

Ecostress con Dobutamina: predittività del risultato della PCI in CAD Stabile

Rossella Gottilla Cardiologia con Utic ed Emodinamica AORN A Cardarelli



# The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

APRIL 12, 2007

VOL. 356 NO. 15

## Optimal Medical Therapy with or without PCI for Stable Coronary Disease

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#### ABSTRACT

#### BACKGROUND

In patients with stable coronary artery disease, it remains unclear whether an initial management strategy of percutaneous coronary intervention (PCI) with intensive pharmacologic therapy and lifestyle intervention (optimal medical therapy) is superior to optimal medical therapy alone in reducing the risk of cardiovascular events.

#### METHODS

We conducted a randomized trial involving 2287 patients who had objective evidence of myocardial ischemia and significant coronary artery disease at 50 U.S. and Canadian centers. Between 1999 and 2004, we assigned 1149 patients to undergo PCI with optimal medical therapy (PCI group) and 1138 to receive optimal medical therapy alone (medical-therapy group). The primary outcome was death from any cause and non-fatal myocardial infarction during a follow-up period of 2.5 to 7.0 years (median, 4.6).

#### RESULTS

There were 211 primary events in the PCI group and 202 events in the medicaltherapy group. The 4.6-year cumulative primary-event rates were 19.0% in the PCI group and 18.5% in the medical-therapy group (hazard ratio for the PCI group, 1.05; 95% confidence interval [CI], 0.87 to 1.27; P=0.62). There were no significant differences between the PCI group and the medical-therapy group in the composite of death, myocardial infarction, and stroke (20.0% vs. 19.5%; hazard ratio, 1.05; 95% CI, 0.87 to 1.27; P=0.62); hospitalization for acute coronary syndrome (12.4% vs. 11.8%; hazard ratio, 1.07; 95% CI, 0.84 to 1.37; P=0.56); or myocardial infarction (13.2% vs. 12.3%; hazard ratio, 1.13; 95% CI, 0.89 to 1.43; P=0.33).

#### CONCLUSIONS

As an initial management strategy in patients with stable coronary artery disease, PCI did not reduce the risk of death, myocardial infarction, or other major cardiovascular events when added to optimal medical therapy. (ClinicalTrials.gov number, NCT00007657.)

Affiliations for all authors are listed in the Appendix. Address reprint requests to Dr. Boden at the Division of Cardiology, Buffalo General Hospital, 100 High St., Buffalo, NY 14201, or at wboden@kaleidahealth. org.

Members of the Clinical Outcomes Utilizing Revascularization and Aggressive Drug Evaluation (COURAGE) trial are listed in the Appendix and in the Supplementary Appendix, available with the full text of this article at www.nejm.org.

This article (10.1056/NEJMoa070829) was published at www.nejm.org on March 26, 2007.

N Engl J Med 2007;356:1503-16.
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## COURAGE





# The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

SEPTEMBER 13, 2012

VOL. 367 NO. 11

#### Fractional Flow Reserve–Guided PCI versus Medical Therapy in Stable Coronary Disease

Bernard De Bruyne, M.D., Ph.D., Nico H.J. Pijls, M.D., Ph.D., Bindu Kalesan, M.P.H., Emanuele Barbato, M.D., Ph.D., Pim A.L. Tonino, M.D., Ph.D., Zsolt Piroth, M.D., Nikola Jagic, M.D., Sven Möbius-Winkler, M.D., Gilles Rioufol, M.D., Ph.D., Nils Witt, M.D., Ph.D., Petr Kala, M.D., Philip MacCarthy, M.D., Thomas Engström, M.D., Keith G. Oldroyd, M.D., Kreton Mavromatis, M.D., Ganesh Manoharan, M.D., Peter Verlee, M.D., Ole Frobert, M.D., Nick Curzen, B.M., Ph.D., Jane B. Johnson, R.N., B.S.N., Peter Jüni, M.D., and William F. Fearon, M.D., for the FAME 2 Trial Investigators\*



#### Fractional Flow Reserve-Guided PCI for Stable Coronary Artery Disease

Bemard De Bruyne, M.D., Ph.D., William F. Fearon, M.D., Nico H.J. Pijls, M.D., Ph.D., Emanuele Barbato, M.D., Ph.D., Pim Tonino, M.D., Ph.D., Zsolt Piroth, M.D., Nikola Jagic, M.D., Sven Mobius-Winckler, M.D., Gilles Riouffol, M.D., Ph.D., Nils Witt, M.D., Ph.D., Petr Kala, M.D., Philip MacCarthy, M.D., Thomas Engström, M.D., Keith Oldroyd, M.D., Kreton Mavromatis, M.D., Ganesh Manoharan, M.D., Peter Verlee, M.D., Ole Frobert, M.D., Nick Curzen, B.M., Ph.D., Jane B. Johnson, R.N., B.S.N., Andreas Limacher, Ph.D., Eveline Nüesch, Ph.D., and Peter Jüni, M.D., for the FAME 2 Trial Investigators\*



#### ABSTRACT

# A Polessay Each Point Dept. Manual crass, 2.49 (EVS. C., 6.73–6.65) Plan series action, 0.19 (EVS. C., 6.73–6.65) Point series action, 0.19 (EVS. C., 6.73–6.65) Plan series action, 0.19 (EVS. C., 6.73–6.65

Shon are the conductor action can be primary and point (a composite of death from any cause, notifical response betterfereinne, require transcriptions) (Part of ) and of which or responsed in derivation (Part of ) in the see subgroups, stratified on the hast of a landment point of 2 days after randomization (united deathed leng), these disclosed superaging from each that accurred which in 24 years of the set that occurred between 6 days and the end of follow-up at 2 years. Chals for the first 2 days are not included in the period after 2 days. The insents whose the data for days 0 to 2 on an expanded year, Pushes for internation were calculated from text of heterogeneity between time periods. Hazard rasion below 100 denote a lower incidence of the primary and point in the CP group than the medical-theory group.

