

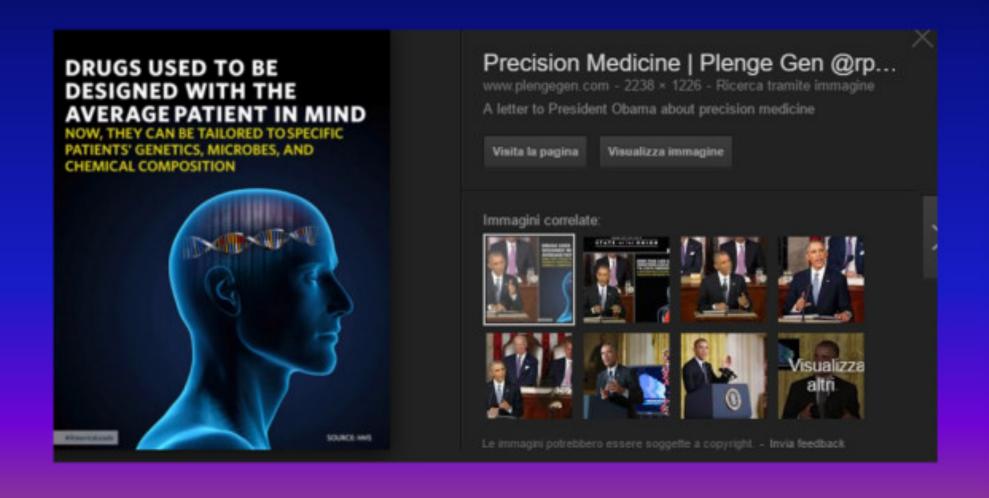


The adoption of the Framingham risk score was waived many years ago as an innovative population-based method to assess the median risk of developing CAD.

