CARDIOVASCULAR INNOVATIONS DIGITAL 2020

Updates in Post PCI Pharmacology Roxana Mehran, MD, FACC, FAHA, MSCAI, FESC

Mount Sinai Professor Of Cardiovascular Clinical Research and Outcomes Professor of Medicine (Cardiology), and Population Health Science and Policy Director of Interventional Cardiovascular Research and Clinical Trials, Icahn School of Medicine at Mount Sinai, New York, NY, USA

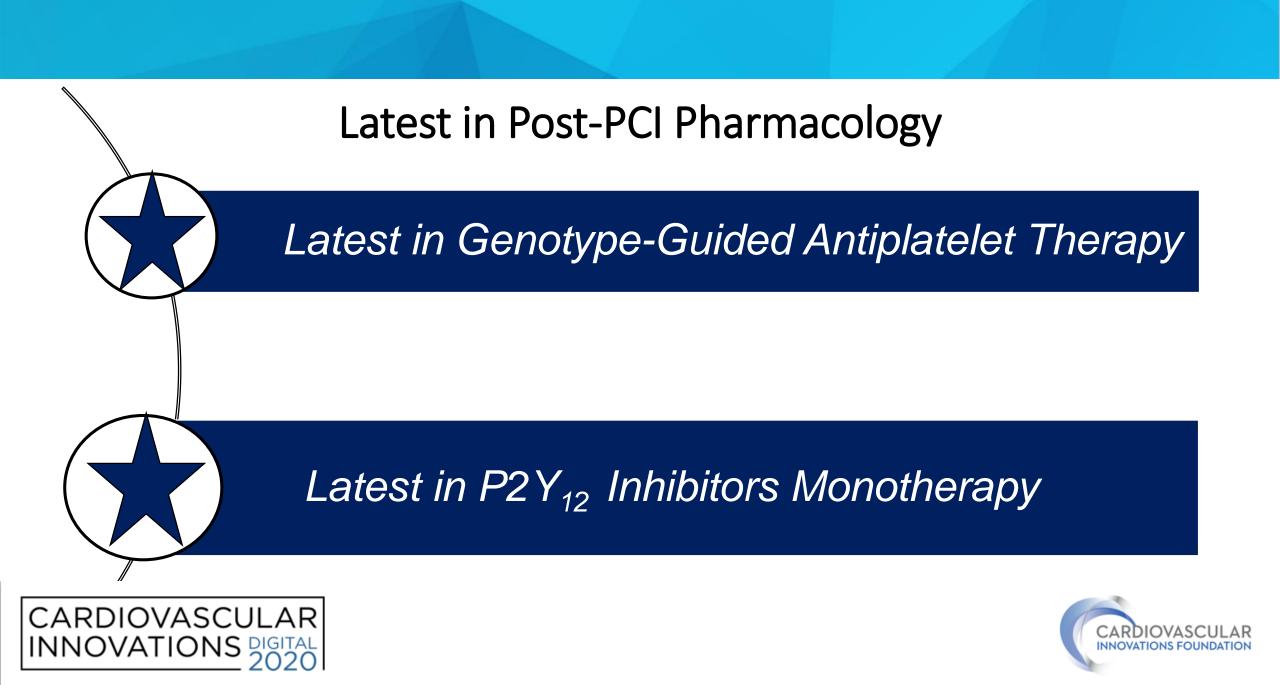


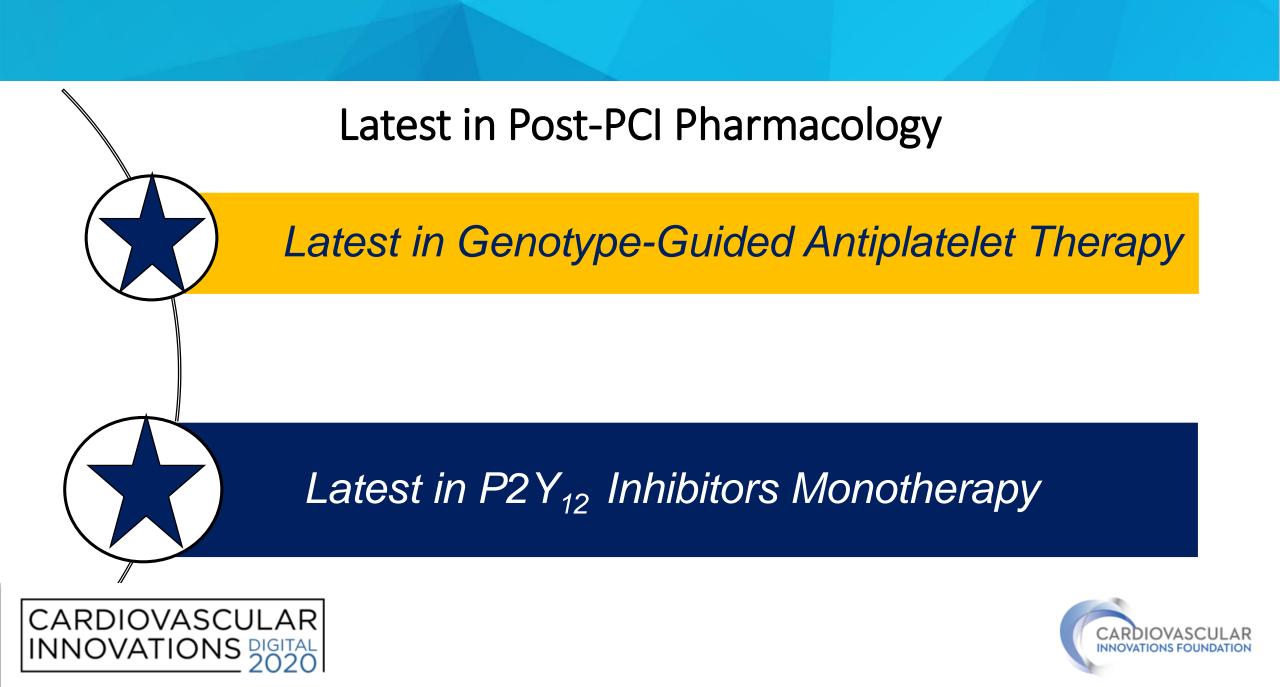
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Consultant/Advisory/Speaking Engagements	Boston