#### CONCLUSIONS

In patients with stable coronary artery disease, FFR-guided PCI, as compared with medical therapy alone, improved the outcome. Patients without ischemia had a favorable outcome with medical therapy alone. (Funded by St. Jude Medical; FAME 2 ClinicalTrials.gov number, NCT01132495.)



## Ecostress Dobutamina in CAD cronica

- Prima della PTCA
- per stabilire il significato emodinamico di una stenosi
- il miocardio ibernato cosi da pianificare l'eventuale intervento.
- Buon valore predittivo pos (93%) e neg (80%) per il miglioramento dell'ischemia precocemente post PTCA.
- Riduzione dell'ischemia e recupero del miocardio ibernato.

Kao H-L Am J Cardiol 1995;76:652-656

Davila-Roman VG Am J Cardiol 1995;76:245-249

# Potere diagnostico per restenosi post PTCA

- ▶ Nel 95% dei pz avviene nei primi 6 mesi post PTCA
- Sensibilità 74% (Cl 69-79%).
- Specificità 87% (CI 84-89%).
- Predittivo positivo (83%)
- Predittivo negativo (97%)

## Restenosi Post PTCA

- ► ECG da sforzo sensibilità 54% specificità 70%
- ▶ VPpos 64% VPN 61%
- Scintigrafia sensibilità 75-80%, specificità 79%
- ▶ VP pos 80% VPN 85%





European Society doi:10.1093/eurhearti/ehz425



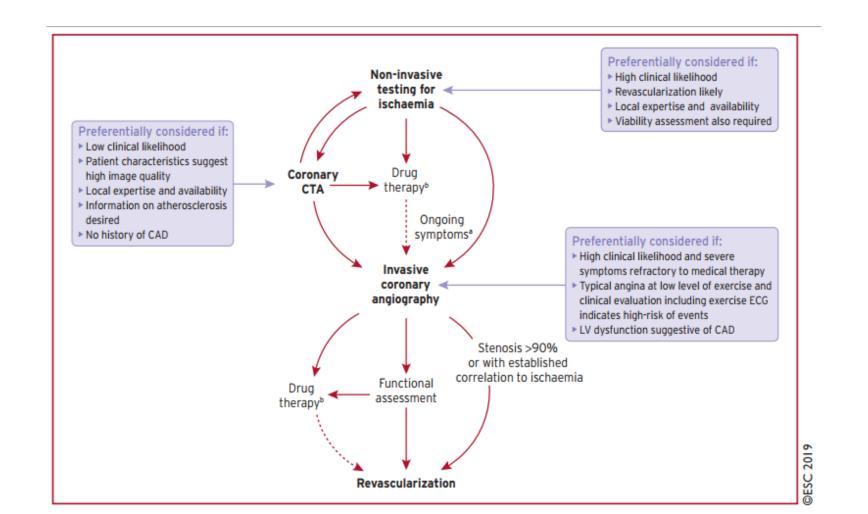
# 2019 ESC Guidelines for the diagnosis and management of chronic coronary syndromes

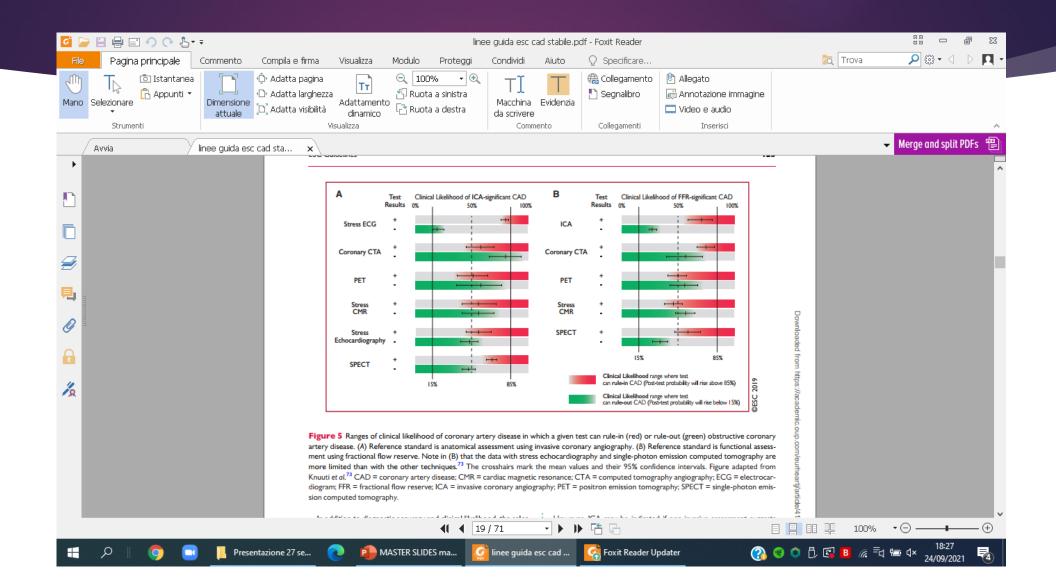
The Task Force for the diagnosis and management of chronic coronary syndromes of the European Society of Cardiology (ESC)

Authors/Task Force Members: Juhani Knuuti\* (Finland) (Chairperson), William Wijns\* (Ireland) (Chairperson), Antti Saraste (Finland), Davide Capodanno (Italy), Emanuele Barbato (Italy), Christian Funck-Brentano (France), Eva Prescott (Denmark), Robert F. Storey (United Kingdom), Christi Deaton (United Kingdom), Thomas Cuisset (France), Stefan Agewall (Norway), Kenneth Dickstein (Norway), Thor Edvardsen (Norway), Javier Escaned (Spain), Bernard J. Gersh (United States of America), Pavel Svitil (Czech Republic), Martine Gilard (France), David Hasdai (Israel), Robert Hatala (Slovak Republic), Felix Mahfoud (Germany), Josep Masip (Spain), Claudio Muneretto (Italy), Marco Valgimigli (Switzerland), Stephan Achenbach (Germany), and Jeroen J. Bax (Netherlands)