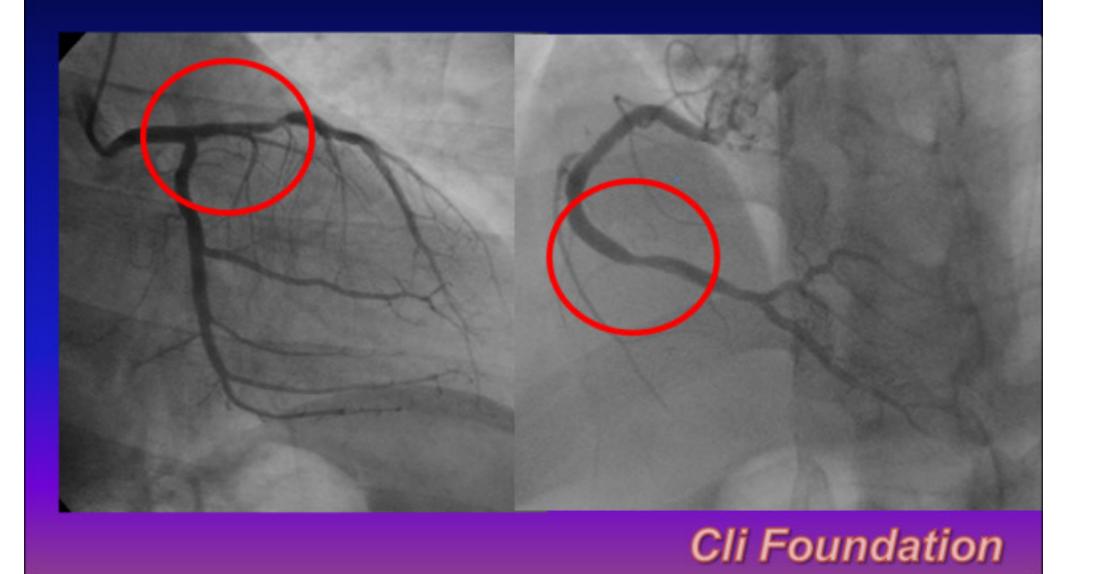
Inherent limitations

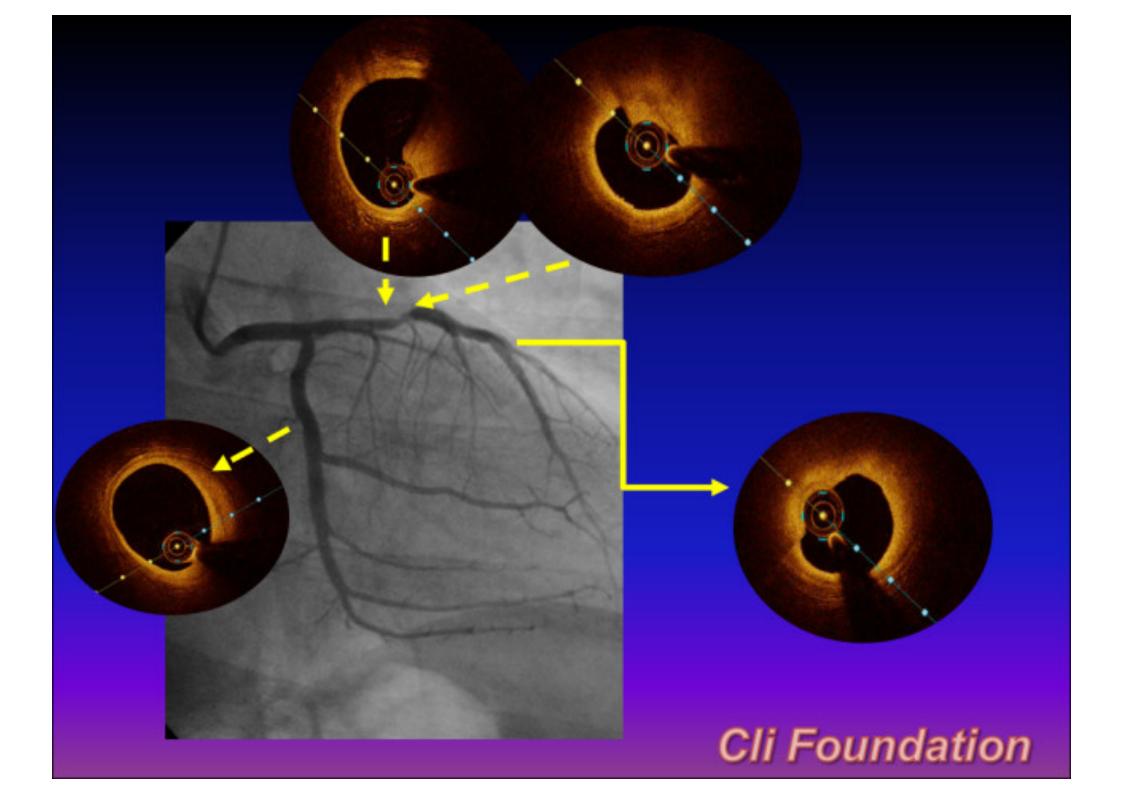
- ☐ Clinical scores do not allow discernment of the individual's risk
- ☐ Enable identification of a small increase in cardiovascular events (3- to 5-fold) in subject categories considered at risk.
- □ Evel when more liberal treatment criteria are applied, as suggested in the recent AFCAPS/TexCAPS trial, only 37% of the AMIs are prevented.

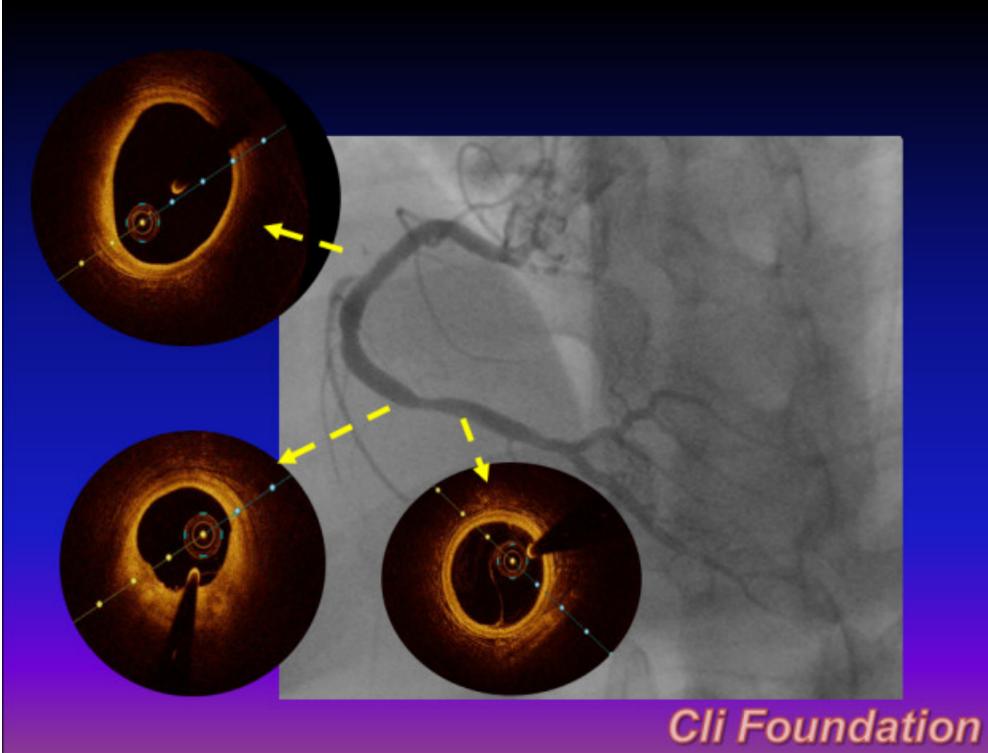
The search for patients at high risk after and ACS



