



# Antithrombotic therapy in the ACS patient with atrial fibrillation

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# Disclosures

## DISCLOSURE STATEMENT OF FINANCIAL INTEREST

Kurt Huber, MD, FESC, FACC

**Research Grants from Bristol-Myers Squibb, Eli Lilly, Medtronic, Sanofi-Aventis**

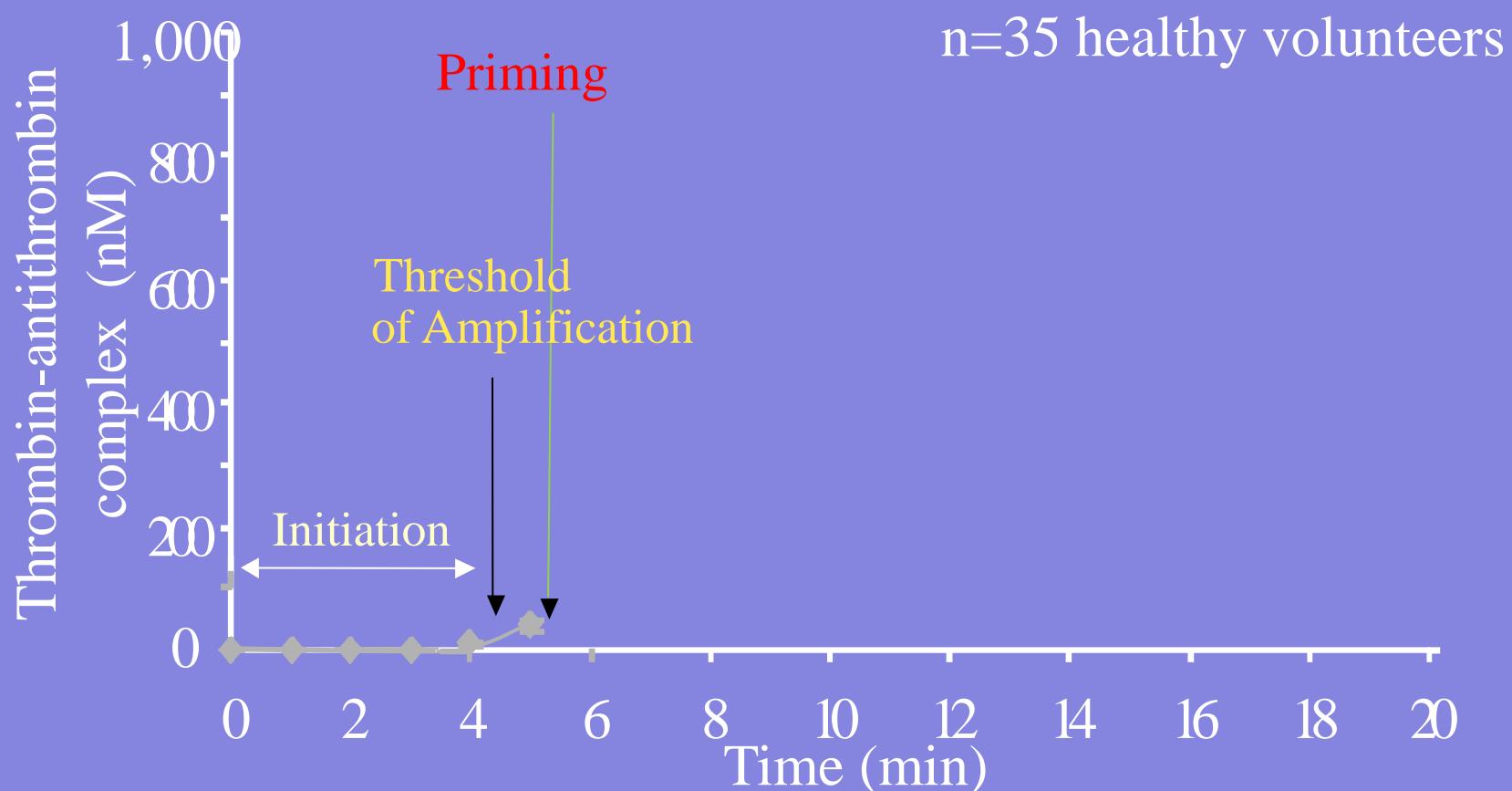
**Consulting Fees from AstraZeneca, Bayer, Boehringer-Ingelheim, Bristol-Myers Squibb, Daiichi Sankyo, Fibrex, Eli Lilly, Portola, Sanofi-Aventis, Schering-Plough, The Medicines Company**

**Lecture Fees from AstraZeneca, Boehringer-Ingelheim, Boston Scientific, Bristol-Myers Squibb, Cordis / Johnson&Johnson, Daiichi Sankyo, Eli Lilly, GlaxoSmithKline, Pfizer, and Sanofi-Aventis.**



# BACKGROUND

# Thrombin generation in thrombosis

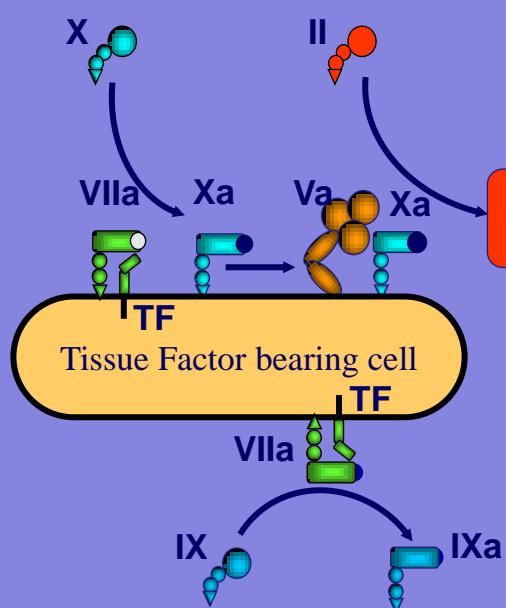


Thrombin activated during initiation phase is only used for platelet activation in the priming phase. Platelets in the Priming Phase are almost exclusively activated by thrombin induced during the initiation phase

