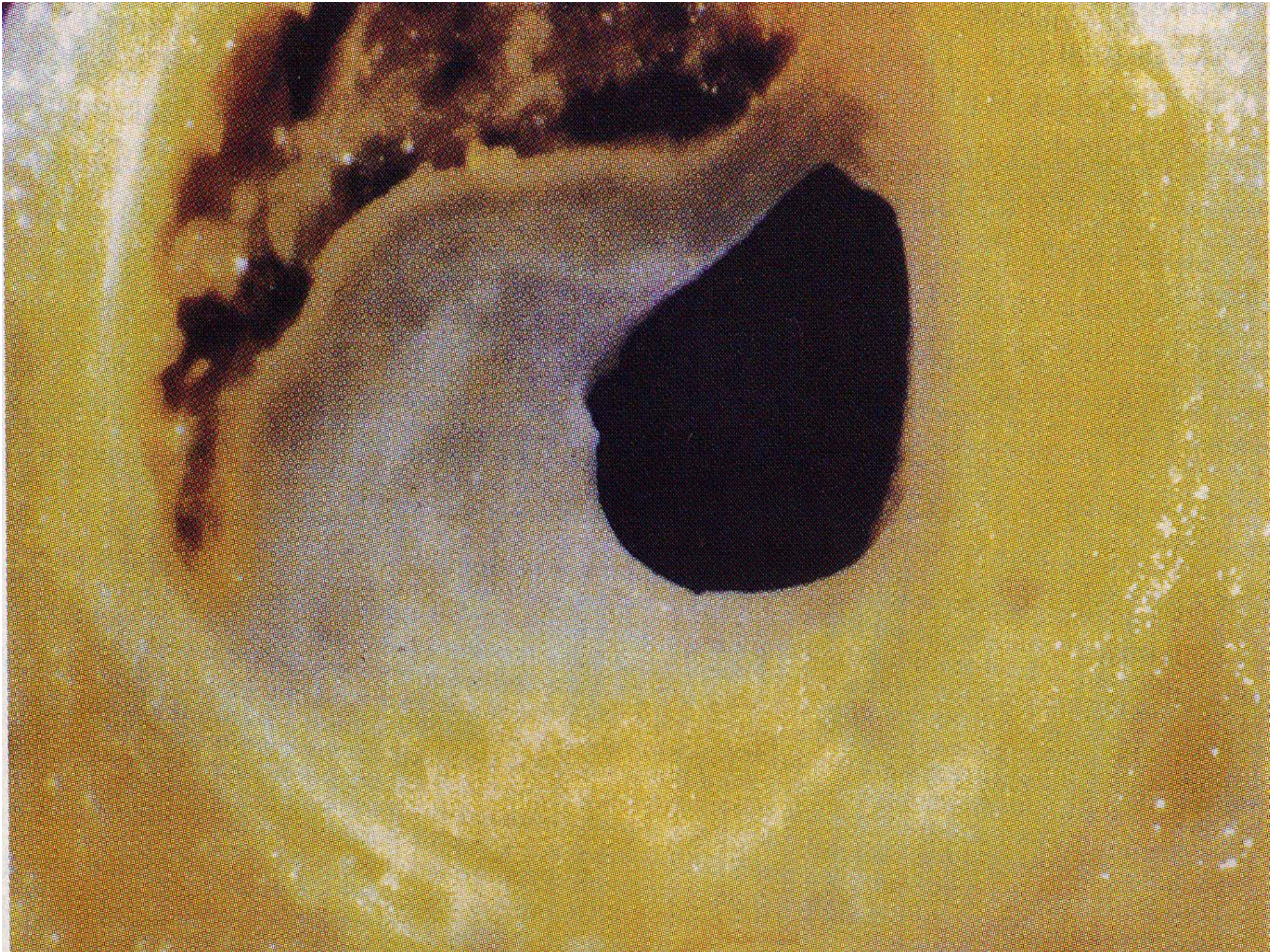
- **Rottura**: più frequente in siti senza stenosi o con stenosi lieve
- **Erosione**: più frequente se pre-esistente stenosi e in arterie più piccole

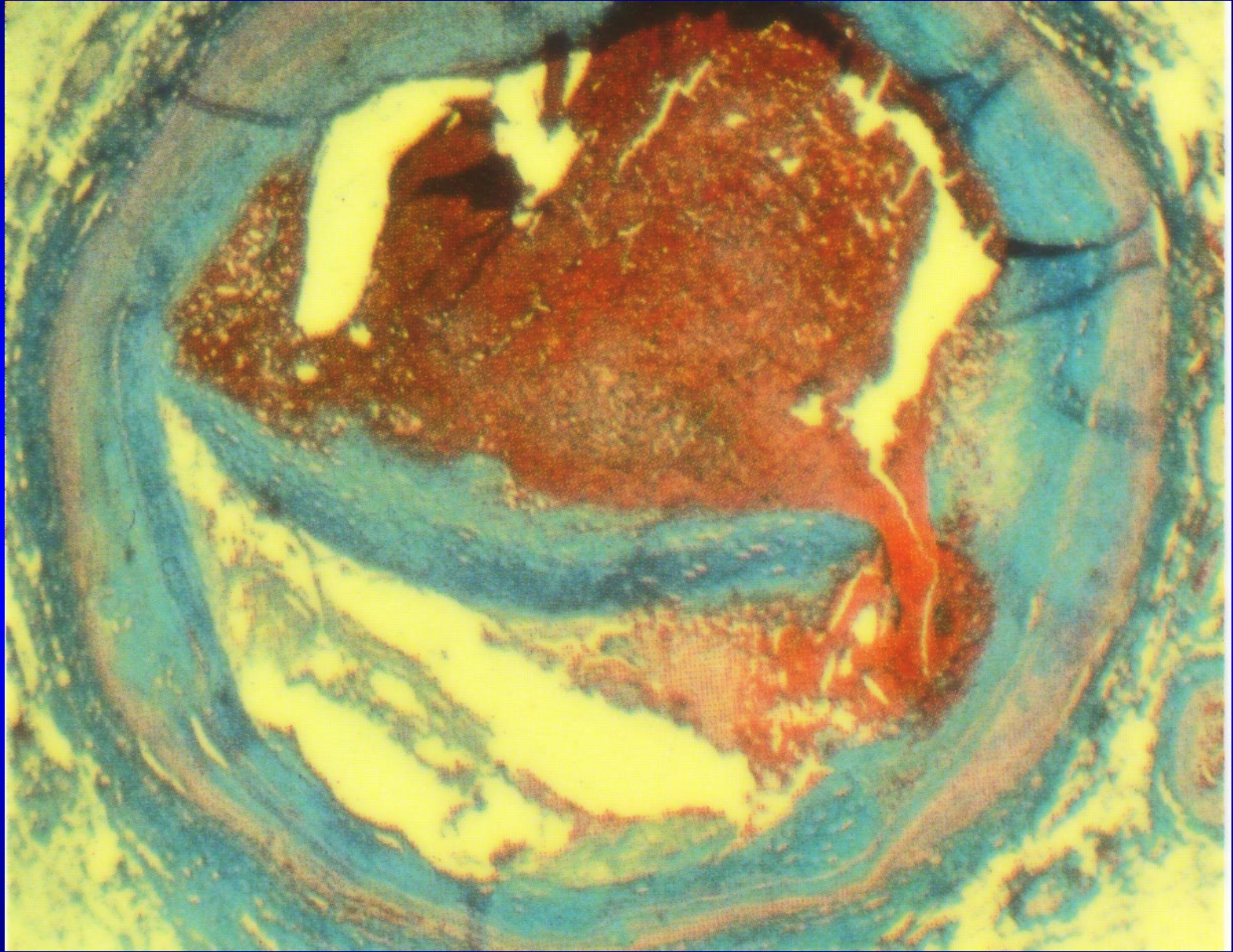


Trombosi causata da erosione (←)

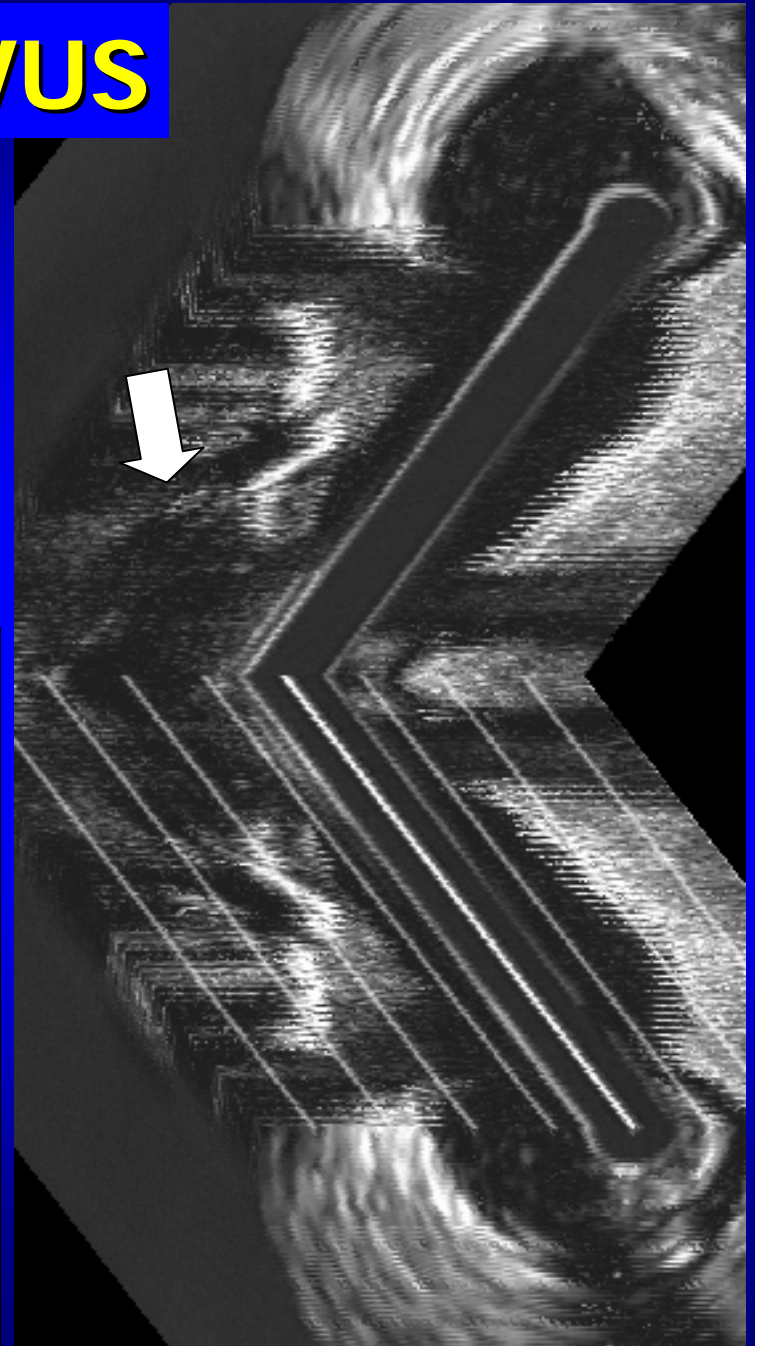
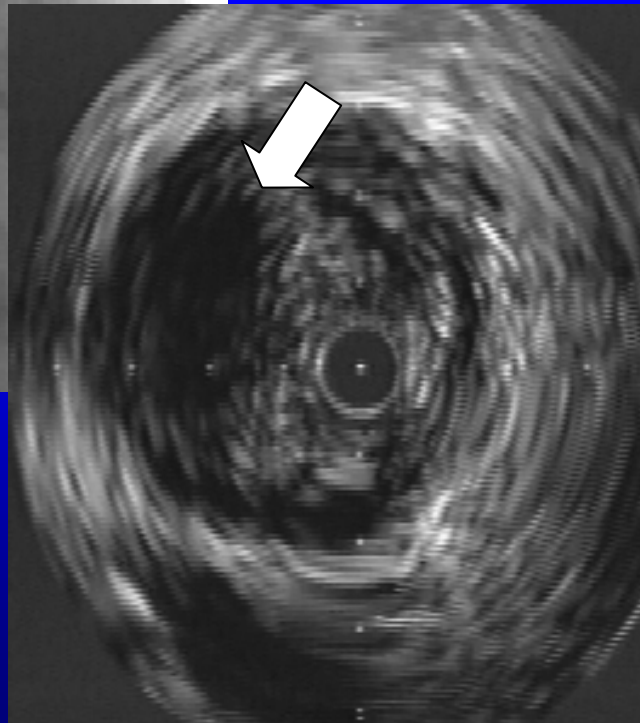
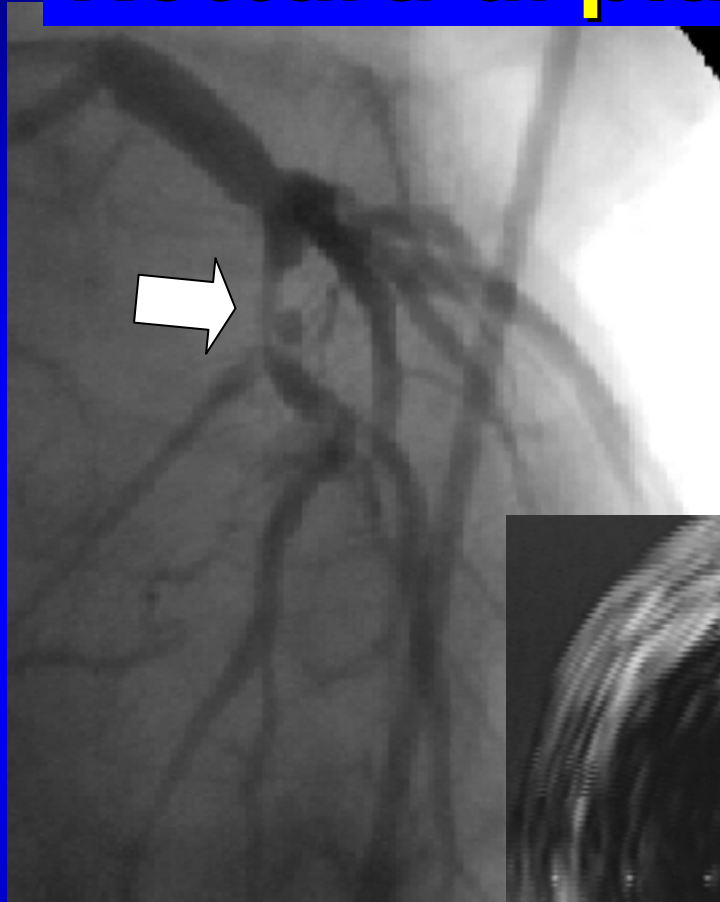
Trombosi causata da rottura



