



LA THROMBOSE DE STENT EST-ELLE TOUJOURS D'ACTUALITE ?

Les thromboses de stent sont-elles toutes identiques ?

Mercredi 06 juin 2012
14h40 – 15h20



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déclare n'avoir aucun conflit d'intérêt concernant les données de sa communication

Definition of Stent Thrombosis

Cutlip D et al. *Circulation* 2007, in press

- **Definite Stent Thrombosis**

- Angiographic or pathologic confirmation of partial or total thrombotic occlusion within the peri-stent region

AND at least ONE of the following, additional criteria:

- Acute ischemic symptoms
 - Ischemic ECG changes
 - Elevated cardiac biomarkers

- **Probable Stent Thrombosis**

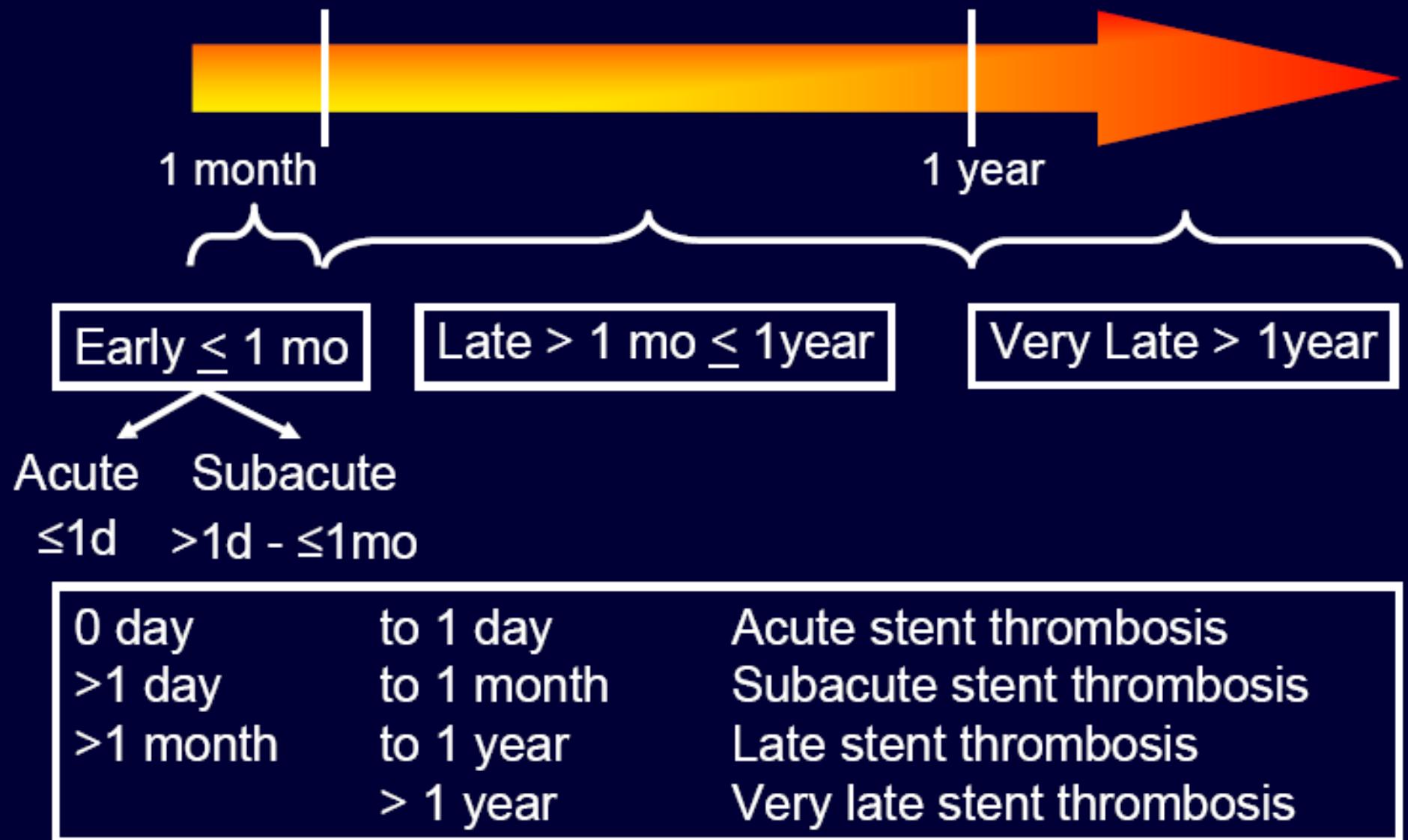
- Any unexplained death within 30 days of stent implantation
 - Any myocardial infarction, which is related to documented acute ischemia in the territory of the implanted stent without angiographic confirmation of stent thrombosis and in the absence of any other obvious cause

- **Possible Stent Thrombosis**

- Any unexplained death beyond 30 days

Time Frame of Stent Thrombosis

Cutlip D et al. *Circulation* 2007, in press

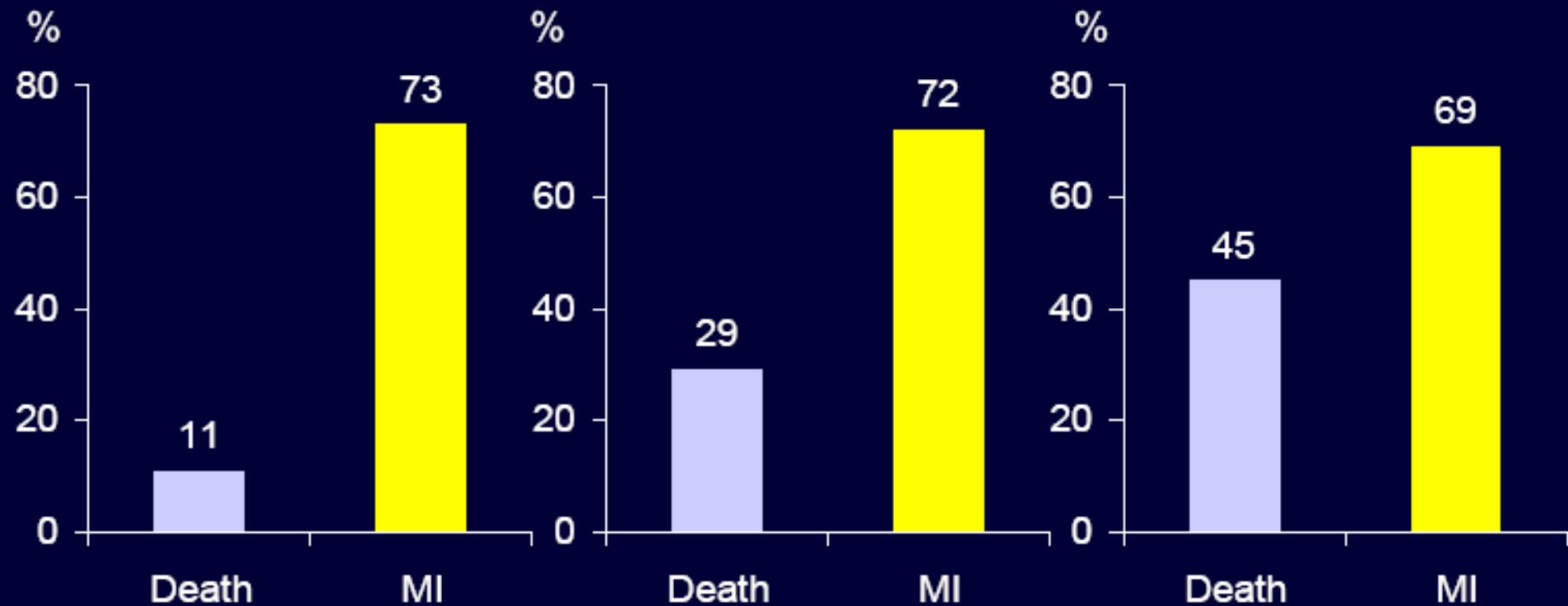


Outcome of Patients With DES Stent Thrombosis at 6 – 9 Months

Definite ST
Bern/Rotterdam (N=8146)
Daemen et al. *Lancet* 2007

Definite ST
Washington (N=2974)
Kuchulakanti et al. *Circulation* 2006

Definite, Probable,
and Possible ST
Milan/Siegburg (N=2229)
Iakovou et al. *JAMA* 2005



