

AAS na prevenção primária ainda existe lugar?

IAGO REINEL DE CASTRO

MÉDICO ESPECIALISTA EM CLÍNICA MÉDICA, CARDIOLOGIA E
ECOCARDIOGRAFIA

Declaração de Conflitos de Interesses

O apresentador declara **não** apresentar conflitos de interesses que possam ser relacionados à sua apresentação.

FEATURE

FEATURE

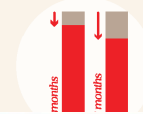
Aspirin was one of the first drugs to come into common usage and is still one of the most researched drugs in the world, with an estimated 700 to 1,000 clinical trials conducted each year

Acetylsalicylic acid is named Aspirin by Bayer. The letter 'A' stands for acetyl, 'spir' is derived from the plant known as *Spiraea ulmaria* (meadowsweet), which yields salicin, and 'in' was a common suffix used for drugs at the time of the first stable synthesis of acetylsalicylic acid.

1899

1974

Data from the first randomised controlled trial of aspirin in the secondary prevention of death from heart attack show a reduction in total mortality of 12% at 6 months and 25% at 12 months but the results are statistically significant (BMJ 1974;1:169).



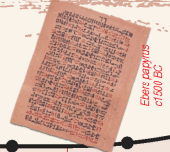
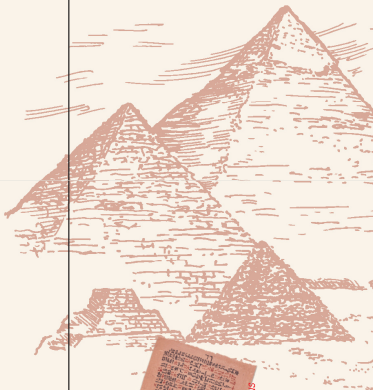
2014

A meta-analysis suggests that long-term prophylactic use of aspirin has a favourable benefit-harm profile and leads to a dramatic reduction in the incidence of bowel, stomach and oesophageal cancer (*Annals of Oncology*, online 5 August 2014).

A history of aspirin

Aspirin Evidence: Secondary Prevention

By Dawn Connelly



c3000 – 1500 BC

Willow is used as a medicine by ancient civilisations like the Sumerians and Egyptians. The Ebers papyrus, an ancient Egyptian medical text, refers to willow as an anti-inflammatory or pain reliever for non-specific aches and pains.



White Willow (Spir. alba)



Helping Cardiovascular Professionals Learn. Advance. Heal.



Hippocrates administers Willow leaf tea to a sick patient.

Salicin is also found in the meadowsweet flower by Swiss pharmacist Johann Pagenstecher and later by German researcher Karl Jacob Löwig.

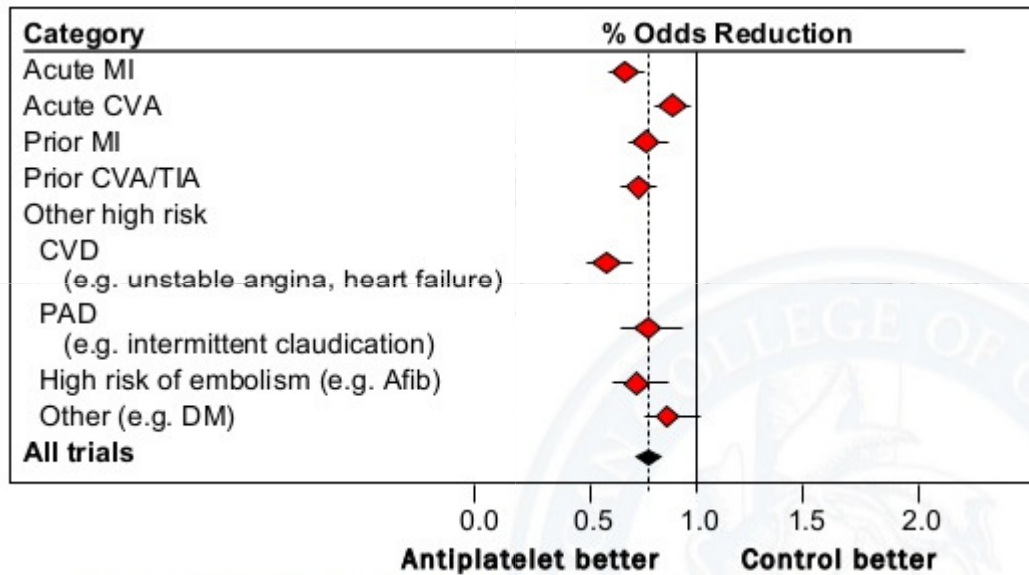


hypertensive patients, with the greatest benefit seen in preventing heart attacks. The incidence of non-fatal major bleeds was twice as common (*Lancet* 1998;351:1755).