Young 28 years old asiatic male Anterior NSTEMI Risk factor: mild smoher







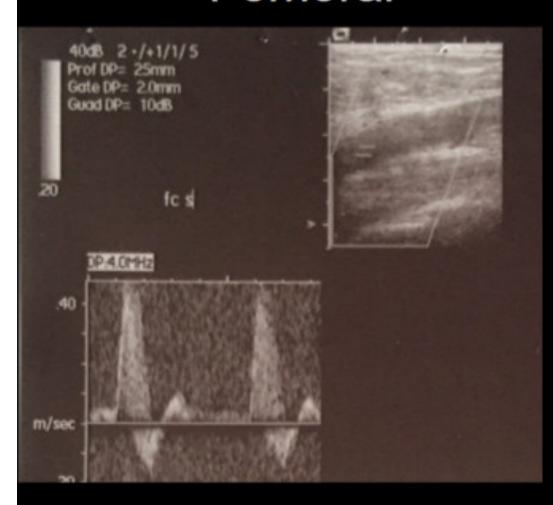
Sliding Doors

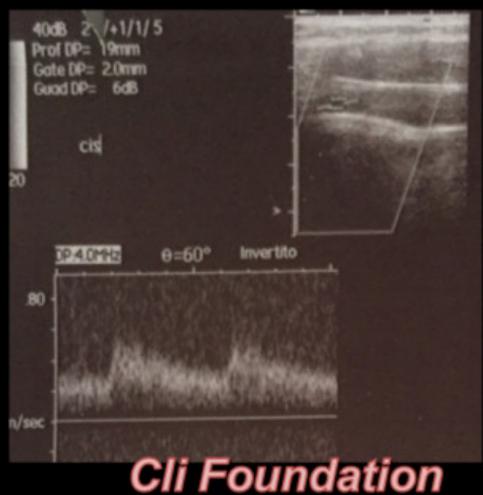


Undiseased carotid and femoral arteries at Echo-doppler

Femoral

Carotid



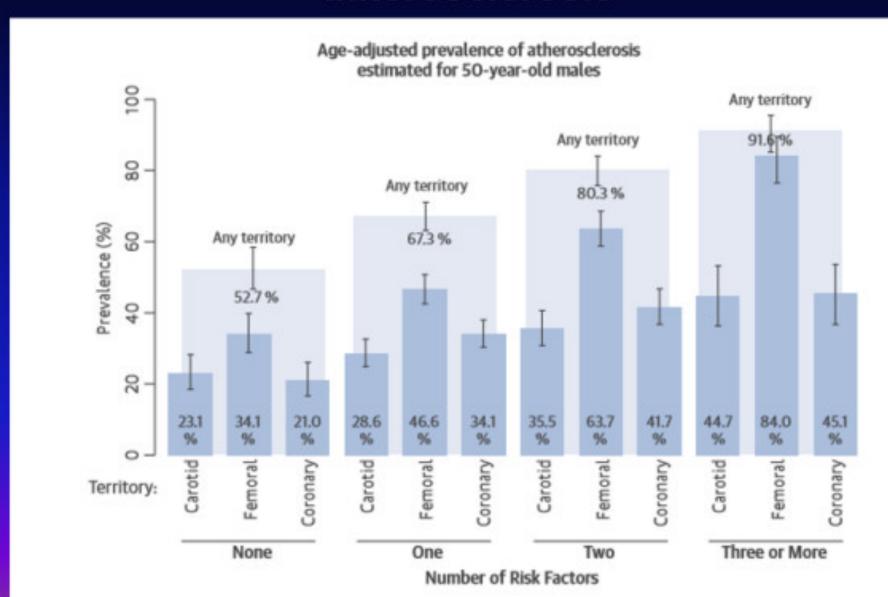


Femoral and subclinical carotid atherosclerosis association with risk factors and coronary calcium. The AWHS study

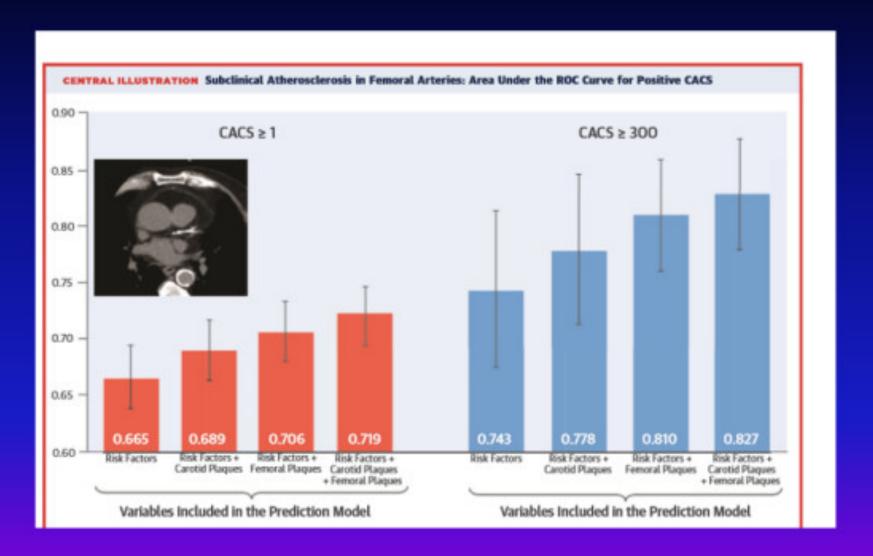
Laclaustra et al. JACC 2016

- Population: 1.423 Middle aged men
- Methods: Risk factors assessment plus Carotid, femoral ultrasound and coronary calcium score

Risk factors are better correlated with femoral atherosclerosis



Association with coronary calcium score stronger in femoral than carotid subclinical atherosclerosis