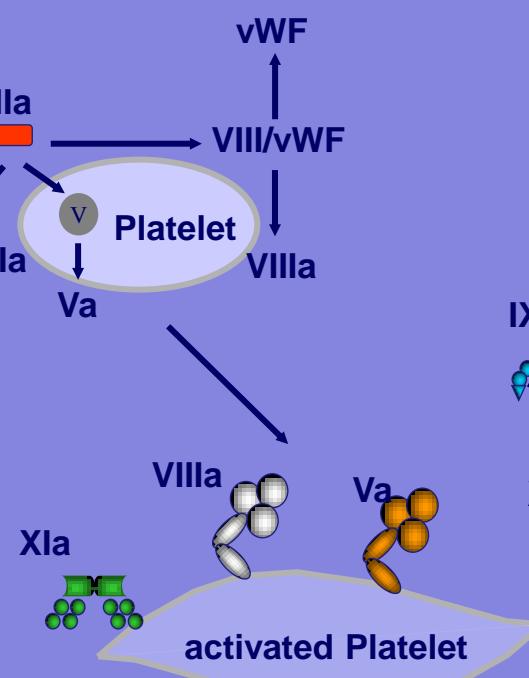


# Cell-based model of coagulation

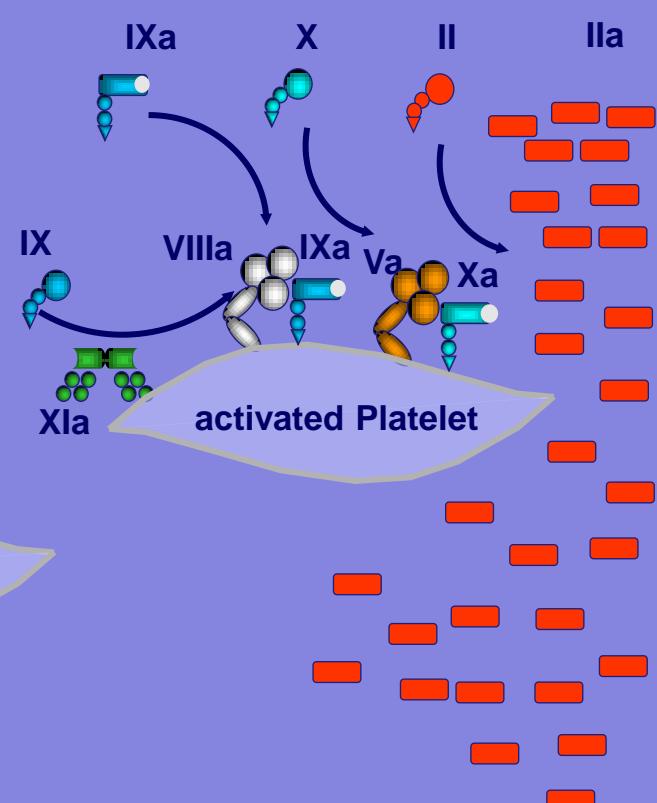
## Initiation



## Priming

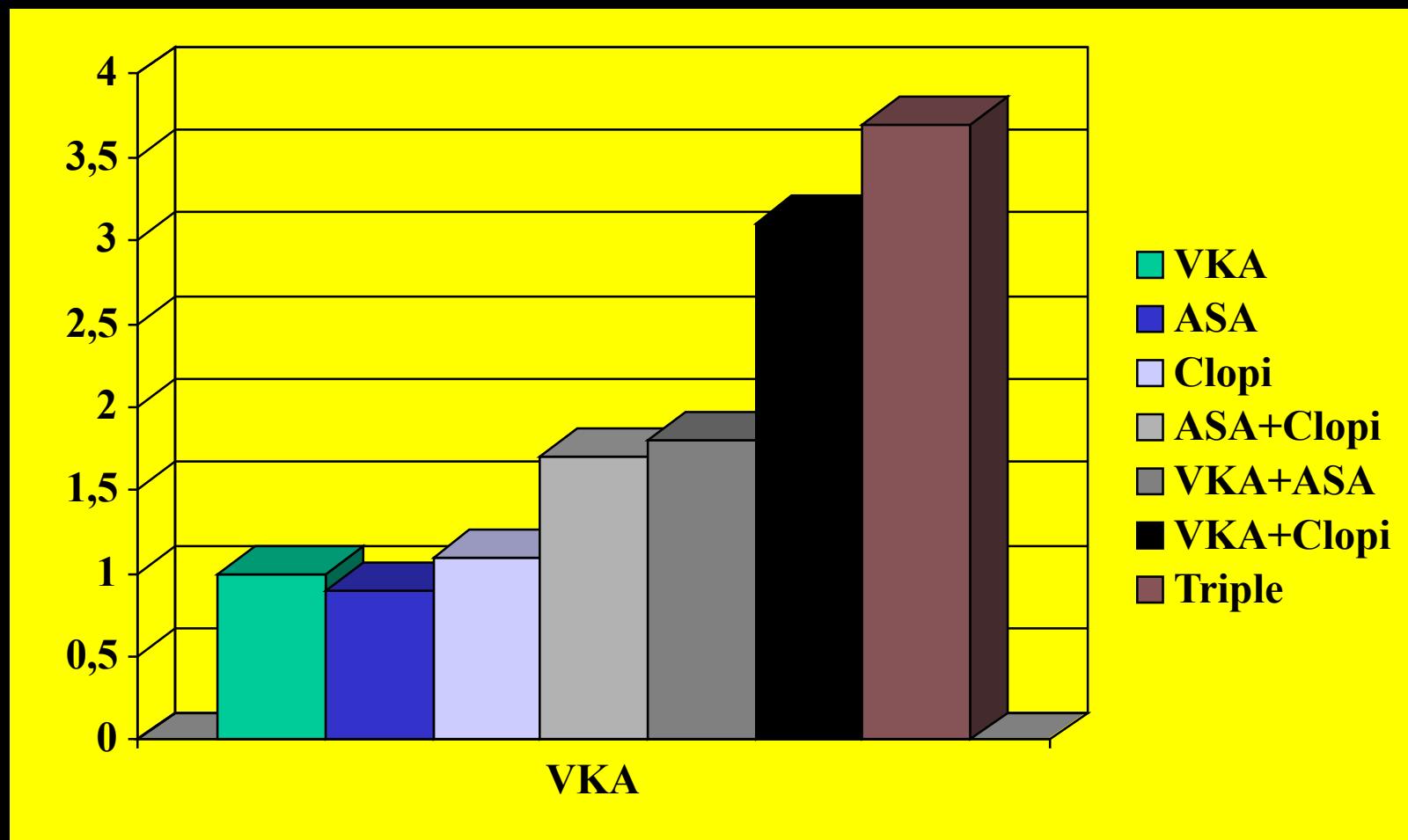


## Propagation





# Major Bleeding\* per Year Odds Ratios



\* non-fatal and fatal

Hansen et al. Arch Intern Med 2010;170:1433-1441



# **“triple”-Therapy in pts. with Atrial Fibrillation after ACS or Elective Stent Implantation**



# ESC Guidelines in AF patients at moderate to high thromboembolic risk in whom OAC is required

## Low bleeding risk

Elective BMS\*



VKA (INR 2.0-3.0)

Elective DES (-olimus)



VKA (INR 2.0-2.5) +  
Clopidogrel (or ASA)