2011

A meta-analysis of eight clinical trials finds that, after five years of follow-up, trial participants who took aspirin daily for a mean of four years have a 44% reduced risk of dying from cancer compared with participants who took a placebo (*Lancet* 2011;377:31).

Effect of antiplatelet treatment* on vascular events**



Aspirin reduces the risk of adverse cardiovascular events

*Aspirin was the predominant antiplatelet agent studied
**Include MI, stroke, or death

Source: Antithrombotic Trialists' Collaboration. *BMJ* 2002;324:71-86

- 2005**: WHS (Women's Health Study) results.
- 2009**: Results from the ARRIVE study.
- 2011**: Results from the ASPREE study.
- 2013**: Follow-up results of the WHS confirm that long-term use of alternate day low-dose aspirin results in a 42% reduction in colorectal cancer incidence, with benefits starting to appear after 10 years.
- 2015**: Results expected from the ASPREE study.
- 2018**: Results expected from the ASPREE study.

Ainda **existe** lugar na prevenção 1^a?



Ainda funcionam? Sim.

Ainda **existe lugar** na prevenção 1^a?



Ainda funcionam? Sim.
Mas ainda existe lugar?

Diretrizes



STF 2016: Adults aged 50 to 59
rs with a $\geq 10\%$ 10-year CVD risk
A ADA 2015: Reasonable for
viduals with a $\geq 10\%$ 10-year CVD
and without increased bleeding



ESC 2016: Antiplatelet therapy
is not recommended in
individuals free from CVD



I Diretriz Brasileira de Prevenção
Cardiovascular 2013

Diretrizes Brasileiras de
Antiagregantes Plaquetários e
Anticoagulantes em
Cardiologia 2013

Benefícios em IAM não fatal AVC não fatal

Study, Year (Reference)	Aspirin Dose, mg/d	Follow-up, mo	Population Description	RR (95% CI)	Events/IG
Nonfatal MI					
PPP, 2001 (38)	100	43.2	Men and women with ≥1 risk factor for CVD	0.69 (0.36–1.33)	15/2226
HOT, 1998 (34)	75	45.6	Men and women with hypertension	0.60 (0.45–0.81)	68/9399
JPAD, 2008 (35)	100	52.4	Men and women with diabetes	1.35 (0.57–3.19)	12/1262
JPPP, 2014 (39)	100	60.2	Men and women with ≥1 risk factor for CVD	0.53 (0.31–0.91)	20/7220
PHS I, 1989 (30)					
BMD, 1988 (36)					
POPADAD, 2008 (31)					
TPT, 1998 (24)					
AAA, 2010 (33)	100	98.4	Men and women with ABI ≤0.95	0.91 (0.65–1.28)	62/1675
WHS, 2005 (37)	50	121.2	Women health professionals	1.01 (0.83–1.24)	184/1993
Overall: ($I^2 = 61.9\%$; $P = 0.005$)				0.78 (0.71–0.87)	
Nonfatal stroke					
PPP, 2001 (38)	100	43.2	Men and women with ≥1 risk factor for CVD	0.84 (0.42–1.07)	15/2220
JPAD, 2008 (35)	100	52.4	Men and women with diabetes	1.01 (0.60–1.72)	27/1262
ETDRS, 1992 (32)	50	60	Men and women with diabetes and diabetic retinopathy	1.35 (0.89–1.90)	67/1855
JPPP, 2014 (39)					
PHS I, 1989 (30)					
BMD, 1988 (36)					
POPADAD, 2008 (31)					
TPT, 1998 (24)	75	81.6	Men at high risk for ischemic heart disease	0.64 (0.34–1.20)	18/1280
AAA, 2010 (33)	100	98.4	Men and women with ABI ≤0.95	0.97 (0.62–1.52)	37/1675
WHS, 2005 (37)	50	121.2	Women health professionals	0.81 (0.67–0.97)	198/1993
Overall: ($I^2 = 25.1\%$; $P = 0.212$)				0.95 (0.85–1.06)	