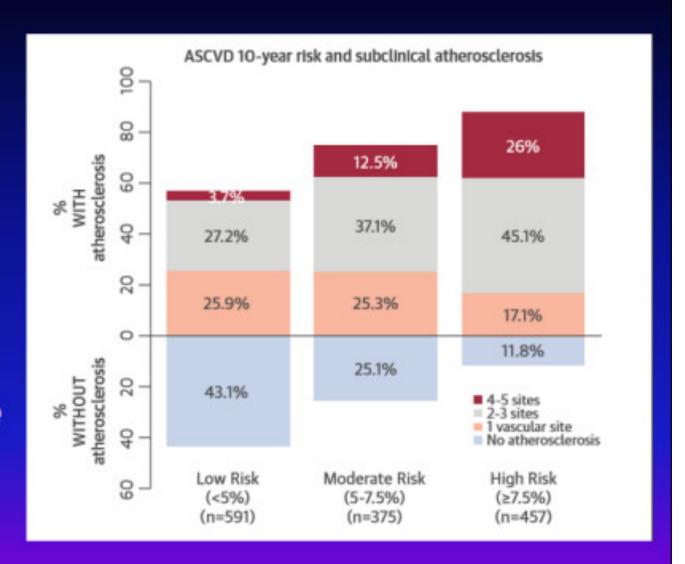


Femoral screening...... A new appealing strategy

Laclaustra et al. JACC 2016

67% of low risk patients have atherosclerosis

12 % of high risk patients don't have atherosclerosis

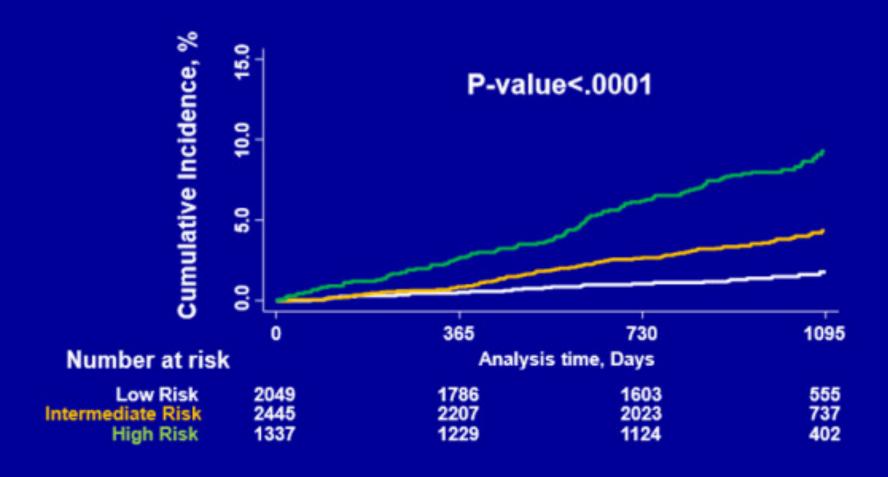


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Carotid Plaque Area Quartiles