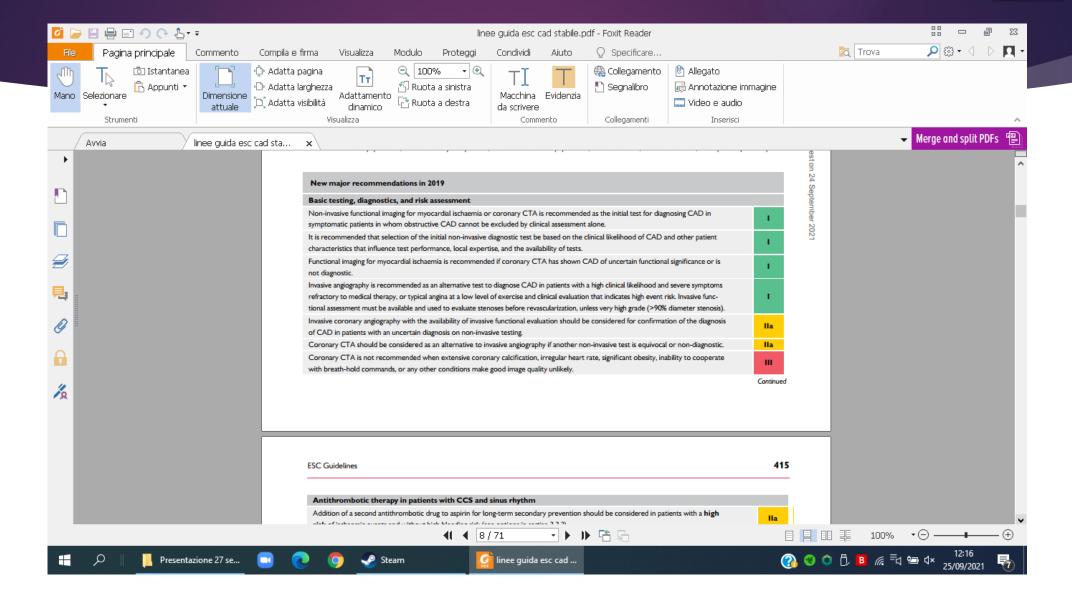
Document Reviewers: Franz-Josef Neumann (Germany) (CPG Review Coordinator), Udo Sechtem (Germany) (CPG Review Coordinator), Adrian Paul Banning (United Kingdom), Nikolaos Bonaros (Austria), Héctor Bueno (Spain), Raffaele Bugiardini (Italy), Alaide Chieffo (Italy), Filippo Crea (Italy), Martin Czerny (Germany), Victoria Delgado (Netherlands), Paul Dendale (Belgium),

# Linee guida ESC 2019 CAD Cronica

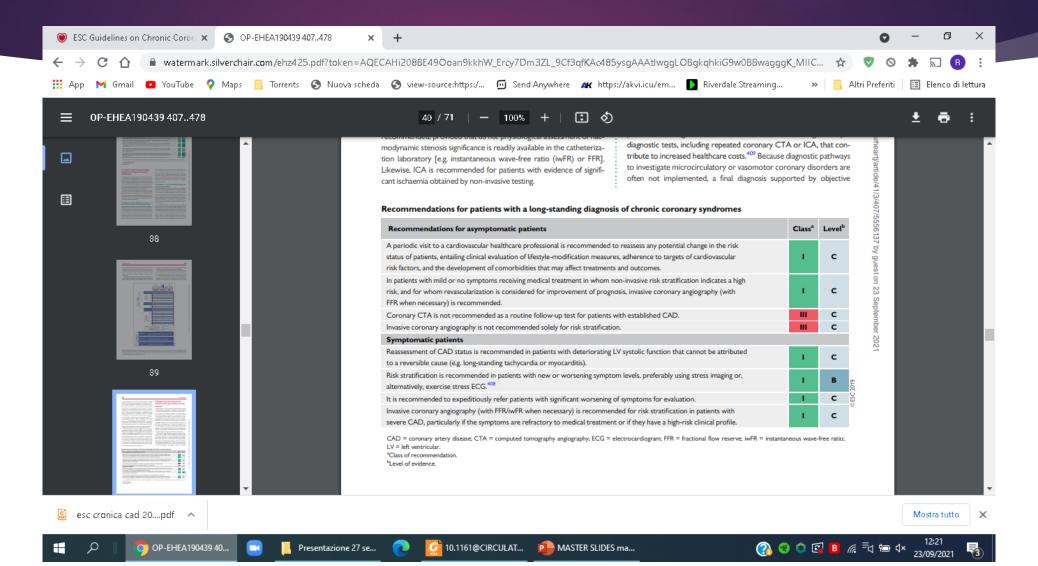


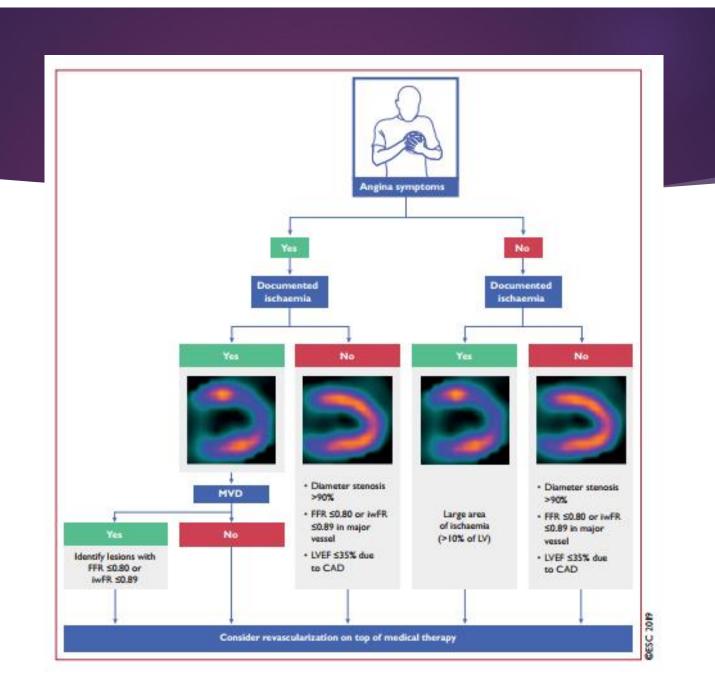


# Linee guida ESC 2019 CAD Cronica



# Lineeguida ESC 2019 CAD cronica









Echocardiography at rest

Early (e.g. I-3 months) after revascularization to set as a reference and/or periodically (e.g. at I year if previously abnormal and/or every 3-5 years) to evaluate LV function, valvular status and haemodynamic status.