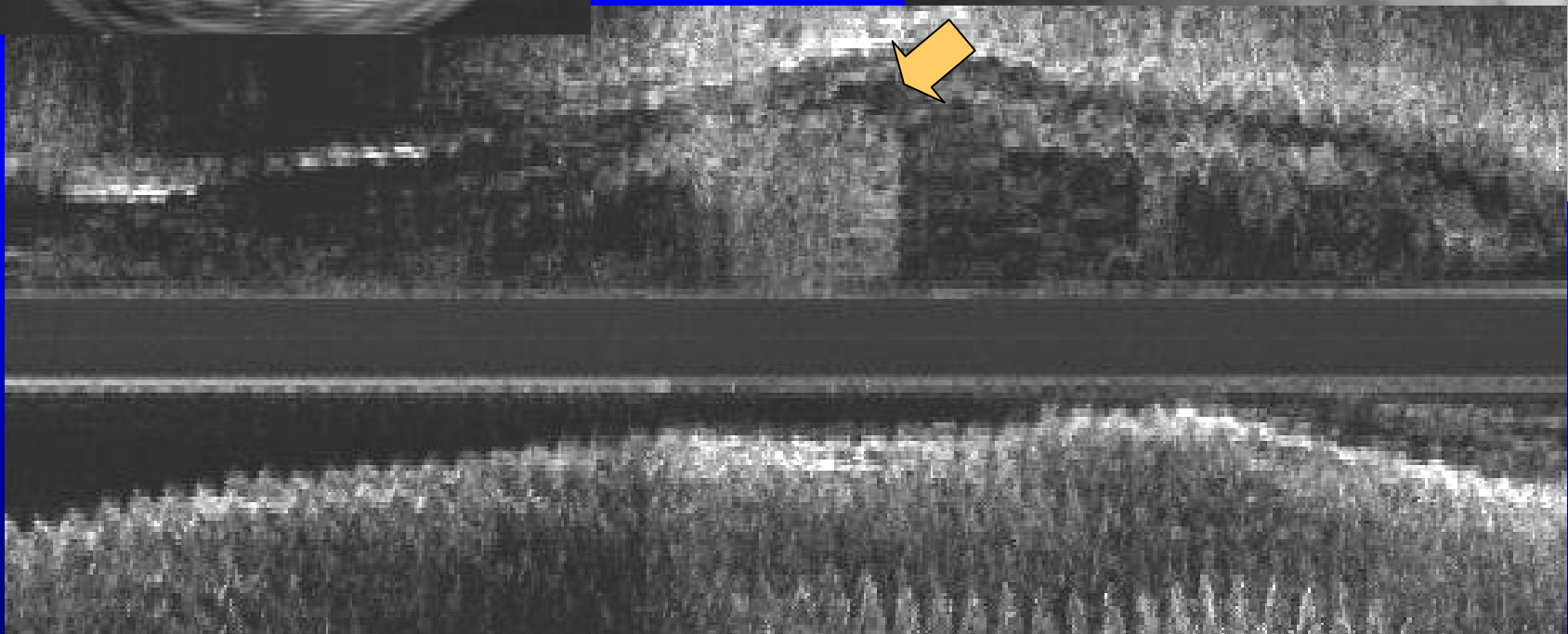
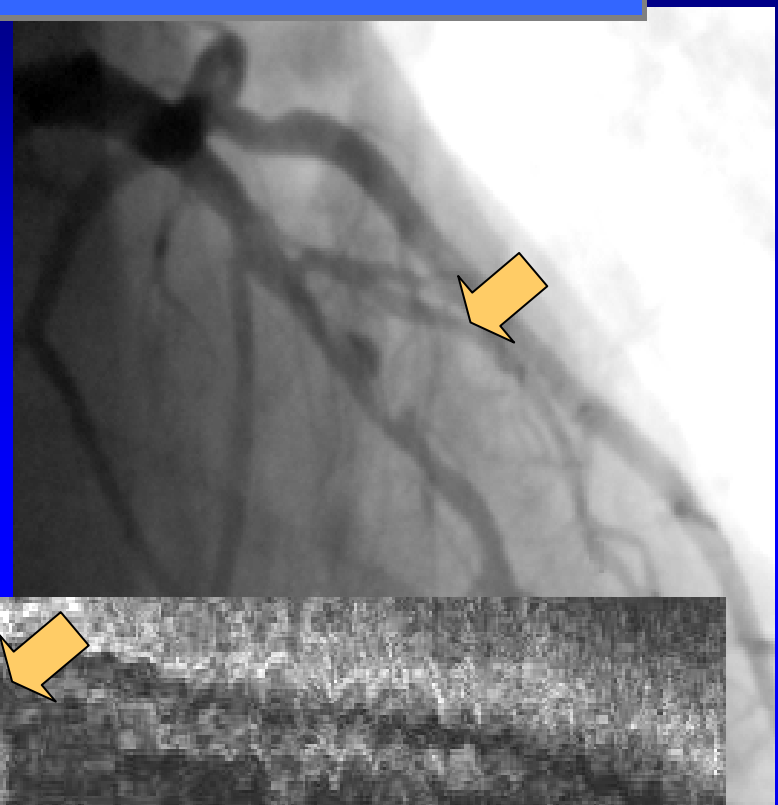
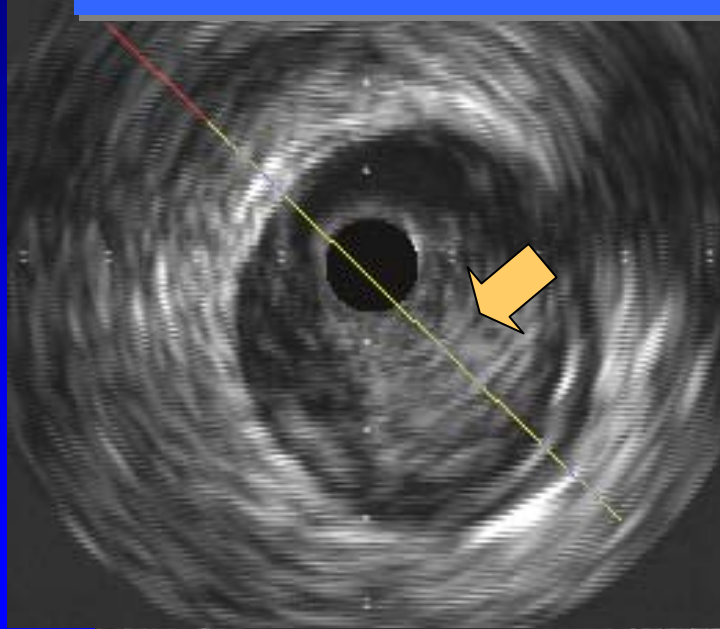




Rottura di placca all'IVUS

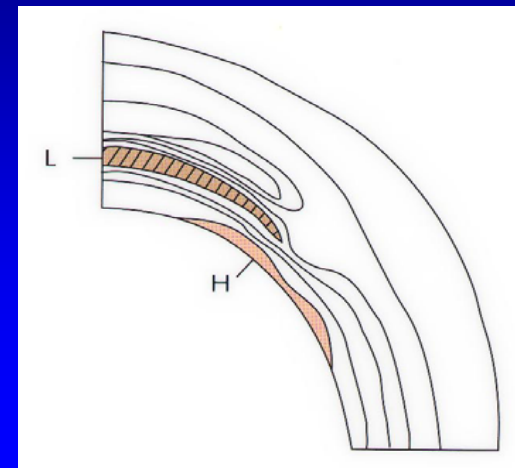
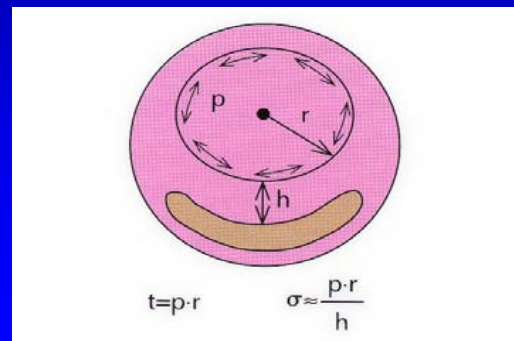
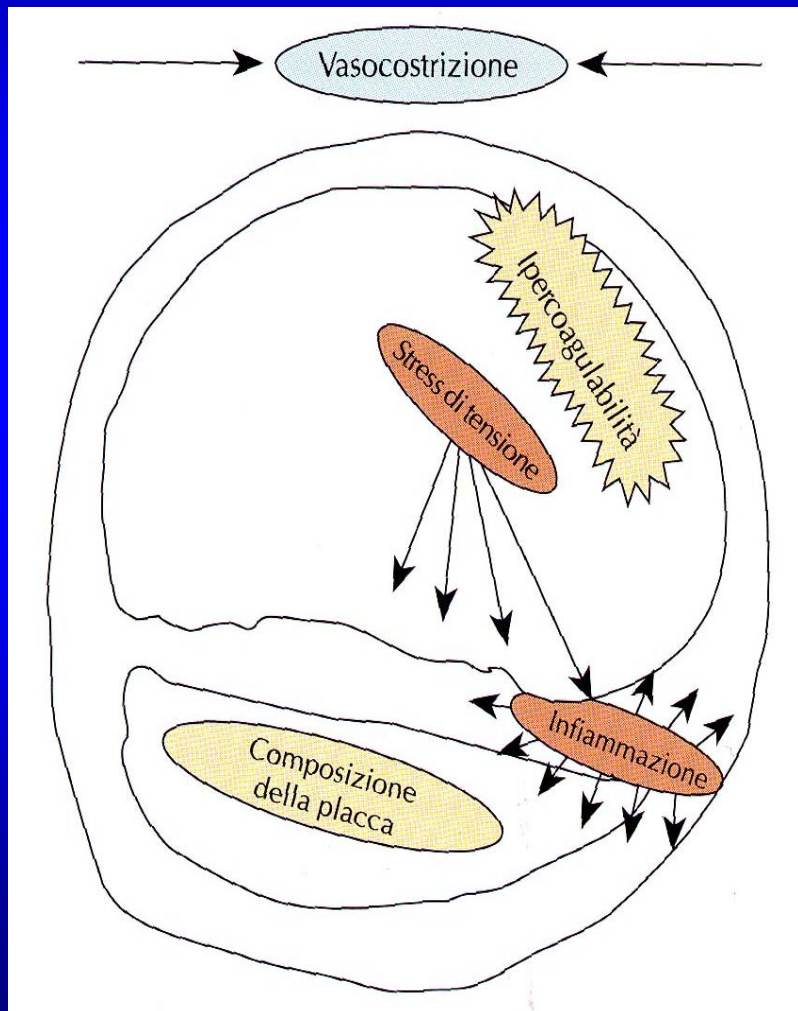


Trombo intracoronarico all' IVUS



LA PLACCA

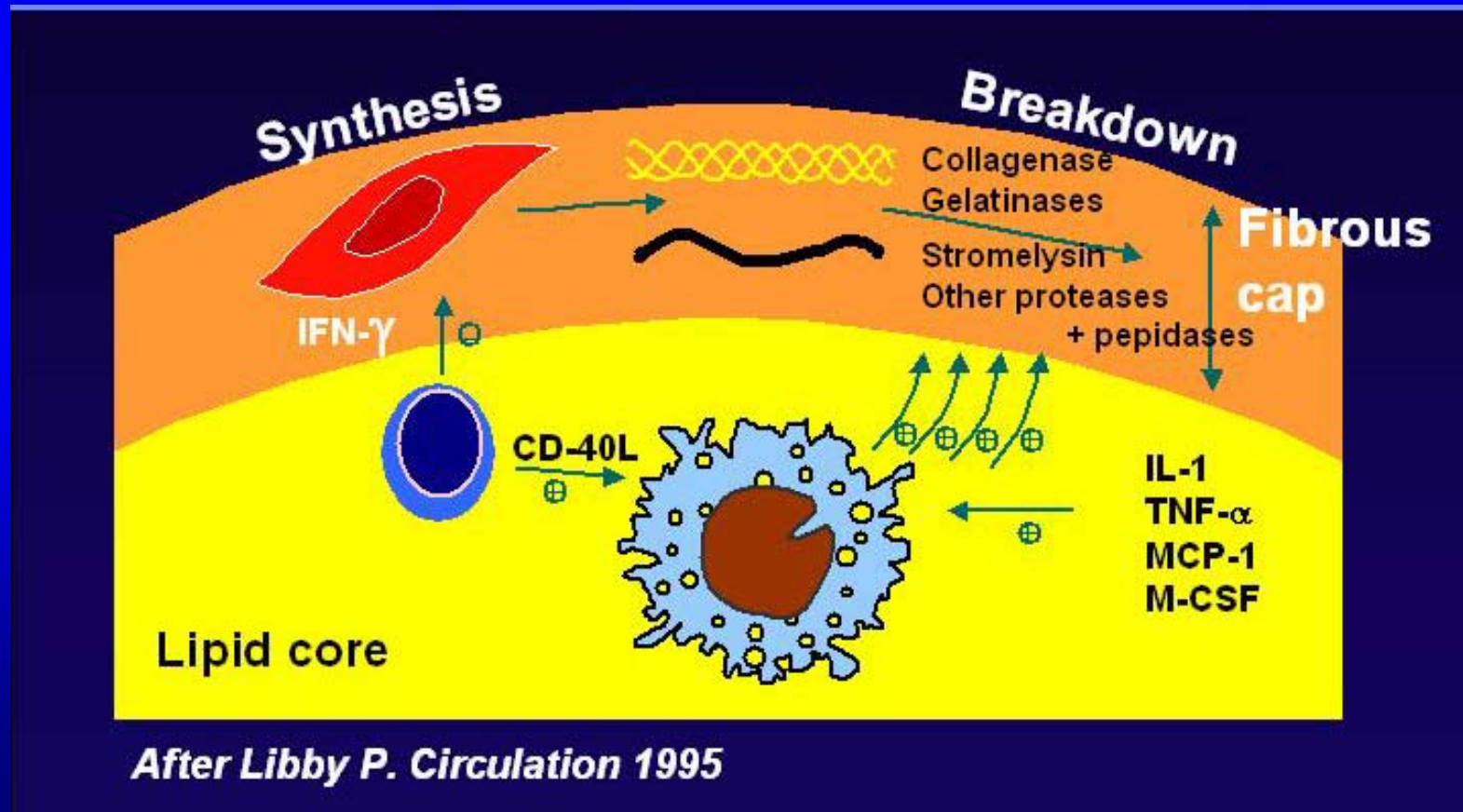
Patogenesi della rottura



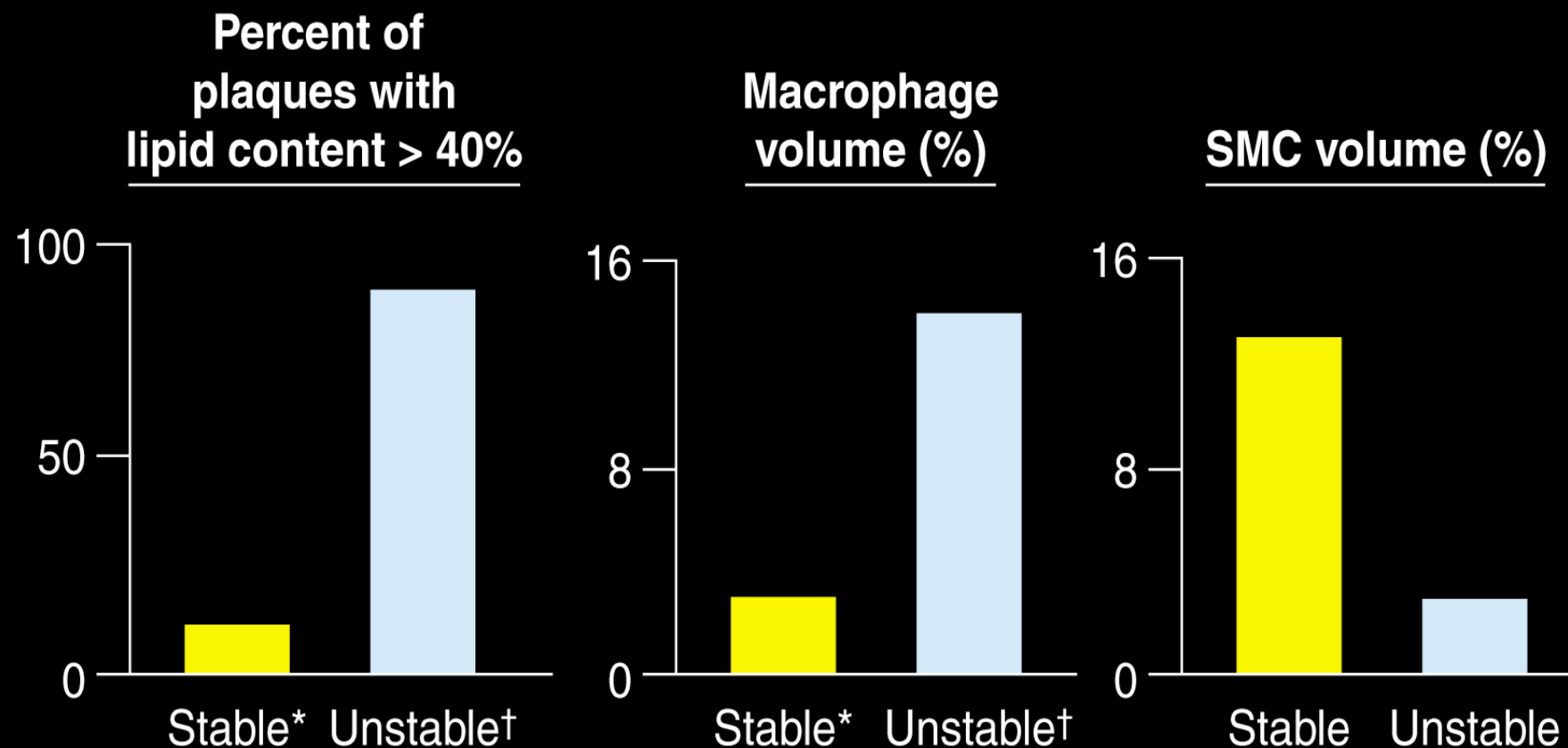
LA PLACCA

Patogenesi della rottura

Fenomeno autodistruttivo



Patients dying from stable vs unstable coronary syndromes: Atheromas have varying composition