Elective DES (paclitaxel)



ACS + BMS/DES



VKA (INR 2.0-2.5) +  
ASA + Clopidogrel

## High bleeding risk\*\*

Elective BMS



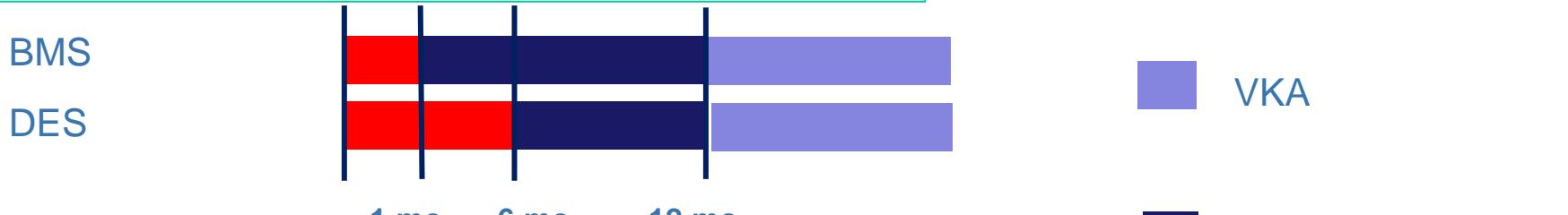
ACS + BMS



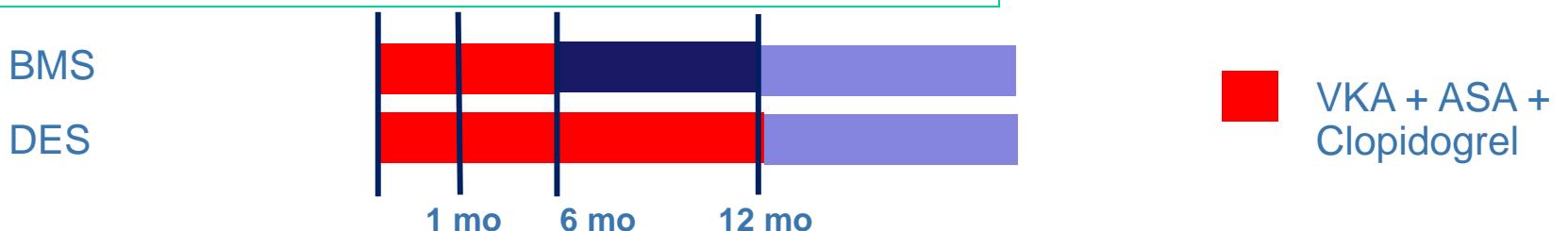


# A North-American consensus document on antithrombotic therapy in AF patients and a coronary stent with moderate/high stroke risk (CHADS<sub>2</sub> ≥2)

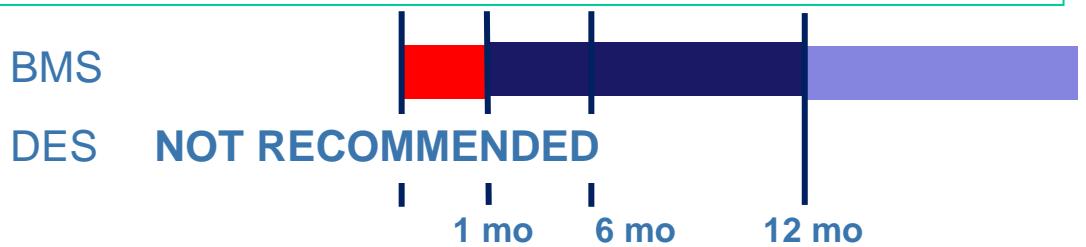
## Low stent thrombosis risk and low bleeding risk



## High stent thrombosis risk and low bleeding risk



## Any stent thrombosis risk and high bleeding risk



Dabigatran 2x100 mg was discussed for the first time as possible replacement of VKAs