Stress test for inducible ischaemia

As necessary, to investigate changes in symptoms level, and/or early (e.g. I-3 months) after revascularization to set as a reference and/or periodically (e.g. every 3-5 years) to reassess ischaemia.



Invasive coronary angiography

As necessary, for patients at high risk based on noninvasive ischaemia testing, or severe angina symptoms (e.g. CCS class 3-4). Not recommended solely for risk stratification.

- ► Fisico
- ▶ Dipiridamolo/adenosina
- ▶ Dobutamina



No, non sono stressato, sto solo attraversando un periodo difficile.

BESTI.IT

#### **GUIDELINES AND STANDARDS**

Guidelines for Performance, Interpretation, and Application of Stress Echocardiography in Ischemic Heart Disease: From the American Society of Echocardiography



Patricia A. Pellikka, MD, FASE, Chair, Adelaide Arruda-Olson, MD, PhD, FASE,
Farooq A. Chaudhry, MD, FASE,\* Ming Hui Chen, MD, MMSc, FASE, Jane E. Marshall, RDCS, FASE,
Thomas R. Porter, MD, FASE, and Stephen G. Sawada, MD, Rochester, Minnesota; New York, New York; Boston,
Massachusetts; Omaha, Nebraska; Indianapolis, Indiana

Journal of the American Society of Echocardiography January 2020

Table 12 Recommendations for stress echocardiography in patients with symptoms or suspected stable coronary artery disease

Recommendations for non-invasive testing for IHD	Class of recommendation	Level of evidence
In patients with suspected stable CAD, intermediate pretest probability and preserved ejection fraction, stress imaging, such as stress echocardiography, is preferred as the initial test option.	1	В
In patients without typical angina, an imaging stress test, such as stress echocardiography, is recommended as the initial test for diagnosing stable CAD if the pretest probability is high or if LVEF is reduced.	I	В
In patients with suspected CAD and with resting ECG abnormalities, which prevent accurate interpretation of ECG changes during stress, an imaging stress test, such as stress echocardiography, is recommended.	I	В
In patients with LBBB and symptoms consistent with IHD, stress echocardiography (either ESE or DSE) is preferred over SPECT imaging because of its greater specificity and because of its versatility for detecting other cardiac conditions associated with LBBB	I	В
Stress echocardiography is the preferred test for women with an indication for an noninvasive imaging test for known or suspected CAD because of its safety (absence of radiation to the breasts), and greater specificity (absence of breast attenuation artifact)	I	В
ESE is the preferred imaging stress test for children with suspected IHD because of the absence of radiation to developing tissues and absence of need for an intravenous line, and the provision of the prognostically important assessment of exercise capacity	I	В
A pharmacologic stress test, such as DSE, is recommended for patients with the above indications for a stress imaging test who are unable to exercise.	1	В
Stress echocardiography is the preferred test in patients with exertional dyspnea of uncertain etiology. In these patients, in addition to assessment of regional wall motion, tricuspid regurgitation velocity and diastolic function should be assessed at rest and with stress	I	В
An imaging stress test, such as stress echocardiography, should be considered in patients with prior coronary artery revascularization (PCI or CABG) and new cardiac symptoms.	lla	В
An imaging stress test, such as stress echocardiography, should be considered to assess the functional severity of intermediate lesions on coronary arteriography.	lla	В

regurgitation velocity and diastolic function should be assessed at rest and with stress		
An imaging stress test, such as stress echocardiography, should be considered in patients with prior coronary artery revascularization (PCI or CABG) and new cardiac symptoms.	lla	В
An imaging stress test, such as stress echocardiography, should be considered to assess the functional severity of intermediate lesions on coronary arteriography.	lla	В
Recommendations for risk stratification using ischemia testing	Class of recommendation	Level of evidence
A stress imaging test such as stress echocardiography for risk stratification is recommended in patients with an inconclusive exercise ECG	1	В
A stress imaging test, such as stress echocardiography, is recommended for risk stratification in patients with known stable CAD and a deterioration in symptoms if the site and extent of ischemia would influence clinical decision making	I	В
In asymptomatic adults with diabetes, peripheral vascular disease, or a strong family history of CAD, or when previous risk assessment testing suggests high risk of CAD, such as a coronary artery calcium score of ≥400, a stress imaging test, such as stress echocardiography, may be considered for advanced cardiovascular risk assessment. <sup>208</sup>	Ilb	В
Recommendation for re-assessment in patients with stable CAD	Class of recommendation	Level of evidence
Recommendation for re-assessment in patients with stable CAD  An exercise ECG or stress imaging test such as stress echocardiography is recommended in the presence of recurrent or new symptoms once instability has been ruled out.		
An exercise ECG or stress imaging test such as stress echocardiography is recommended in the		evidence
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An exercise ECG or stress imaging test such as stress echocardiography is recommended in the presence of recurrent or new symptoms once instability has been ruled out.  In symptomatic patients with revascularized stable CAD, a stress imaging test, such as stress echocardiography, is indicated rather than stress ECG.  Reassessment of prognosis using a stress test, such as stress echocardiography, may be considered in asymptomatic patients after the expiration of the period for which the previous	recommendation	evidence C
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\_\_\_



# Percutaneous coronary intervention in stable angina (ORBITA): a double-blind, randomised controlled trial



Rasha Al-Lamee, David Thompson, Hakim-Moulay Dehbi, Sayan Sen, Kare Tang, John Davies, Thomas Keeble, Michael Mielewczik,
Raffi Kaprielian, Iqbal S Malik, Sukhjinder S Nijjer, Ricardo Petraco, Christopher Cook, Yousif Ahmad, James Howard, Christopher Baker,
Andrew Sharp, Robert Gerber, Suneel Talwar, Ravi Assomull, Jamil Mayet, Roland Wensel, David Collier, Matthew Shun-Shin, Simon A Thom,
Justin E Davies, Darrel P Francis, on behalf of the ORBITA investigators\*

#### Summary

Background Symptomatic relief is the primary goal of percutaneous coronary intervention (PCI) in stable angina and is commonly observed clinically. However, there is no evidence from blinded, placebo-controlled randomised trials to show its efficacy.