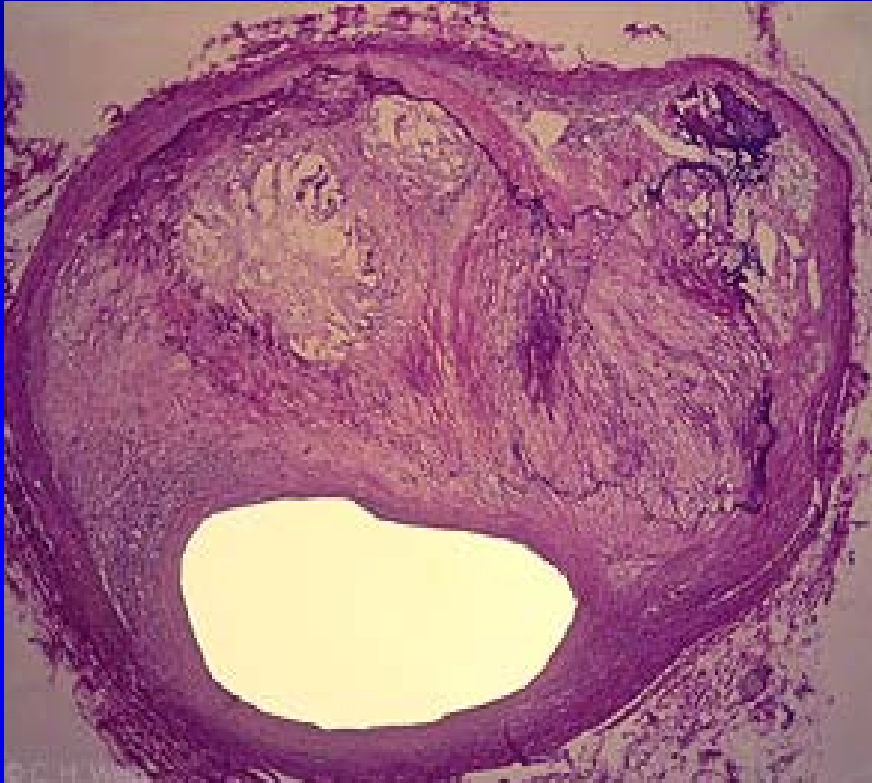


*Stable plaque = smooth surface

†Unstable plaque = fissured

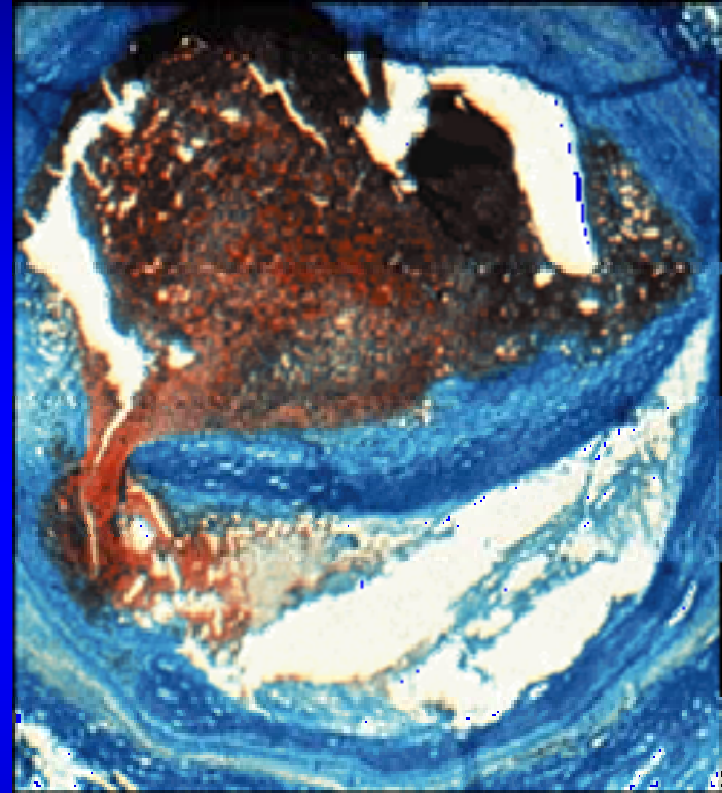
Davies MJ, et al. *Br Heart J.* 1993;69:377-381.

Placca aterosclerotica stabile



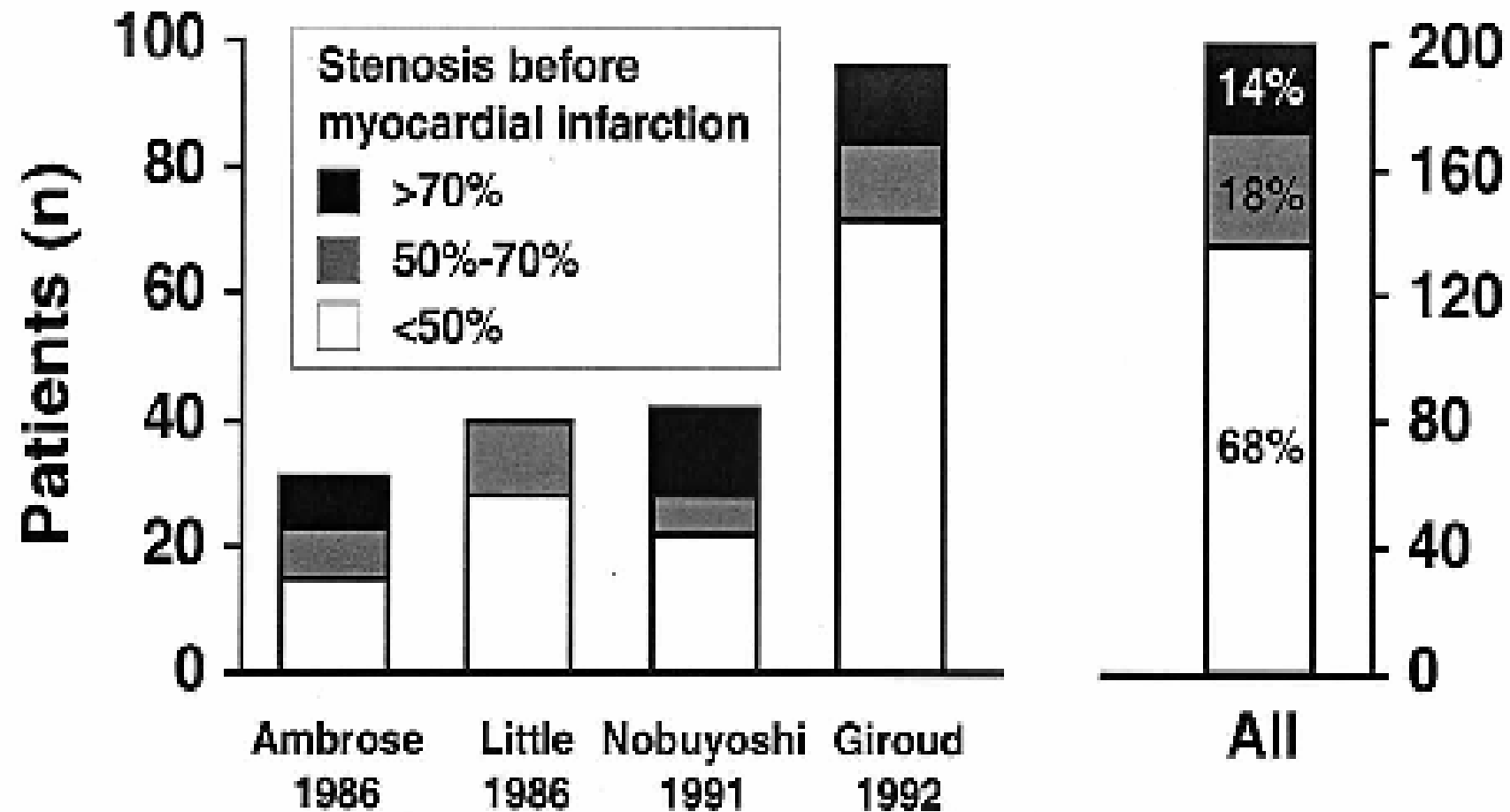
- **Paziente asintomatico**
- **Angina stabile**

Placca complicata



Sindrome coronarica acuta

Severity of coronary stenosis detected by angiography before myocardial infarction



Perché stenosi moderate causano sindromi coronariche acute?

Lo stato funzionale dell'ateroma, non soltanto le sue dimensioni o il grado di restringimento del lume, determina la propensione allo sviluppo di sindromi coronariche acute.

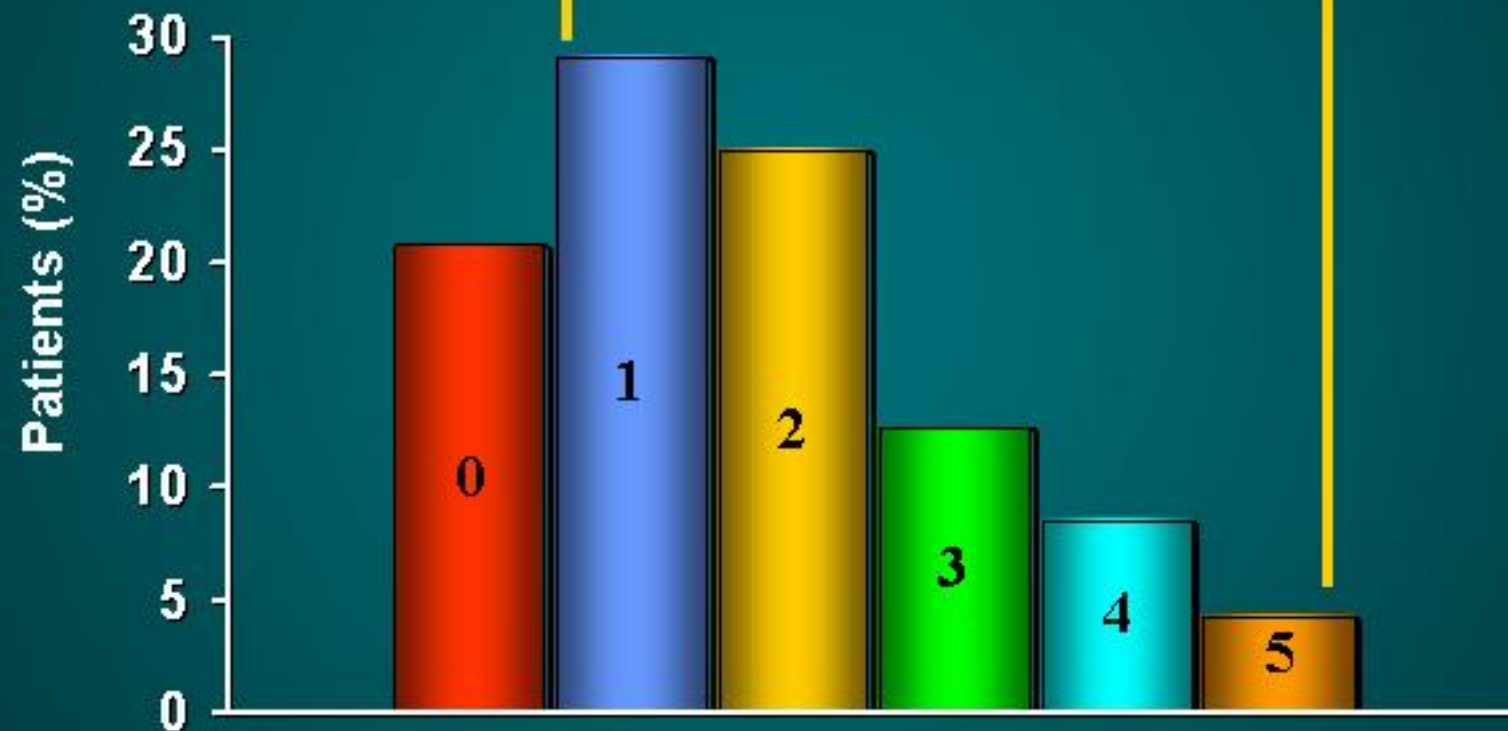
Placca Vulnerabile

Un Concetto Fuori Moda?