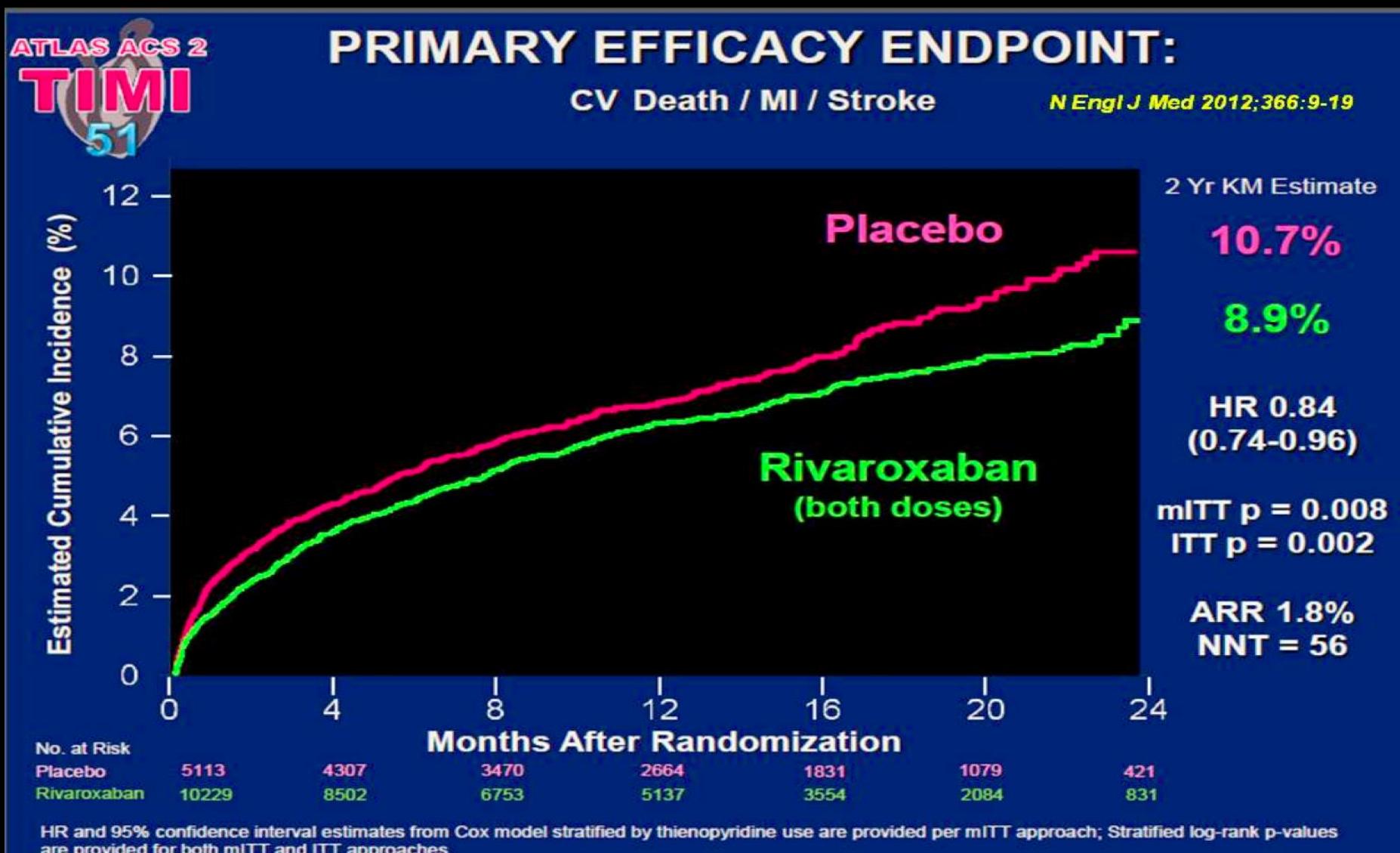


# **„Triple“ Therapy in Secondary Prevention after ACS with a NOAC added to DAPT**



# ATLAS-2 Studie

## (Rivaroxaban)

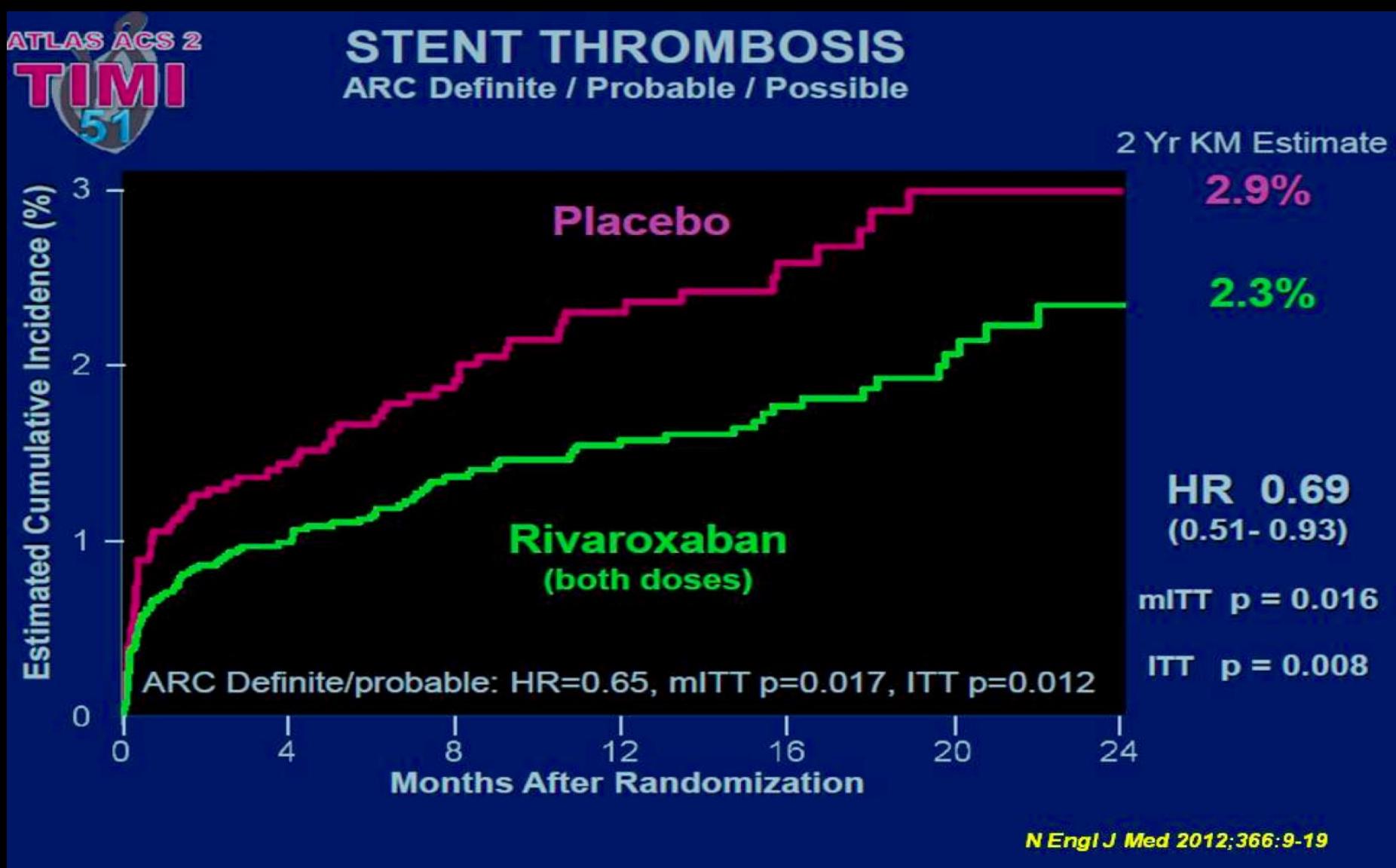