Methods ORBITA is a blinded, multicentre randomised trial of PCI versus a placebo procedure for angina relief that was done at five study sites in the UK. We enrolled patients with severe (≥70%) single-vessel stenoses. After enrolment, patients received 6 weeks of medication optimisation. Patients then had pre-randomisation assessments with

#### Lancet 2018; 391: 31-40

Published Online November 2, 2017 http://dx.doi.org/10.1016/ 50140-6736(17)32714-9

This online publication has been corrected. The corrected version first appeared at the lancet.com

Interpretation In patients with medically treated angina and severe coronary stenosis, PCI did not increase exercise time by more than the effect of a placebo procedure. The efficacy of invasive procedures can be assessed with a placebo control, as is standard for pharmacotherapy.

Findings ORBITA enrolled 230 patients with ischaemic symptoms. After the medication optimisation phase and between Jan 6, 2014, and Aug 11, 2017, 200 patients underwent randomisation, with 105 patients assigned PCI and 95 assigned the placebo procedure. Lesions had mean area stenosis of 84·4% (SD 10·2), fractional flow reserve of 0·69 (0·16), and instantaneous wave-free ratio of 0·76 (0·22). There was no significant difference in the primary endpoint of exercise time increment between groups (PCI minus placebo 16·6 s, 95% CI =8·9 to 42·0, p=0·200). There were no deaths. Serious adverse events included four pressure-wire related complications in the placebo group, which required PCI, and five major bleeding events, including two in the PCI group and three in the placebo group.

Interpretation In patients with medically treated angina and severe coronary stenosis, PCI did not increase exercise time by more than the effect of a placebo procedure. The efficacy of invasive procedures can be assessed with a placebo control, as is standard for pharmacotherapy.

Funding NIHR Imperial Biomedical Research Centre, Foundation for Circulatory Health, Imperial College Healthcare Charity, Philips Volcano, NIHR Barts Biomedical Research Centre.

R Petraco MRCP, C Cook MRCP, Y Ahmad MRCP, J Howard MRCP, Prof J Mayet FRCP, M Shun-Shin MRCP, Prof S A Thorn FRCP, J E Davies MRCP, Prof D P Francis FRCP); Imperial College Healthcare NHS Trust, London, UK (R Al-Lamee, D'Thompson, S Sen, R Kaprielian FRCP, I S Malik FRCP, S S Niger MRCP, R Petraco, C Cook, Y Ahmad, J Howard, C Baker FRCP, R Assomuli MRCP, Prof J Mayet, M Shun-Shin, Prof SThorn, J.E. Davies, Prof D P Francis FRCP); Cancer

## <u>Circulation</u>

#### ORIGINAL RESEARCH ARTICLE

6

Dobutamine Stress Echocardiography Ischemia as a Predictor of the Placebo-Controlled Efficacy of Percutaneous Coronary Intervention in Stable Coronary Artery Disease

The Stress Echocardiography–Stratified Analysis of ORBITA

## **ORBITA** trial

(Objective Randomised Blinded Investigation With Optimal

Medical Therapy of Angioplasty in Stable Angina).

- ▶ 183 pazienti
- Randomizzato Doppio cieco
- Pz con angina stabile
- Lesione coronarica singola severa
- PCI contro terapia medica
- ► Il grado di ischemia all'ecostress predice l'efficacia della PTCA sulla frequenza di angina del paziente.



- Non vi è relazione significativa tra la frequenza di angina e la PTCA
- ▶ Il miglioramento sintomatico non è relazionato ai parametri invasivi.

# Orbita

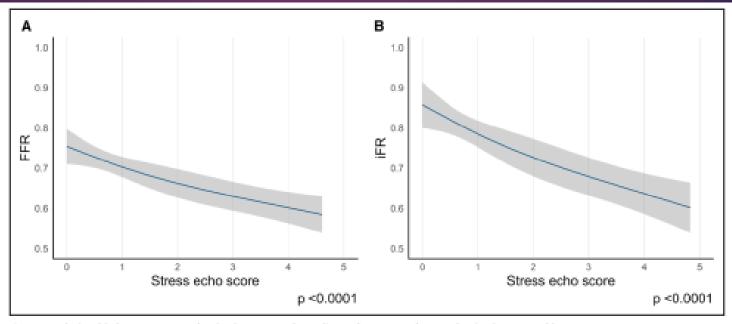
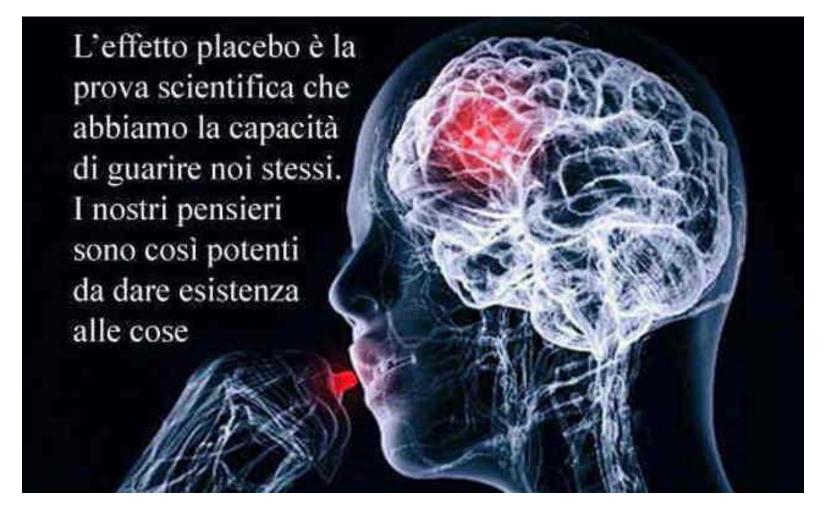


Figure 1. Relationship between prerandomization stress echocardiography score and prerandomization FFR and iFR.