- ✓ Arteria vulnerabile
- ✓ Letto vascolare vulnerabile
- ✓ Paziente vulnerabile

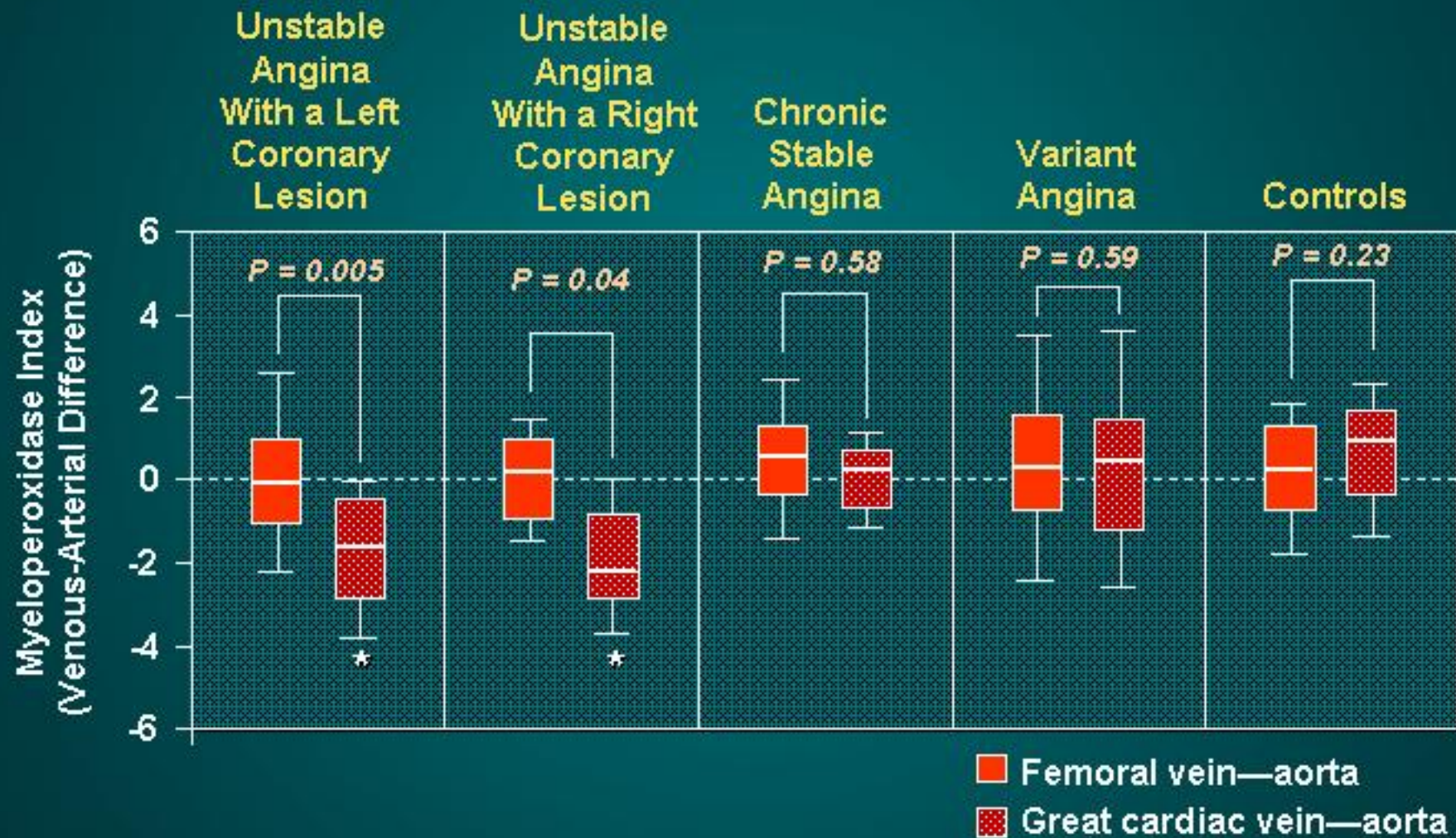
Vulnerable Patients

80% of ACS patients have > 1 ruptured plaque



ruptured plaques in addition to culprit lesion

Widespread Coronary Inflammation in Unstable Angina



Sindromi Coronariche Acute

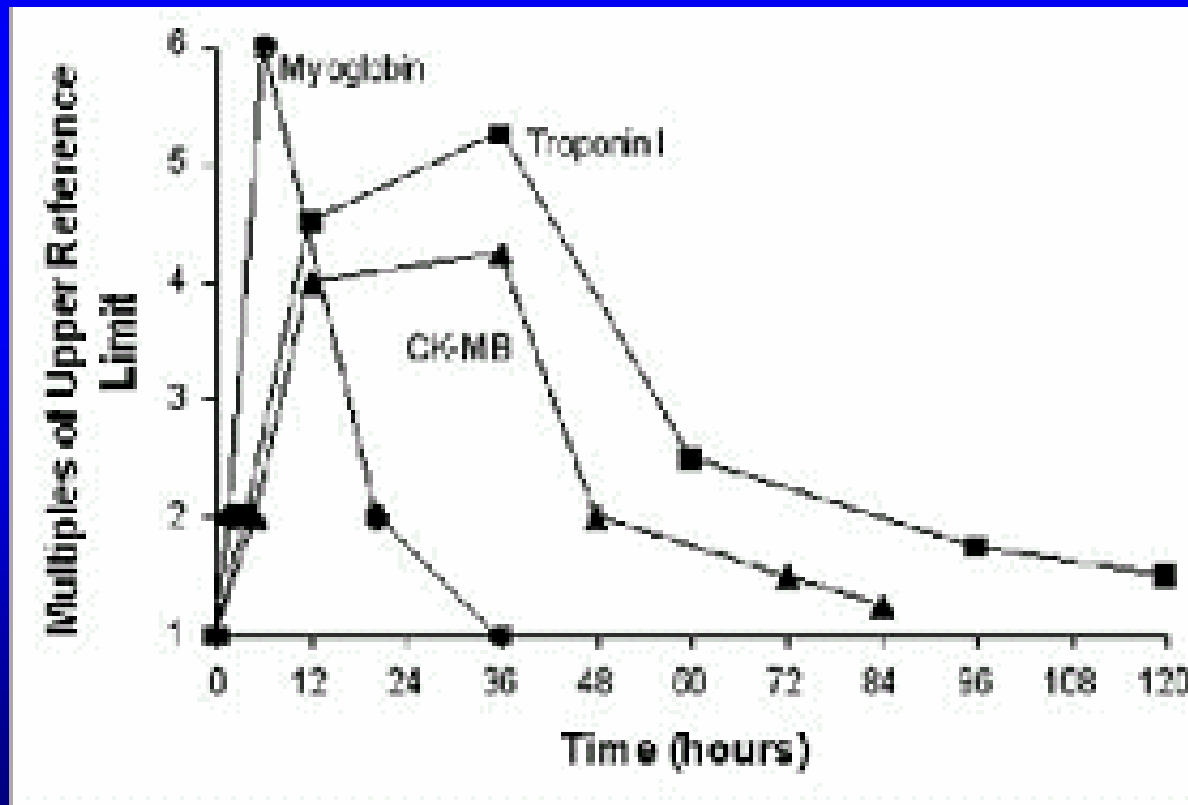
Diagnosi

SCA: quadri clinici

- Dolore toracico “tipico” o “atipico”
- “Equivalenti anginosi”:
 - dispnea-EPA-shock (classificazione di Killip)
 - lipotimia/sincope
 - astenia
 - dolore epigastrico (IMA inferiore)
 - sudorazione algida, nausea, vomito
- SCA asintomatica o paucisintomatica (diabetici, anziani)

SCA: diagnosi di laboratorio

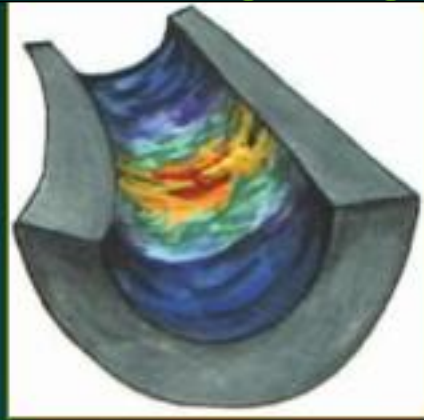
- Enzimi di citonecrosi miocardica:
 - mioglobina, troponine, CK-Mb (SGOT, LDH)



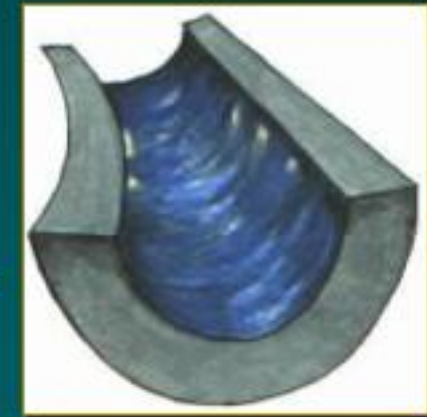
SCA: diagnosi strumentale

- ECG:
 - alteraz. tratto ST e/o onda T;
 - onda Q
- Ecocardiogramma:
 - diagnosi precoce (ECG normale)
 - diagnosi di sede ed estensione
 - valutazione della funzione VS
 - diagnosi delle complicanze meccaniche
- Coronarografia:
 - sede ed estensione della malattia coronarica

Imaging di morfologia vs "attività"



Termografia,
Spettroscopia,
RM con mdc



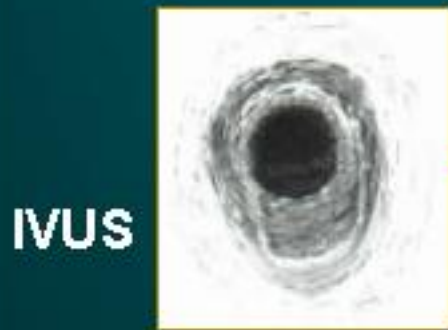
Attività

Placca
"attiva"
e
infiammata



Placca
"inattiva"
e non
infiammata

Morfologia

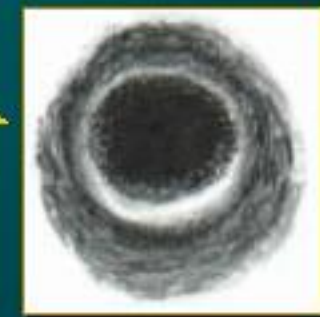


IVUS

OCT



RM
senza
mdc



SCA: principi di terapia

- Terapia medica:
 - Antitrombotica:
 - ASA, fibrinolitici, eparina (non frazionata/ a basso PM), inibitori GP IIb/IIIa, clopidogrel
 - Antiischemica:
 - nitroderivati, beta-bloccanti, Ca-antagonisti
 - Prevenzione secondaria:
 - ACE-inibitori, statine
- Terapia interventistica:
 - Angioplastica coronarica
- Terapia chirurgica:
 - By-pass aorto-coronarico

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SCA senza ST sopraslivellato

Epidemiologia

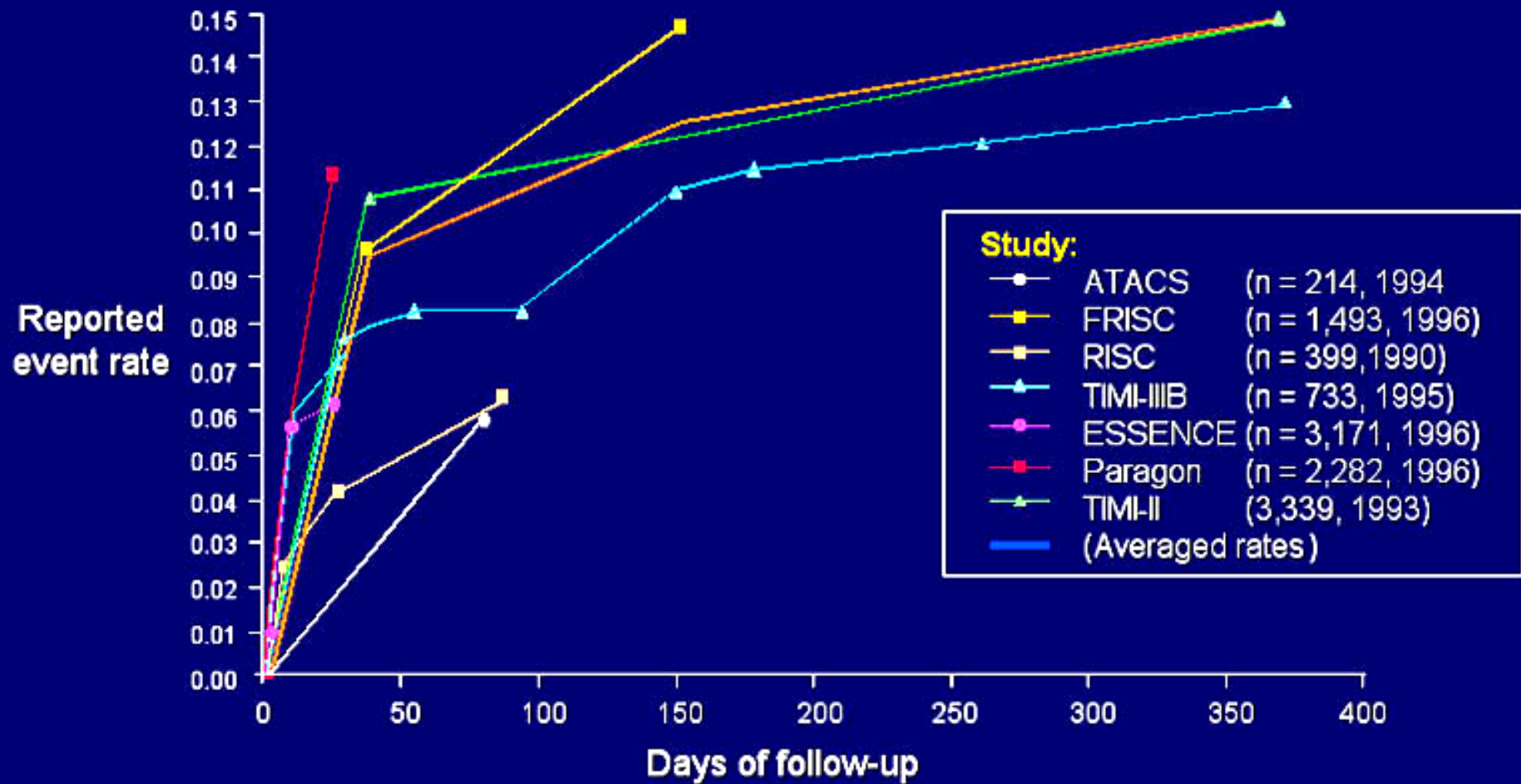
SCA senza persistente ST sopraslivellato

- Uno dei maggiori problemi di salute pubblica
- 1.43 milioni di ospedalizzazioni/anno negli Stati Uniti
- Ancora oggi la prognosi rimane sfavorevole

**Rischio di Morte o Infarto Miocardico:
8-16% ad 1 mese di follow-up***

*da recenti trial clinici con antiplastrinici, antitrombotici o di interventistica

Studi sull'Angina Instabile: percentuale di morte e di infarto



**ACC/AHA Guidelines Update
Ottobre 2002
Sindromi Coronariche Acute NSTEMI
Nuove direttive “evidence-based”**

- **Importanza della stratificazione del rischio**
- **Beneficio del Clopidogrel a breve e a lungo termine**
- **Meglio le LMWH (?)**
- **Inibitori IIb/IIIa e strategia invasiva per pz ad alto rischio**

SCA senza ST sopraslivellato

Stratificazione del rischio

SCA senza persistente ST sopraslivellato

Valutazione del rischio

- Angor/ischemia ricorrente
- Depressione del tratto ST
- Alterazioni dinamiche ST
- Alterazioni troponina
- Diabete
- Instabilità emodinamica
- Aritmie maggiori (TV, FV)

Rischio acuto

- Markers clinici:
Età
Storia di IM, BPAC
Diabete
- Markers biologici:
CRP
- Markers angiografici:
EF
Estensione della malattia

Rischio a lungo termine

Stratificazione del rischio

TIMI Risk Score nelle SCA NSTE

7 Independent Predictors

- Age > 65
- Multiple risk factors
- Known CAD (>50% stenosis)
- Aspirin therapy
- Recurrent angina < 24 hours
- ST-segment deviation
- Elevated CK-MB or troponin

SCA senza ST sopraslivellato

Terapia:
beneficio del clopidogrel

The New York Times

TUESDAY, MARCH 20, 2001

Drug Hailed As a Heart And Stroke Protector

In the syndrome, not enough blood can flow to nourish the heart because the coronary arteries are blocked by fatty deposits from the underlying condition, atherosclerosis. A heart attack can result unless a blood-thinning drug, aspirin, is given. In the syndrome, people experience at least one episode of acute coronary syndrome a year, Dr. Yusuf said.

Dr. Yusuf's study, the largest and longest of acute coronary syndrome, involved more than 12,500 patients in 428 hospitals in 25 countries. Half the participants received aspirin, which is given to most patients with the syndrome. In those who received aspirin, death occurred at a rate of 10.5 percent, compared with 8.5 percent in those receiving clopidogrel, a reduction of about 20 percent — a statistically significant reduction. Heart

Clopidogrel, like aspirin, helps keep components of the blood known as platelets from sticking together. But the two drugs act by different

The drug is clopidogrel ...

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Dr. Christopher P. Cannon, a cardiologist at Brigham and Women's Hospital in Boston, said in an interview: "This is the best news since aspirin. This is really a super aspirin that lives up to its name." But like aspirin, clopidogrel can cause life-threatening bleeding into the brain to cause a stroke, or elsewhere.