Multifactorial Nature of Stent Thrombosis

Stent factors

- Surface
- Drugs
- Polymer

Procedural factors

- Dissection
- Incomplete stent apposition

Lesion factors

- Vessel size/length
- Thrombus
- Plaque characteristics
- Bifurcation

STENT THROMBOSIS

Blood factors

- Coagulation activity
- Platelet inhibition

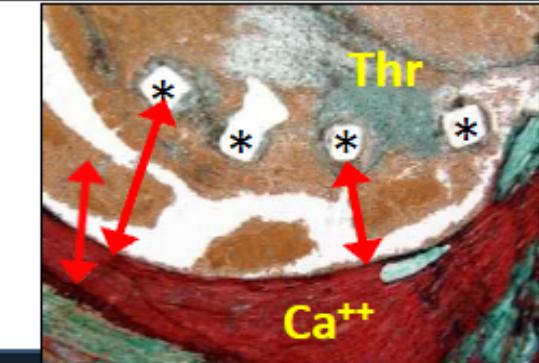
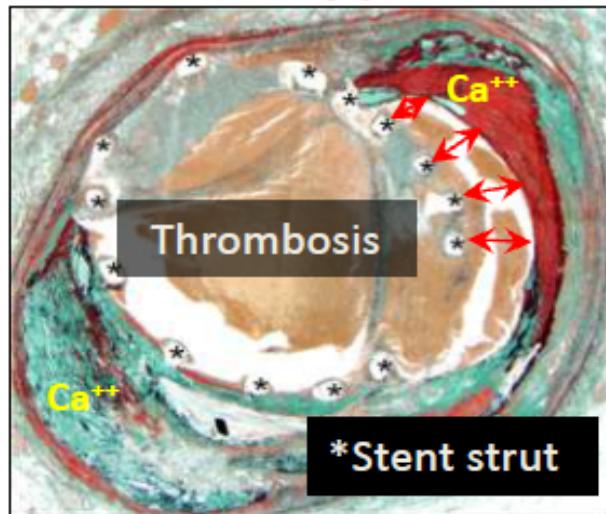
Antithrombotic and anticoagulation therapy

Patient factors

- Drug response/interactions
- Gene polymorphism
- LV function
- ACS
- Renal failure
- Diabetes

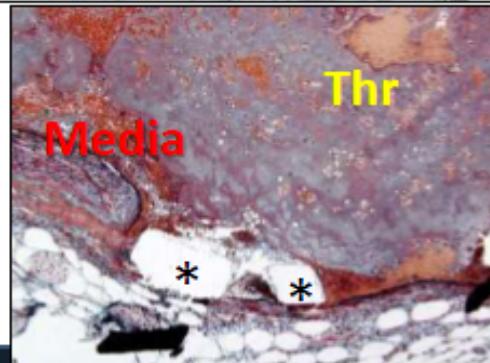
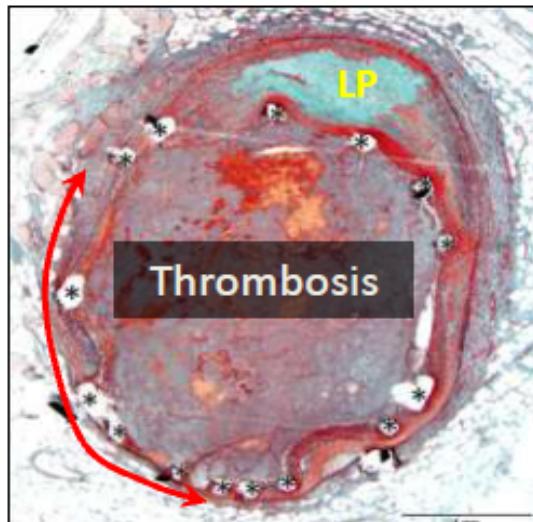
Speculative Predictors of Early ST in Histology

Strut malapposition



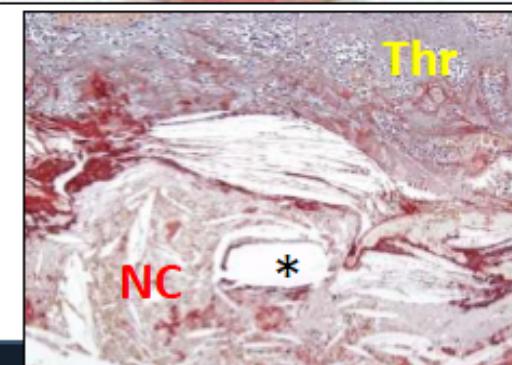
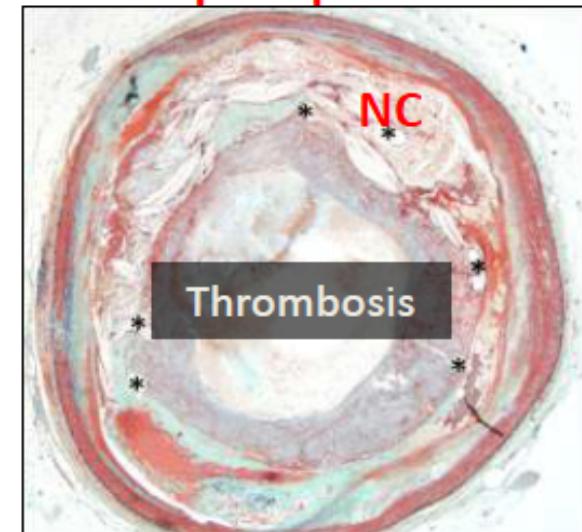
BMS implanted in a heavily calcified plaque (fibrocalcific). Struts are partially malapposed.

Media disruption



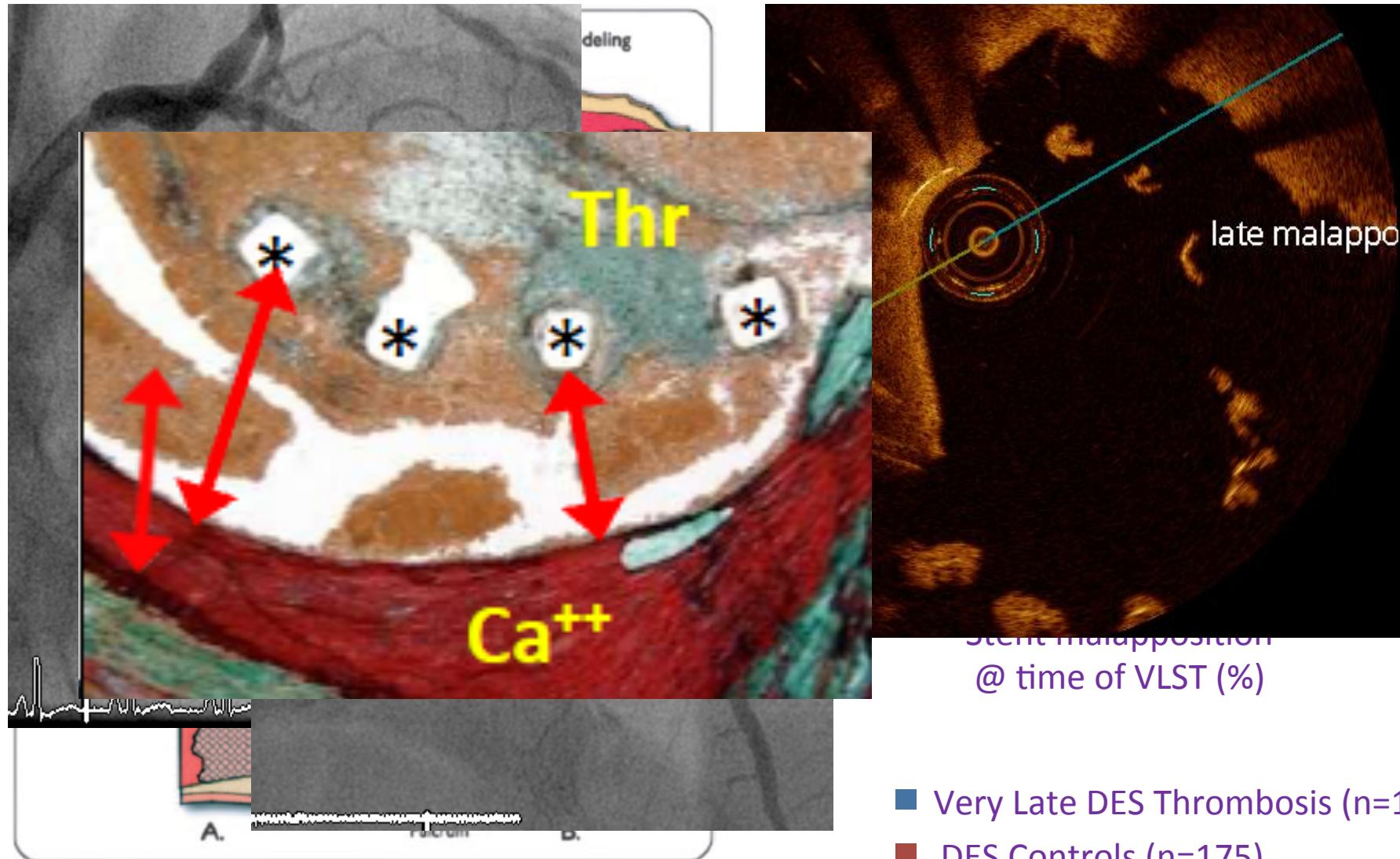
DES implanted in an eccentric pathological intimal thickening plaque (PIT). Media is tore at the opposite side of plaque.