A, Relationship between prerandomization stress echocardiography score and prerandomization FFR. B, Relationship between prerandomization stress echocardiography score and prerandomization iFR. echo indicates echocardiography; FFR, fractional flow reserve; and iFR, instantaneous wave-free ratio.

Tuttavia non vi è relazione significativa tra parametri invasivi e sintomi o tempo di esercizio





# Grado di ischemia-frequenza di angina

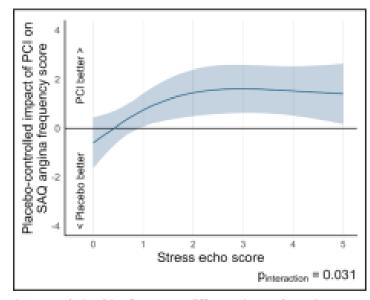


Figure 2. Relationship of treatment difference in Seattle Angina Questionnaire (SAQ) angina frequency score at follow-up to prerandomization stress echocardiography score by randomization arm.

There is a significant interaction between stress echocardiography score and Seattle Angina Frequency score with a progressive tendency for larger effects on angina frequency score with higher stress echocardiography score (P<sub>Interation</sub>=0.031), echo indicates echocardiography; and PCI, percutaneous coronary intervention.

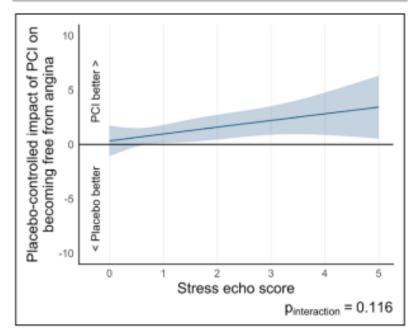


Figure 3. Relationship of treatment difference in freedom from angina and prerandomization stress echocardiography by randomization arm. There is no discernible dependence on prerandomization stress echocardiography score. echo indicates echocardiography; and PCI, percutaneous coronary intervention.

# Orbita

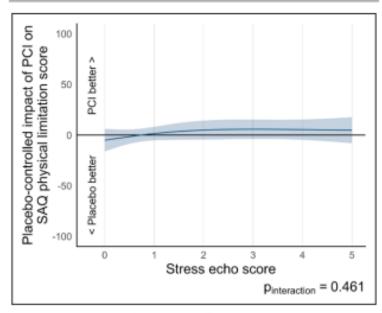


Figure 4. Relationship of treatment difference in Seattle Angina Questionnaire (SAQ) physical limitation score and prerandomization stress echocardiography by randomization arm.

There is no discernible dependence on prerandomization stress echocardiography score. echo indicates echocardiography; and PCI, percutaneous coronary intervention.

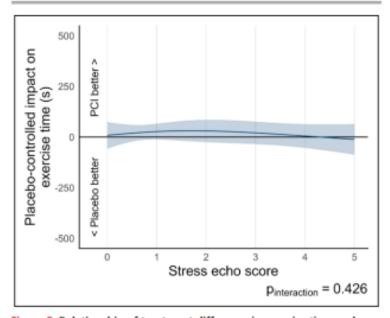


Figure 5. Relationship of treatment difference in exercise time and prerandomization stress echocardiography by randomization arm. There is no discernible dependence on prerandomization stress echocardiography score. echo indicates echocardiography; and PCI, percutaneous coronary intervention.

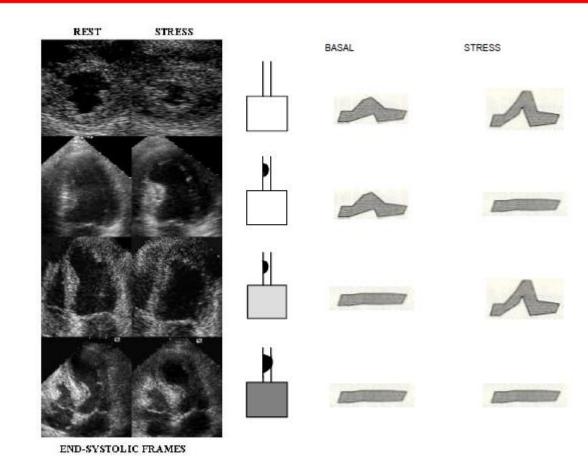
## LA DIAGNOSI

NORMALE

ISCHEMIA

VITALITA'

NECROSI





## Evidenziare/escludere anomalie della cinetica

#### **ESCURSIONE**

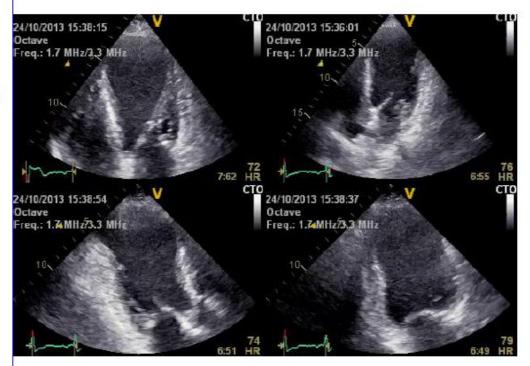
- . normale: normocinesia→1
- . ridotta: ipocinesia→2
- . assente: acinesia →3
- . espansione sistolica: discinesia→4
- . espansione fissa: aneurisma→5

#### SPESSORE

- . diminuito spessore diastolico
- . ridotto/assente ispessimento sistolico
- . assottigliamento sistolico