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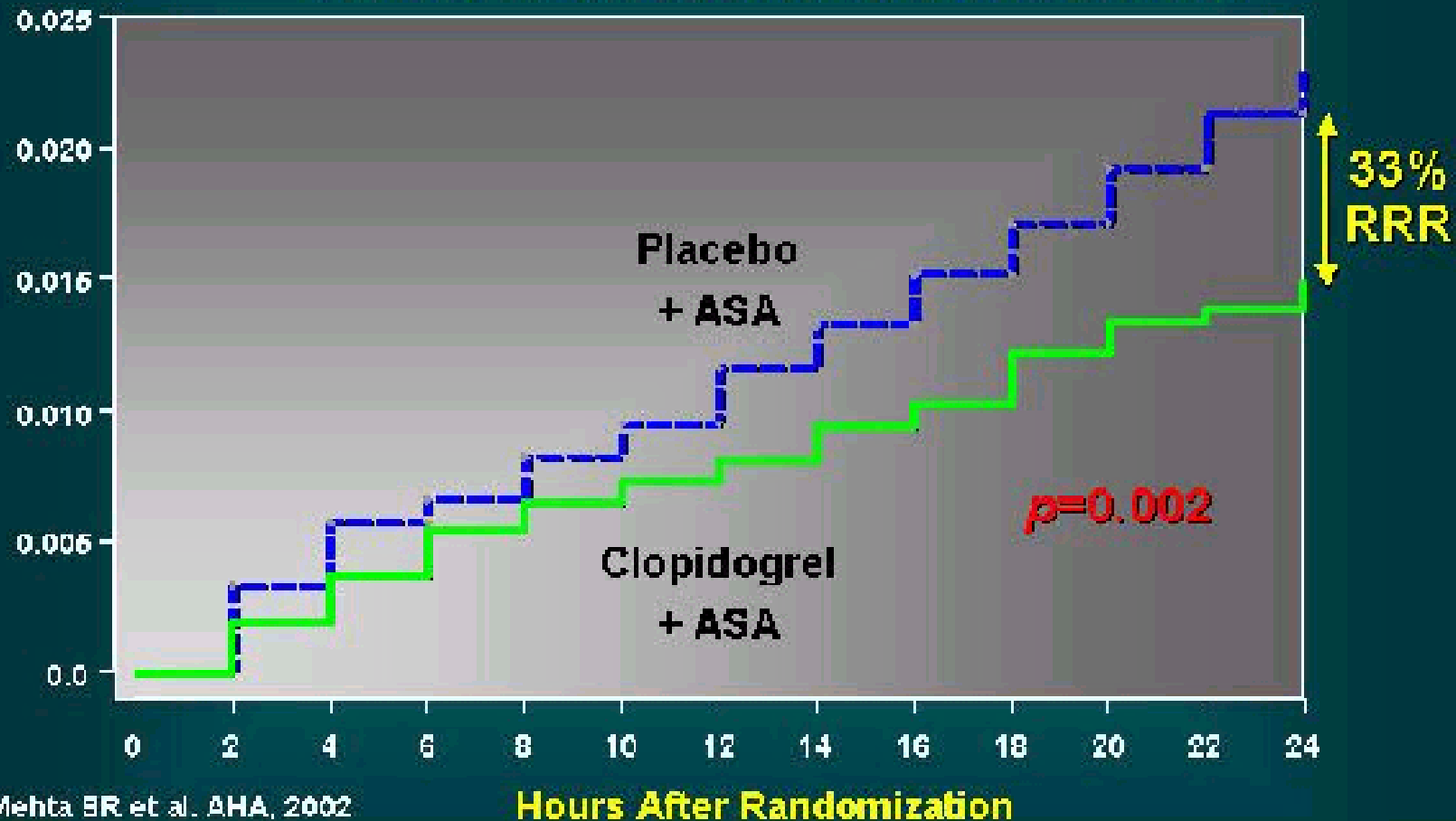
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Clopidogrel: beneficio a breve..

CURE

Clopidogrel in the 1st 24 Hours

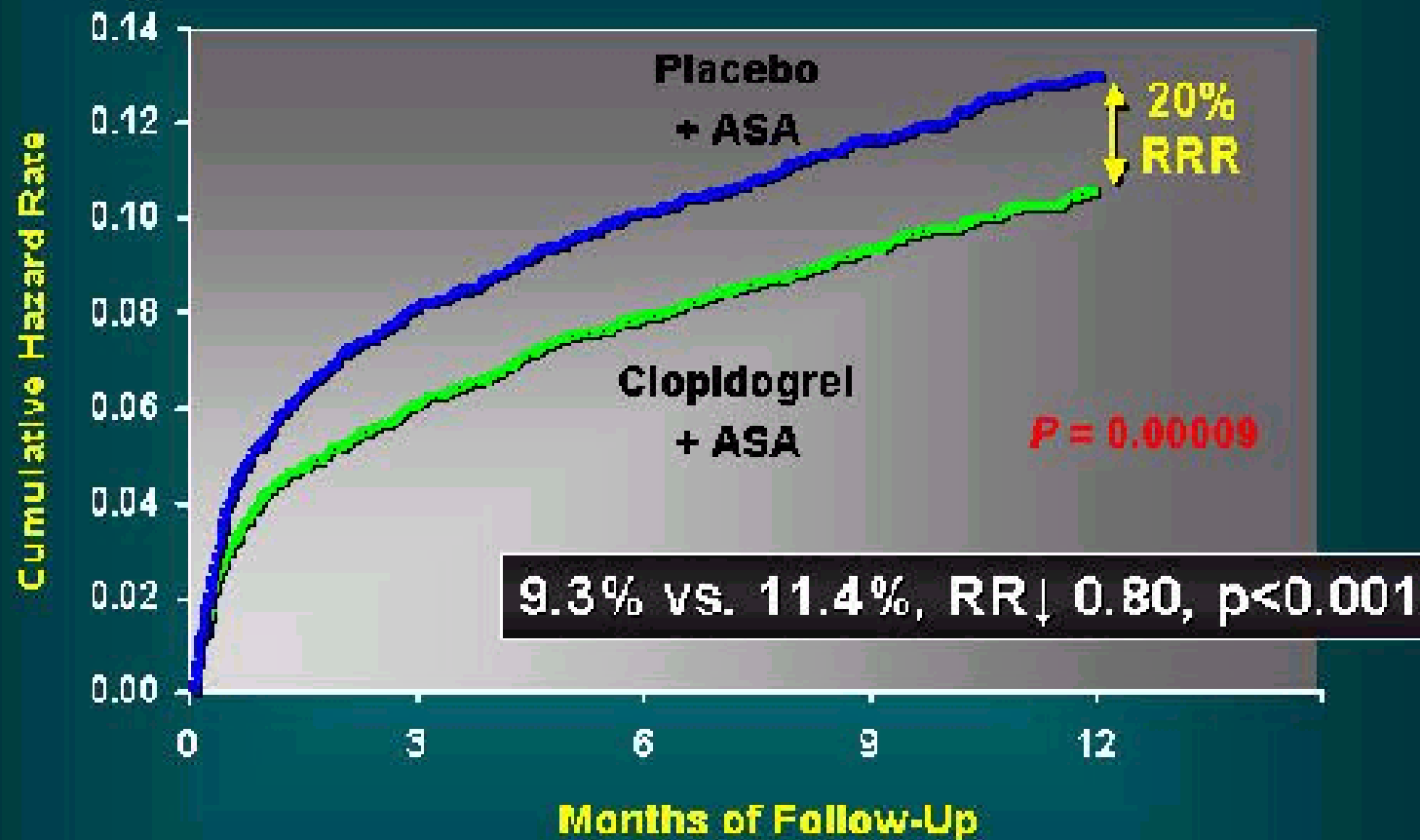
CV Death, MI, Stroke, Severe Ischemia



.. e a lungo termine

CURE

1° End Point: CV Death, MI, Stroke



SCA senza ST sopraslivellato

Eparina non frazionata o a
basso peso molecolare?

SCA senza ST sopraslivellato

Inibitori GP IIb/IIIa

Inibitori IIb/IIIa nelle SCA

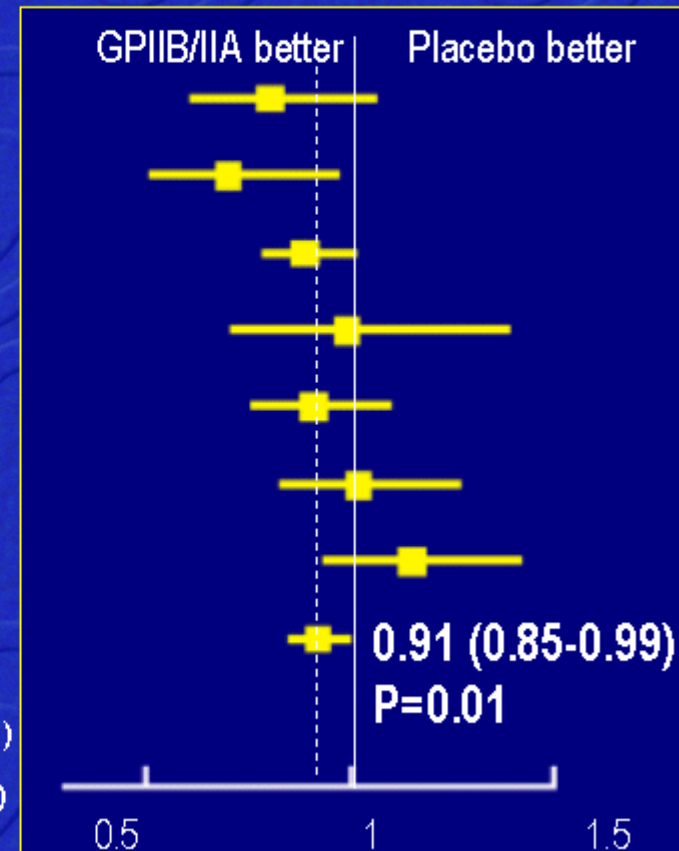
Seven Trials comparing GPIIb/IIIa vs Placebo in ACS

Both groups received Heparin +ASA

31,402 patients

Death and MI at 30-day FU

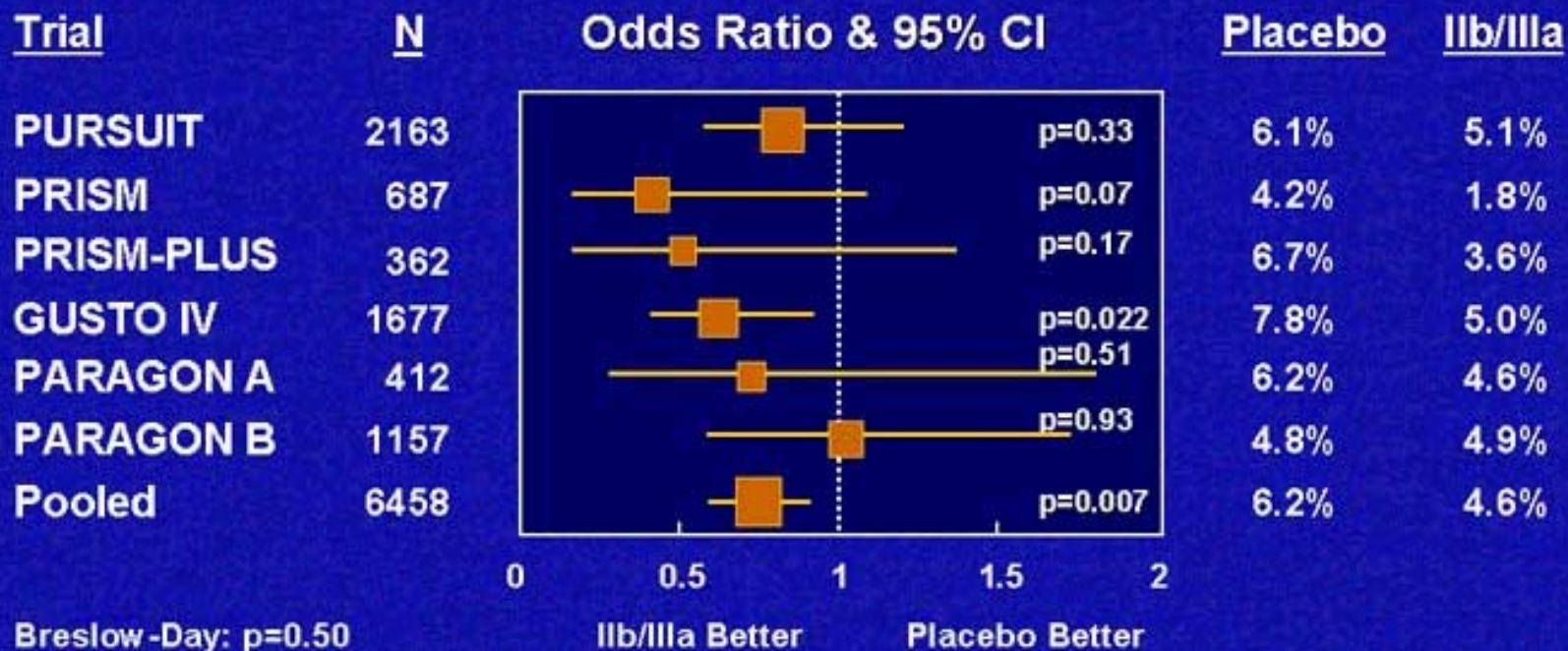
	Drug	Placebo
PRISM	5.8%	7.1%
PRISM-PLUS	8.7%	11.9%
PURSUIT	14.2%	15.7%
PARAGON A	11.6%	11.7%
PARAGON B	10.6%	11.5%
GUSTO IV 24h	8.2%	8%
GUSTO IV 48h	9.1%	8%
Total	10.3%	11.9%
Major bleedings	2.5%	1.4% (0.001)
ICH	0.08%	0.05% (NS)



Inibitori IIb/IIIa nei pz diabetici

GP IIb/IIIa Inhibitors Reduce Mortality in Diabetic Patients with non-ST Segment Elevation Acute Coronary Syndromes

30-Day Mortality Diabetic Patients

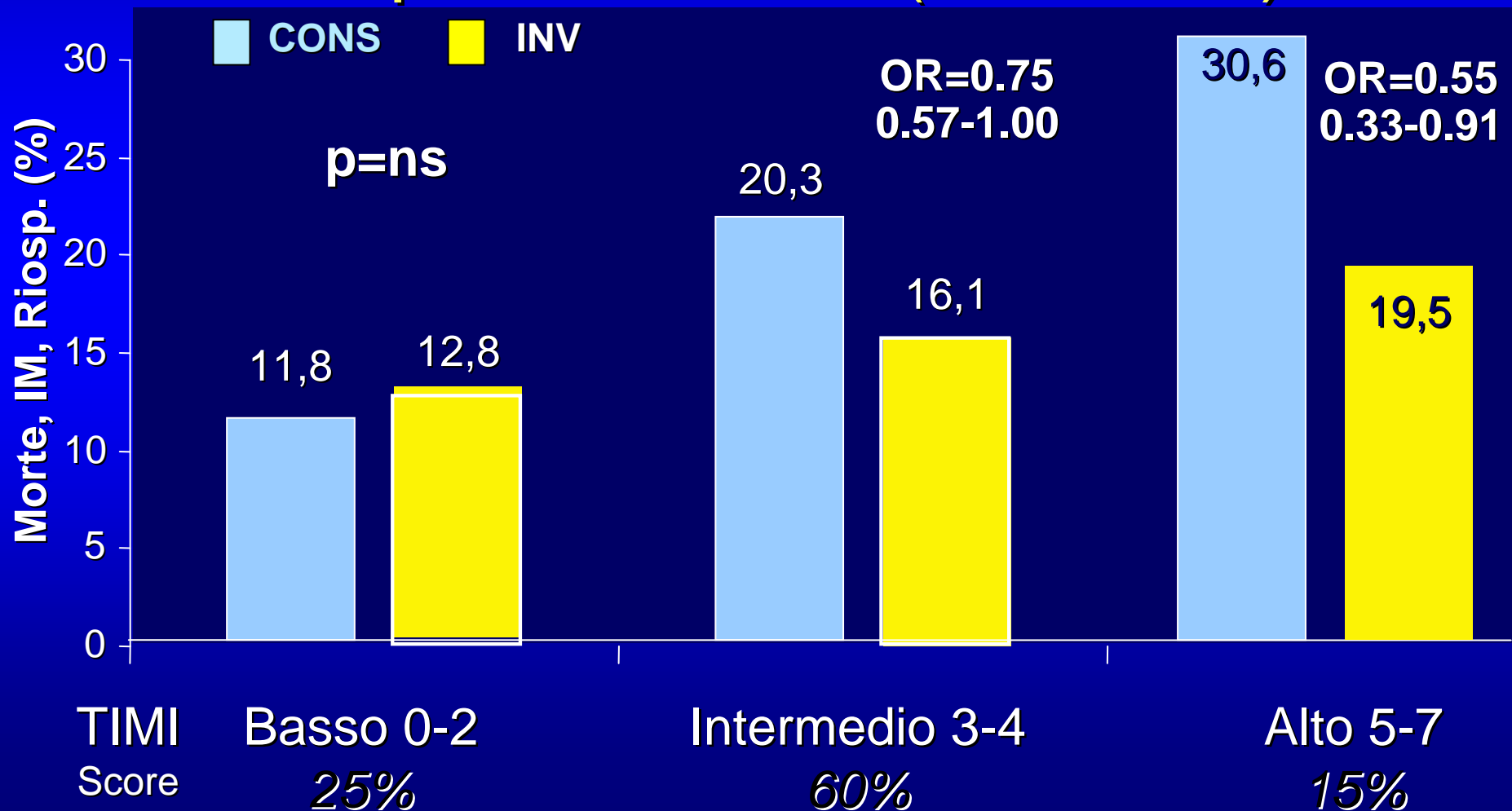


SCA senza ST sopraslivellato

Strategia invasiva o
conservativa?