Scientific, Medscape, Siemens Medical Solutions, Roivant Sciences Inc, Sanofi, Janssen, Abbott Laboratories, Medtelligence (Janssen Scientific Affairs), Idorsia Pharmaceuticals Ltd. (no fee), Regeneron Pharmaceuticals (no fee), Abiomed (spouse), The Medicines Company (spouse), Abbott Laboratories (to institution), Spectranetics/Philips/Volcano Corp (to institution)
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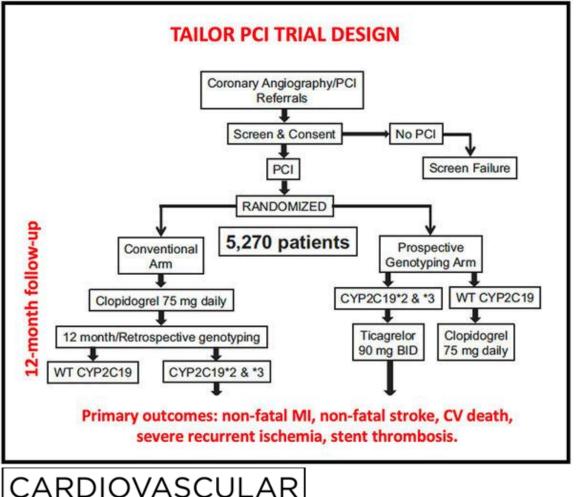








TAILOR-PCI: Genotype-guided Antiplatelet Therapy Post PCI



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Background:

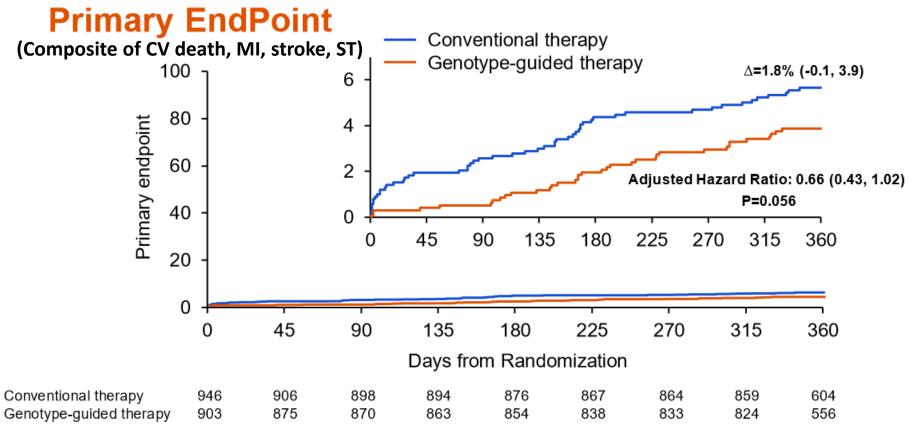
Current clinical guidelines do not recommend testing patients for the CYP2C19 genetic abnormality before prescribing clopidogrel.

Hypothesis:

Guiding the choice of post-PCI DAPT according to CYP2C19 loss-of-function (LOF) status will improve outcomes in CYP2C19 LOF carriers versus prescribing clopidogrel for all.



TAILOR-PCI: Genotype-guided Antiplatelet Therapy Post PCI

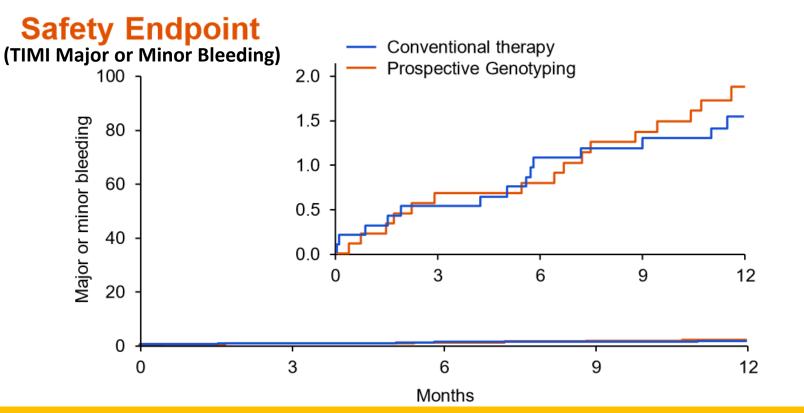






Pereira NL et al., ACC 2020 Virtual Session.