U Baber, R Mehran, V Fuster et al, 2013

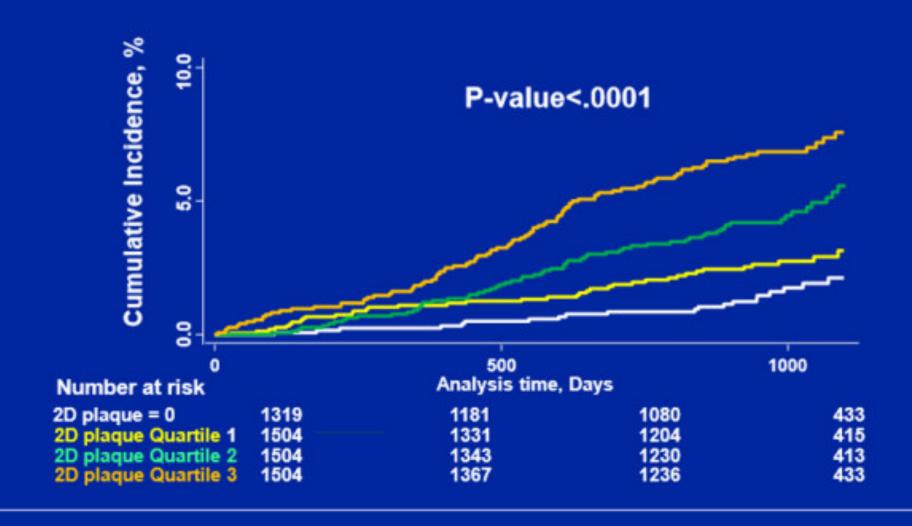
Cumulative MACE by Framingham Score



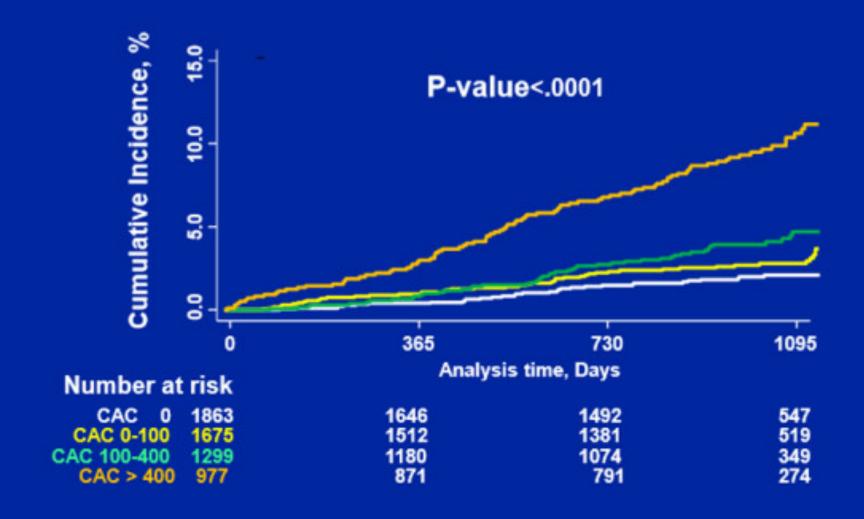




MACE by 2D US Carotid Plaque



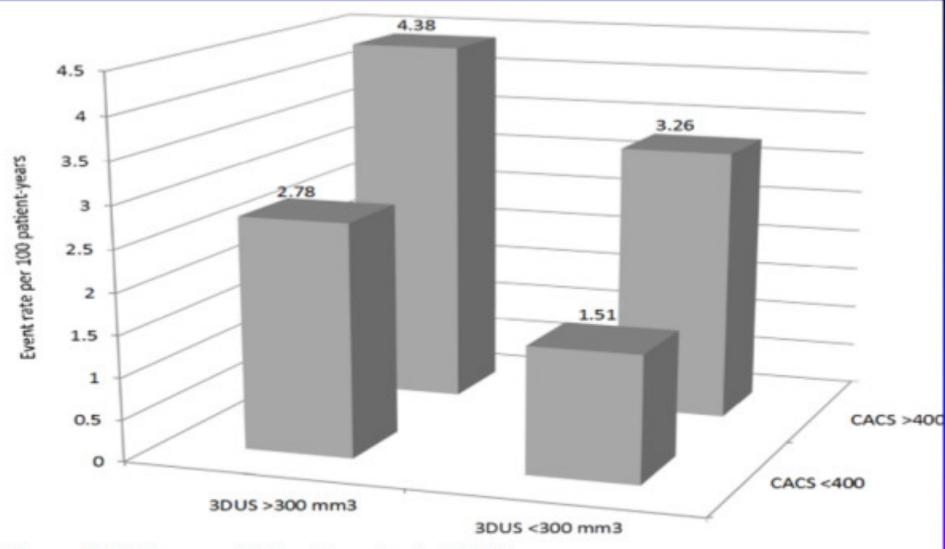
MACE by Coronary Calcium Score



U Baber, R Mehran, V Fuster et al, 2013

3D-US Transducer + CAC

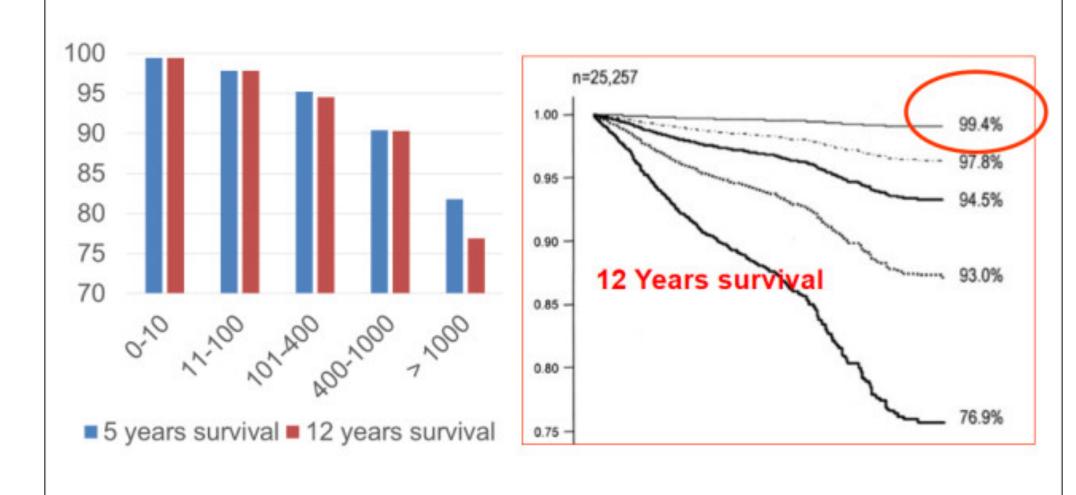
3D-US Transducer + CAC Impact on Events (Intermediate FRS Group)



U Baber, R Mehran, V Fuster et al, 2013

Hecht et al. JACC 2010

Survival according to calcium score



Treatment of Asymptomatic Adults With Elevated Coronary Calcium Scores With Atorvastatin, Vitamin C, and Vitamin E

The St. Francis Heart Study Randomized Clinical Trial

Yadon Arad, MD, FACC, Louise A. Spadaro, MD, FACC, Marguerite Roth, RN, David Newstein, DRPH, Alan D. Guerci, MD, FACC