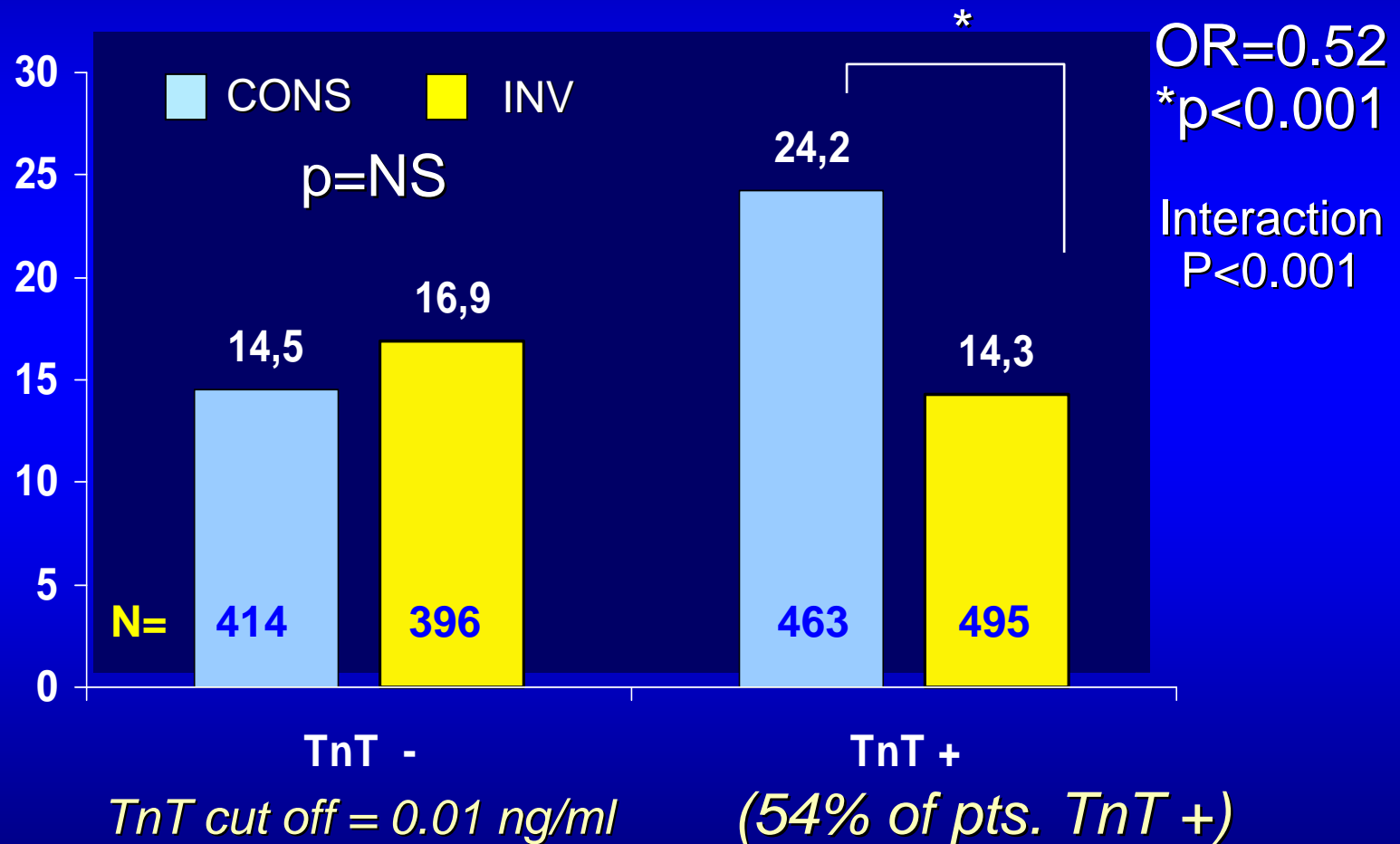
Strategia invasiva per pz. ad alto rischio

Stratificazione del rischio:
Endpoint a 6 Mesi (TACTICS)



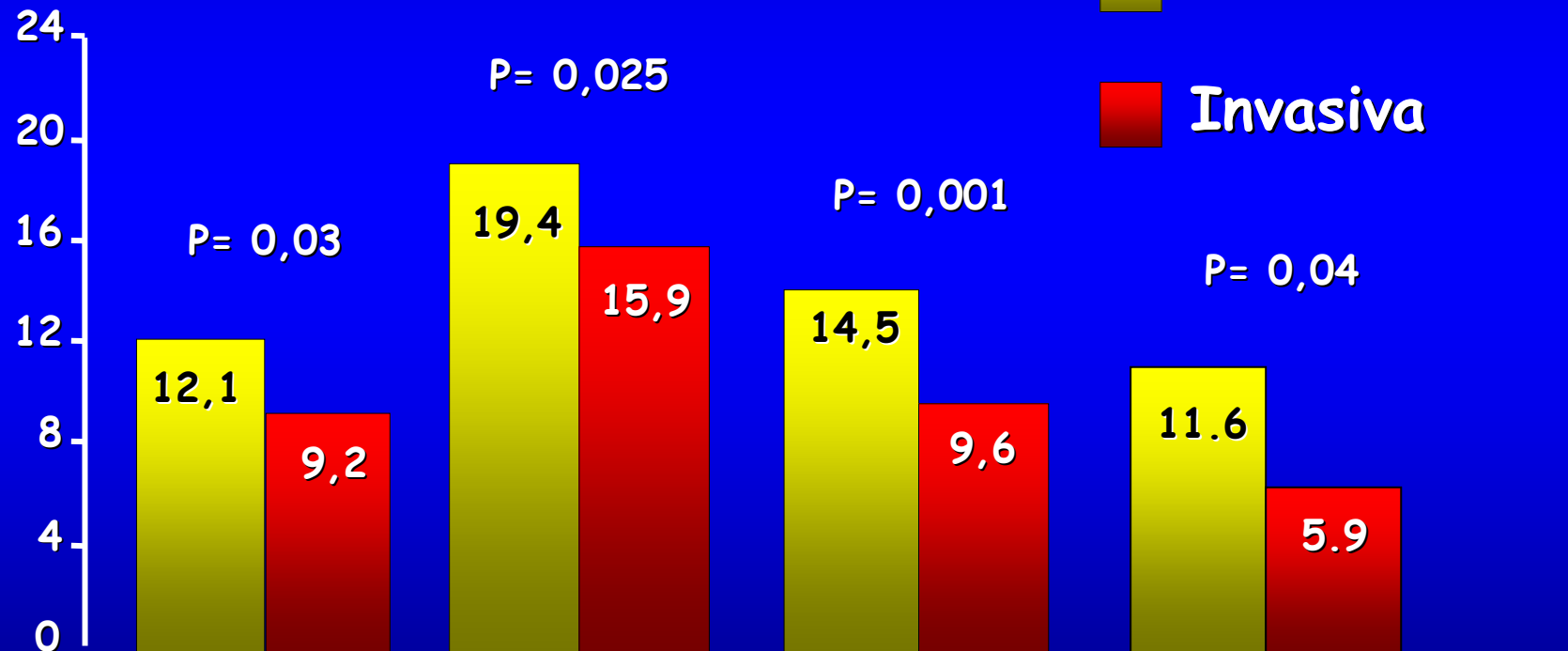
Pazienti ad alto rischio (TACTICS): Primary Endpoint a 6 Mesi

morte, IM, riospedalizzazione per SCA



SCA "NSTE" 2003: Evidenze per una Strategia Invasiva

Primary Endpoint (%)



FRISC II

TACTICS

RITA 3

ISAR-COOL

rischio pz

alto

medio

medio

alto

t.m. alla PCI

5 gg

1 g

3 gg

2 h

SCA senza ST sopraslivellato

Linee Guida

European Heart Journal (2002) **23**, 1809–1840
doi:10.1053/euhj.2002.3385, available online at <http://www.idealibrary.com> on IDEAL®



Task Force Report

Management of acute coronary syndromes in patients
presenting *without* persistent ST-segment elevation

The Task Force on the Management of Acute Coronary Syndromes of the
European Society of Cardiology*

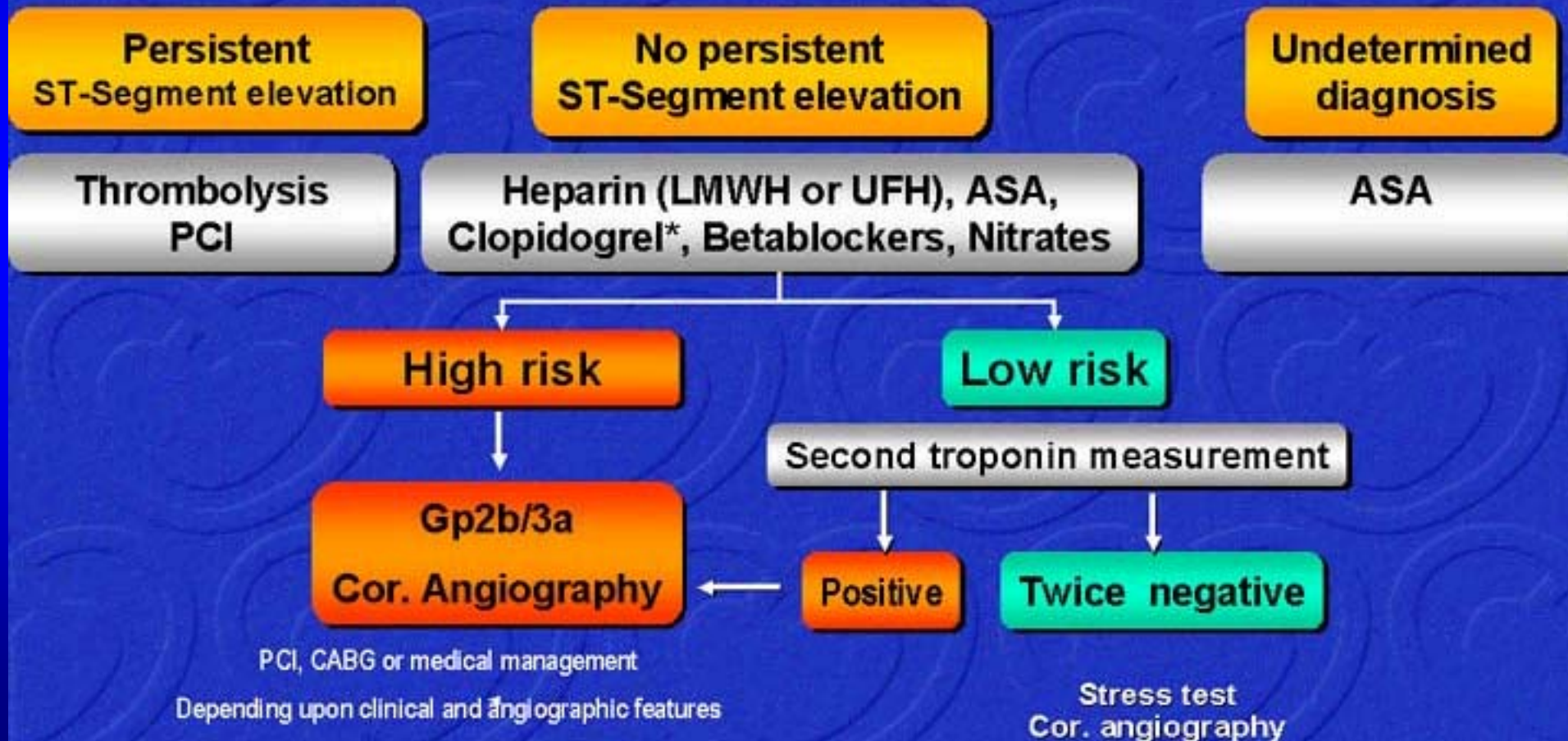
Michel E. Bertrand, Chair, Maarten L. Simoons, Keith A. A. Fox, Lars C. Wallentin,
Christian W. Hamm, Eugene McFadden, Pim J. De Feyter,
Giuseppe Specchia, Witold Ruzyllo

ACS ESC guidelines



Clinical suspicion of ACS

Physical examination, (Echocardiogram)
ECG monitoring, Blood samples



* omit clopidogrel if the patient is likely to go to CABG within 5 days

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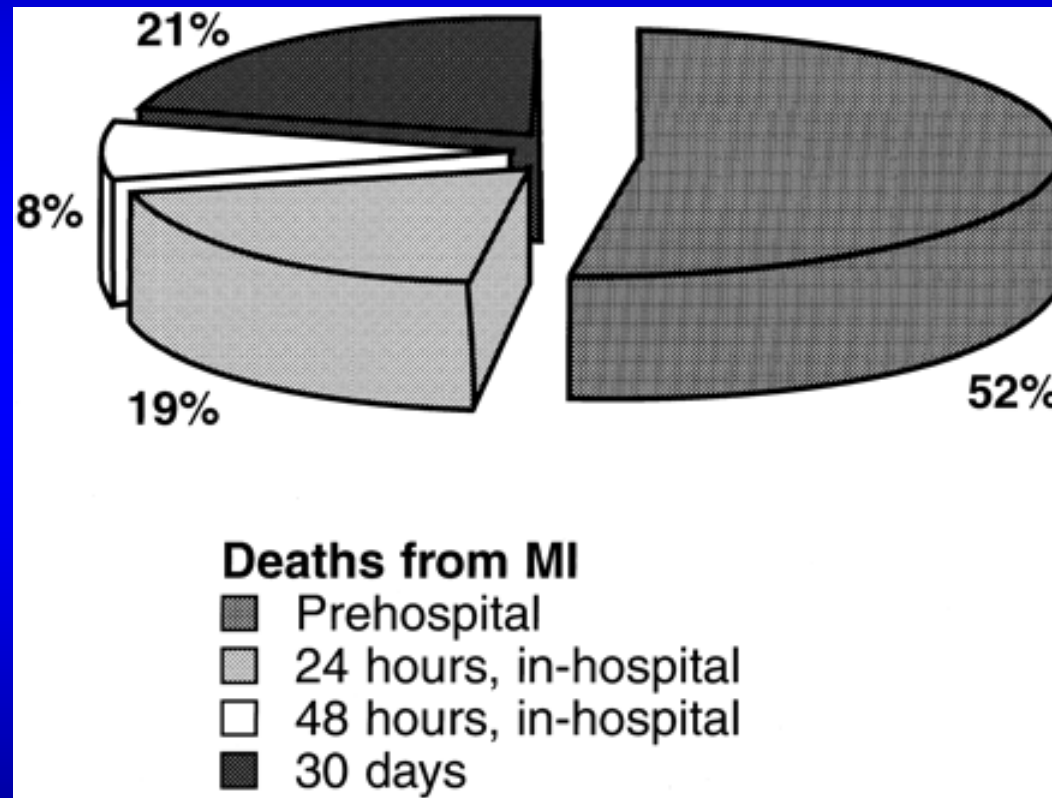
Epidemiologia

Incidenza: 1.1 milione persone/anno negli Stati Uniti; di questi:

- circa la metà muore entro 1 ora dall'esordio dei sintomi o prima di raggiungere l'ospedale;
- 24% degli uomini e il 42% delle donne muore entro 1 anno;
- 21% degli uomini e il 30% delle donne sviluppa CHF entro 6 anni.

L'1% di riduzione della mortalità salverebbe 3.400 vite/anno.

