



Stents and anticoagulation therapy

Malcolm R. Bell, MBBS, FRACP, FACC
Vice Chair, Department of Cardiovascular Medicine
Mayo Clinic, Rochester MN, USA

Conflicts and disclosures – none



+





Oh no!
Now what?
Triple therapy?



Pre-PCI



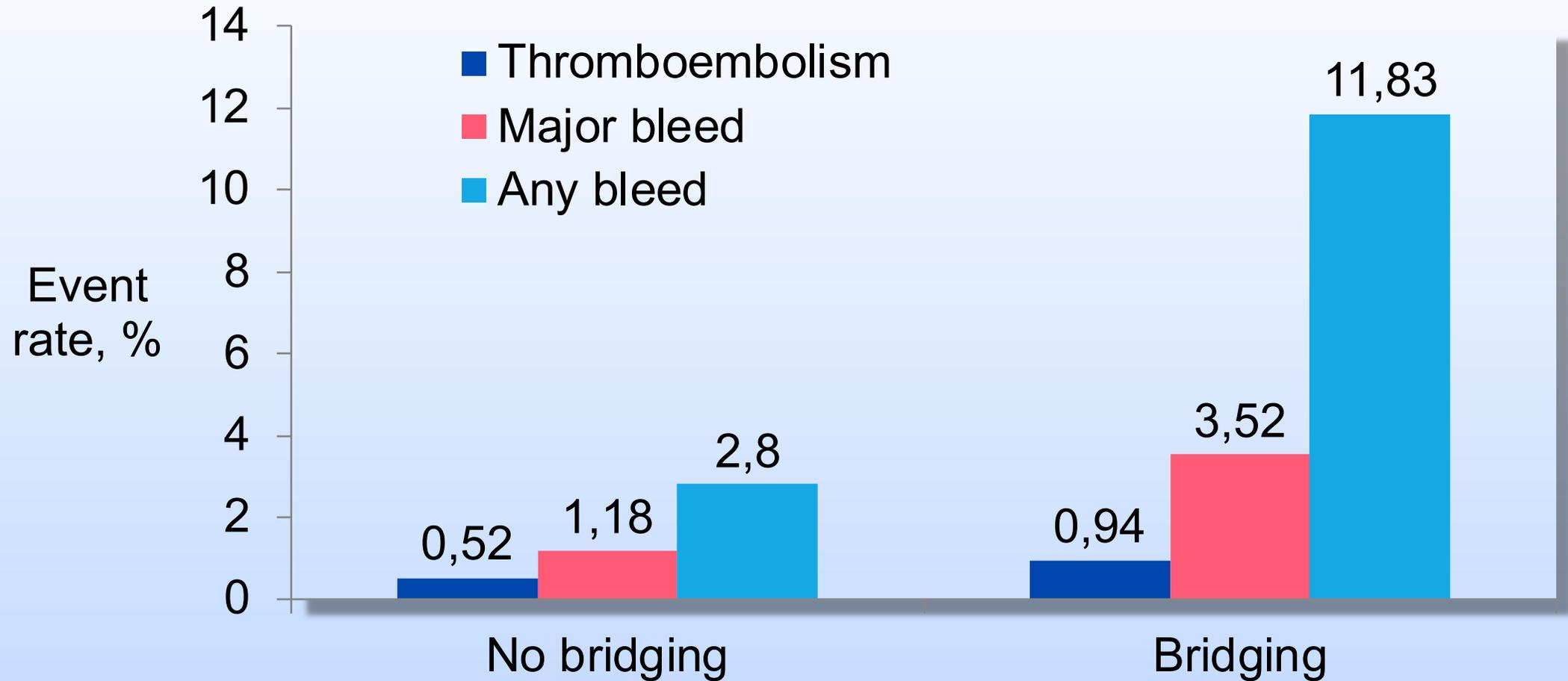
Bridging with heparin is over-utilized and highly variable

Bleeding 13x more common and bigger threat than clotting

No bridging for most AF

OAC interruptions unnecessary for most endovascular procedures

Periprocedural complications (pooled data)



PCI procedure

Radial access for PCI



Feasible and safe with therapeutic INR
Uninterrupted DOACs – need more data

Choice of stent should no longer be a question:

Nothing to lose with DES and
which are safer and more effective than BMS

DES should be default stent

Post PCI

Timeline of contemporary RCTs

2013 – WOEST trial (63% DES)

2016 – PIONEER AF-PCI trial (63% DES)

2017 – RE-DUAL PCI trial (83% DES)