# ATLAS-2 Studie

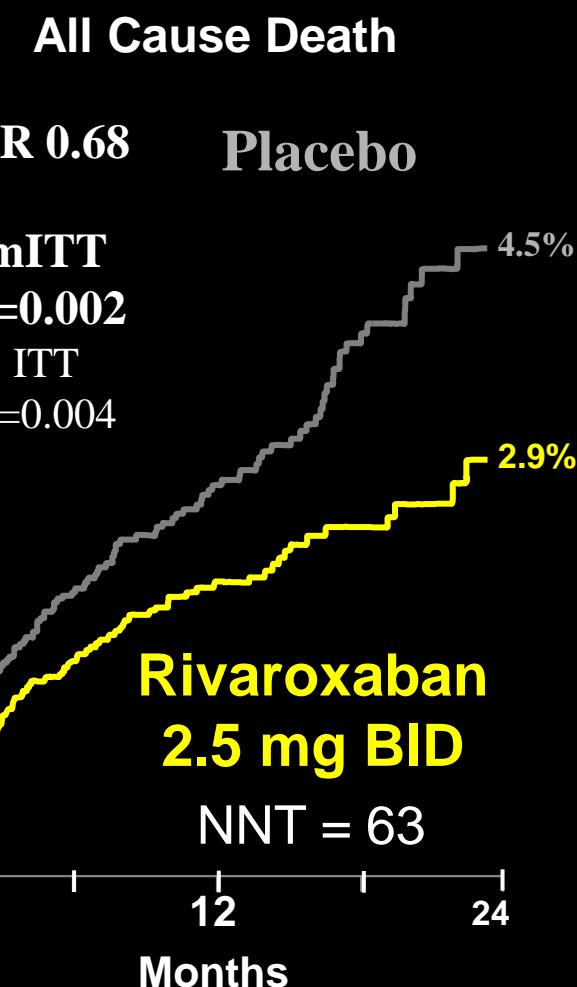
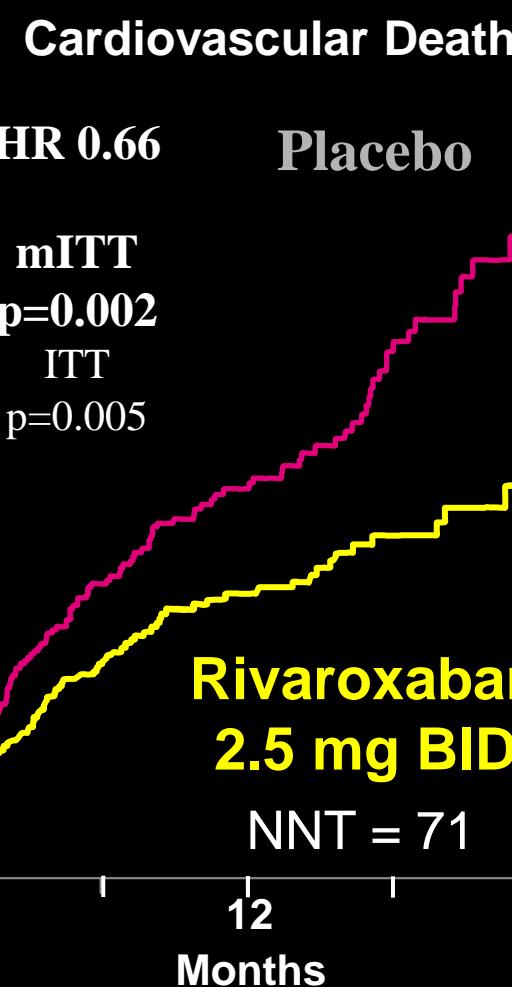
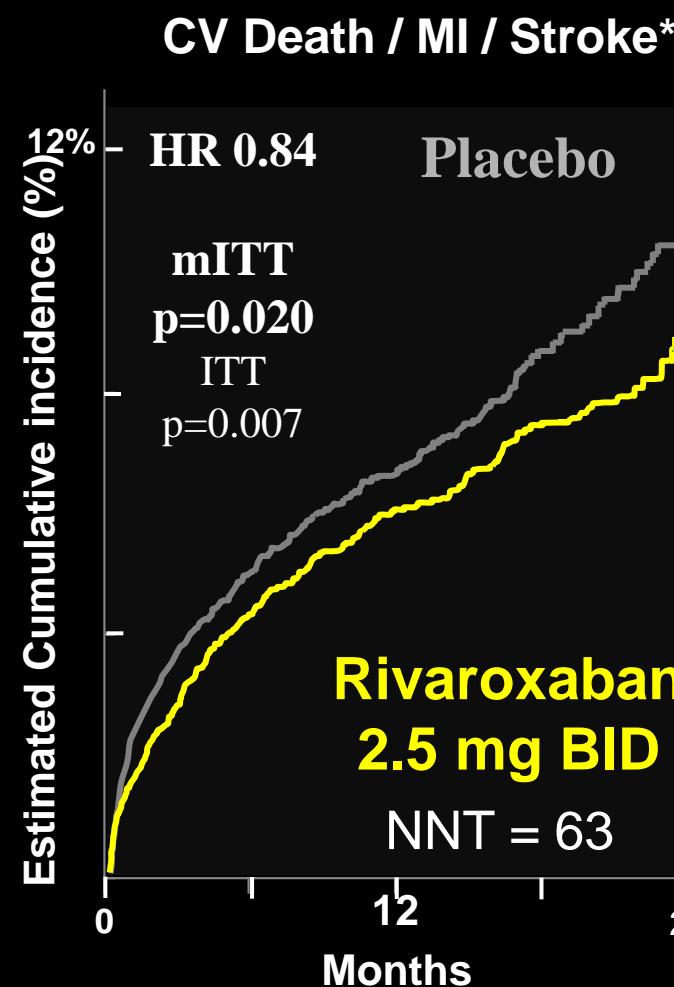
## (Rivaroxaban)