TAILOR-PCI: Genotype-guided Antiplatelet Therapy Post PCI

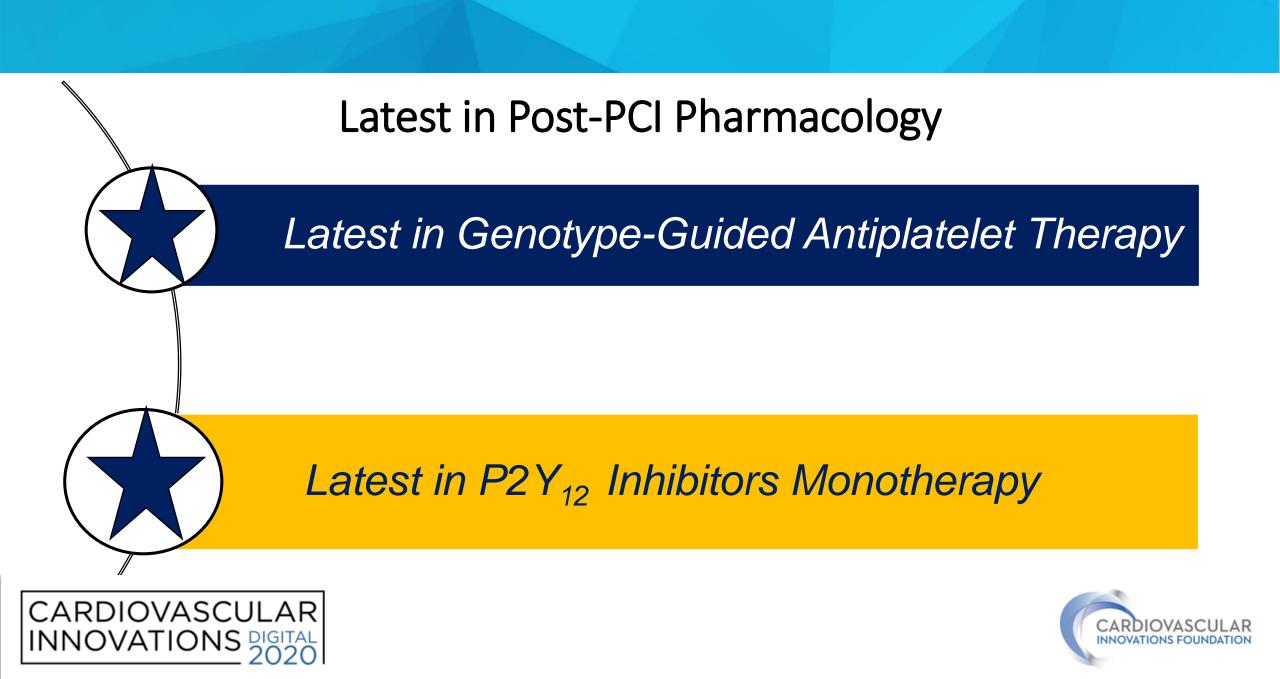


Bleeding rates were similar between the two strategies





Pereira NL et al., ACC 2020 Virtual Session.



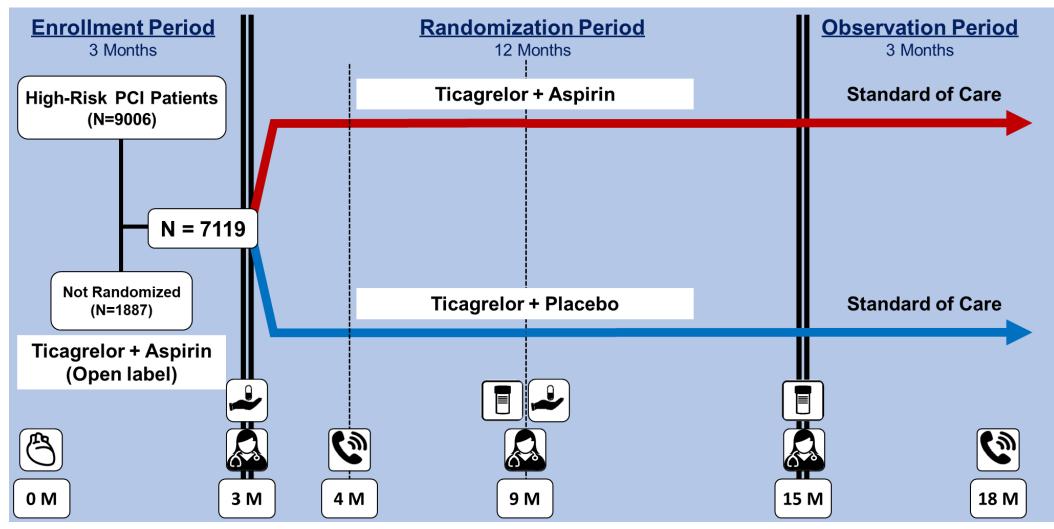
P2Y12 Monotherapy: A New Paradigm

- Many issues with Aspirin (Aspirin resistance, bleeding risk, gastrotoxicity, etc.).
- Aspirin might <u>abolish</u> or <u>decrease</u> the proposed anti-thrombotic effect of P2Y12 receptor blockers in vivo that is mediated by the amplification of anti-platelet effect of inhibitory prostaglandins such as PGI₂.
- In the presence of strong P2Y12 receptor blockade, aspirin provides little additional inhibition of platelet aggregation.





