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# Anti-Platelet Therapy : who needs it & what to prescribe?

**Koen Deloose, MD**

# Anti-platelet therapies : targets

## THIENOPYRIDINES

### 1<sup>st</sup> generation : irreversible

Ticlopidine  
Clopidogrel  
Prasugrel

### 2nd generation : reversible – non competitive

Cangrelor  
Ticagrelor

### 3rd generation : reversible – competitive

Elinogrel

phosphodiesterase

ADP

## PHOSPHO-DIESTERASE INHIBITOR

Cilostazol - Dipyridamole

Collagen

Thrombin

Activation

## GP IIB/IIIA INHIBITOR

ReoPro - Abciximab

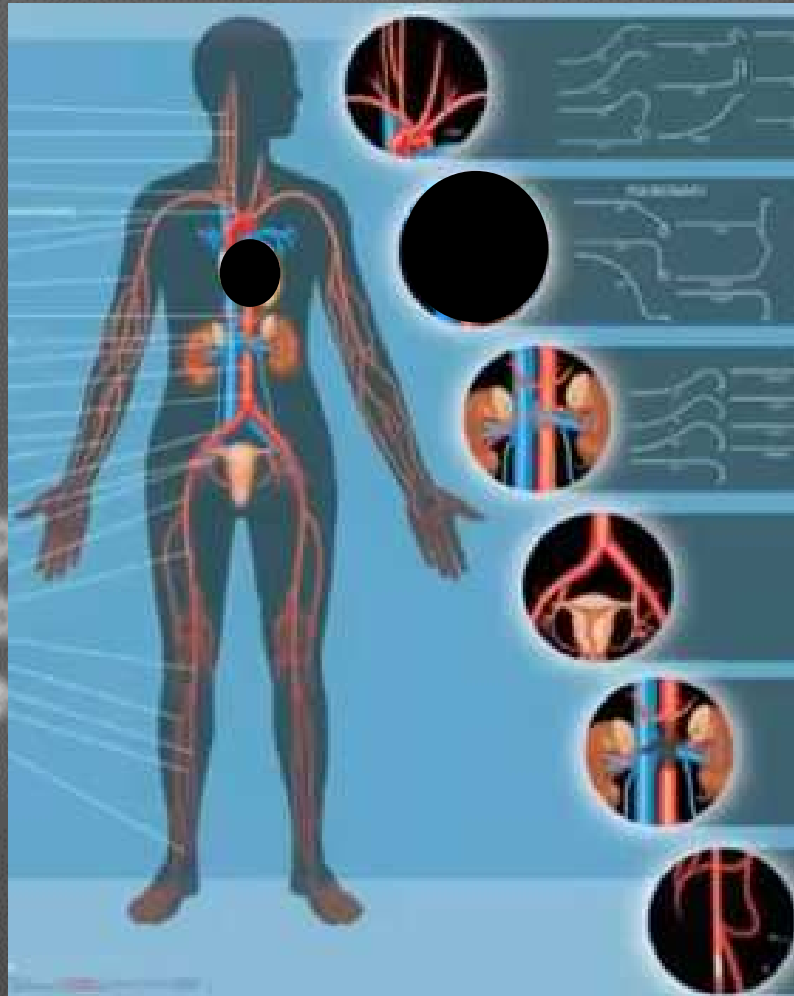
## PAR1 thrombin receptor ANTAGONIST

Voraxapar (Zontivity<sup>®</sup>) - Atopaxar

## CYCLO-OXYGENASE INHIBITOR

Aspirin

# Anti-platelet therapies : targets



CYCLO-OXYGENASE INHIBITOR  
Aspirin

THIENOPYRIDINES  
**1<sup>st</sup> generation : irreversible**  
Ticlopidine  
Clopidogrel

PHOSPHO-DIËSTERASE INHIBITOR  
Cilostazol – Dipyridamole

# Role of Anti-platelet therapies

- **Primary prevention**
- **Secondary prevention**
  - **The cardiovascular patient**
  - **(Endo)vascular therapy**

# Role of Anti-platelet therapies

- **Primary prevention**

Physicians' Health Study Research Group. *NEJM* 1989;321:129-35

22,071 men randomized to aspirin (325mg every other day) followed for an average of 5 years

**End point**

**Relative**

Myocardial infarction

**Aspirin significantly reduces the MI-risk but not the stroke-risk in men**

0.43
0.20 (0.91-1.59)
1.22 (0.93-1.60)
0.15

# Role of Anti-platelet therapies

- **Primary prevention**

**W**omens' **H**ealth **S**tudy. *Ridker P et al. NEJM 2005;352:1293-304*

39,876 women randomized to aspirin (100 mg every other day) or placebo for an average of 10 years