Necrotic core prolapse

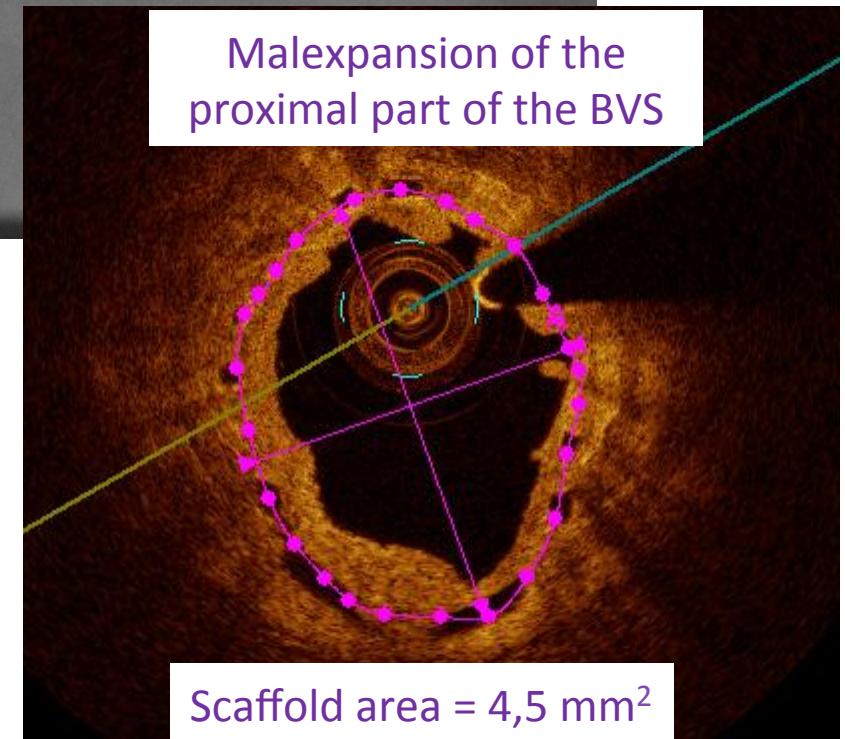
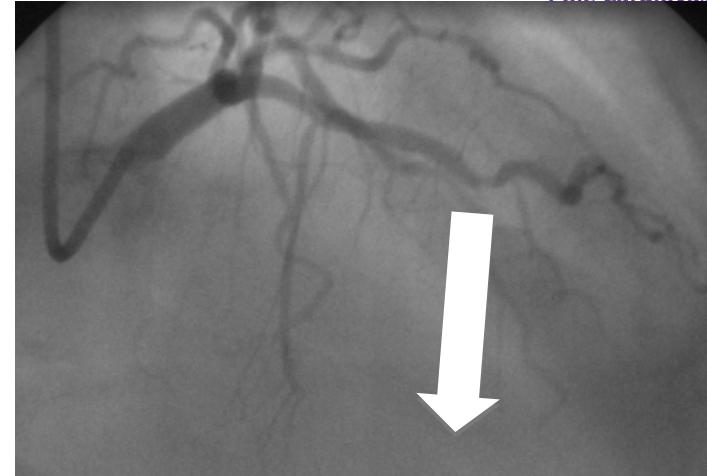
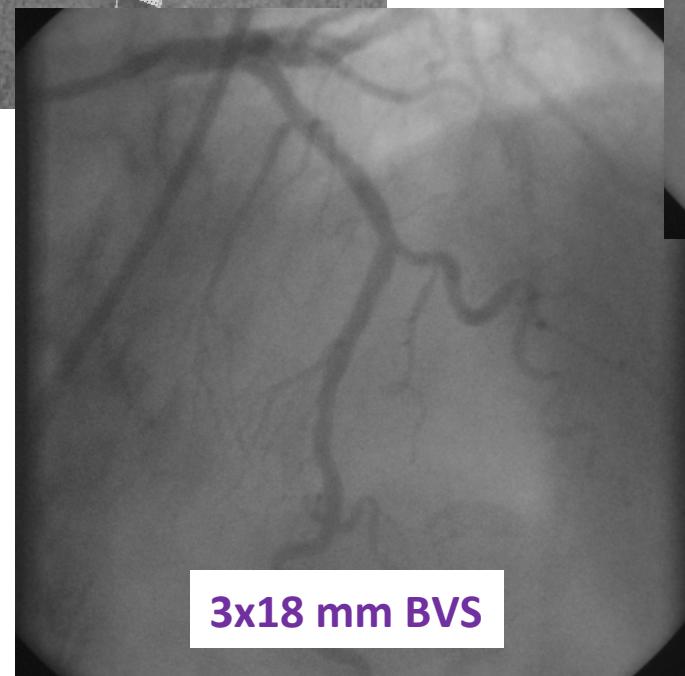


BMS implanted in ruptured plaque. Struts are penetrating into necrotic core (NC).

Causes mécaniques, évitables ?



Causes mécaniques, évitables ?



Les TS sont-elles toutes identiques ?



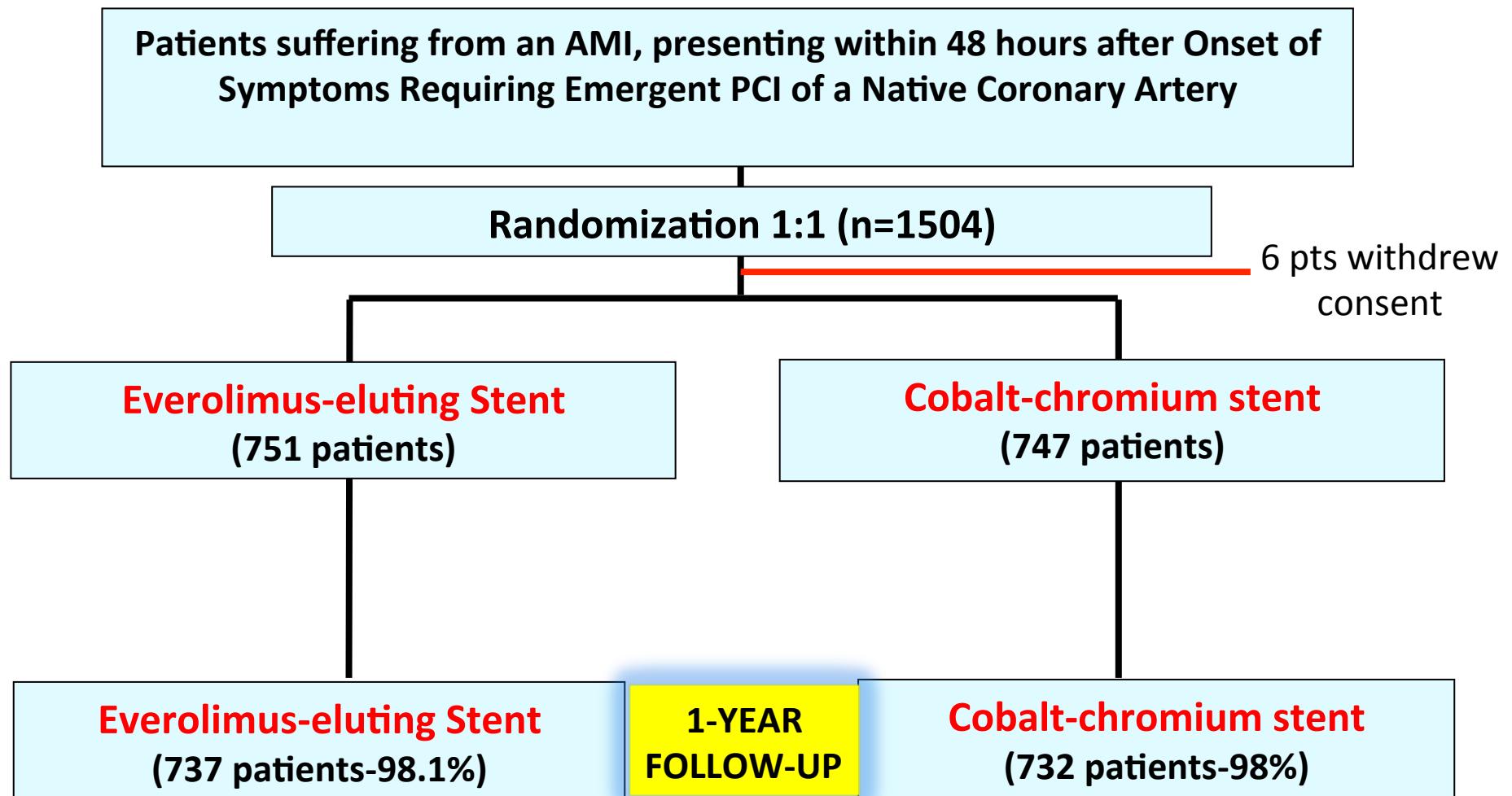
- Il existe bien certains facteurs, pour certains évitables (mécaniques, pharmacologiques) permettant d'individualiser certaines TS.
- Une fois éliminés ces différents facteurs, y-a-t-il des différences de TS entre les différents stents à notre disposition, utilisés à large échelle ?
- BMS vs DES 1^{ère} vs DES 2^{ème} génération ?