TWILIGHT Trial: Study Design



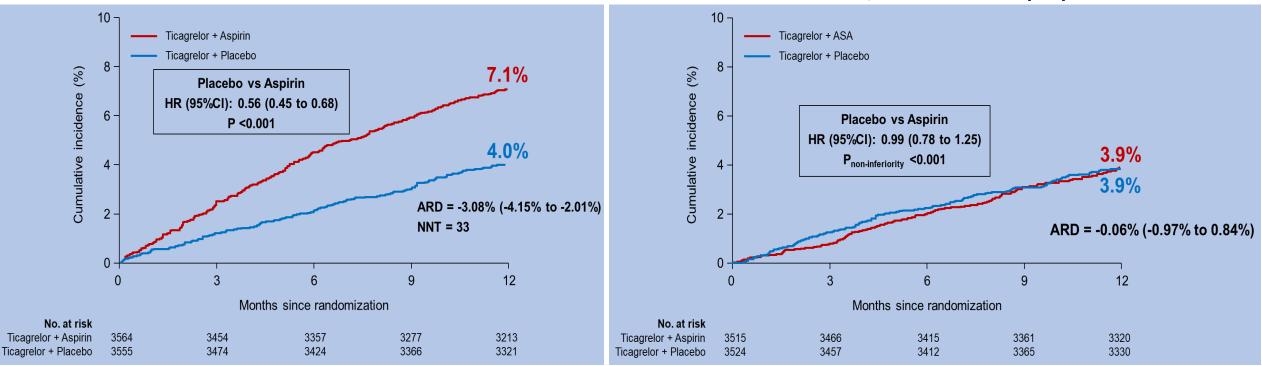
Mehran R et al., NEJM, 2019

TWILIGHT Trial: Results

Primary Endpoint BARC 2, 3 or 5 bleeding (ITT)

Key Secondary Endpoint

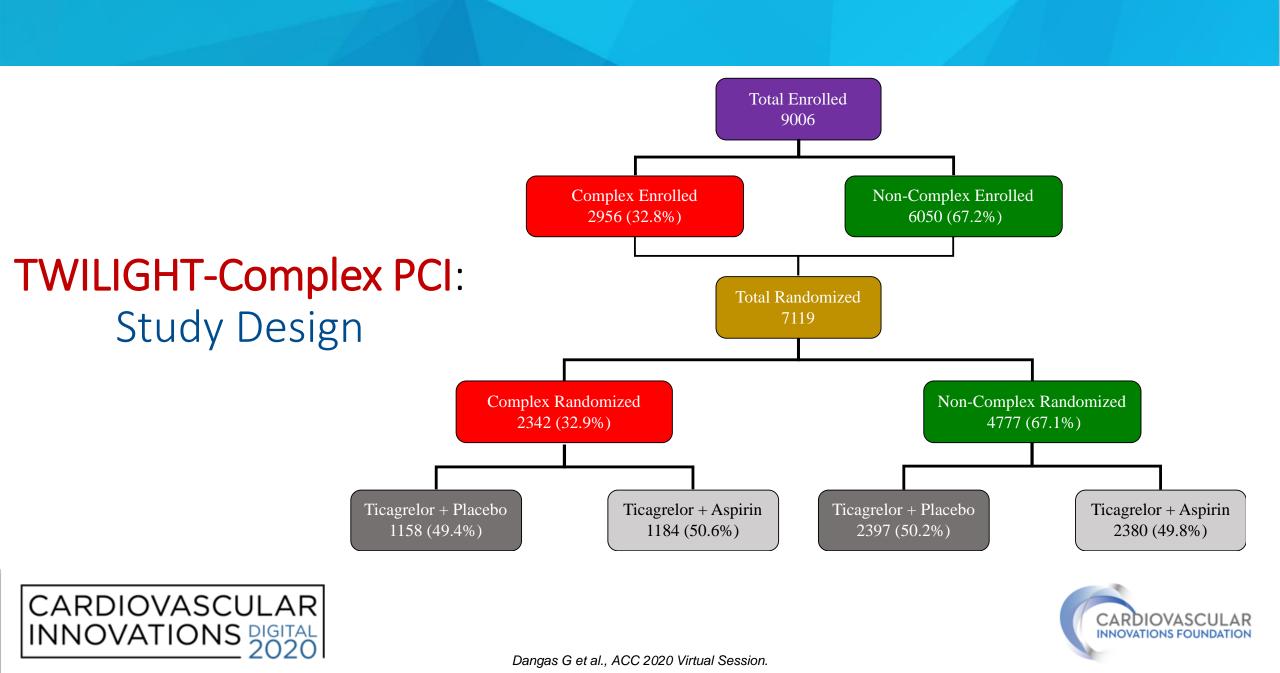
Death, MI or Stroke (PP)