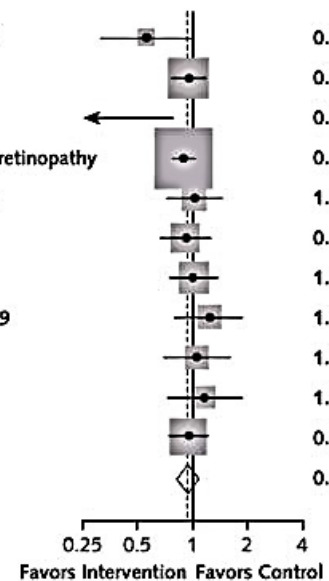
A significant 22 percent reduction in nonfatal MI (relative risk [RR] 0.78, 95% CI 0.71–0.87).

No significant benefit on nonfatal stroke (RR 0.95, 95% CI 0.85–1.06).

Benefícios em mortalidade por DCV

mortalidade total

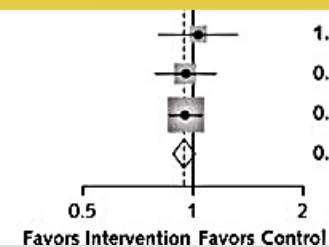
Study, Year (Reference)	Aspirin Dose, mg/d	Follow-up, mo	Population Description	RR (95% CI)	IG
CVD mortality					
PPP, 2001 (38)	100	43.2	Men and women with ≥ 1 risk factor for CVD	0.56 (0.31–1.01)	17/22
HOT, 1998 (34)	75	45.6	Men and women with hypertension	0.95 (0.75–1.20)	133/93
JPAD, 2008 (35)	100	52.4	Men and women with diabetes	0.10 (0.01–0.79)	1/126
ETDRS, 1992 (32)	650	60	Men and women with diabetes and diabetic retinopathy	0.89 (0.76–1.04)	244/18
JPPP, 2014 (39)	100	60.2	Men and women with ≥ 1 risk factor for CVD	1.02 (0.71–1.47)	58/72
PHS I, 1989 (30)	162.5	60.2	Men physicians	0.92 (0.66–1.28)	66/110
BMD, 1988 (36)	500	72	Men physicians	1.01 (0.74–1.37)	119/34
POPADAD, 2008 (31)	100	80.4	Men and women with diabetes and ABI ≤ 0.99	1.23 (0.80–1.89)	43/63
TPT, 1998 (24)	75	81.6	Men at high risk for ischemic heart disease	1.05 (0.69–1.61)	42/12
AAA, 2010 (33)	100	98.4	Men and women with ABI ≤ 0.95	1.17 (0.72–1.89)	35/16
WHS, 2005 (37)	50	121.2	Women health professionals	0.95 (0.74–1.22)	120/19
Overall: ($I^2 = 8.8\%$; $P = 0.360$)				0.94 (0.86–1.03)	



All-cause mortality					
PPP, 2001 (38)	100	43.2	Men and women with ≥ 1 risk factor for CVD	0.81 (0.58–1.13)	62/22
HOT, 1998 (34)	75	45.6	Men and women with hypertension	0.93 (0.79–1.09)	284/93
JPAD, 2008 (35)	100	52.4	Men and women with diabetes	0.91 (0.57–1.43)	34/12

A significant 6 percent reduction in all-cause mortality (RR 0.94, 95% CI 0.89-0.99)

TPT, 1998 (24)	75	81.6	Men at high risk for ischemic heart disease	1.03 (0.80–1.32)	113/12
AAA, 2010 (33)	100	98.4	Men and women with ABI ≤ 0.95	0.95 (0.78–1.15)	176/16
WHS, 2005 (37)	50	121.2	Women health professionals	0.95 (0.85–1.06)	609/19
Overall: ($I^2 = 0.0\%$; $P = 0.996$)				0.94 (0.89–0.99)	