EXAMINATION trial

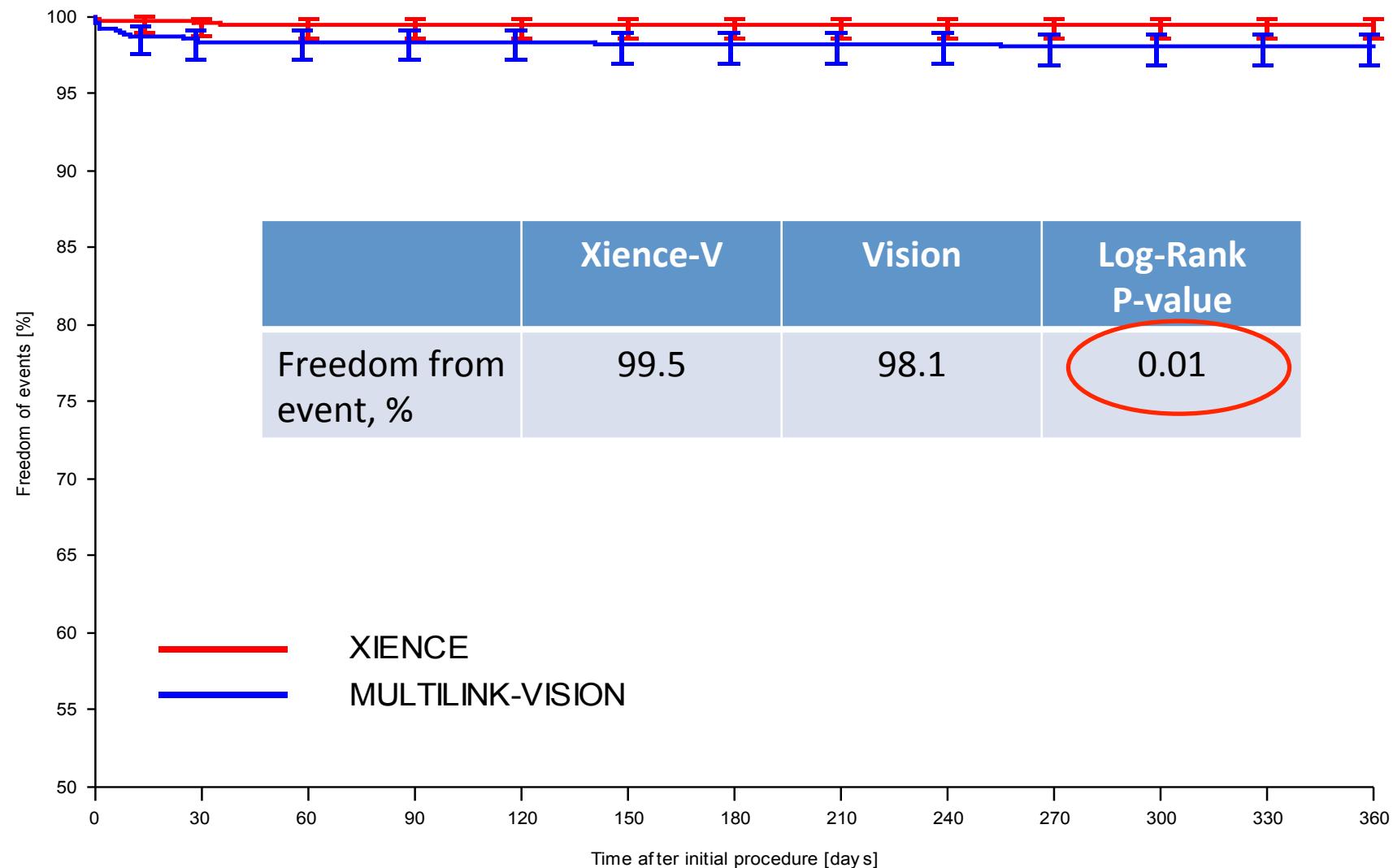
*The EXAMINATION (a clinical
Evaluation of Xience-V stent in
Acute Myocardial INfArcTION)
trial*

*Manel Sabaté
Hospital Clínic, Barcelona
(On behalf of the Examination Investigators)*

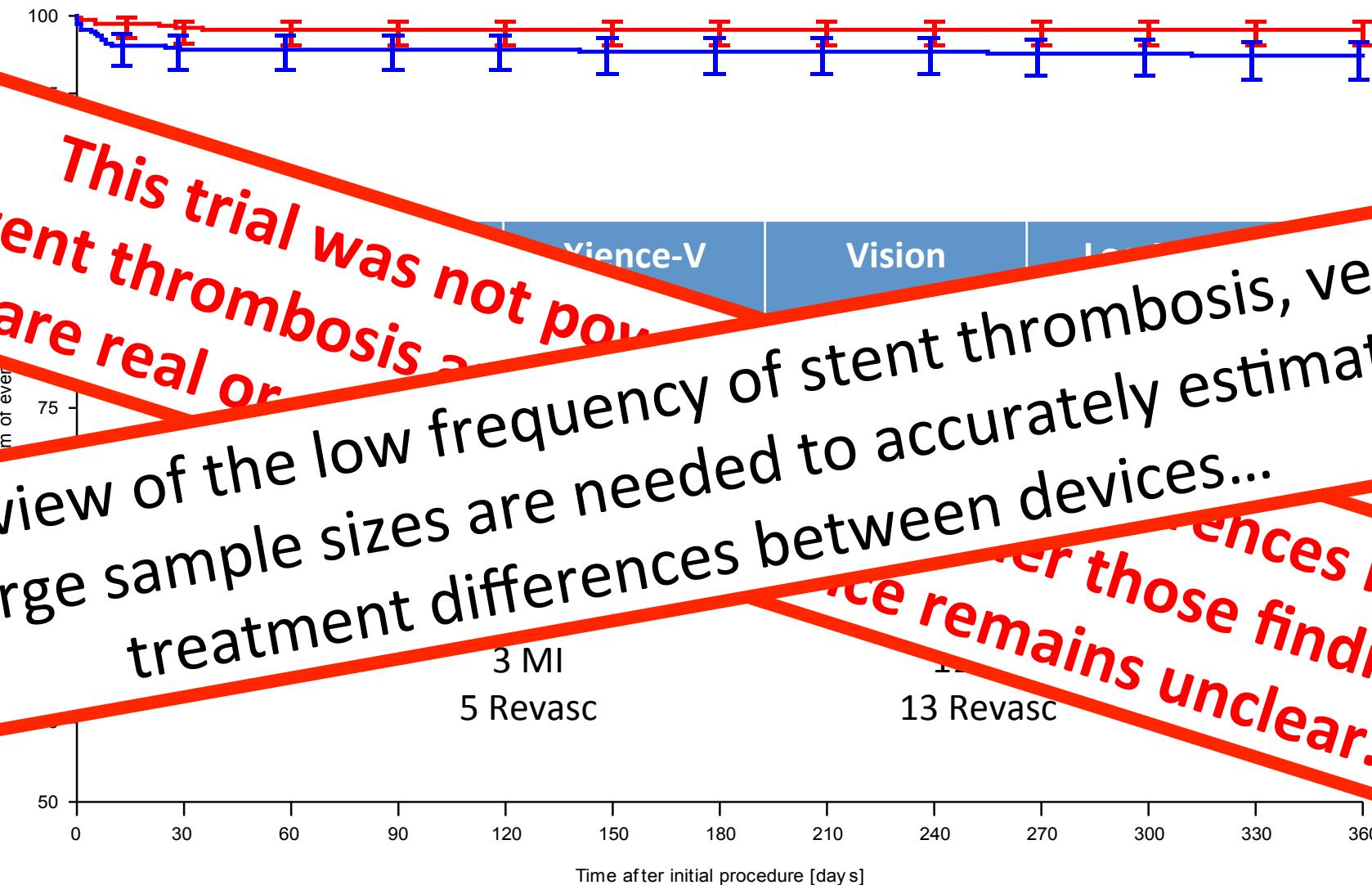
Study Design = All-comer RCT



Secondary Endpoints: Definite Stent Thrombosis



Secondary Endpoints: Definite/Probable Stent Thrombosis



Stent thrombosis with drug-eluting and bare-metal stents: evidence from a comprehensive network meta-analysis



Tullio Palmerini, Giuseppe Biondi-Zoccai, Diego Della Riva, Christoph Stettler, Diego Sangiorgi, Fabrizio D'Ascenzo, Takeshi Kimura, Carlo Briguori, Manel Sabaté, Hyo-Soo Kim, Antoinette De Waha, Elvin Kedhi, Pieter C Smits, Christoph Kaiser, Gennaro Sardella, Antonino Marullo, Ajay J Kirtane, Martin B Leon, Gregg W Stone

Palmerini T et al. *Lancet* 2012:On-line

DOI:10.1016/S0140-6736(12)60324-9

Stent Thrombosis Network Meta-analysis

Primary EP: ARC Definite ST (FU to 2 years)

2602 potentially relevant articles

Review of title
and abstract

2441 excluded
2117 not a comparison between DES
324 post-hoc, subgroup, follow-up, or
pooled analyses of included trials