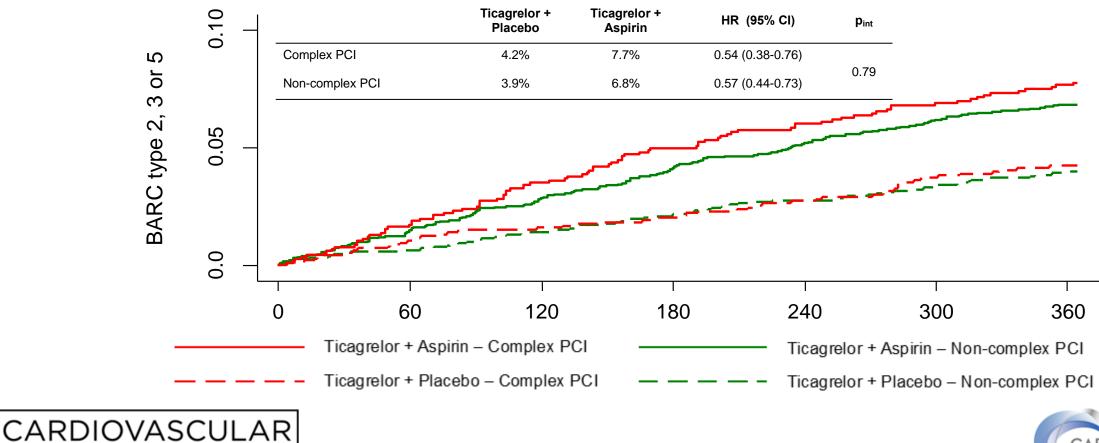


Mehran R et al., NEJM, 2019



TWILIGHT-Complex PCI: BARC 2, 3, or 5 Bleeding

Intention-To-Treat Cohort





Dangas G et al., ACC 2020 Virtual Session.

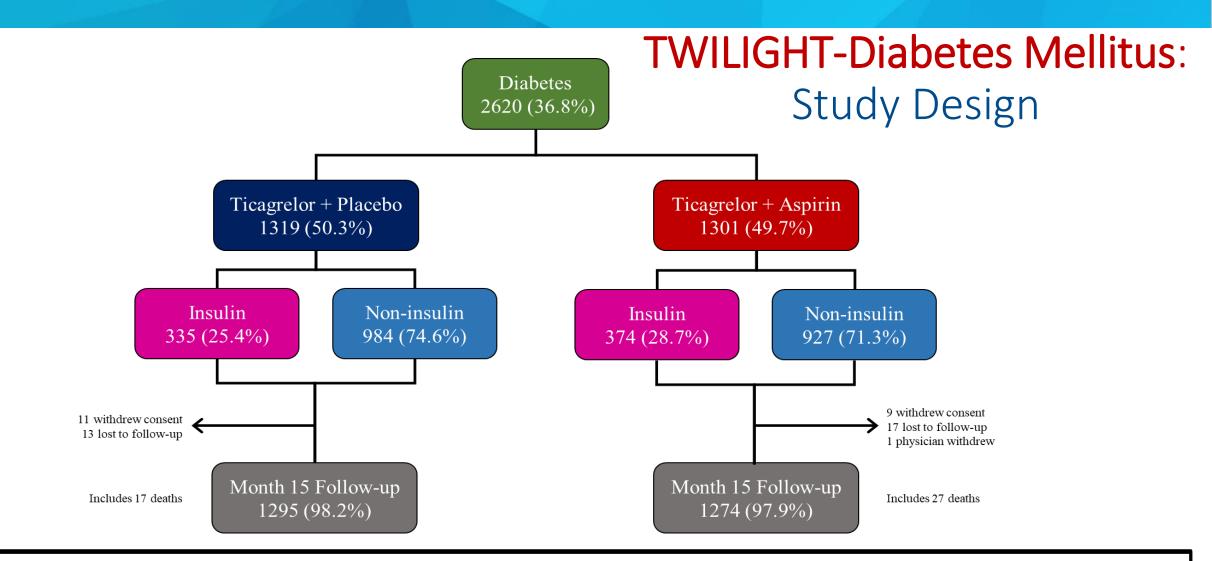
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TWILIGHT-Complex PCI: Death, MI, or Stroke