#### **ECOGENICITA**'

. Aumentata



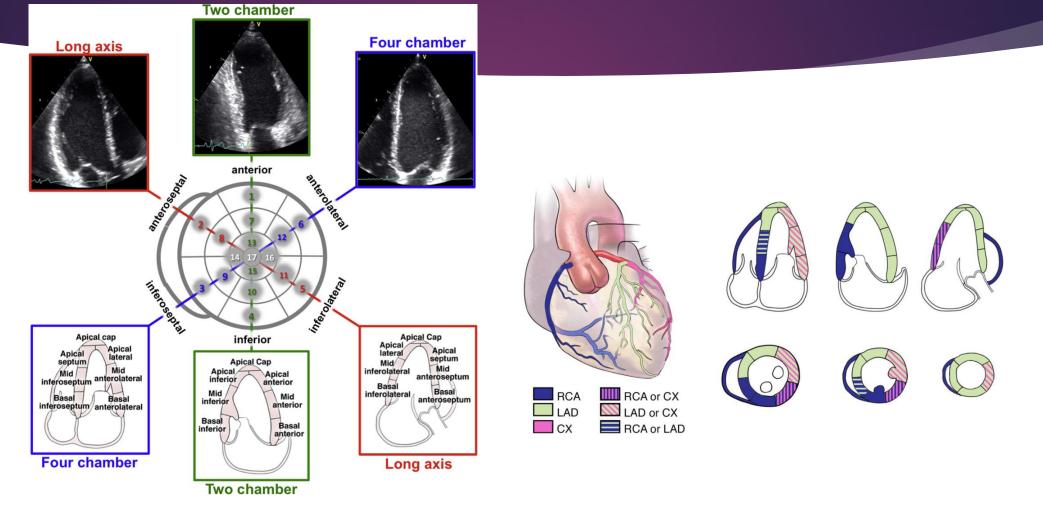
## Ecostress con dobutamina

- Sicuro
- Senza radiazioni ionizzanti
- Disponibile
- Basso costo
- Risultato immediato
- Accuratezza paragonabile a ecostress fisico e tomoscintigrafia miocardica
- Operatore dipendente
- ▶ Limitato da finestra acustica subottimale

# Dobutamina sconsigliata

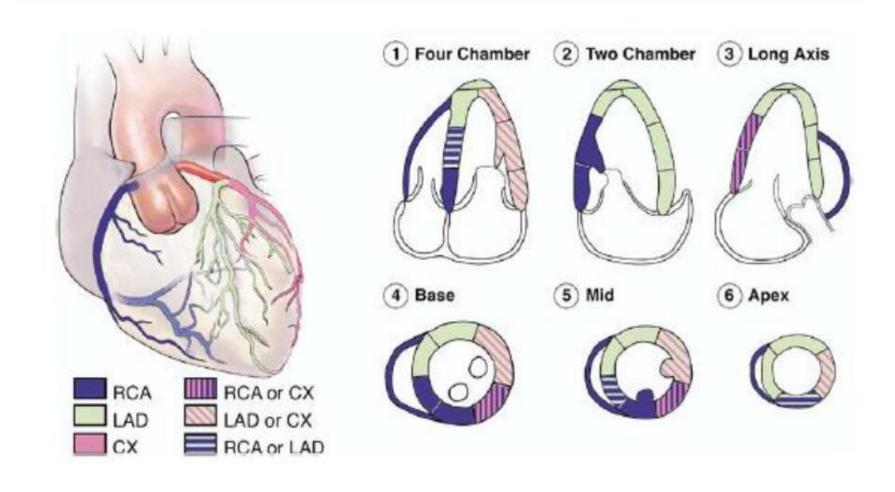
- Severa Ipertensione non controllata
- Aneurismi cerebrali noti
- ► Importanti aneurismi addominali