161 articles needing full review

Full-text
review

112 excluded
84 not an RCT
13 DES not FDA approved
11 no ARC definition
4 DES pooled

49 articles meeting criteria

FDA
approved
stents
(BMS, SES, PES,
End-ZES, Res-ZES,
CoCr-EES, PtCr-EES)

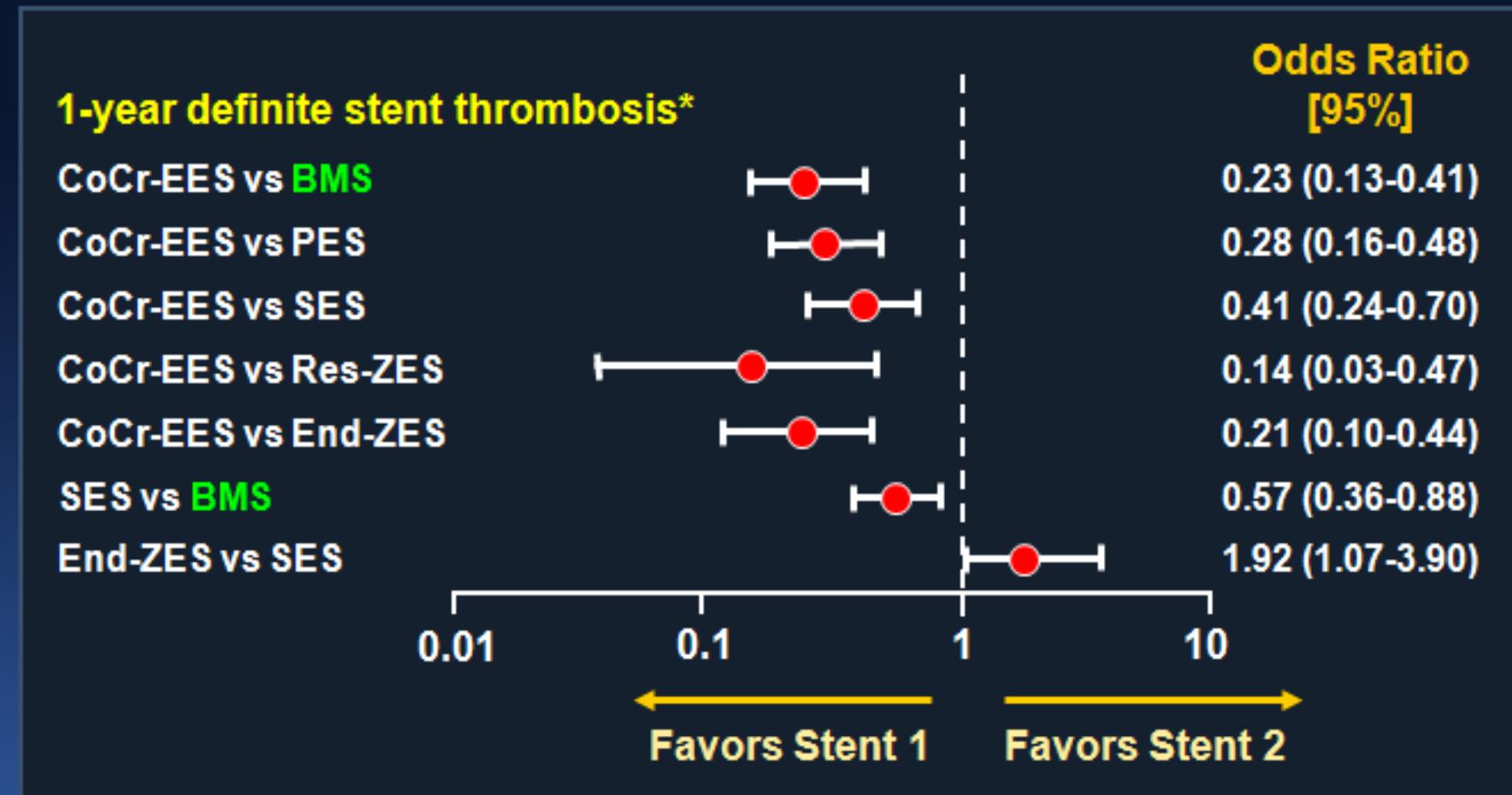
49 RCTs
50,844 pts

Palmerini T et al. *Lancet* 2012:On-line

Stent Thrombosis Network Meta-analysis

Primary EP: ARC Definite ST (FU through 2 years)

49 RCTs, 50,844 pts



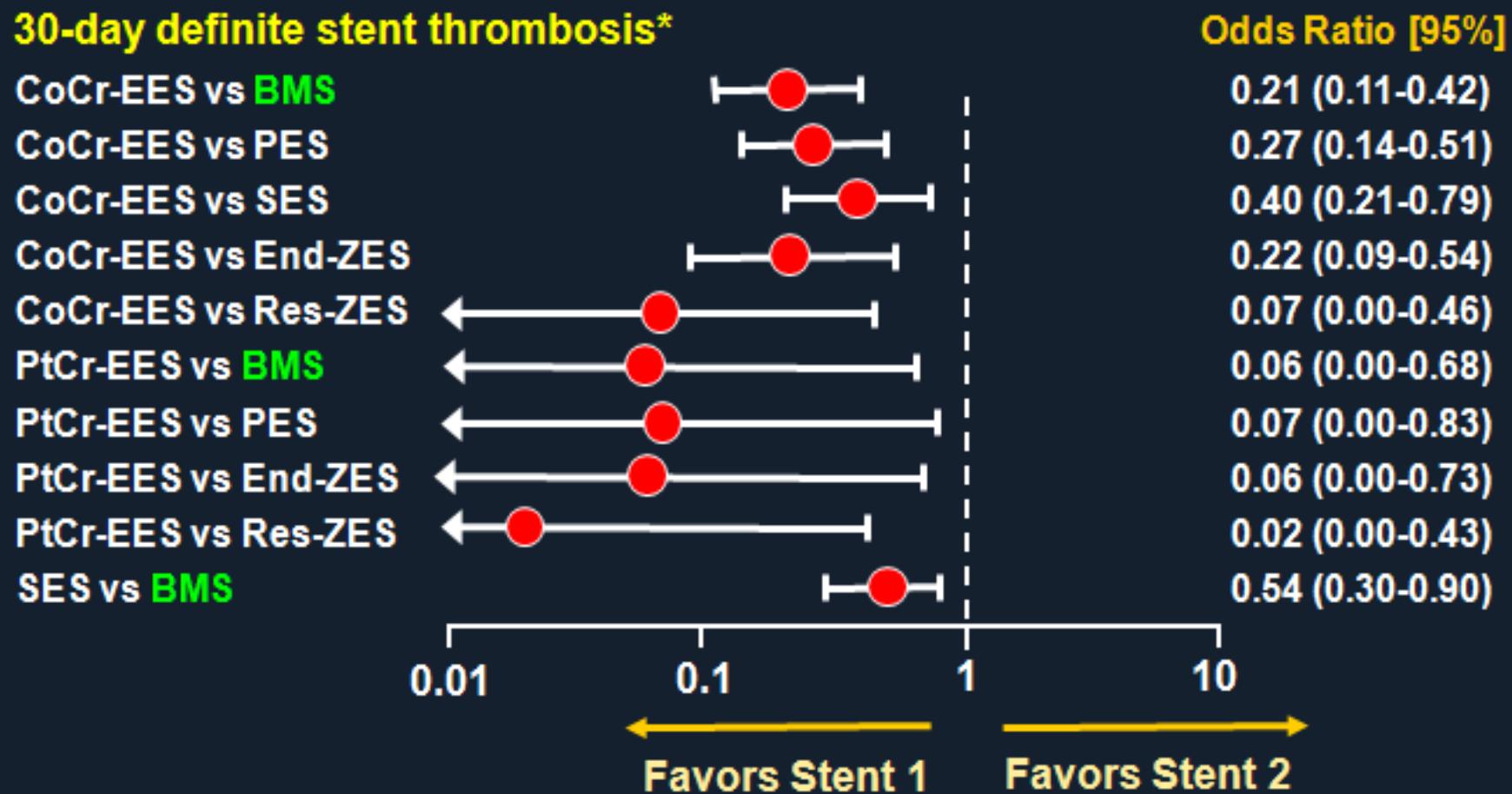
*Only statistically significant results are shown