0.10 All-cause death, MI or stroke Ticagrelor + Ticagrelor + HR (95% CI) **p**int Placebo Aspirin Complex PCI 3.8% 4.9% 0.77 (0.52-1.15) 0.13 Non-complex PCI 3.9% 3.5% 1.13 (0.84-1.53) 0.05 0.0 60 120 180 240 300 360 0 Days after randomization Ticagrelor + Aspirin – Complex PCI Ticagrelor + Aspirin – Non-complex PCI Ticagrelor + Placebo - Complex PCI Ticagrelor + Placebo - Non-complex PCI CARDIOVASCULAR CARDIOVASCULAR INNOVATIONS DIGITAL INNOVATIONS FOUNDATION

Per-Protocol Cohort

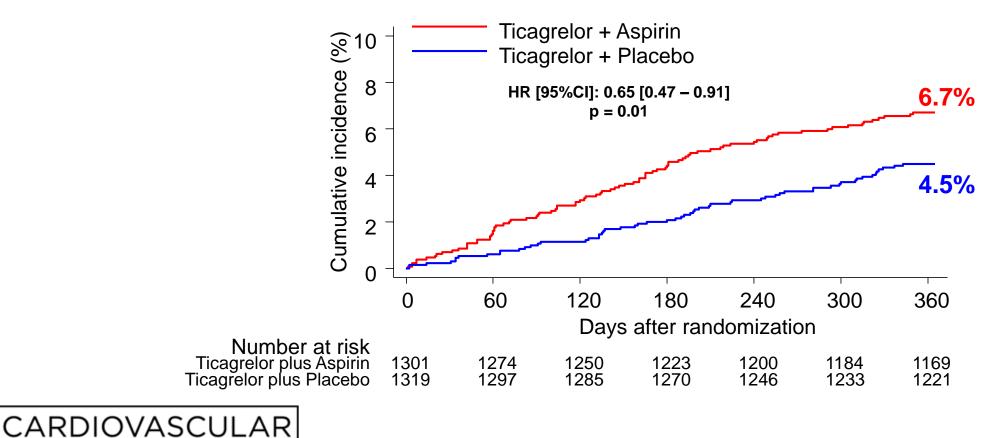
Dangas G et al., ACC 2020 Virtual Session.



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TWILIGHT-DM: BARC 2, 3, or 5 Bleeding

Intention-To-Treat Cohort



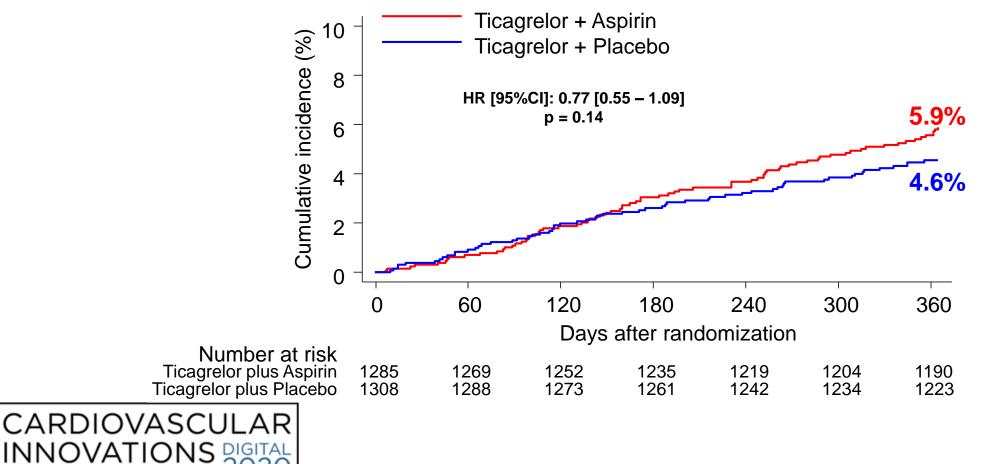


Angiolillo D et al., ACC 2020 Virtual Session.

