

STROKE AND MEDICATIONS

From Acute Care to Primary Care

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Speaker Disclosures

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- Clinical practice in Neurosciences and Mental Health Programs at KHSC

No conflicts of interest to disclose

Objectives



At the end of this session the participant will be able to:

- Apply evidence for dual-antiplatelet therapy post-ischemic stroke in the discharged patient
- Incorporate drug-related factors into monitoring of anticoagulant therapy for prevention of cardioembolic stroke
- Apply evidence for new diabetes medications to the post-ischemic stroke patient
- Evaluate risks and benefits of ASA for primary prevention of cardiovascular disease

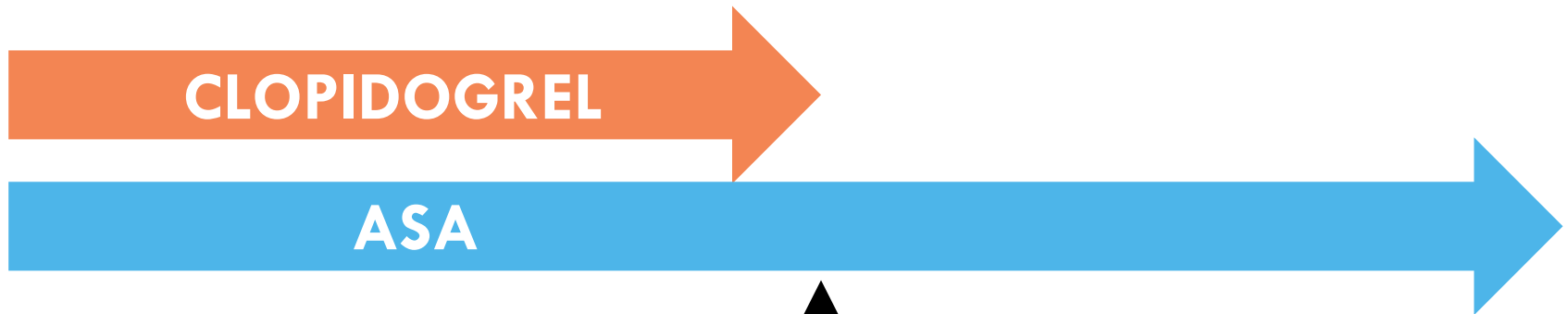


Dual Antiplatelet Therapy

Who, How Long, and Why

Dual Antiplatelets

High-risk TIA/Minor Stroke



**Day 21- 30
Post-Event**

Early DAPT prevents
recurrent events
(NNT 29 – 56)^{1,2}

Late DAPT increases risk
of major hemorrhage
(NNH = 200)²

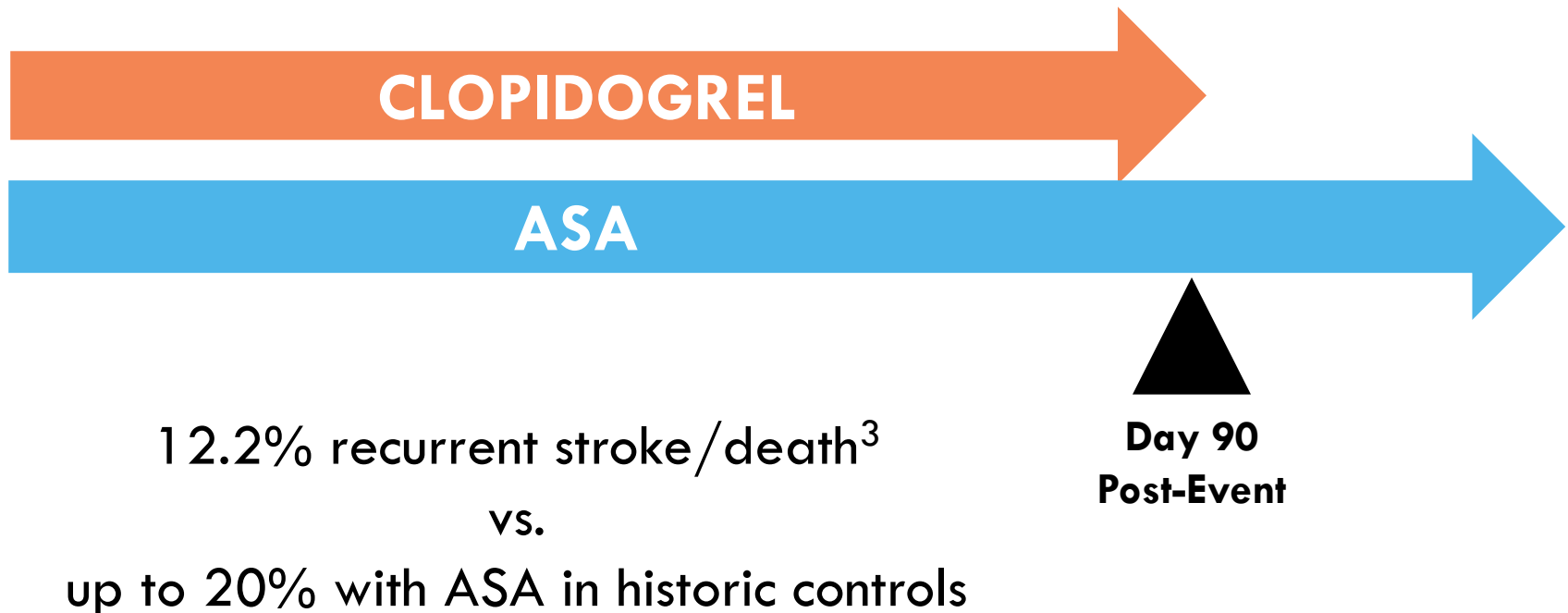
¹CHANCE Trial, 2013

²POINT Trial, 2018

Dual Antiplatelets

Intracranial Atherosclerotic Stenosis

- 70 – 99% stenosis without surgical intervention





Anticoagulants

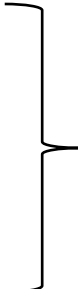
Maximizing Benefit over Risk

Anticoagulants

Meds and bleed risk

- Antiplatelets: stop UNLESS compelling indication
- NSAIDs: consider alternate analgesics, dose reduction, change to cox-2 selective, addition of PPI

Meds and fall risk

- Sedatives/hypnotics
 - Gabapentinoids
 - Anticholinergics
 - Antipsychotics
- 
- Avoid starting
 - Monitor use
 - Lowest effective dose
 - Taper off if risks > benefits

Anticoagulants