Palmerini T et al. Lancet 2012:On-line

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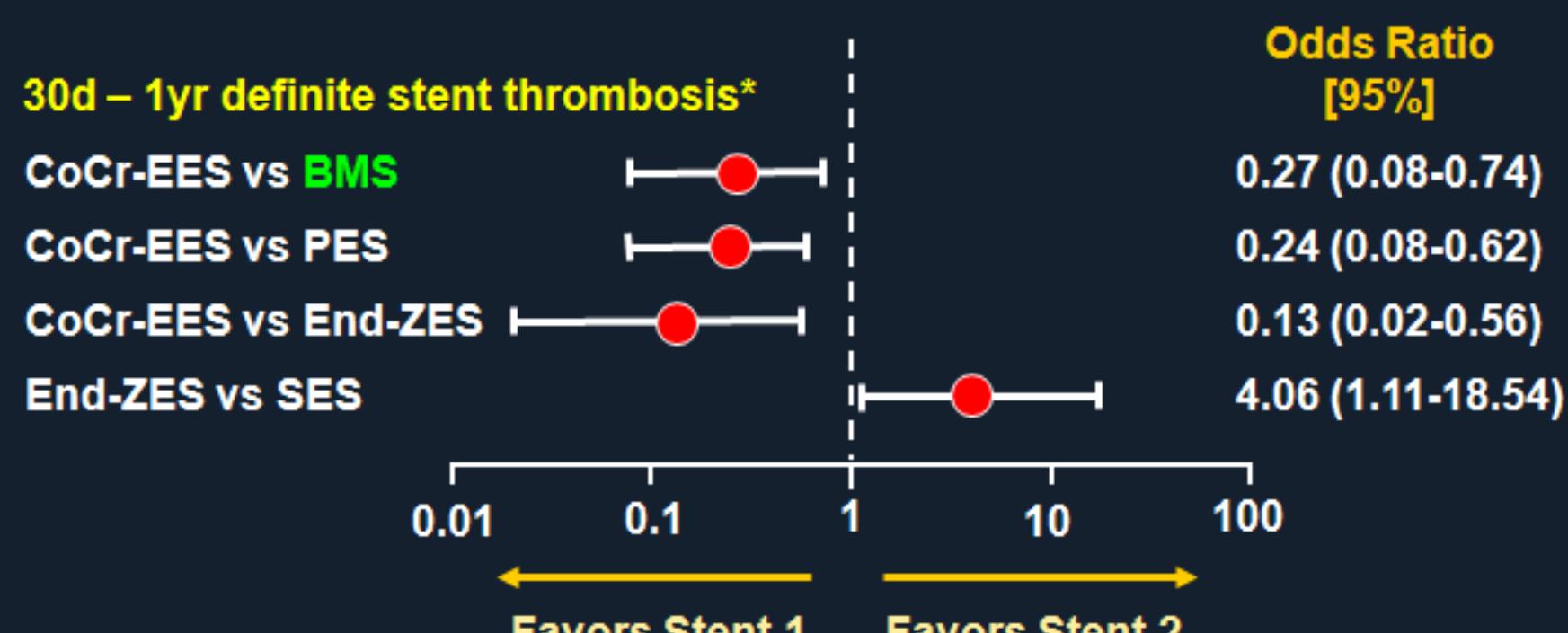
AP2936556-OUS Rev. A. Information contained herein for use outside of the U.S. only.

Palmerini T et al. *Lancet* 2012;On-line

Stent Thrombosis Network Meta-analysis

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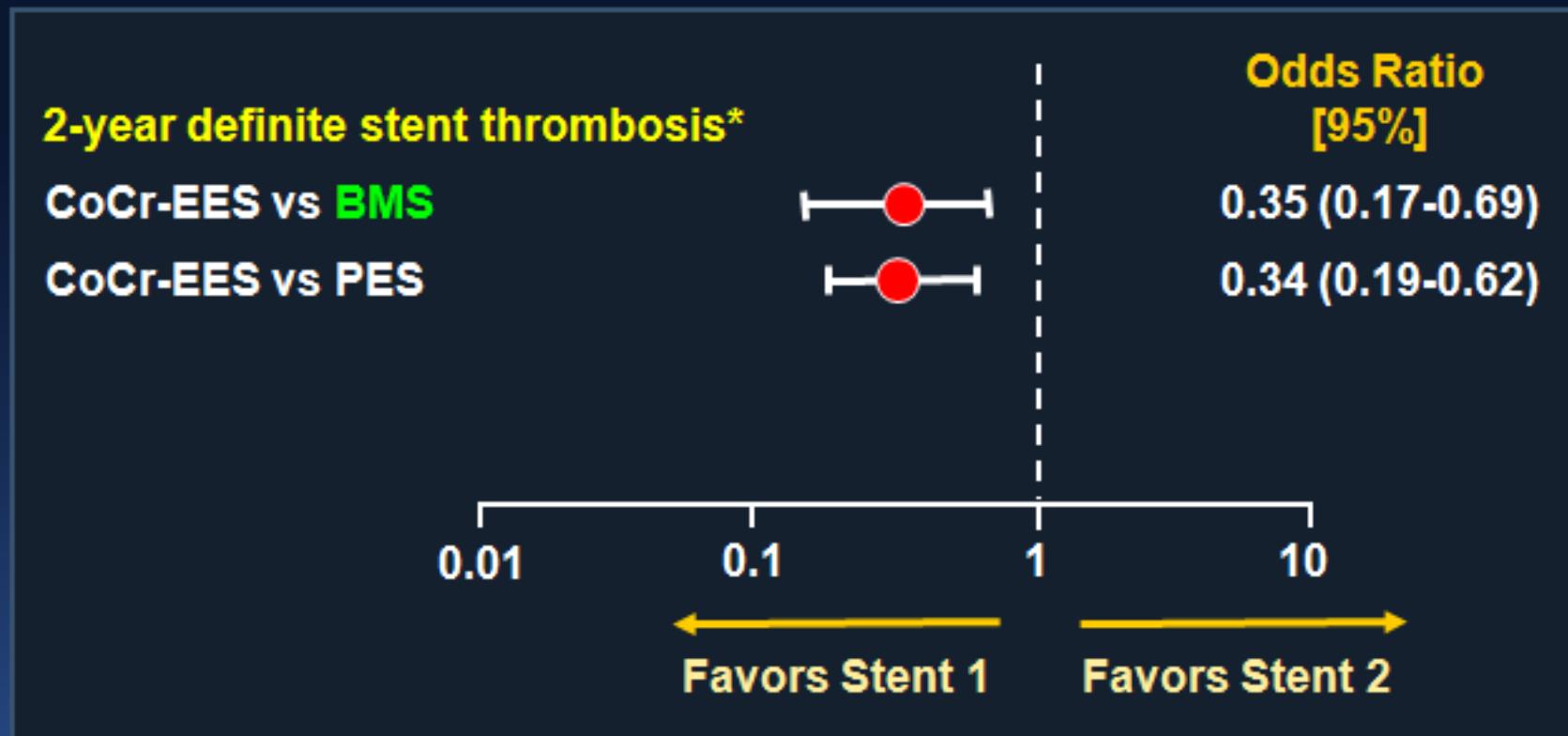
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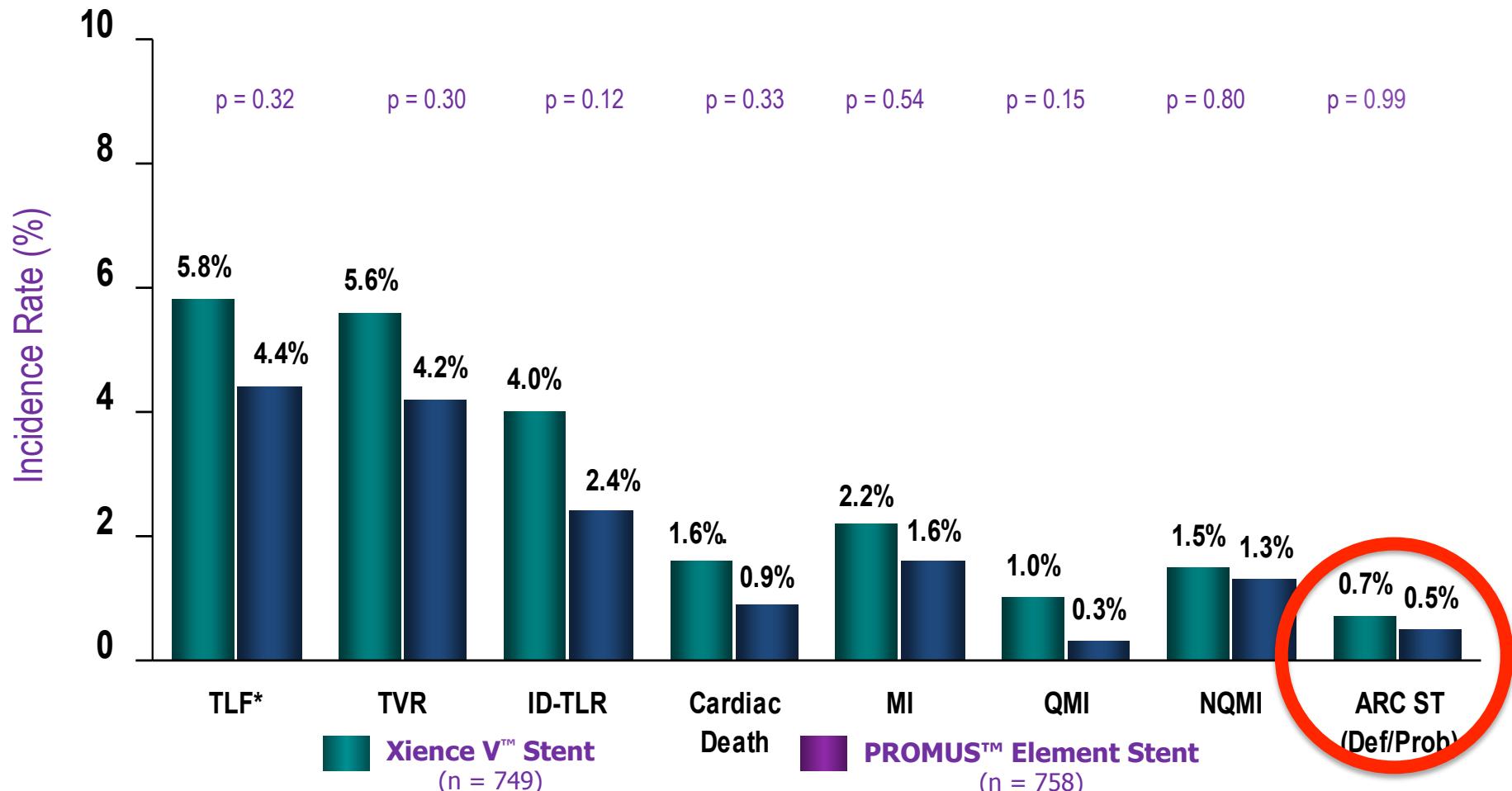
Palmerini T et al. *Lancet* 2012:On-line