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TWILIGHT-DM: Death, MI, or Stroke

Per-Protocol Cohort



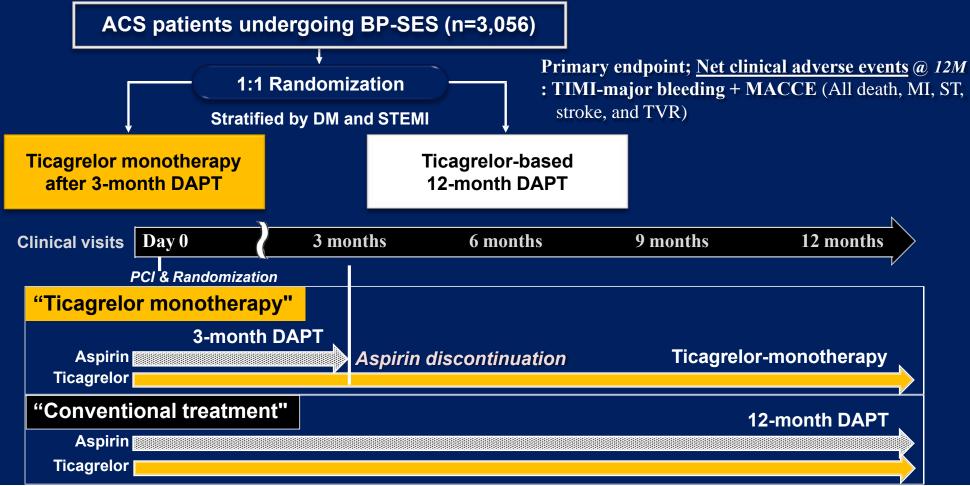


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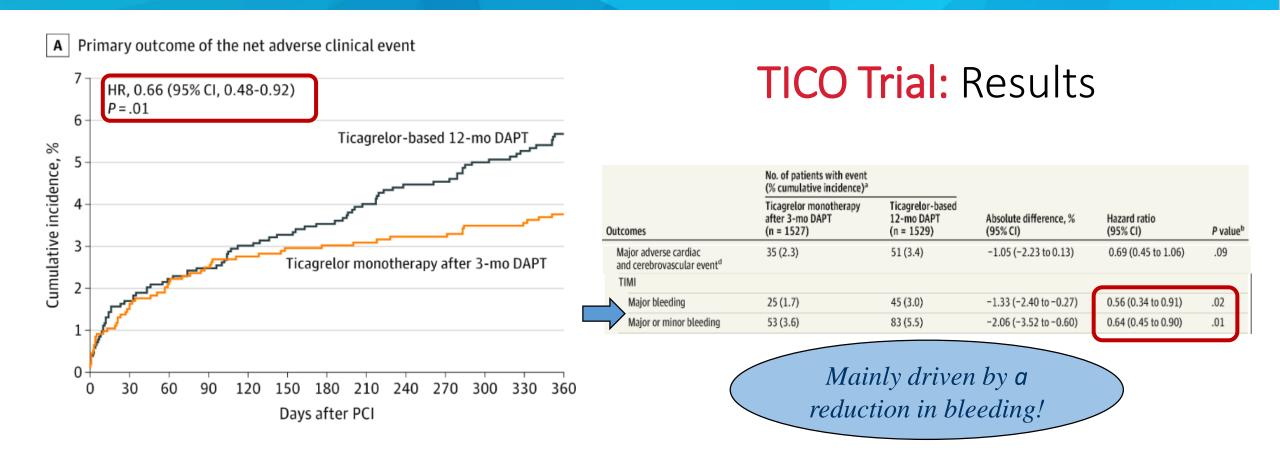
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Ticagrelor Monotherapy Vs DAPT in Patients with ACS:

TICO Randomized Trial



Kim BK et al., ACC, 2020



Among patients with ACS treated with DES, ticagrelor monotherapy after 3 months of DAPT, compared with ticagrelor-based 12month DAPT, resulted in <u>a modest but statistically significant reduction</u> in a composite outcome of major bleeding and CV events at 1 year.

Kim BK et al., JAMA, 2020

Take-Home Messages

A genotype-guided strategy was NOT superior at reducing adverse cardiovascular events compared with standard therapy after PCI.

 P2Y12 Inhibitor Monotherapy after a short course of DAPT reduces risk of bleeding without compromising ischemic protection in different subsets of high-risk patients who have undergone PCI.





Thank You!