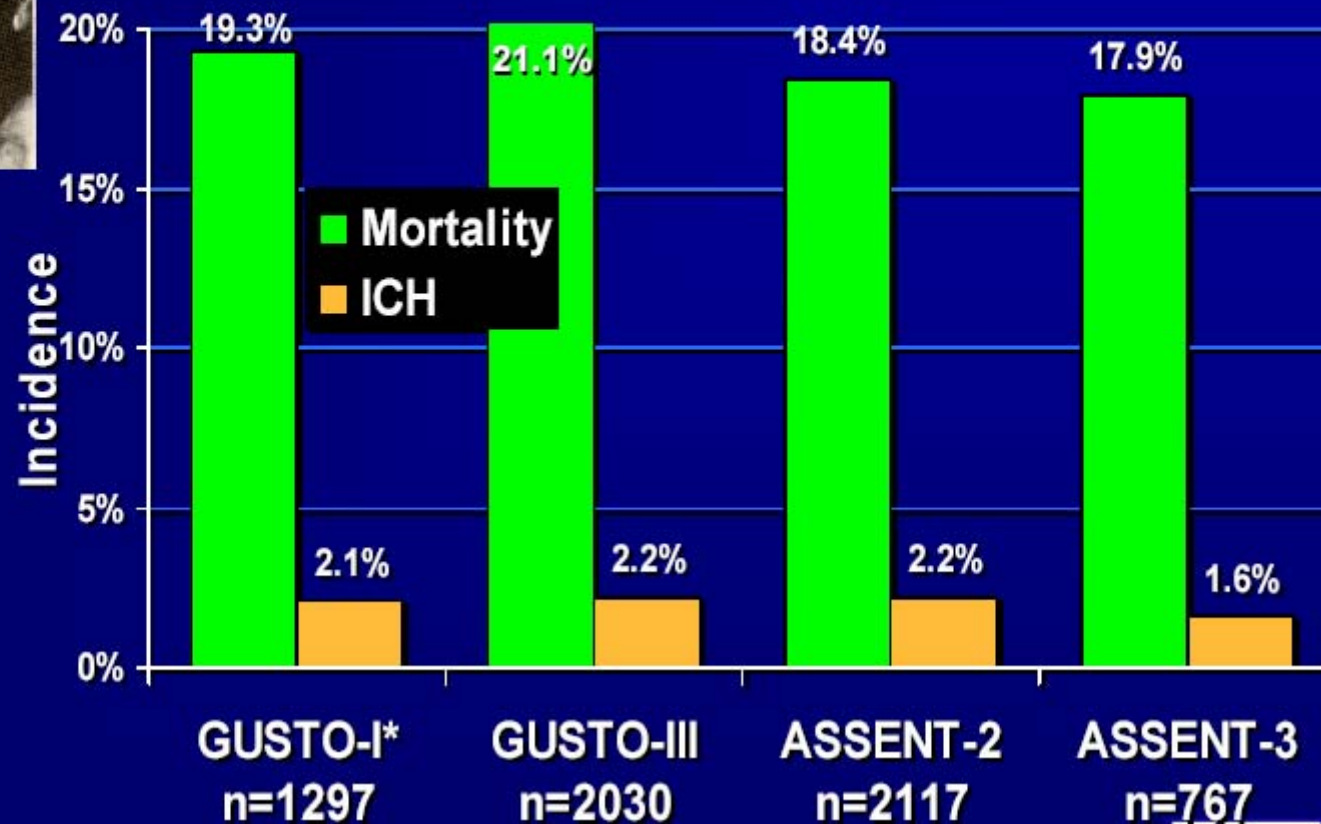
Distribuzione della mortalità in pazienti con IMA nei primi 30 gg



Gli ultra75enni: una popolazione ad alto rischio nell'IMA



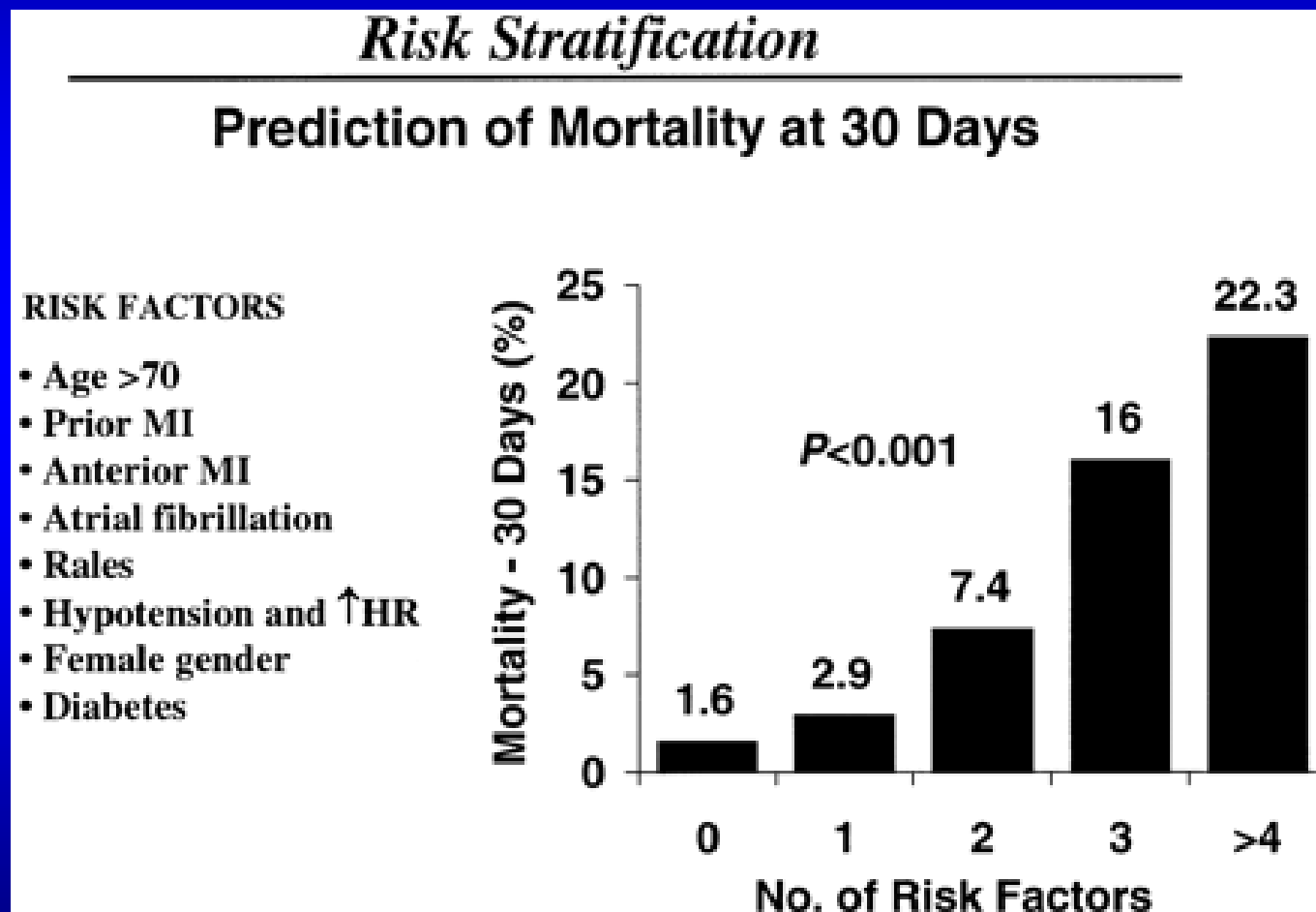
The Elderly in AMI trials: >75 Years Old



SCA con ST sopraslivellato

Stratificazione del rischio

Predittori di mortalità a 30 giorni in pazienti con IMA (indipendentemente dai valori di troponina)



SCA con ST sopraslivellato

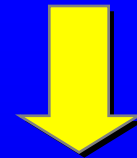
Terapia:
rapida riperfusione

Vantaggi della riperfusione precoce

RIPERFUSIONE PRECOCE



SALVATAGGIO DEL MIOCARDIO



PRESERVAZIONE DELLA FUNZIONE VS



AUMENTO SOPRAVVIVENZA

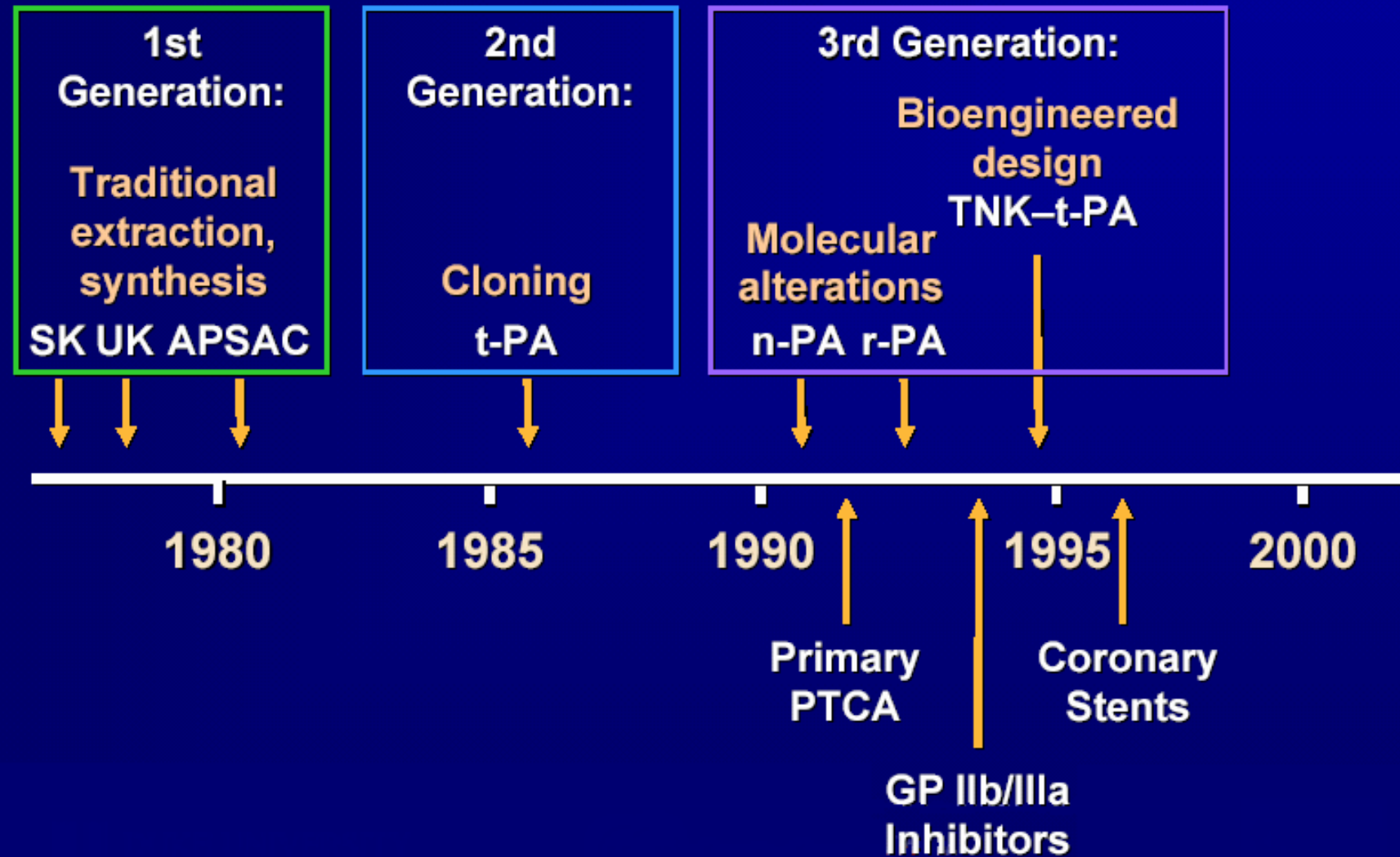
Terapia riperfusiva

Fibrinolisi: - streptochinasi - reteplase (rPA)
- urochinasi - lanoteplase (nPA)
- alteplase (rtPA) - tenecteplase (TNK)

PTCA: - primaria (di prima intenzione)
- rescue (dopo fibrinolisi inefficace)
- facilitata (dopo pretrattamento con mezza dose di fibrinolitico e/o inib. GP IIb/IIIa)

BPAC d'emergenza: - stenosi TC (?)
- PTCA inefficace o complicata
- complicanze meccaniche

L'evoluzione della terapia per la riperfusione coronarica

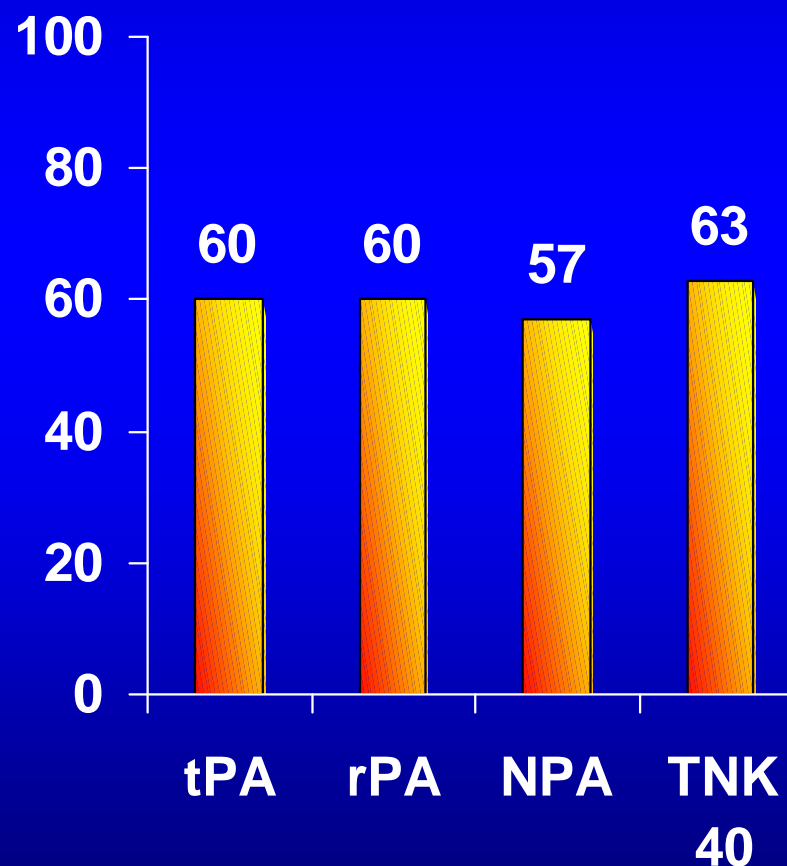


La trombolisi ha i suoi limiti

Nessun trombolitico riesce a superare il muro del 60% di flusso TIMI 3 a 90 minuti