2018.....more ongoing

All showed
substantial decrease
in bleeding compared
to triple therapy

Use of clopidogrel with or without aspirin in patients taking oral anticoagulant therapy and undergoing percutaneous coronary intervention: an open-label, randomised, controlled trial



Willem J M Dewilde, Tom Oirbans, Freek W A Verheugt, Johannes C Kelder, Bart J G L De Smet, Jean-Paul Herrman, Tom Adriaenssens, Mathias Vrolix, Antonius A C M Heestermans, Marije M Vis, Jan G P Tijssen, Arnoud W van 't Hof, Jurriën M ten Berg, for the WOEST study investigators

Summary

Background If percutaneous coronary intervention (PCI) is required in patients taking oral anticoagulants, antiplatelet therapy with aspirin and clopidogrel is indicated, but such triple therapy increases the risk of serious bleeding. We investigated the safety and efficacy of clopidogrel alone compared with clopidogrel plus aspirin.

Lancet 2013; 381: 1107-15

Published Online
February 13, 2013
<http://dx.doi.org/10.1016/>

Clopidogrel without aspirin associated with significant reduction in bleeding complications and no increase in rate of thrombotic events

therapy (hazard ratio [HR] 0·36, 95% CI 0·26–0·50, $p < 0·0001$). In the double-therapy group, six (2·2%) patients had multiple bleeding events, compared with 34 (12·0%) in the triple-therapy group. 11 (3·9%) patients receiving double therapy required at least one blood transfusion, compared with 27 (9·5%) patients in the triple-therapy group (odds ratio from Kaplan-Meier curve 0·39, 95% CI 0·17–0·84, $p = 0·011$).

Interpretation Use of clopidogrel without aspirin was associated with a significant reduction in bleeding complications and no increase in the rate of thrombotic events.

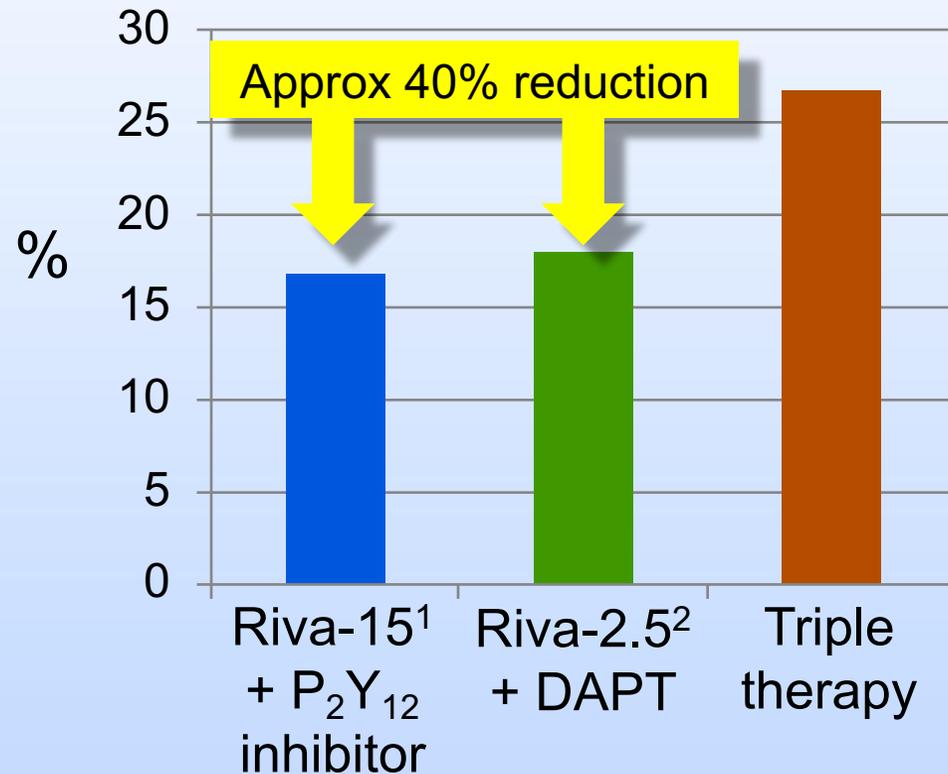
Funding Antonius Ziekenhuis Foundation, Strect Foundation.