Summary

- (1) CoCr-EES were associated with significantly lower rates of 1-year and 2-year definite stent thrombosis than were BMS, a result not present with other DES
- (2) the reduction in stent thrombosis with CoCr-EES compared with BMS was apparent both early and late (occurring before 30 days and between 31 days and 1 year)
- (3) CoCr-EES were also associated with significantly lower 1-year rates of definite stent thrombosis than were other first and second generation DES, including PES, SES, PC-ZES, and Re-ZES.

PLATINUM Workhorse Trial

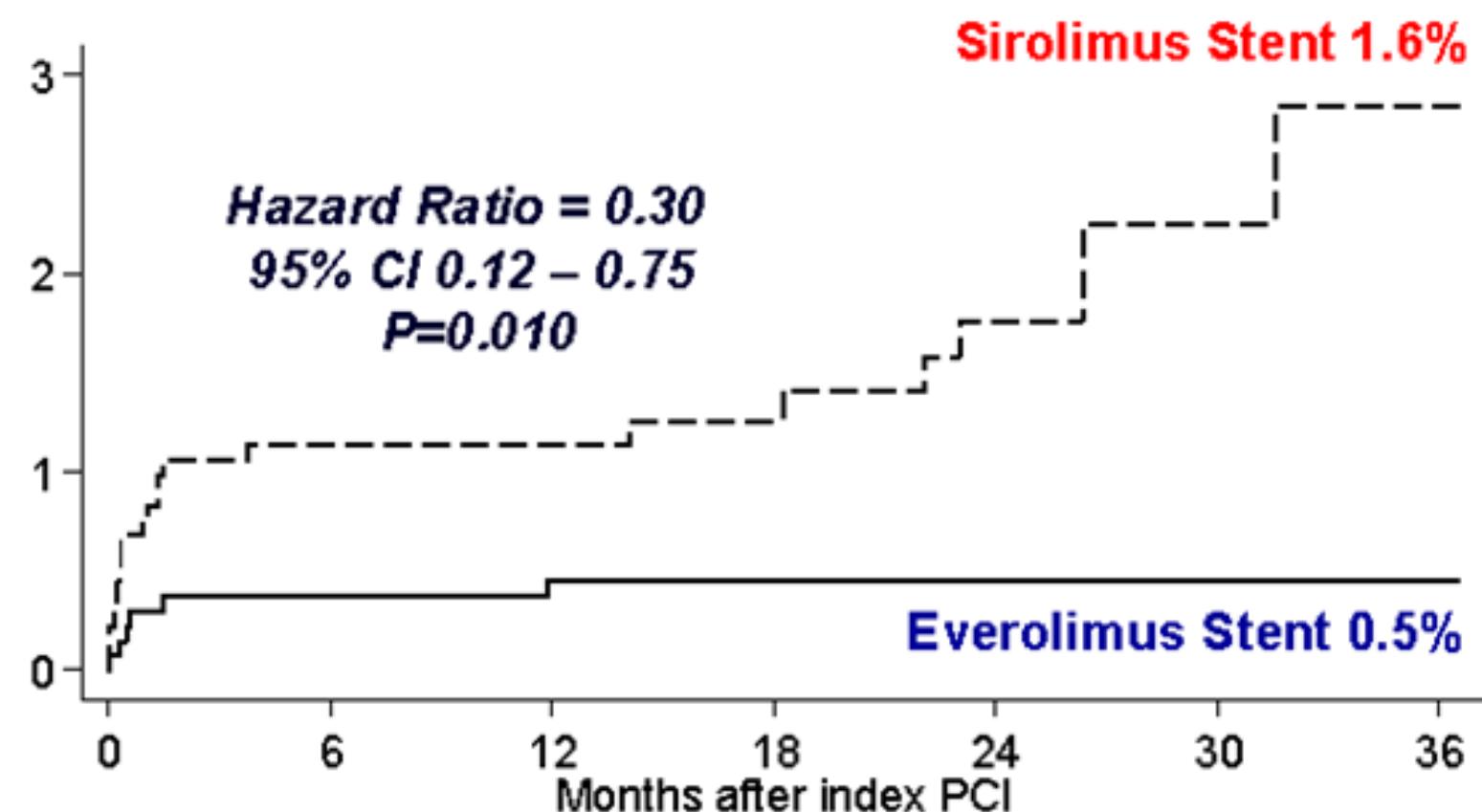
2-Year Clinical Results



*TLF=Ischemia-driven (ID) target lesion revascularization (TLR), cardiac death related to the target vessel (TV) or myocardial infarction (MI) related to the TV.
TVR: target vessel revascularization. ST: stent thrombosis.

Presented by Gregg W. Stone, MD, ACC 2012.