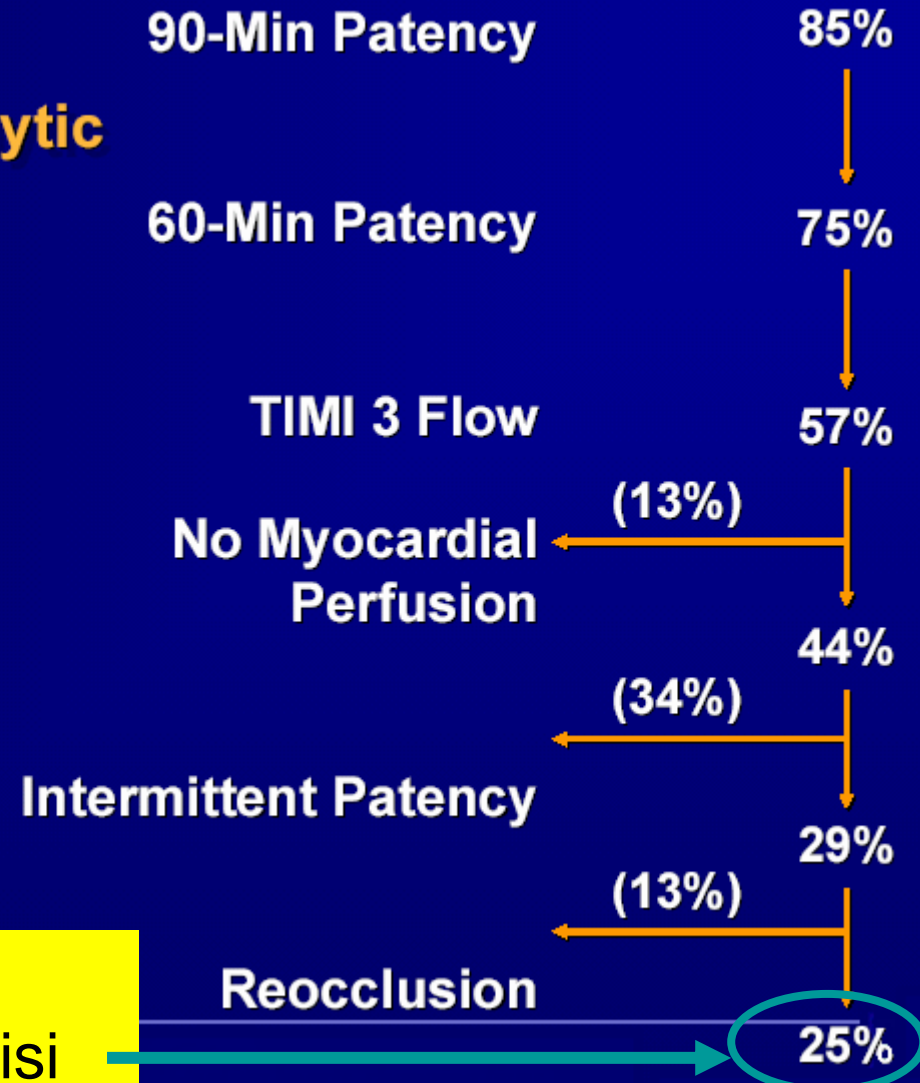


% TIMI 3 Flow



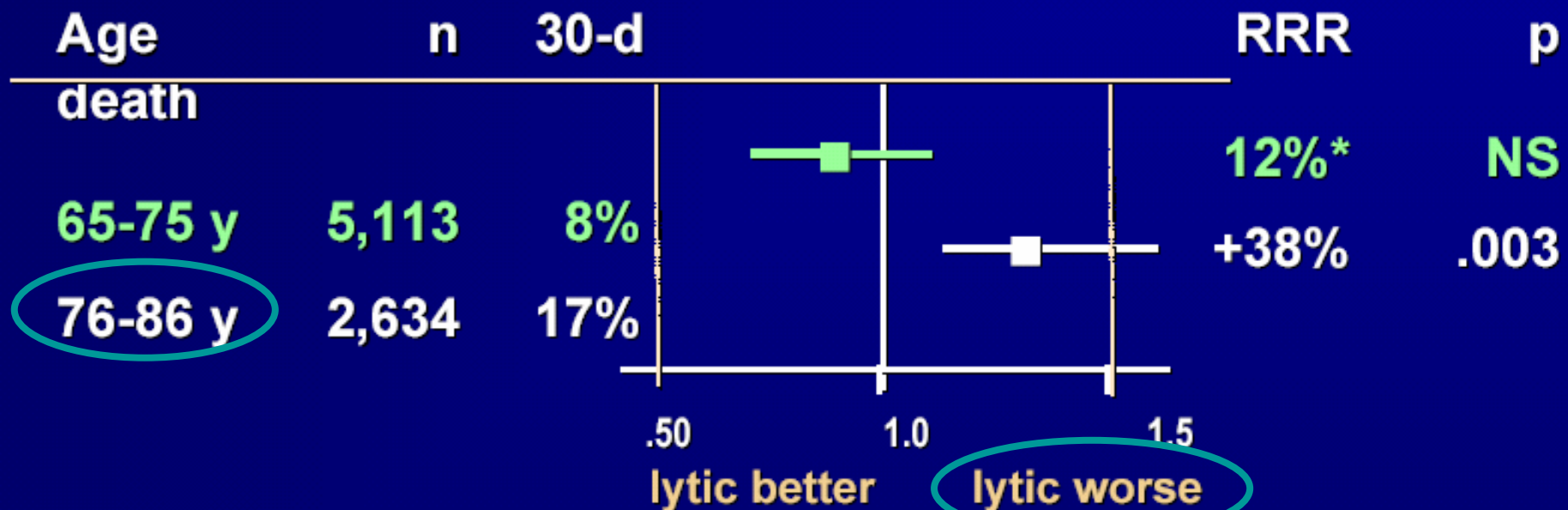
La trombolisi ha i suoi limiti

Limitations of Current Fibrinolytic Strategies: "The Illusion of Reperfusion"



Gli ultra75enni: la trombolisi è meno efficace

Cooperative Cardiovascular Project Database

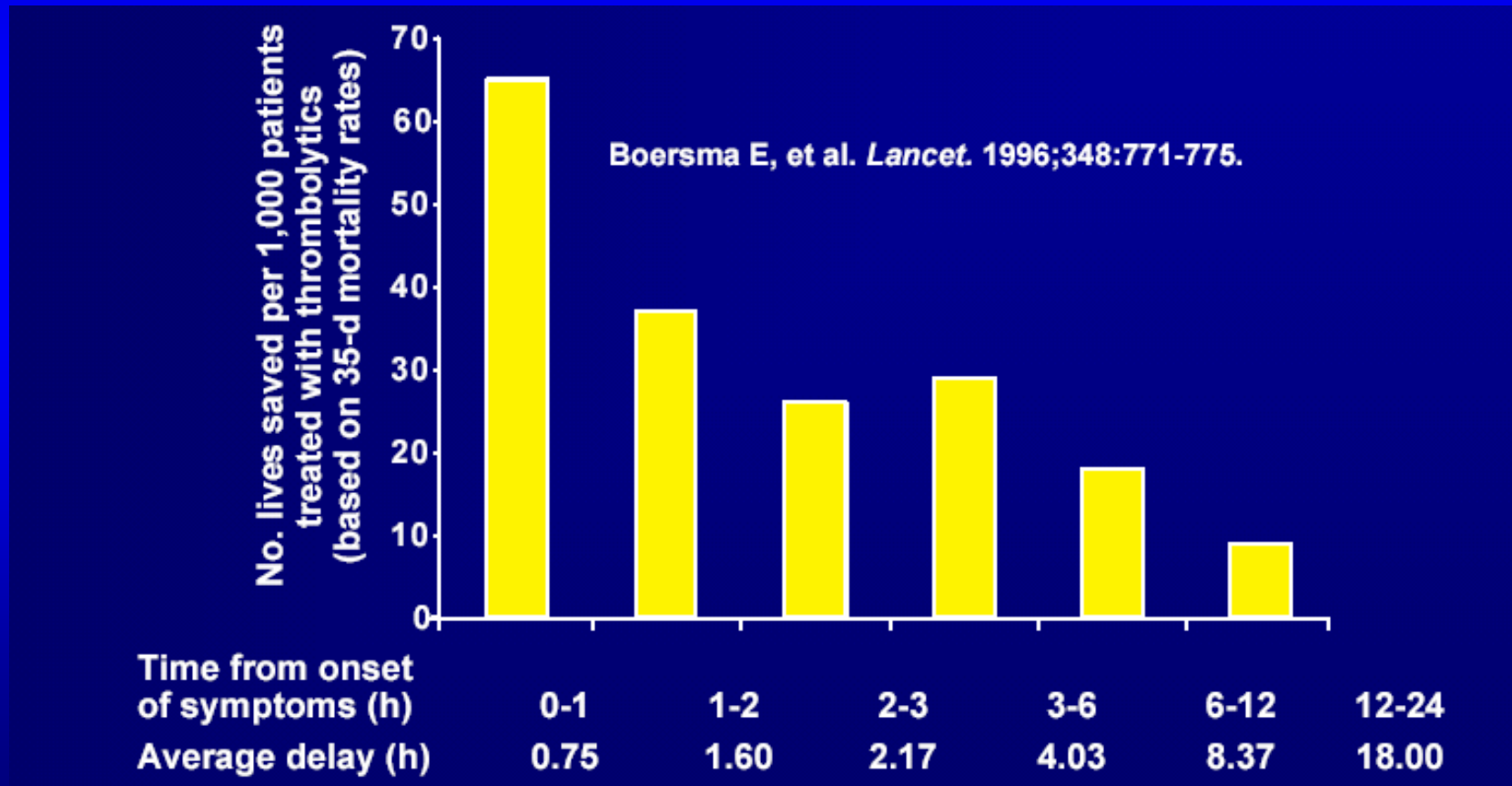


*no benefit for first 4 days

-- Thiemann, Circulation, 2000

Trombolisi: “time is muscle” (and survival)

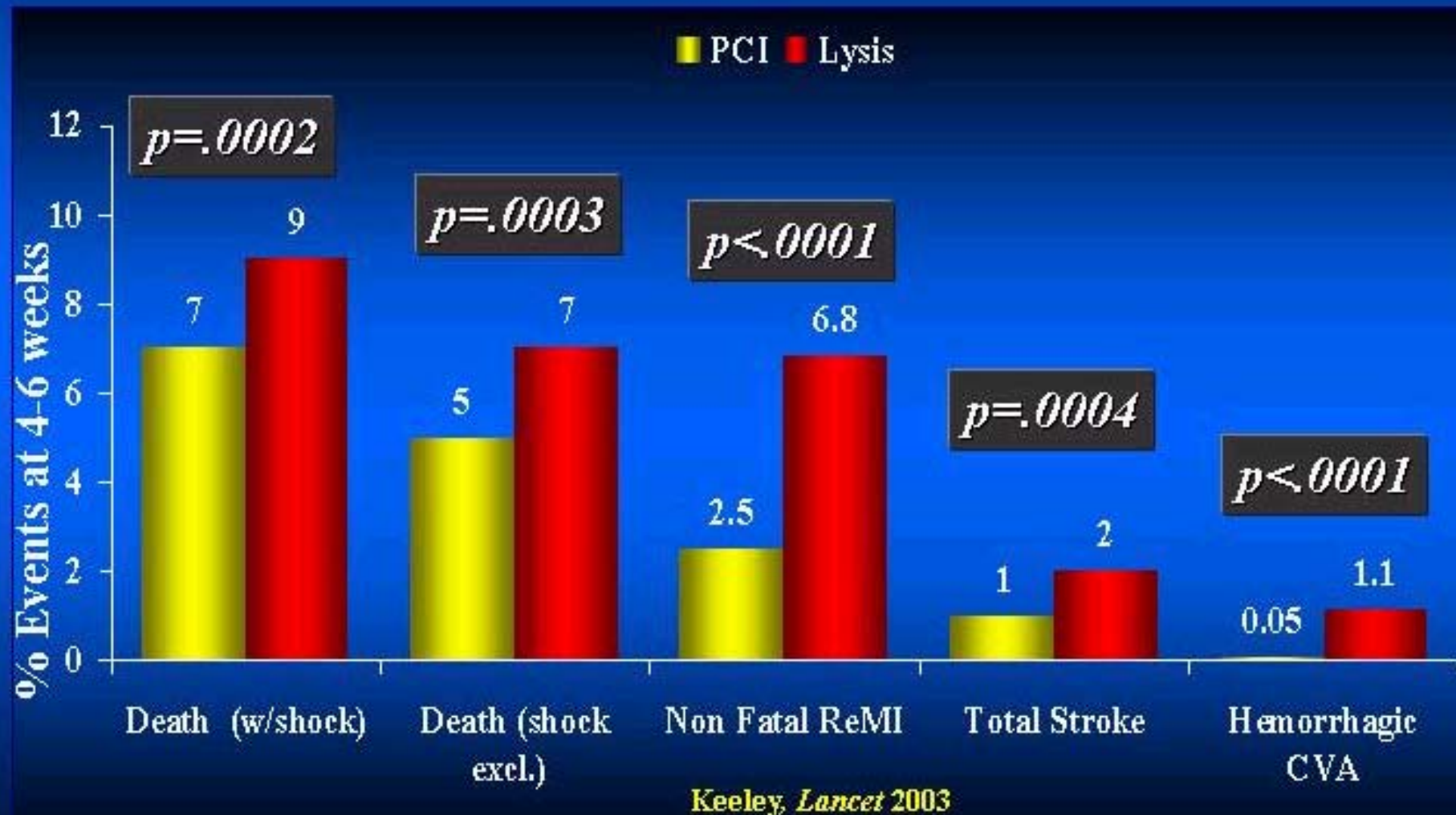
Tempo precoronarico e mortalità



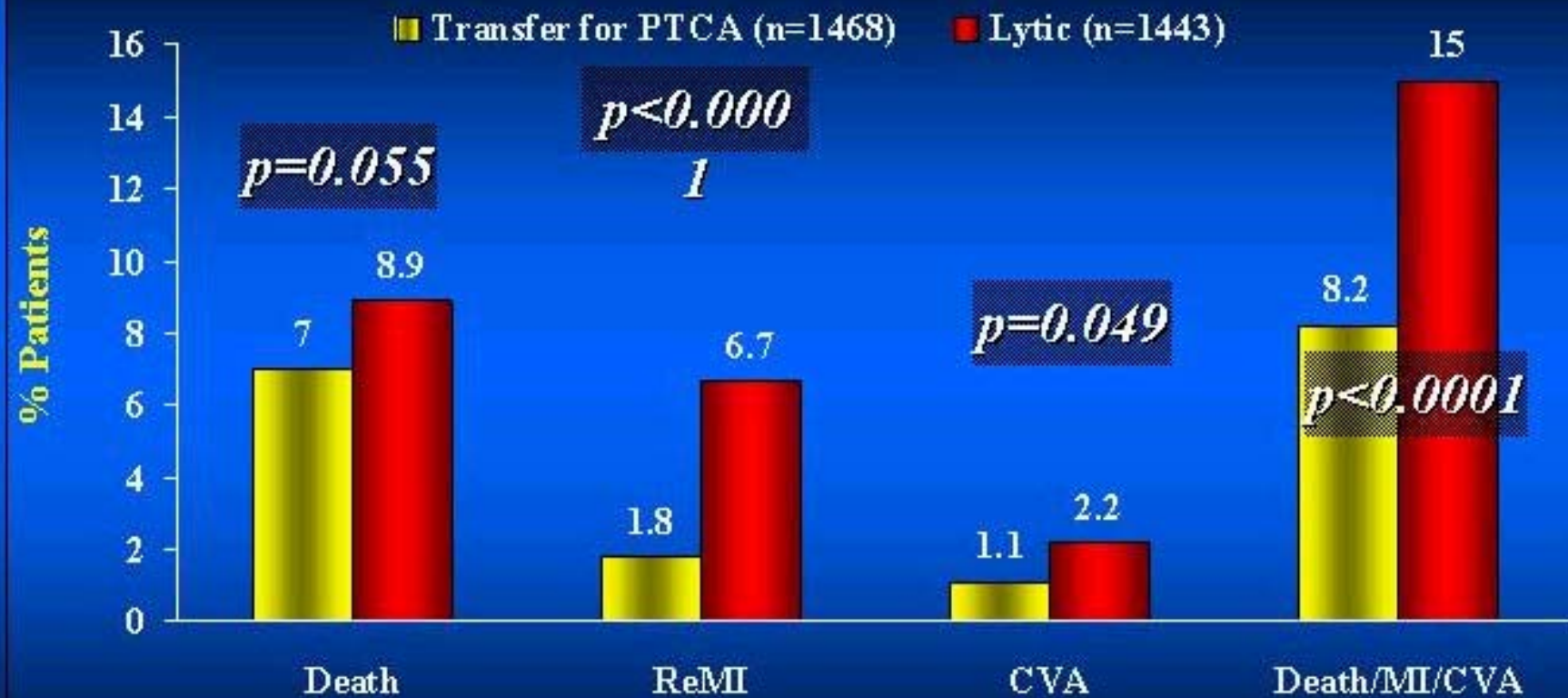
SCA con ST sopraslivellato

Fibrinolisi o angioplastica?

PCI vs Lysis: metanalisi di 23 trial randomizzati (7739 pz)



Trasferimento per pPTCA vs lisi “on-site” (dati ricavati da 5 RCT*)



*LIMI, Prague I & II, Air PAMI, DANAMI-II trials

Keeley & Grines, in press

SCA con ST sopraslivellato

Linee Guida



2003 ESC Guidelines for STEMI

Task Force Report

Management of acute myocardial infarction in patients presenting with ST-segment elevation

The Task Force on the Management of Acute Myocardial Infarction of the European Society of Cardiology,

Frans Van de Werf, Chair, Diego Ardissino, Amadeo Betriu, Dennis V. Cokkinos, Erling Falk, Keith A.A. Fox, Desmond Julian, Maria Lengyel, Franz-Josef Neumann, Witold Ruzyllo, Christian Thygesen, S. Richard Underwood, Alec Vahanian, Freek W.A. Verheugt, William Wijns

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2003 Updated ESC Guidelines

PCI should be considered the treatment of choice

*when it can be performed by an experienced team
within 90min from the first medical contact*