**Aspirin significantly reduces the risk of stroke but not this of MI in women**

# Role of Anti-platelet therapies

- **Primary prevention**

Physicians' Health Study Research Group. *NEJM* 1989;321:129-35

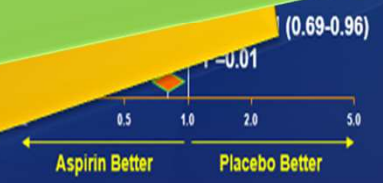
Womens' Health

304

22,071 men randomized to aspirin (22,071) or placebo (22,071) followed for an average of 5 years

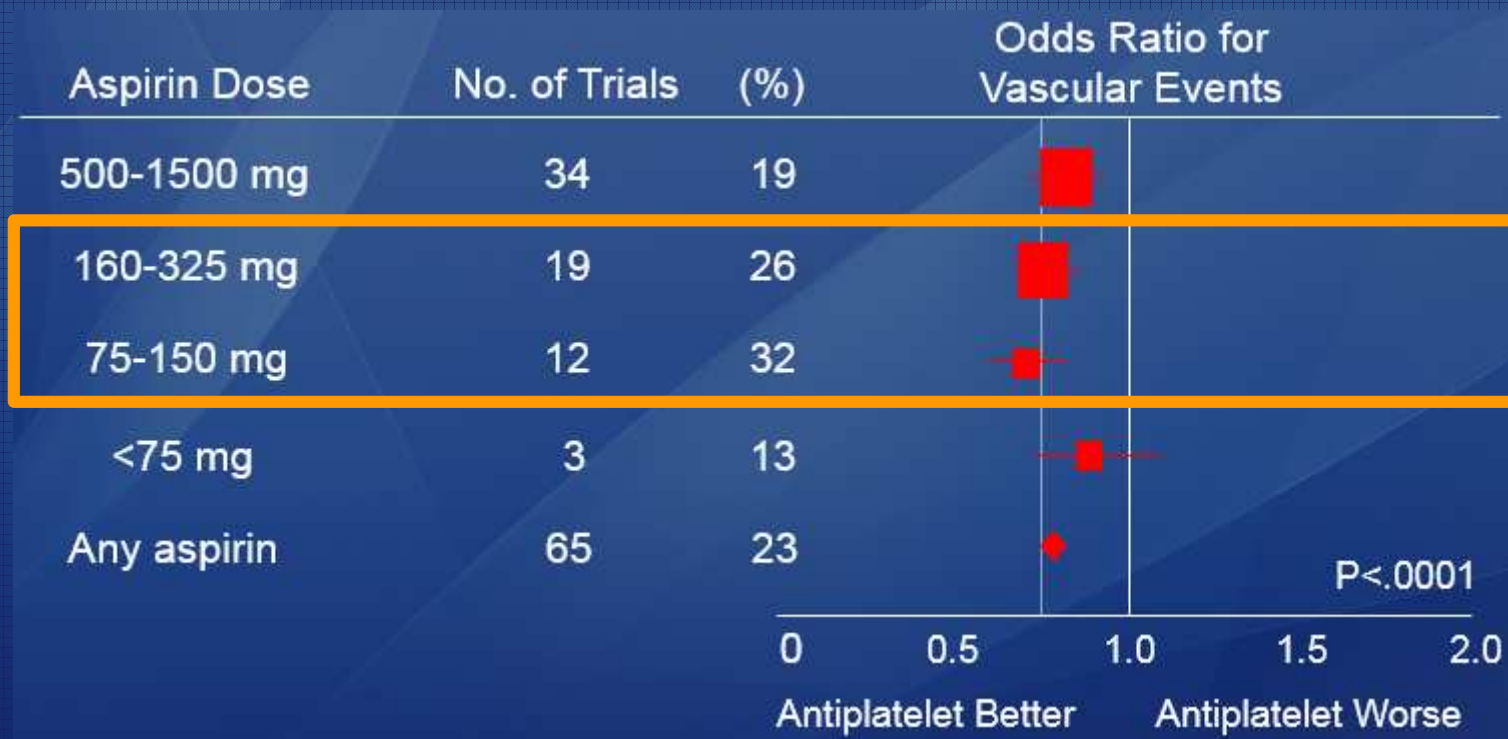
ay) or

**Overall conclusion : Aspirin is recommended for men & women whose 10-year risks for CVD is >10% over 10 years**



# Role of Anti-platelet therapies

- **Primary prevention**  
**Optimal dose**





# Role of Anti-platelet therapies

- **Primary prevention**

**No data to support the role of other anti-platelet agents in primary prevention**



# Role of Anti-platelet therapies

- **Secondary prevention in CV patients**

Effect of Aspirin on vascular

h)