Eating and Swallowing

- Dabigatran bioavailability increased by 30 – 80% if capsules opened
- Rivaroxaban bioavailability reduced by 1/3 in fasted state

Adherence

- Once daily: rivaroxaban, edoxaban
- Cost: 80 – 120\$/month for DOACs vs. 10\$/month for warfarin



Diabetes

New(er) Kids on the Block

SGLT2 Inhibitors

Empagliflozin

EMPA-REG, 2015

Canagliflozin

CANVAS, 2017

Dapagliflozin

DECLARE TIMI 58, 2019

GLP-1 Agonists

Liraglutide
LEADER, 2016

Semaglutide
SUSTAIN 6, 2017

Dulaglutide
REWIND, 2019

SGLT2 & GLP1 Summary

Reasonable choices for post-stroke patients needing additional control

Some caveats:

- Stroke patients less represented vs. heart patients
 - Consider cardiovascular comorbidity + overall risk
- Benefits of SGLT2 inhibitors driven by decreased MI, HF hospitalization
- Affordability, tolerability



ASA for Primary Prevention

What's new since 2018?

ASA for Primary Prevention

2019 ACC/AHA Guidelines:

*“...recent studies have shown that **in the modern era, aspirin should not be used in the routine primary prevention of ASCVD due to lack of net benefit.** Most important is to avoid aspirin in persons with increased risk of bleeding...”*



ASA for Primary Prevention

ARRIVE, 2019

Adults with moderate CV Risk (10-20% over 10 years)

No difference in symptomatic CV events or mortality

More GI bleeds
NNH = 196

ASCEND, 2018

Diabetics over age 40

Decreased symptomatic CV events
NNT = 91

More major bleeds
NNH = 111

ASPREE, 2018

Adults over age 70 living at home

No difference in symptomatic CV events or disability-free survival

More major bleeds
NNH = 100
Higher mortality
NNH = 142

ASA for Primary Prevention

Any patients under-represented in trials who may benefit?