Absolute Risk Reduction With Low-Dose Aspirin Use ≤ 10 Years

Table 2. Absolute Risk Reduction With Low-Dose Aspirin Use ≤ 10 Years

Outcome	Risk Level*	Baseline Risk for Outcome, events per 1000 person-years	RR (95% CI)†	Events Prevented per Person-Years (95% CI)
All-cause mortality (k = 8)	Low	3.19	0.95 (0.89 to 1.01)	0.16 (−0.03 to 0.35)
	Median	8.55		0.43 (−0.09 to 0.94)
	High	13.54		0.68 (−0.14 to 1.49)
CVD mortality (k = 8)	Low	0.63	0.97 (0.85 to 1.10)	0.02 (−0.06 to 0.09)
	Median	2.18		0.07 (−0.22 to 0.33)
	High	4.62		0.14 (−0.46 to 0.69)
Nonfatal stroke (k = 7)	Low	1.21	0.86 (0.76 to 0.98)	0.17 (0.02 to 0.29)
	Median	2.83		0.40 (0.06 to 0.68)
	High	4.84		0.68 (0.10 to 1.16)
Nonfatal MI (k = 8)	Low	0.90	0.83 (0.74 to 0.94)	0.15 (0.05 to 0.23)
	Median	2.69		0.46 (0.16 to 0.70)
	High	8.44		1.43 (0.51 to 2.19)

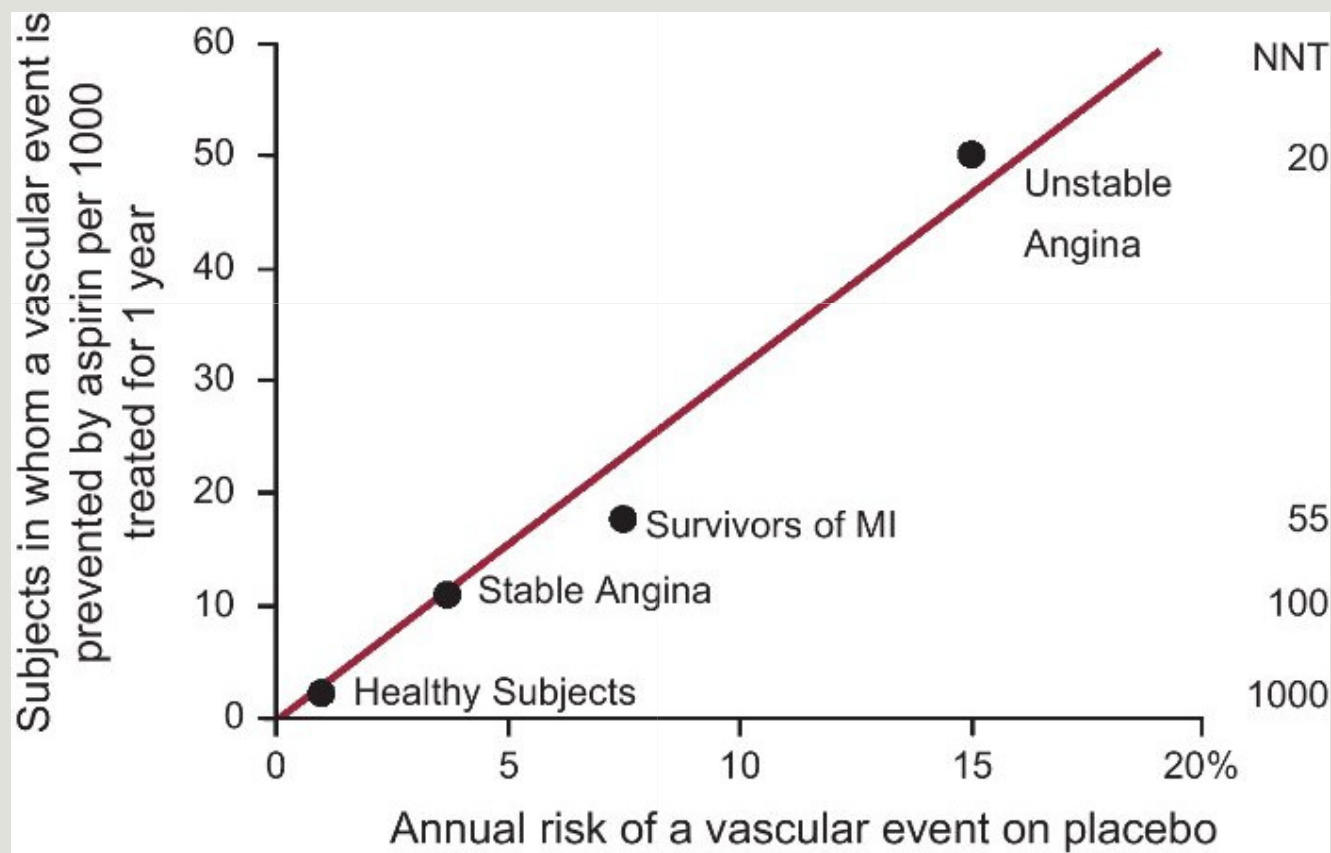
*Low, median, and high (maximum) control group rate for each outcome, excluding zeros and outliers.