Roslyn, New York

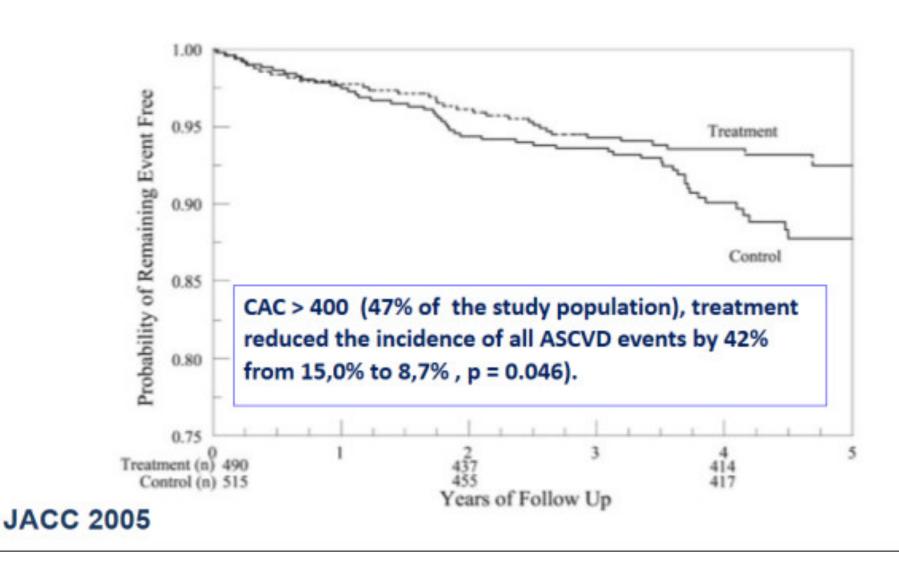
- Double-blind, placebo-controlled randomized clinical trial
- Pupulation. 1,005 asymptomatic subjects with coronary calcium scores > 80th percentile for age and gender.
- Treatment. torvastatin 20 mg daily, vitaminC1g daily, and vitamin E (alpha-tocopherol) 1,000 U daily, versus placebos



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Tailored therapy by means of Calcium Score assessment

Associations between C-reactive protein, coronary artery calcium, and cardiovascular events: implications for the JUPITER population from MESA, a population-based cohort study

Blaha et al. Lancet 2011

Population: 950 participants eligible for JUPITER. 5.8 y FU. Role of CAC for stratifying risk

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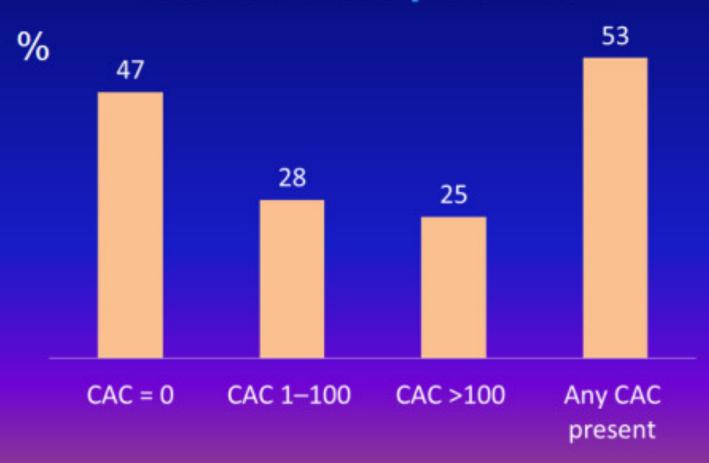
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The MESA study a population-based cohort study

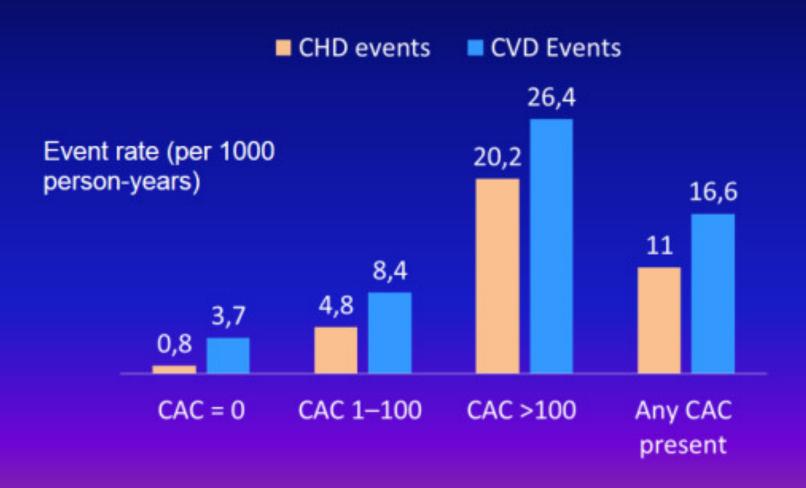
Blaha et al. Lancet 2011

Calcium score prevalence



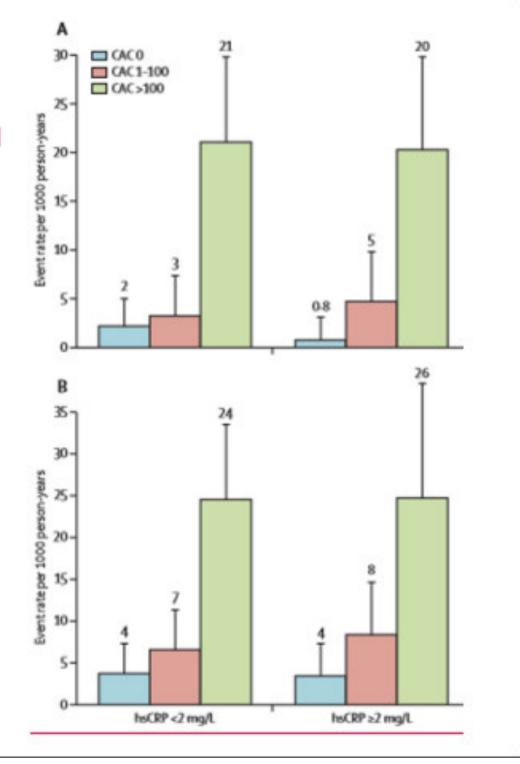
The MESA study a population-based cohort study