**There is strong evidence to recommend Aspirin (75-162 mg daily) if known CHD/ASVD**



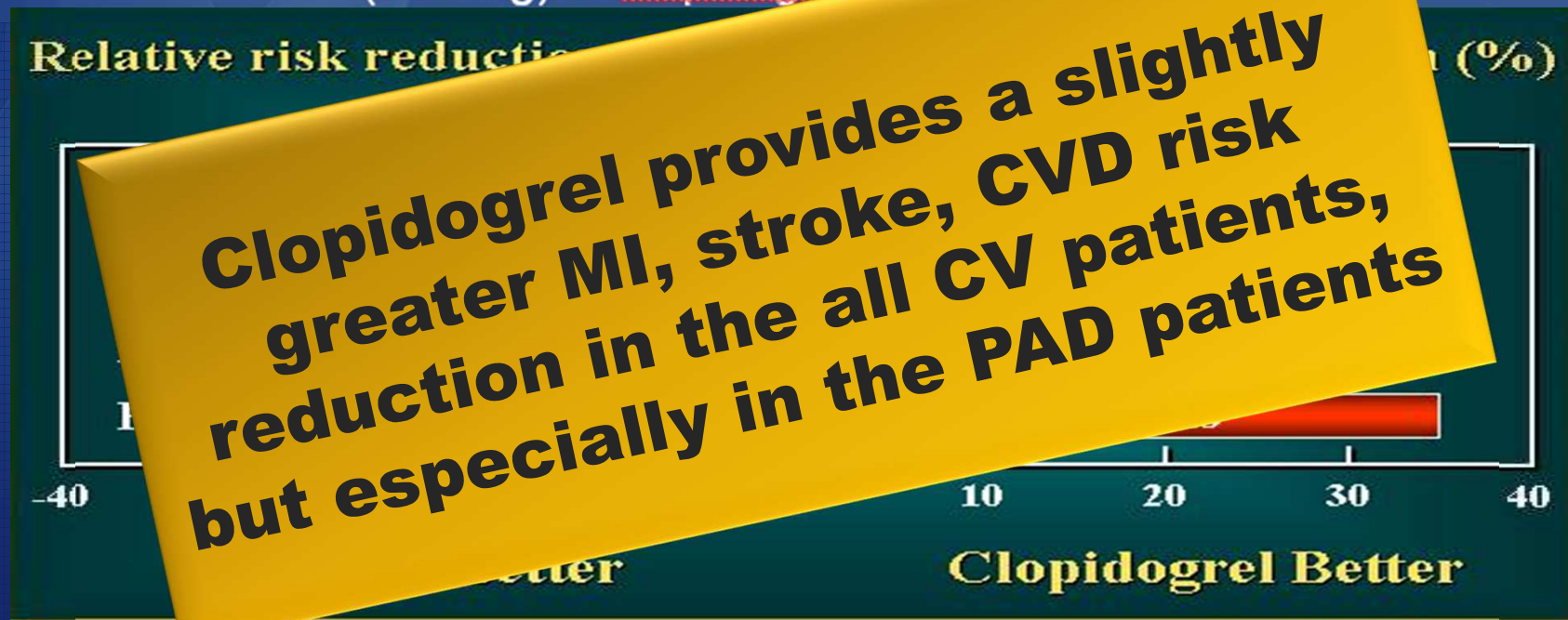
0.0 0.5 1.0 1.5 2.0  
Antiplatelet better Control better

# Role of Anti-platelet therapies

- **Secondary prevention in CV patient @ risk**

## Effect of Clopidogrel on vascular events

19,185 patients with ischemic CVA, MI, or PAD randomized to clopidogrel (75 mg) or aspirin (325 mg)



## Role of Anti-platelet therapies

- **Secondary prevention in CV patient @ risk**

Effect of Aspirin + Clopidogrel on vascular events :

**CHARISMA trial** : 15,603 patients with multiple CV risk factors or known CVD randomized to aspirin (75-162 mg) or aspirin (75-162 mg) & clopidogrel (75 mg) for a mean of 28 months.