*N Engl J Med* 2012;366:9-19

Mega JL et al. Lancet 2009;374:29-3

# PRIMARY EFFICACY ENDPOINT\*: 2.5 mg PO BID

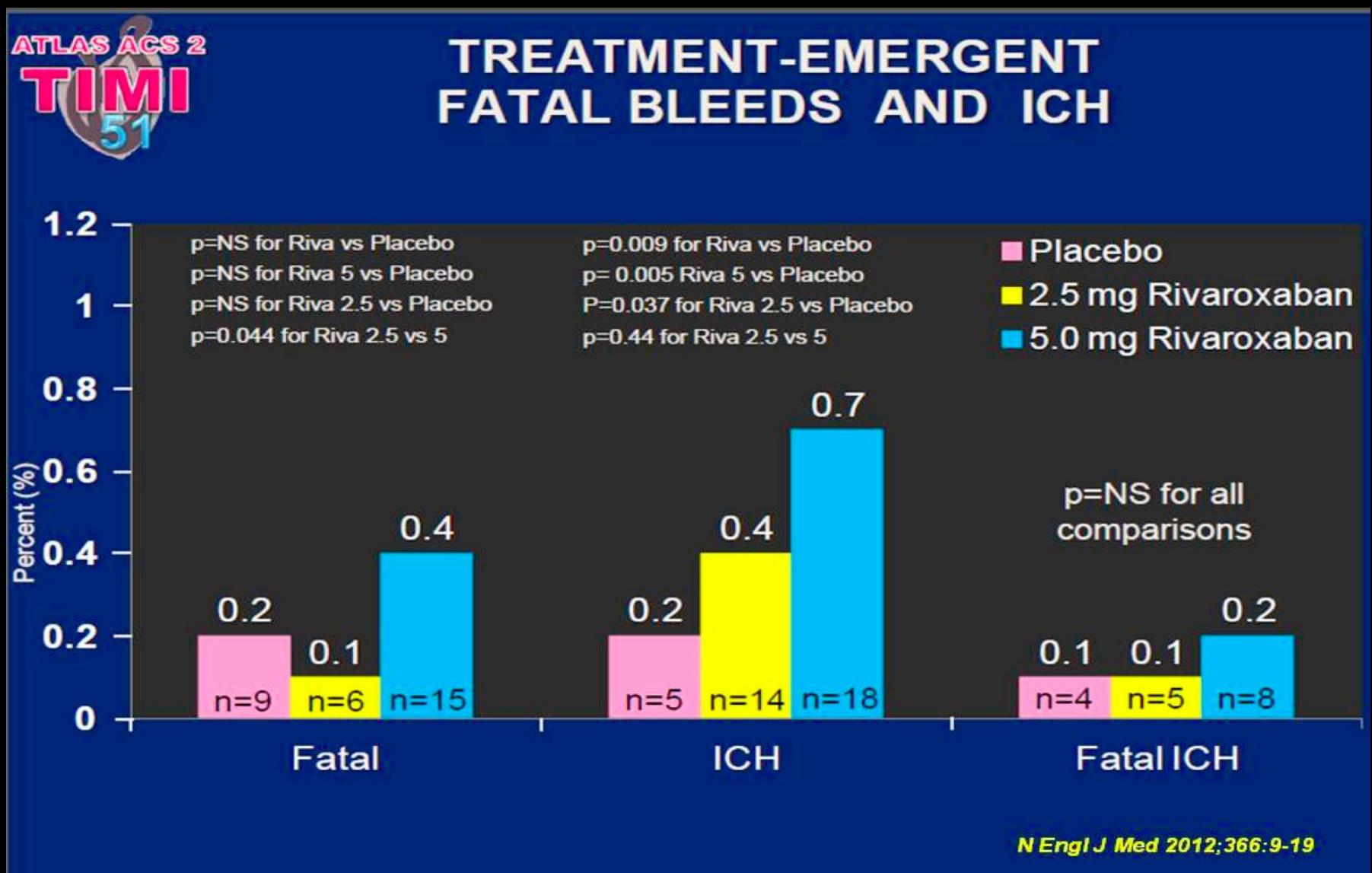


\* First occurrence of cardiovascular death, MI, stroke (ischemic, hemorrhagic, and uncertain) as adjudicated by the CEC across thienopyridine use strata  
Two year Kaplan-Meier estimates, HR and 95% confidence interval estimates from Cox model stratified by thienopyridine use are provided per mITT approach; Stratified log-rank p-values are provided for both mITT and ITT approaches; NNT=Number needed to treat.