# Segmenti e territori di distribuzione coronarica

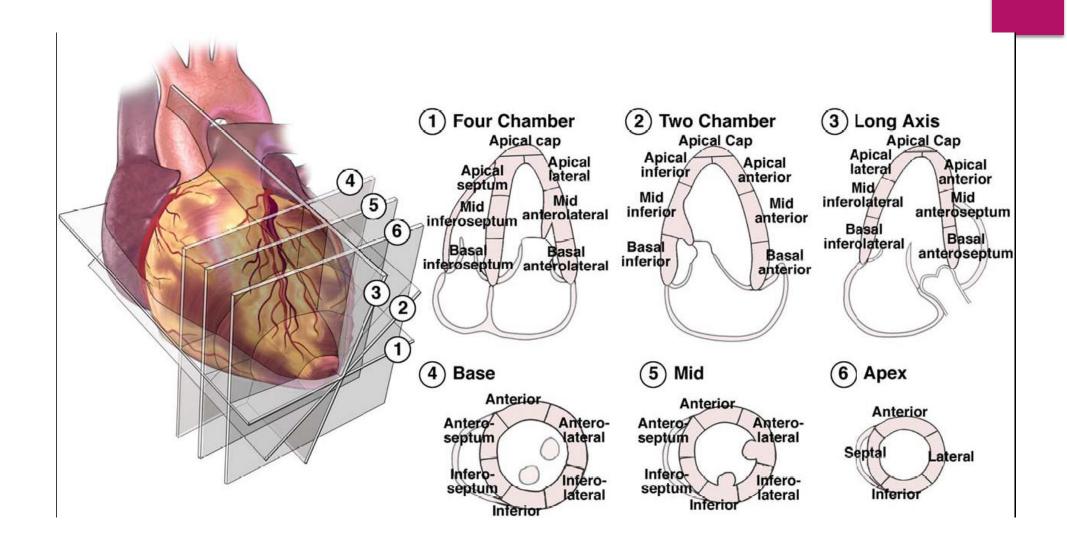


Settore Formazione 06/10/2021

# www.escardio.org ASE-EAE 2006: modello a 17 segmenti (16 utilizzati per lo studio della cinetica)



#### **ANALISI DELLA CINESI SEGMENTARIA-17 SEGMENTI**



Settore Formazione 06/10/2021







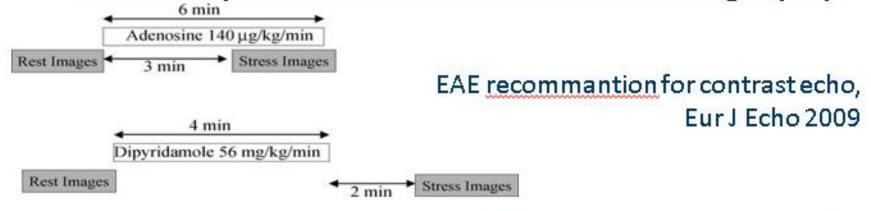
## Ecostress in CAD

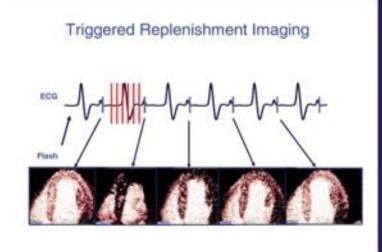
- Diagnosi di CAD
- ➤ Indicazione a PTCA
- Nel follow up
- Valutazione restenosi

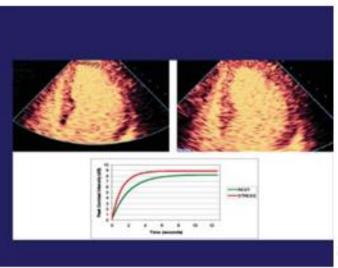




## Stress myocardial contrast echocardiography



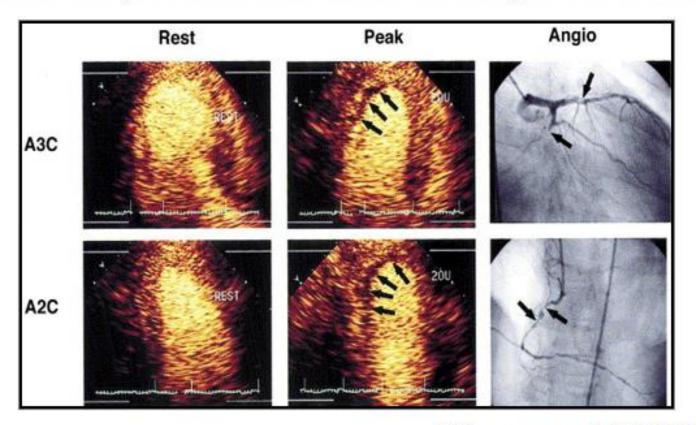








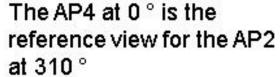
## Reversible perfusion defect during dobutamine



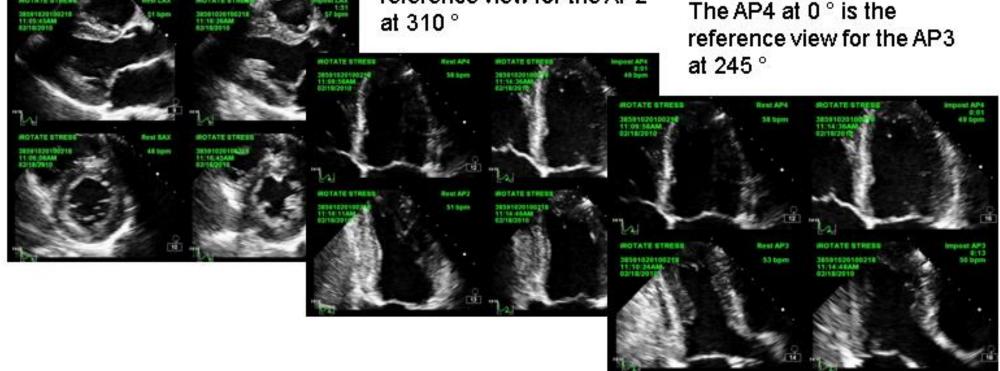
Dijkmans et al JACC 2006

## **IROTATE STRESS**

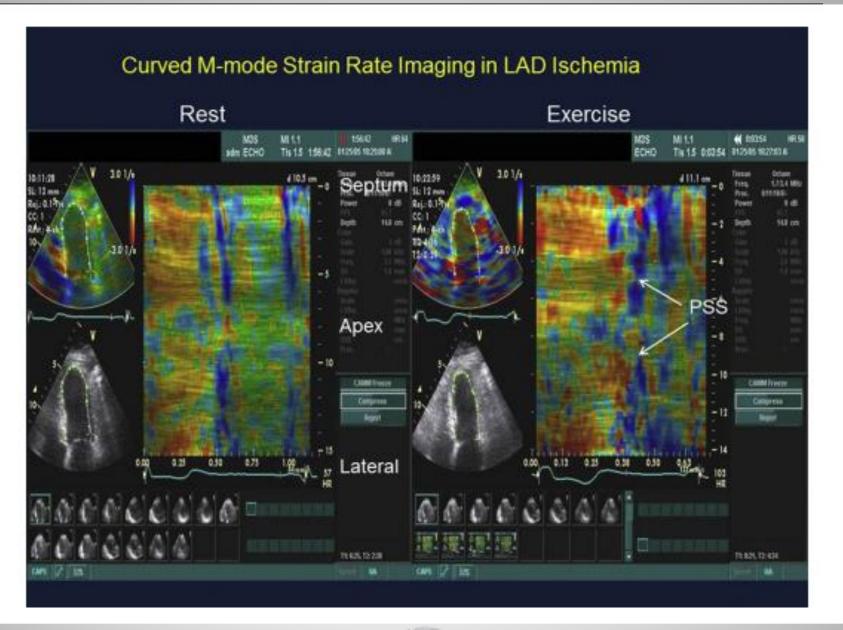
The LAX at 0° is the reference view for the SAX at 90°



The AP4 at 0° is the









### Polar Map Display of Peak Systolic Strain by Speckle Tracking Inferior, Apical Anterior Ischemia



#### Peak Dobutamine Stress



GLS -14.7

GLS - 9.1