**Routine DAP offers no long term benefit**



# Role of Anti-platelet therapies

- **Secondary prevention in CV patient @ risk**

Effect of Aspirin + Clopidogrel on vascular events :

**CHARISMA trial :**



## Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**

In contrast to the coronary field, there is **no evidence** for the optimal anti-platelet therapy and duration after peripheral interventions.

Decision is based on **“good clinical practice”**, mainly derived from the coronaries or device studies.

# Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>
Antiplatelet therapy with aspirin is recommended in all patients with angioplasty for LEAD to reduce the risk of systemic vascular events.	I	C
Dual antiplatelet therapy with aspirin and a thienopyridine for at least one month is recommended after infrainguinal bare-metal-stent implantation.	I	C

## Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**

**MIRROR : 1yr results RCT post endo R/ for PAD**

**80 patients, 1 : 1 randomization DAPT vs ASA mono**

**6 months : significant in favor of DAPT for TLR**

**12 months : no significant difference in TLR anymore**



# Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**



For patients undergoing peripheral artery percutaneous transluminal angioplasty with stenting, *we suggest single rather than dual antiplatelet therapy*

# Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**

**Dual antiplatelet therapy can be considered current state of the art after infrainguinal stent implantation**

because it is ...

- \* supported by a multitude of data from the coronary field
- \* applied in nearly all major endovascular centers and ongoing stenting trials and
- \* propagated by key opinion leaders all over the world

# Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**

## POBA / PTA-DCB / BMS

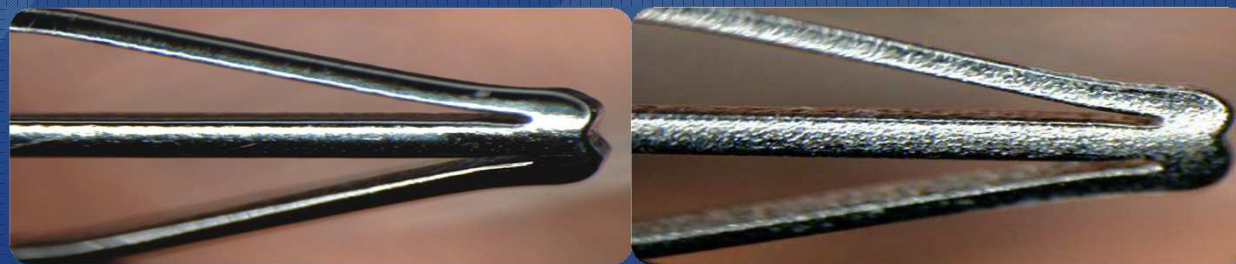


*DAPT 1 month, thereafter ASA lifelong*

## Role of Anti-platelet therapies

- Secondary prevention post (endo) vascular R/

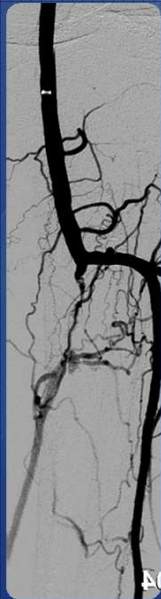
### DES ATK



*DAPT  $\geq$  2 month, thereafter ASA lifelong*

# Role of Anti-platelet therapies

- **Secondary prevention post (endo) vascular R/**



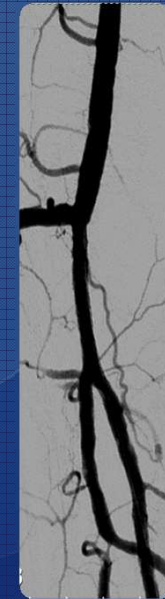
## DES BTK

3 randomized studies:

Achilles (Cypher - Sirolimus): DAPT **6** months

Destiny (Xcience V - Everolimus): DAPT **12** months

Yukon (Sirolimus): DAPT **6** months



*DAPT  $\geq$  6 month, thereafter ASA lifelong*

## Role of Anti-platelet therapies

- Secondary prevention post (endo) vascular R/

### Covered stents PAD



*DAPT  $\geq$  6 month, thereafter ASA lifelong*

## Conclusion

- **Only Aspirin** is proven to play an important role in **primary prevention** of CVD, especially in higher risk patients
- There is a **level Ia evidence** to recommend **Aspirin** as a **secondary prevention** in known CV-patients
- **Longer term DAPT** offers only benefit in **secondary prevention** in severe CV disease
- There is weaker evidence to recommend **DAPT post (endo)vascular intervention** as **secondary prevention**.

## Conclusion

- Based on world-wide (KOL) experiences and coronary/peripheral device trials, **DAPT** is recommended for 1 (POBA; DCB; BMS) up to 6-12 months (DES; Covered stents).