Blaha et al. Lancet 2011



Associations between C-reactive protein, coronary artery calcium, and cardiovascular events: implications for the JUPITER population from MESA, a population-based cohort study

CAC seems to further stratify risk in patients eligible for J CAC can target subgroups of patients to be trated with statin



Lancet 2011

Statins for all?

Mesa trial (LDL-C <130 mg/dL and hsCRP ≥2 mg/L)

- CAC = 0
 NNT to prevent one cardio-vascular disease in 5 years with atorvastatin
- CAC > 100
 NNT to prevent one cardio-vascular disease in 5 years with atorvastatin
 24

Aspirine to whom? Mesa trial (LDL-C <130 mg/dL and hsCRP ≥2 mg/L)

- Pts with elevated CAC scores (>100) were 2-4 times more likely to benefit from aspirin therapy than to be harmed, even if they did not qualify for aspirin use according to current AHA guidelines.
- Conversely, pts with no calcified plaque (CAC score = 0) were 2-4 times more likely to be harmed by aspirin use than to benefit.

Secondary Prevention



Patient Selection



History of cardiovascular disease (MI or ishemic stroke or diagnosis of nonhemorrhagic stroke or symptomatic PAD.

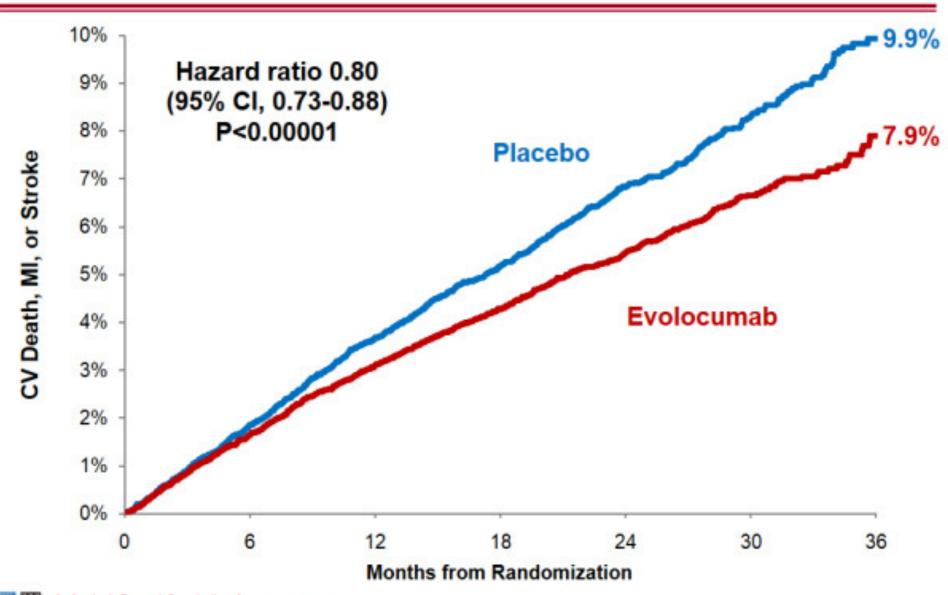
At least 1 Major risk factors : diabetes, age ≥ 65y, MI or nonhemorrhagic stroke, cigarette smoking, symptomatic PAD

Minor risk factors (2 required): • history of non–MIrelated coronary revascularization • residual coronary
artery disease with ≥40% stenosis in ≥2 large vessels,
low HDL, high CRP, LDL-C ≥130 mg/dL (3.4 mmol/L) or
non-HDL-C ≥160 mg/dL (4.1 mmol/L).



Key Secondary Endpoint









Key Inclusion & Exclusion Criteria



KEY INCLUSION

- Age ≥50 years
- At least 1 of the following:
 - Age ≥65 years
 - Diabetes requiring medication
 - 2nd prior MI (>1 year ago)
 - Multivessel CAD
 - CrCl <60 mL/min
- Tolerating ASA and able to be dosed at 75-150 mg/d