Department of Cardiology, Onze Lieve Vrouwe Gasthuis (OLVG), Amsterdam, Netherlands (Prof F W A Verheugt MD, J-P Herrman MD); Department of Cardiology, University Medical Center Groningen, Groningen and Meander Hospital, Amersfoort, Netherlands (B J G L De Smet MD); Department of Cardiology, Catholic University of Leuven

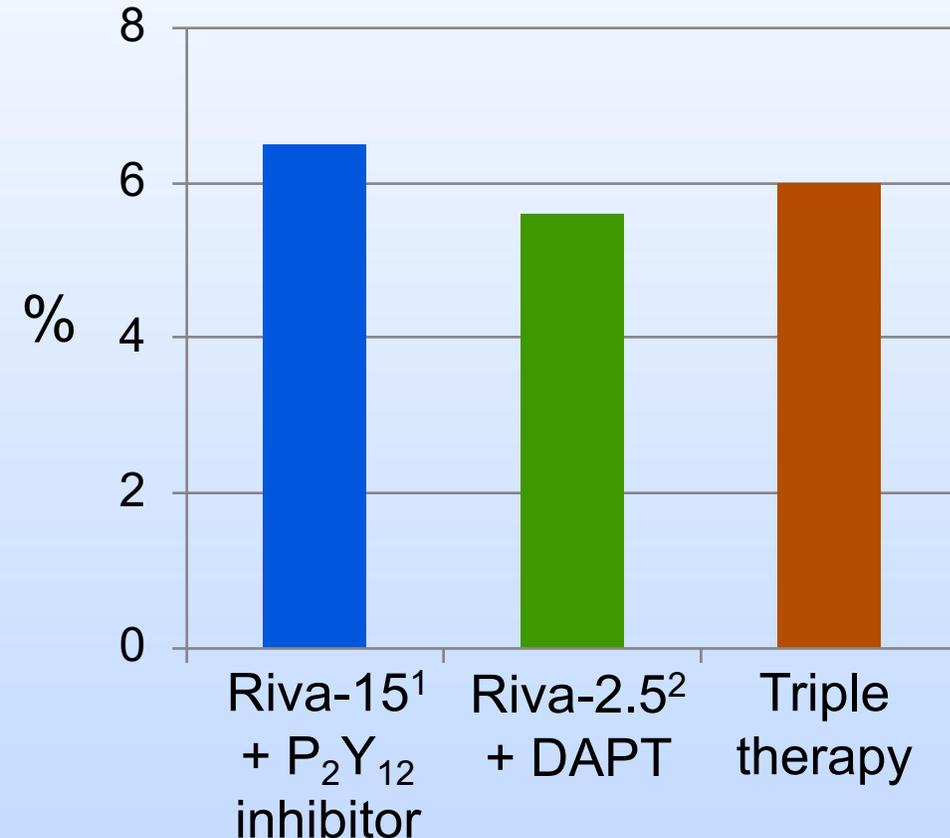
Dewilde WJM: Lancet 381:1107, 2013

PIONEER-AF trial

Clinically significant bleed



CV death, MI, CVA



The NEW ENGLAND JOURNAL *of* MEDICINE

ESTABLISHED IN 1812

OCTOBER 19, 2017

VOL. 377 NO. 16

Dual Antithrombotic Therapy with Dabigatran after PCI in Atrial Fibrillation

Patients with AF who had PCI, risk of major bleeding was significantly lower with dual therapy (dabigatran and a P₂Y₁₂ inhibitor) compared to triple therapy. Adverse ischemic events were similar.

2725 patients
AF (CHA₂DS₂-VASc 3.6)
PCI (DES >82%)

981 pts
Dabigatran 110-mg
P₂Y₁₂ inhibitor

“Dual 110-mg”

763 pts*
Dabigatran 150-mg
P₂Y₁₂ inhibitor

“Dual 150-mg”

*Excluded elderly pts
outside USA

981 pts
Warfarin, P₂Y₁₂ inhibitor
and ASA

“Triple therapy”

RE-DUAL Results at 14 Months

Endpoint	Dual 110-mg	Dual 150-mg	Triple therapy	Non inferior
<i>Primary:</i> Major bleeding	15.4%	20.2%	26.9%	Yes
<i>Efficacy:</i> Composite*	13.7%		13.4%	Yes
CVA	1.7%	1.2%	1.2%	
Def stent thrombosis	1.5%	1.0%	0.9%	

*Thromboembolic event, death or unplanned revascularization

Cannon CP: NEJM 2017

Commentary on trials of OAC after PCI

Dropping ASA results in significant decrease in major bleeding

No signal of increase in ischemic events

Majority were on clopidogrel, not ticagrelor

Caveats:

- No triple therapy using DOAC

- No dual therapy with warfarin

- No data for apixaban

3 Danish Nation-wide Registry studies

Atrial fibrillation and CAD

Triple therapy after MI or PCI:
Immediate high risk of fatal and non fatal bleeding, higher at any time than other combinations
No safe window
No diff in thromboembolic risk

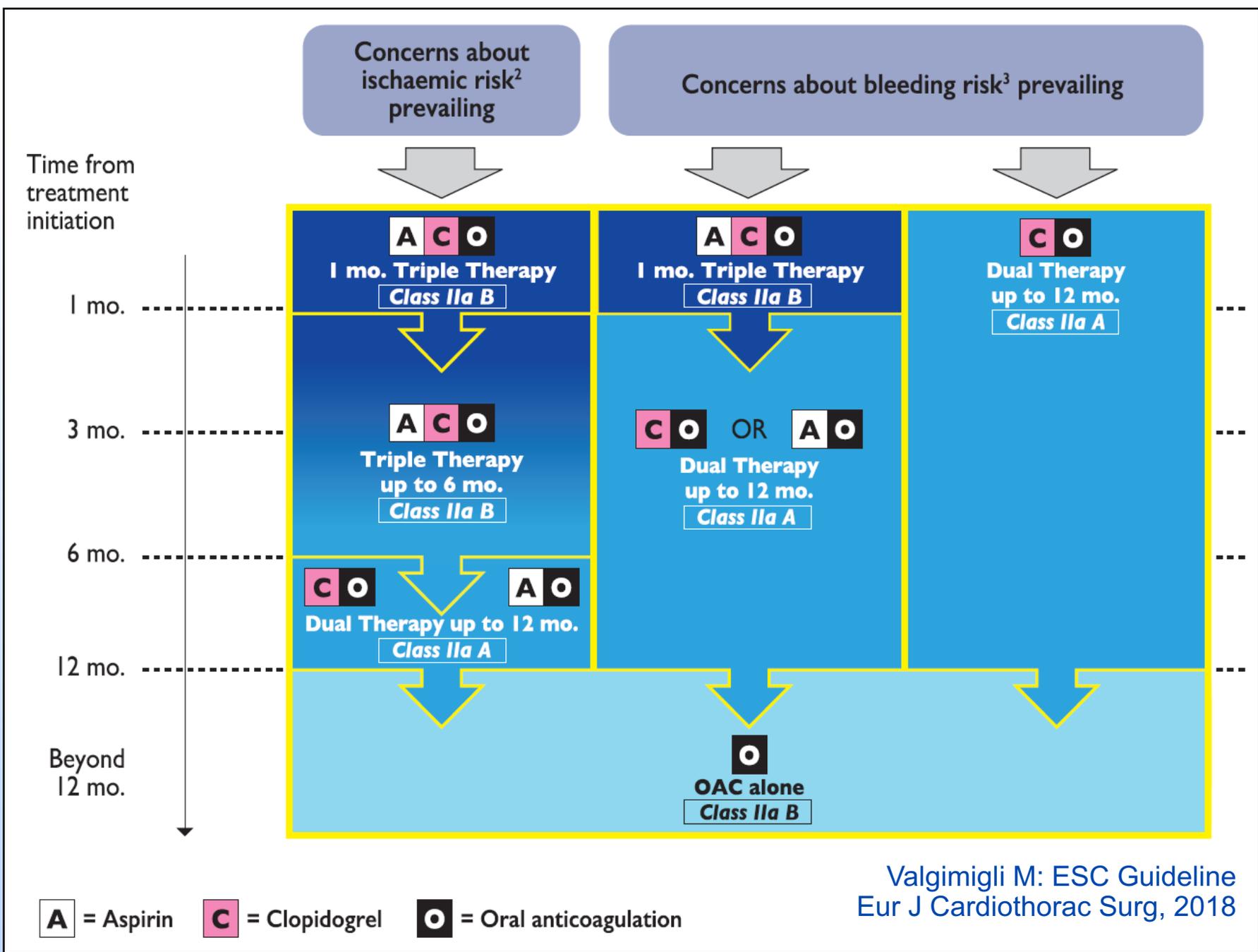
In stable CAD, no combination was better than OAC alone which also had lowest bleeding rate over 3+ yr