- Young, low bleeding risk
- Multiple CV risk factors
- Difficulty optimizing lipids, BP, smoking status
- Known asymptomatic atherosclerosis

References

- 1) Clopidogrel with Aspirin in Acute Minor Stroke or Transient Ischemic Attack. Yongjun Wang, Yilong Wang, Xingquan Zhao, Liping Liu, et al. July 4, 2013. N Engl J Med 2013; 369:11-19 DOI: 10.1056/NEJMoa1215340
- 2) Clopidogrel and Aspirin in Acute Ischemic Stroke and High-Risk TIA. S. Claiborne Johnston, J. Donald Easton, Mary Farrant, William Barsan, et al. July 19, 2018 N Engl J Med 2018; 379:215-225 DOI: 10.1056/NEJMoa1800410
- 3) Stenting versus Aggressive Medical Therapy for Intracranial Arterial Stenosis. Marc I. Chimowitz, Michael J. Lynn, Colin P. Derdeyn, Tanya N. Turan, et al. September 15, 2011 N Engl J Med 2011; 365:993-1003 DOI: 10.1056/NEJMoa1105335
- 4) Empagliflozin, Cardiovascular Outcomes, and Mortality in Type 2 Diabetes. Bernard Zinman, Christoph Wanner, John M. Lachin, David Fitchett, et al. November 26, 2015. N Engl J Med 2015; 373:2117-2128 DOI: 10.1056/NEJMoa1504720
- 5) Canagliflozin and Cardiovascular and Renal Events in Type 2 Diabetes. Bruce Neal, Vlado Perkovic, Kenneth W. Mahaffey, Dick de Zeeuw, et al. August 17, 2017. N Engl J Med 2017; 377:644-657 DOI: 10.1056/NEJMoa1611925
- 6) Dapagliflozin and Cardiovascular Outcomes in Type 2 Diabetes. Stephen D. Wiviott, Itamar Raz, Marc P. Bonaca, Ofri Mosenzon, et al. January 24, 2019 N Engl J Med 2019; 380:347-357 DOI: 10.1056/NEJMoa1812389
- 7) Liraglutide and Cardiovascular Outcomes in Type 2 Diabetes. Steven P. Marso, Gilbert H. Daniels, Kirstine Brown-Frandsen, Peter Kristensen, et al. July 28, 2016 N Engl J Med 2016; 375:311-322 DOI: 10.1056/NEJMoa1603827
- 8) Semaglutide and Cardiovascular Outcomes in Patients with Type 2 Diabetes. Steven P. Marso, Stephen C. Bain, Agostino Consoli, Freddy G. Eliaschewitz, et al. November 10, 2016 N Engl J Med 2016; 375:1834-1844 DOI: 10.1056/NEJMoa1607141
- 9) Dulaglutide and cardiovascular outcomes in type 2 diabetes (REWIND): a double-blind, randomised placebo-controlled trial. Gerstein HC, Colhoun HM, [Dagenais GR](#), Diaz R, [Lakshmanan M](#). Lancet. 2019 Jul 13;394(10193):121-130. doi: 10.1016/S0140-6736(19)31149-3.
- 10) Use of aspirin to reduce risk of initial vascular events in patients at moderate risk of cardiovascular disease (ARRIVE): a randomised, double-blind, placebo-controlled trial. Gaziano JM, Brotons C, Coppolecchia R, Crceci C, et al. [Lancet](#). 2018 Sep 22;392(10152):1036-1046. doi: 10.1016/S0140-6736(18)31924-X
- 11) Effects of Aspirin for Primary Prevention in Persons with Diabetes Mellitus. Bowman L, Mafham M, Wallendszus K, Stevens W, et al. N Engl J Med. 2018 Oct 18;379(16):1529-1539. doi: 10.1056/NEJMoa1804988
- 12) Effect of Aspirin on Cardiovascular Events and Bleeding in the Healthy Elderly. John J. McNeil, Rory Wolfe, Robyn L. Woods, Andrew M. Tonkin, et al. October 18, 2018. N Engl J Med 2018; 379:1509-1518 DOI: 10.1056/NEJMoa1805819