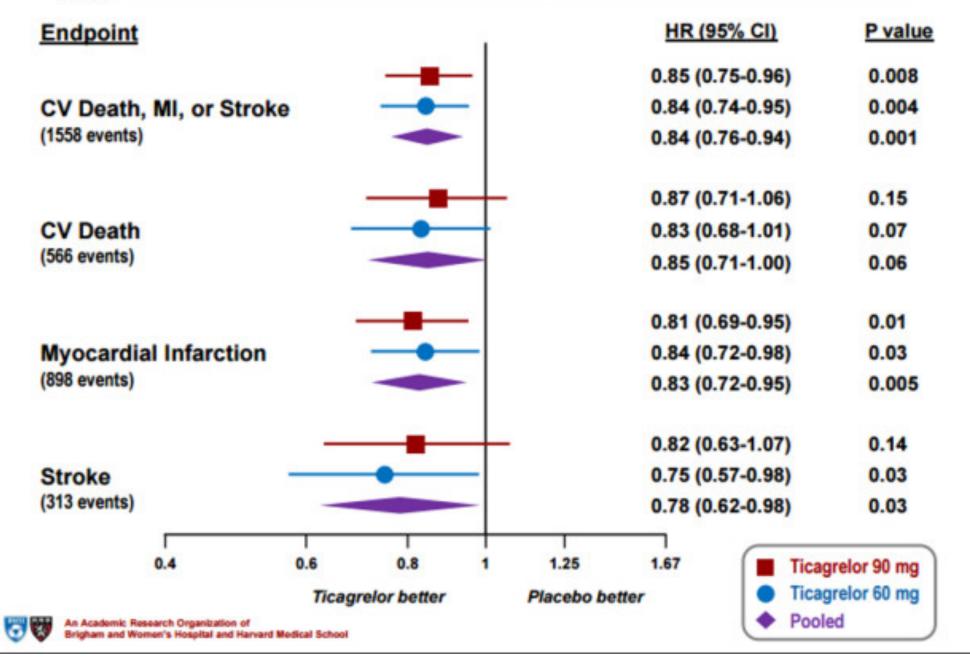
KEY EXCLUSION

- Planned use of P2Y₁₂ antagonist, dipyridamole, cilostazol, or anticoag
- Bleeding disorder
- History of ischemic stroke, ICH, CNS tumor or vascular abnormality
- Recent GI bleed or major surgery
- At risk for bradycardia
- Dialysis or severe liver disease



Components of Primary Endpoint



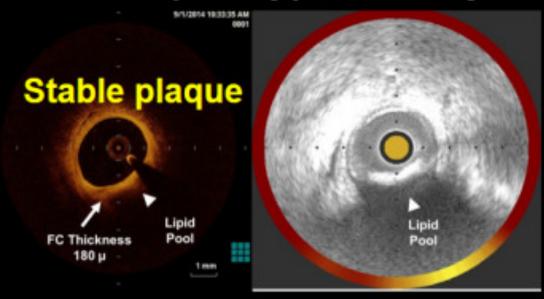




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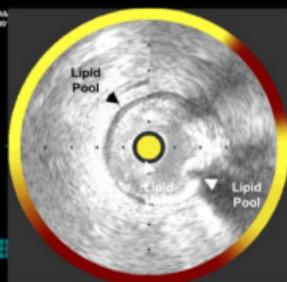
N 99 lesions studied with IVUS-NIRS and OCT

Culprit lesions of ACS pts have small LA (< 4 mm², thin FC (< 80 μ), and superficial inflammation



Submitted Prati et al.



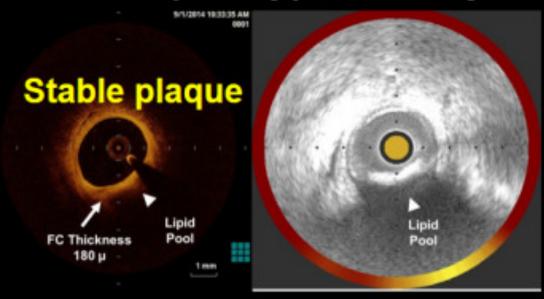




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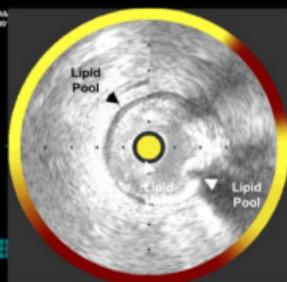
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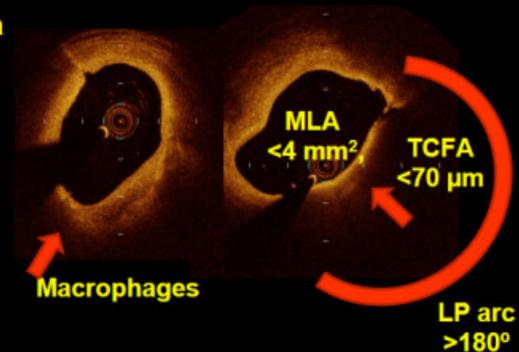
ClinicalTrials.Gov

N 700 pts

OCT vulnerable plaque criteria

Lesion presenting all of the following four features:

- TCFA <65 µm
- Lipid pool arc >180°
- MLA <4.0 mm²
- Macrophages



Clinical outcome

Composite of cardiac death and/or target vessel MI (STEMI/NSTEMI)



Take Home Message

- To discern the individual's risk clinical risk scores should be integrated with the search of arterial atherosclerosis
- Among imaging modalities to detect atherosclerosis,
 the coronary calcium score has an established role
- Absence of detectable calcium score conveys an excellent long term outcome with a ten year risk of cardiovascular events less than 1 / 1000
- For secondary prevention IC imaging modalities potentially better identify patients with a more aggressive atherosclerosis