# ATLAS-2 Studie

## (Rivaroxaban)





# Should we skip aspirin ?

# WOEST Trial - Study Design

## 1:1 Randomisation:

Double therapy group:

**OAC + 75mg Clopidogrel qd**

1 month minimum after BMS

**1 year after DES**

Triple therapy group

**OAC + 75mg Clopidogrel qd + 80mg Aspirin qd**

1 month minimum after BMS

**1 year after DES**

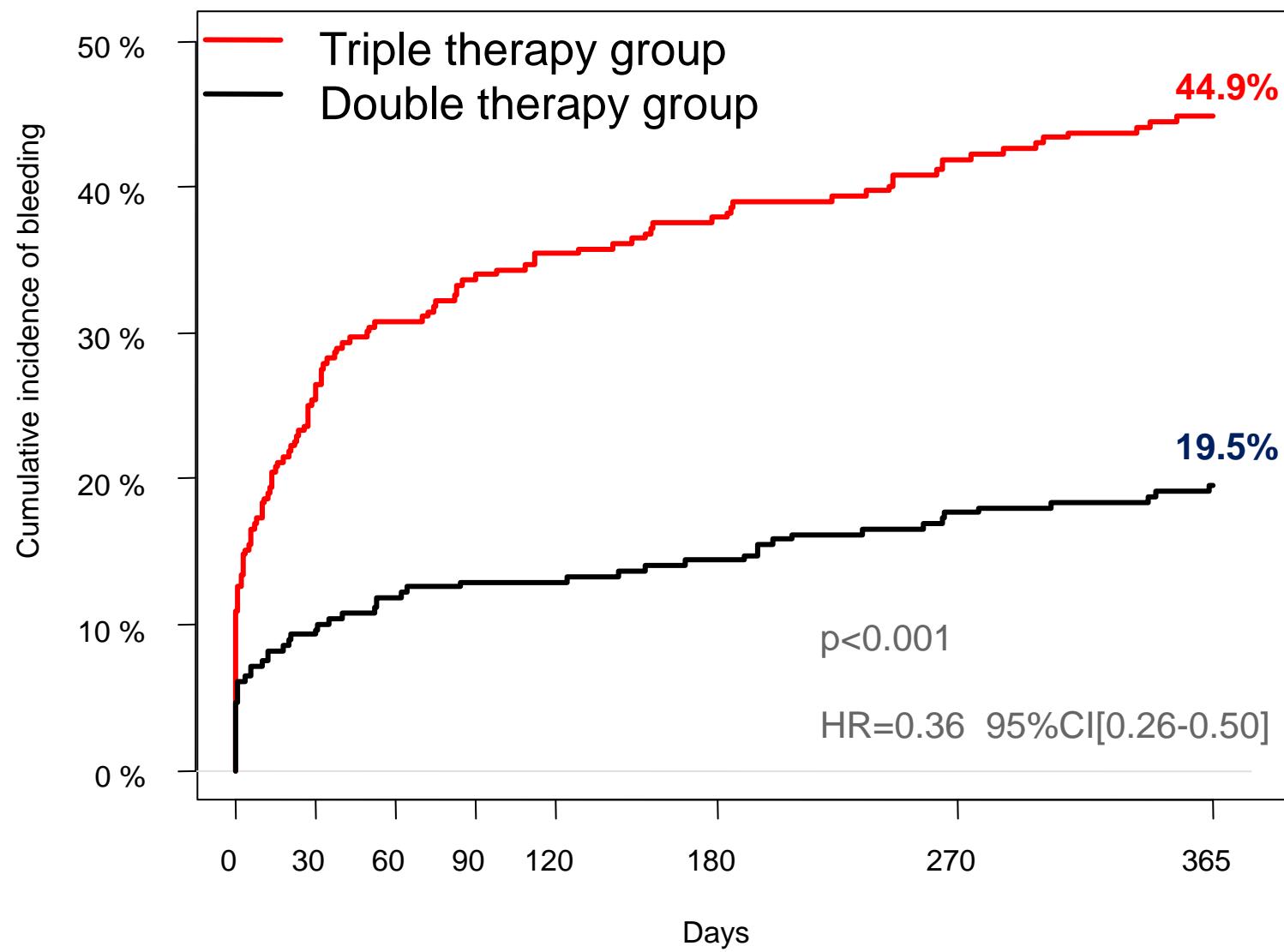
## Follow up: 1 year

**Primary Endpoint:** The occurrence of all bleeding events (TIMI criteria)

## Secondary Endpoints:

- Combination of stroke, death, myocardial infarction, stent thrombosis and target vessel revascularisation
- All individual components of primary and secondary endpoints

## Primary Endpoint: Total number of bleeding events (TIMI criteria)



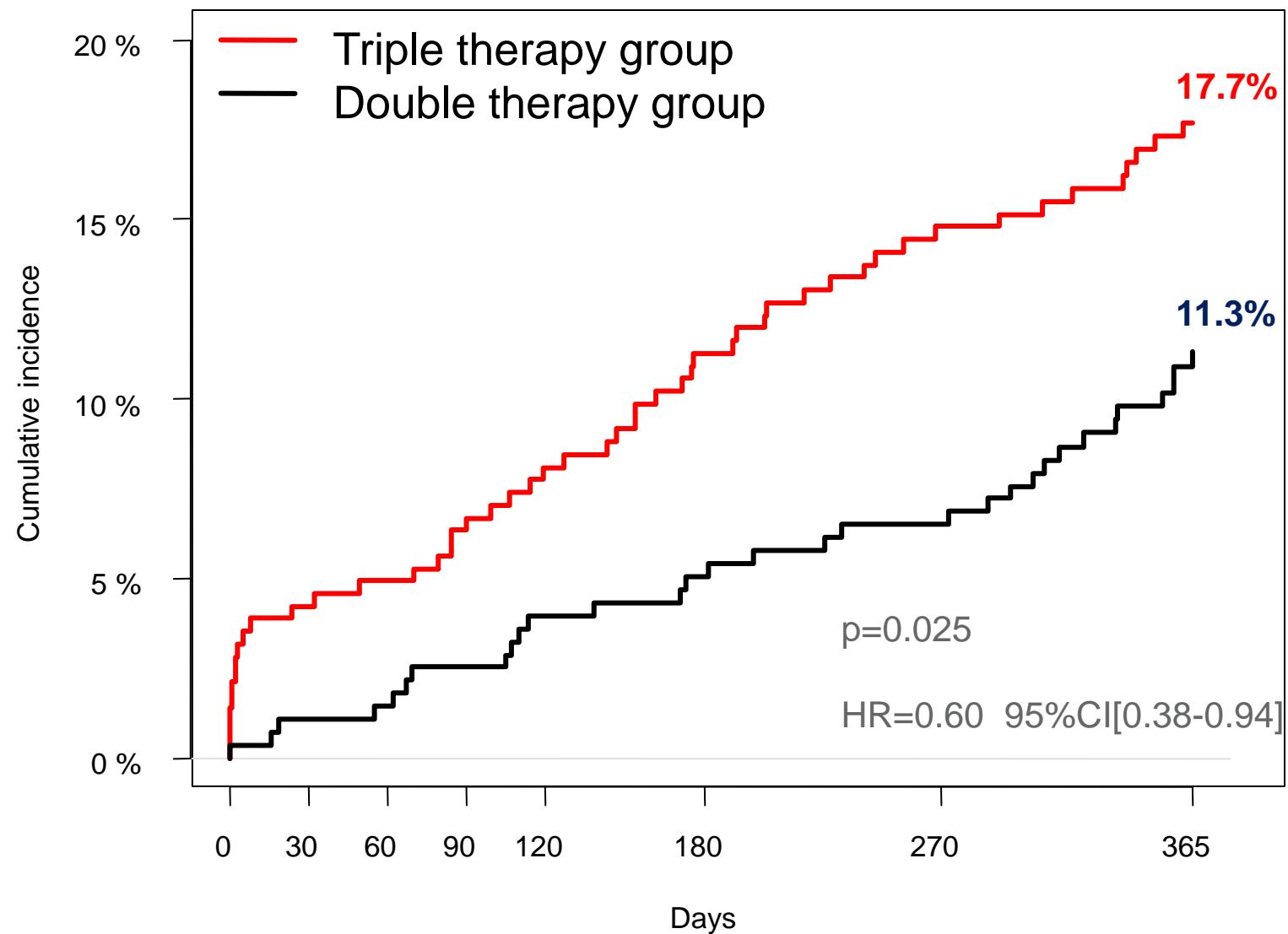
n at risk:

284  
279210  
253194  
244186  
241181  
241173  
236159  
226140  
208



WOEST

## Secondary Endpoint (Death, MI, TVR, Stroke, ST)



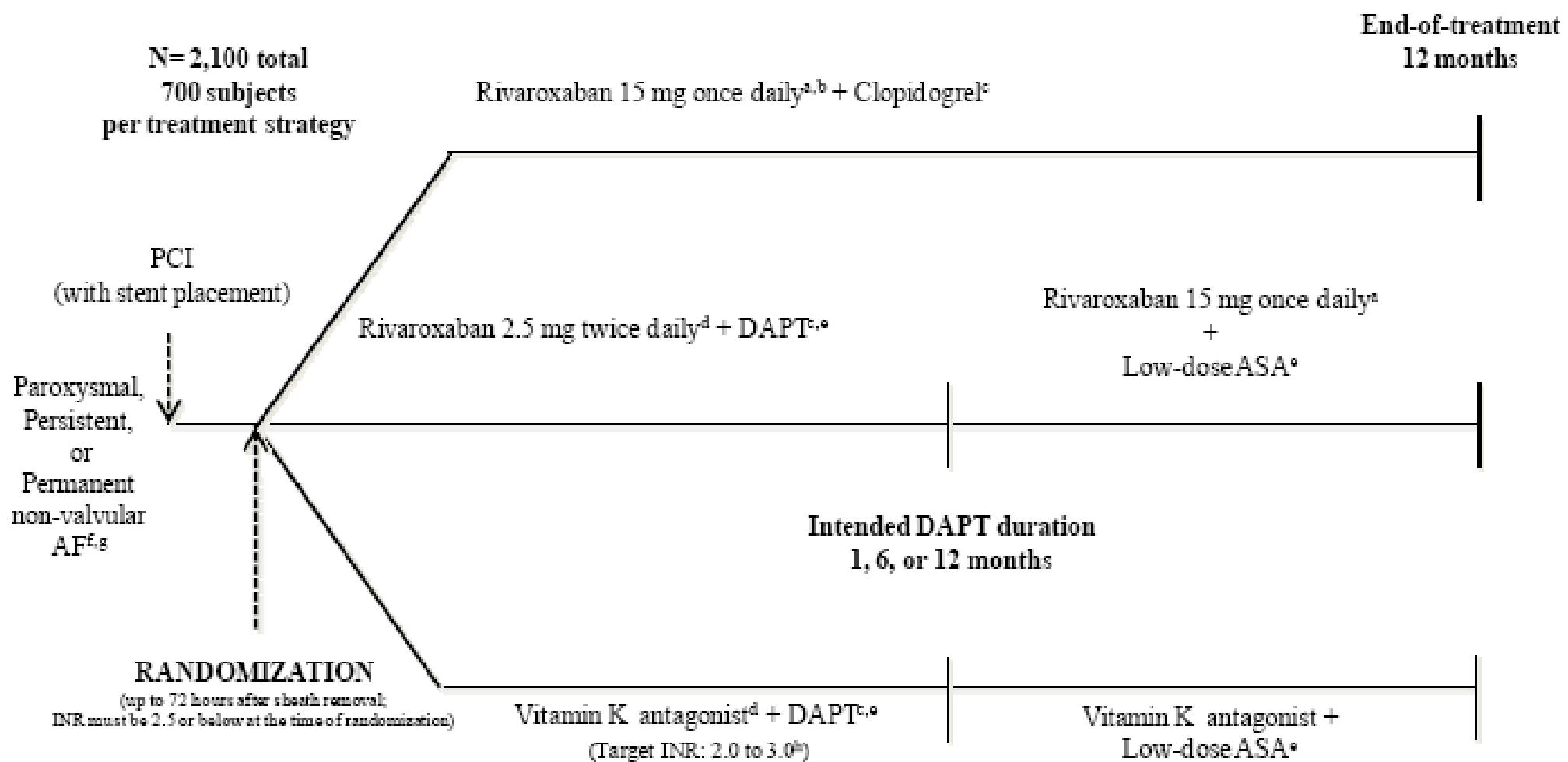
n at risk:

284	272	270	266	261	252	242	223
279	276	273	270	266	263	258	234



# PIONEER AF-PCI

Study Diagram: Clinical Protocol RIVAROXAFL3003





Thank you