12,165 patients on OAC 1 year after MI and PCI

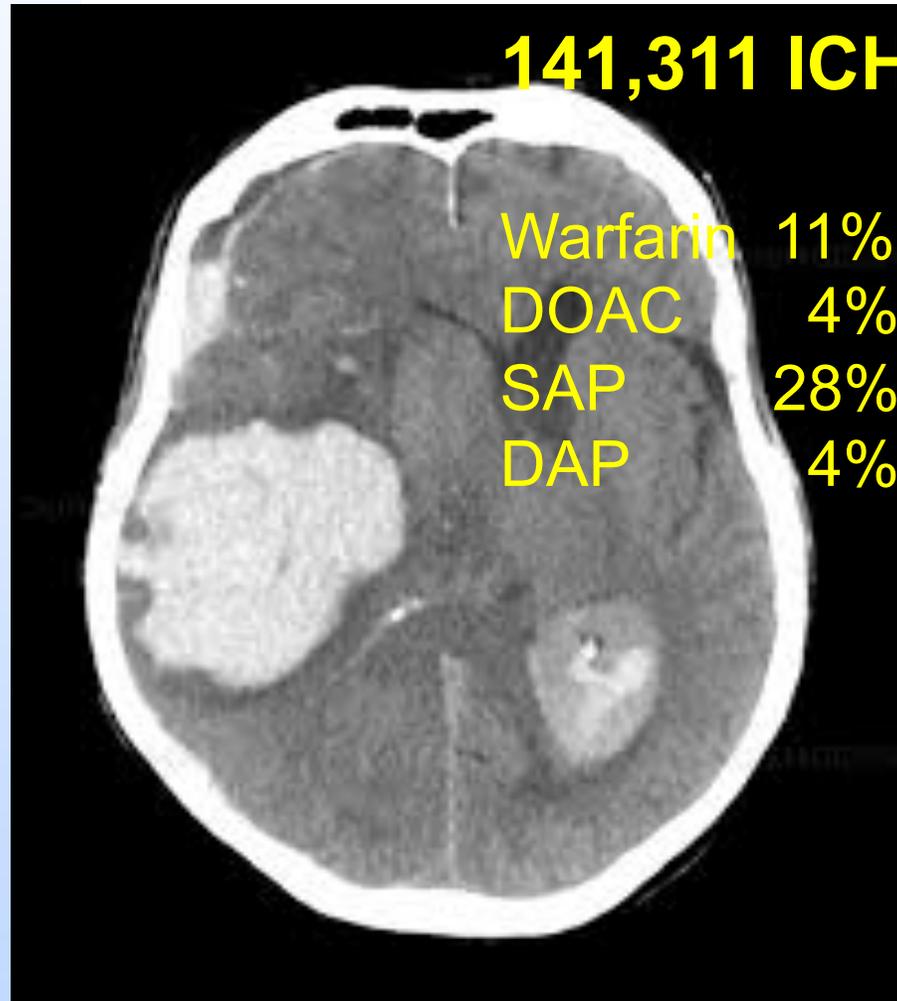


OAC and clopidogrel:
Best and safest combination with lowest all-cause mortality

Lamberts M in: Circ 2012; JACC 2013; Circ 2014



Association of Intracerebral Hemorrhage Among Patients Taking Non-Vitamin K Antagonist vs Vitamin K Antagonist Oral Anticoagulants With In-Hospital Mortality



tsouaka, PhD; Jeffrey L. Saver, MD;
Adrian F. Hernandez, MD, MS;
w, MD

agulators (NOACs) are
are limited data on

Mortality

study of 141 311 patients
1662 Get With

any use of OACs within 7 days

[+ Supplemental content](#)



Influence of Antiplatelet therapy on In-hospital Mortality

	Died	Adjusted risk difference	Adjusted OR
Warfarin			
No antiplatelet agent	32%	Ref	Ref
Single antiplatelet agent	33%	3.2%	1.2
Dual antiplatelet agents	47%	16.5%	2.1

Influence of Antiplatelet therapy on In-hospital Mortality

	Died	Adjusted risk difference	Adjusted OR
Warfarin			
No antiplatelet agent	32%	Ref	Ref
Single antiplatelet agent	33%	3.2%	1.2
Dual antiplatelet agents	47%	16.5%	2.1
DOAC			
No antiplatelet agent	26%	Ref	Ref
Single antiplatelet agent	26%	1.4%	1.1
Dual antiplatelet agents	33%	7%	1.4

Conclusions

Risk-benefit of triple therapy should be seriously questioned

Dropping ASA results in:

- Significant decrease in major bleeding

- No signal of increase in ischemic events

Majority were on clopidogrel

Ideal combination and duration not known

Consider use of OAC alone long-term



bell.malcolm@mayo.edu