Lesson I – Definite Stent Thrombosis @ 3 Years



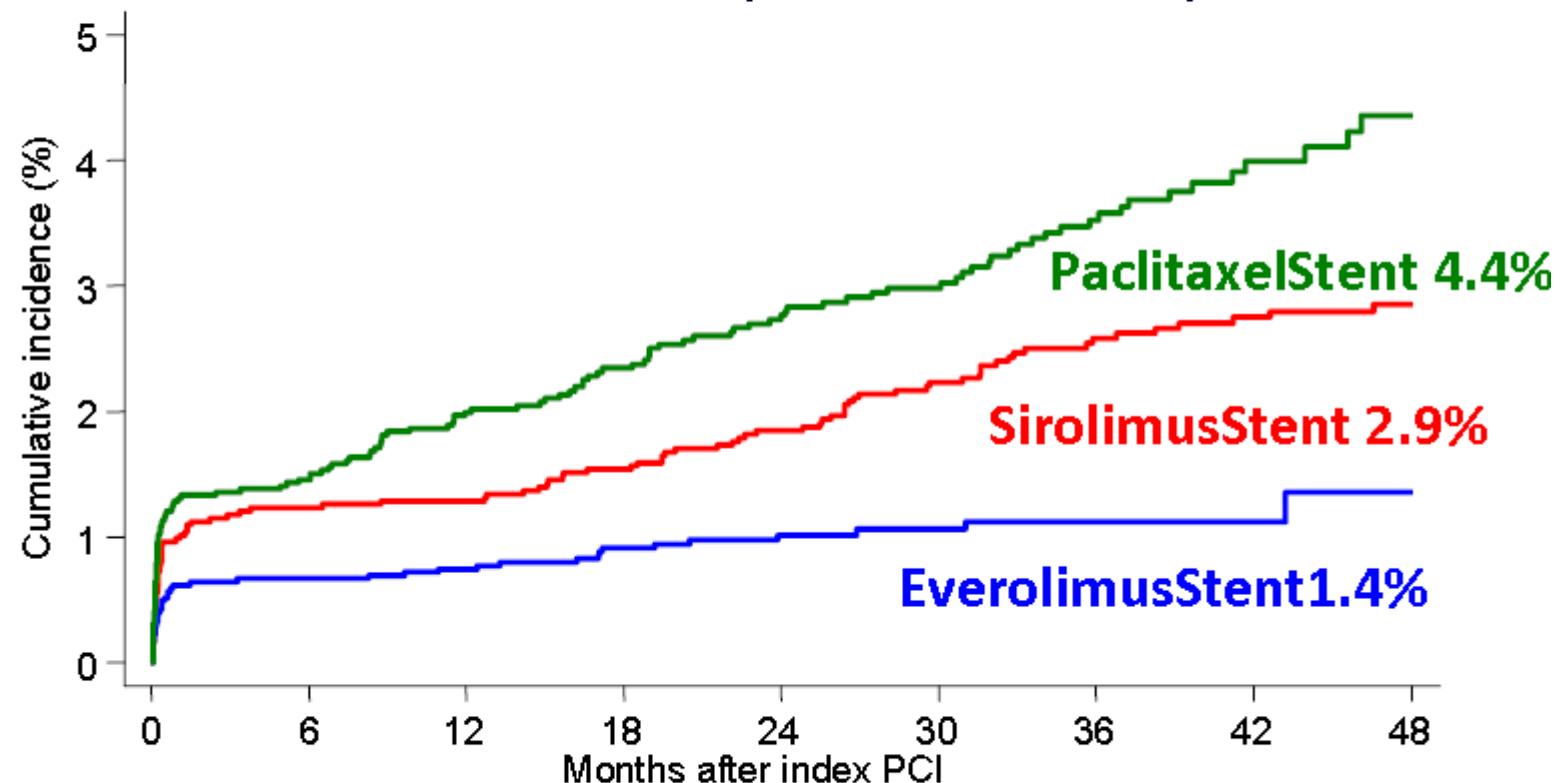
No. at risk

EES	1342	1296	1234	620	543	226	29
SES	1342	1271	1216	619	527	223	28

Primary Endpoint ARC Definite ST @ 4 Years

EES vs. SES Hazard Ratio* = 0.41, 95% CI 0.27–0.62, P<0.0001

EES vs. PES Hazard Ratio* = 0.33, 95% CI 0.23–0.48, P<0.0001



No. at risk

PES	4214	3916	3797	3176	2905	2344	1880	1077	686
SES	3784	3617	3589	3499	3404	3080	2521	2118	1734
EES	4135	3913	3793	3284	2604	1856	1041	514	208

*from Cox proportional hazards model

COMPARE II trial

Pieter Smits

On behalf of all principal COMPARE II investigators:

Ad van Boven, Jean-Jaques Goy, Peter den Heyer,
Antonio Serra, Ton Slagboom, Mario Togni, Ramiro
Trillo Nouche, Mariano Valdés, Andre Vuillomenet,
Jose Vázquez, Vassilis Voudris

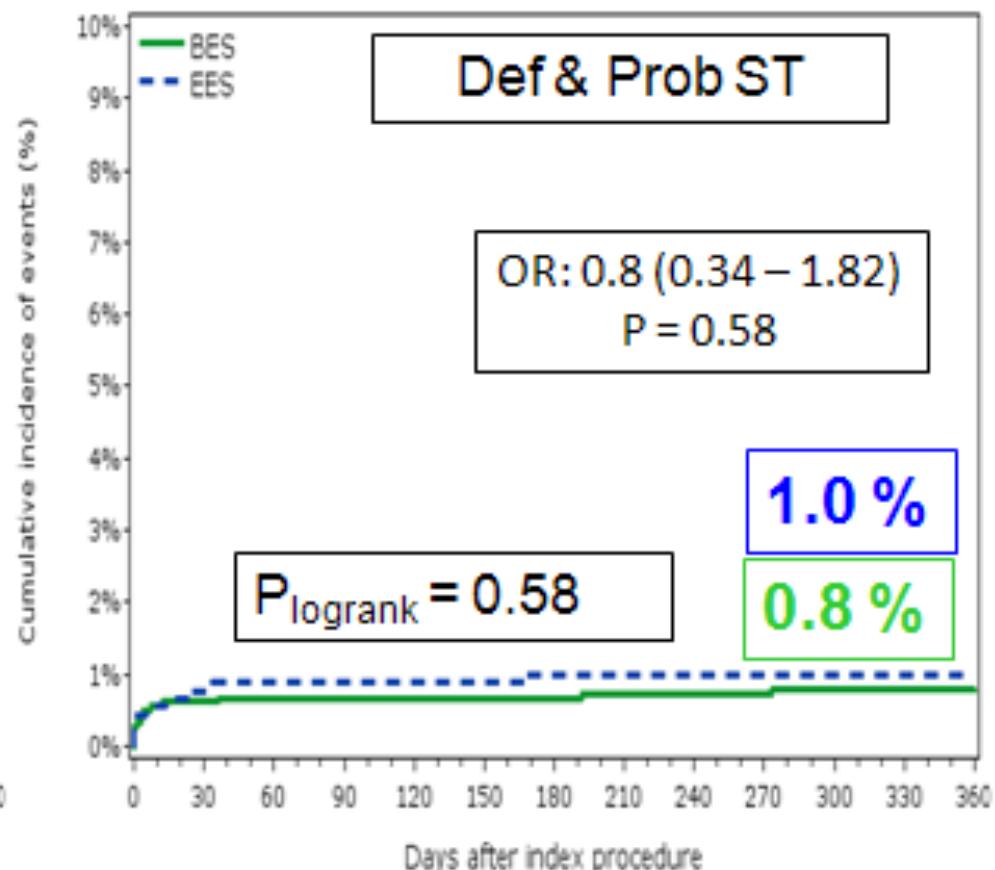
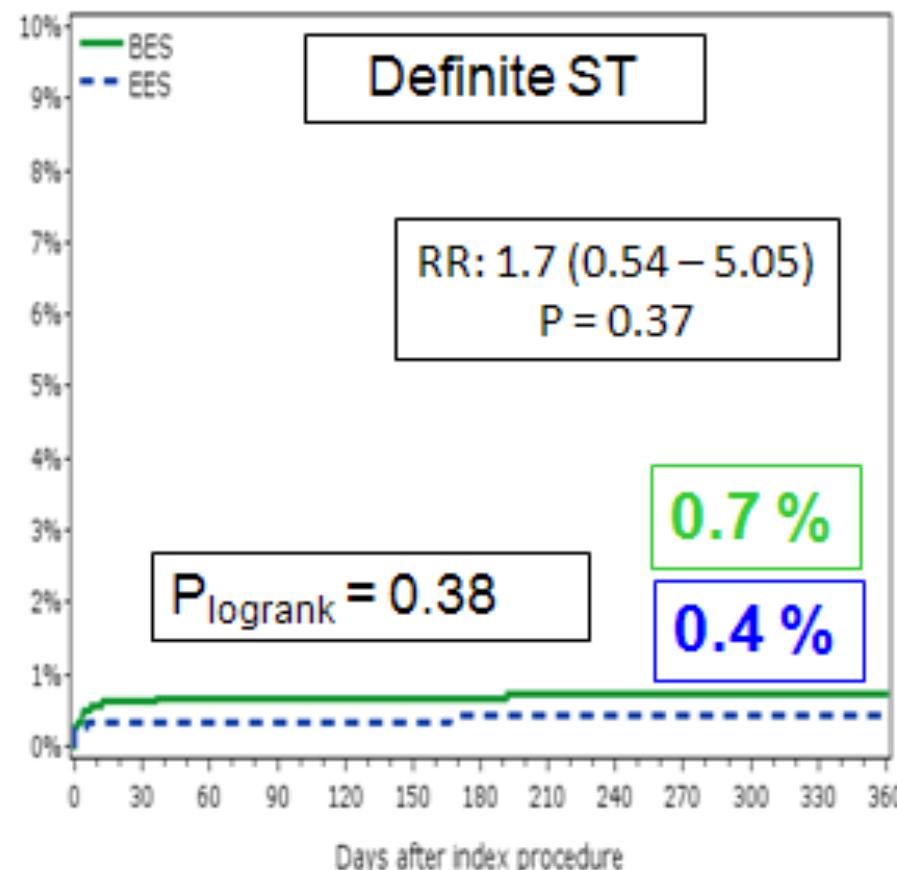
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Purpose

The main objective of the COMPARE II trial is
a head to head comparison of
the everolimus eluting XIENCE-V/PRIME/ PROMUS®(EES)
with
the biolimus eluting NOBORI® stent (BES)
to assess:
whether there is a difference in clinical outcome between
both stent types in a real life situation

Stent Thrombosis (ARC)



Number at Risk:

5	1795	1776	1769	1767	1767	1767	1766	1766	1766	1766	1766	1766	1766	1766	1766	1766	1765	1765	1765
5	912	902	900	900	898	898	897	897	897	897	897	897	897	897	897	897	892	892	892

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Les TS sont-elles toutes identiques ?



- Non, plurifactorielles, certains facteurs potentiellement évitables
- A large échelle, données récentes plaident pour une sécurité supérieure CoCr-EES vs BMS/ DES 1ère et 2ème génération jusqu'à 2 ans
- Mécanismes: rôle du polymère ?
- Quid des PtCr-EES, polymères biodégradables ?

Effect of drug-eluting stents in patients with acute
ST-segment elevation myocardial infarction undergoing
percutaneous coronary intervention:
a meta-analysis of randomised trials and an adjusted indirect
comparison

- Until July 2009
- 13 clinical trials
- N=7224
- **STENT THROMBOSIS** for BMS (2.76%) and DES (2.65%) were similar
- *EuroIntervention.* 2010 Feb;5(7):853-60

TITAX-AMI trial

Definite Stent Thrombosis: 3-Year Follow-Up



TITAX-OCT study

Follow-Up Measurement