†RR = relative risk.

‡Based on aspirin doses ≤ 100 mg/d in primary CVD prevention trials.

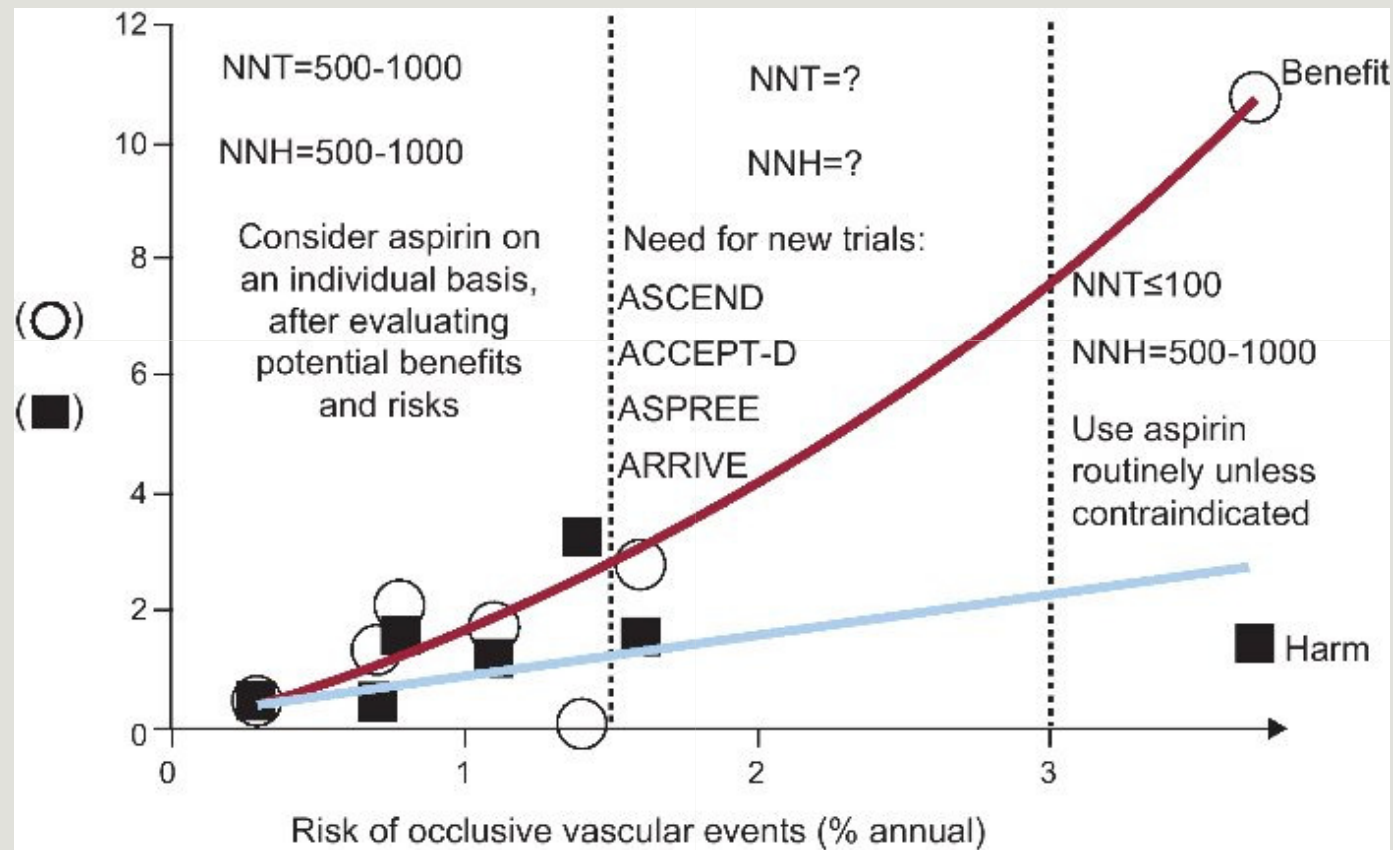
§Boldface data represent clearly prevented events (i.e., 95% CI does not include both caused and prevented events).

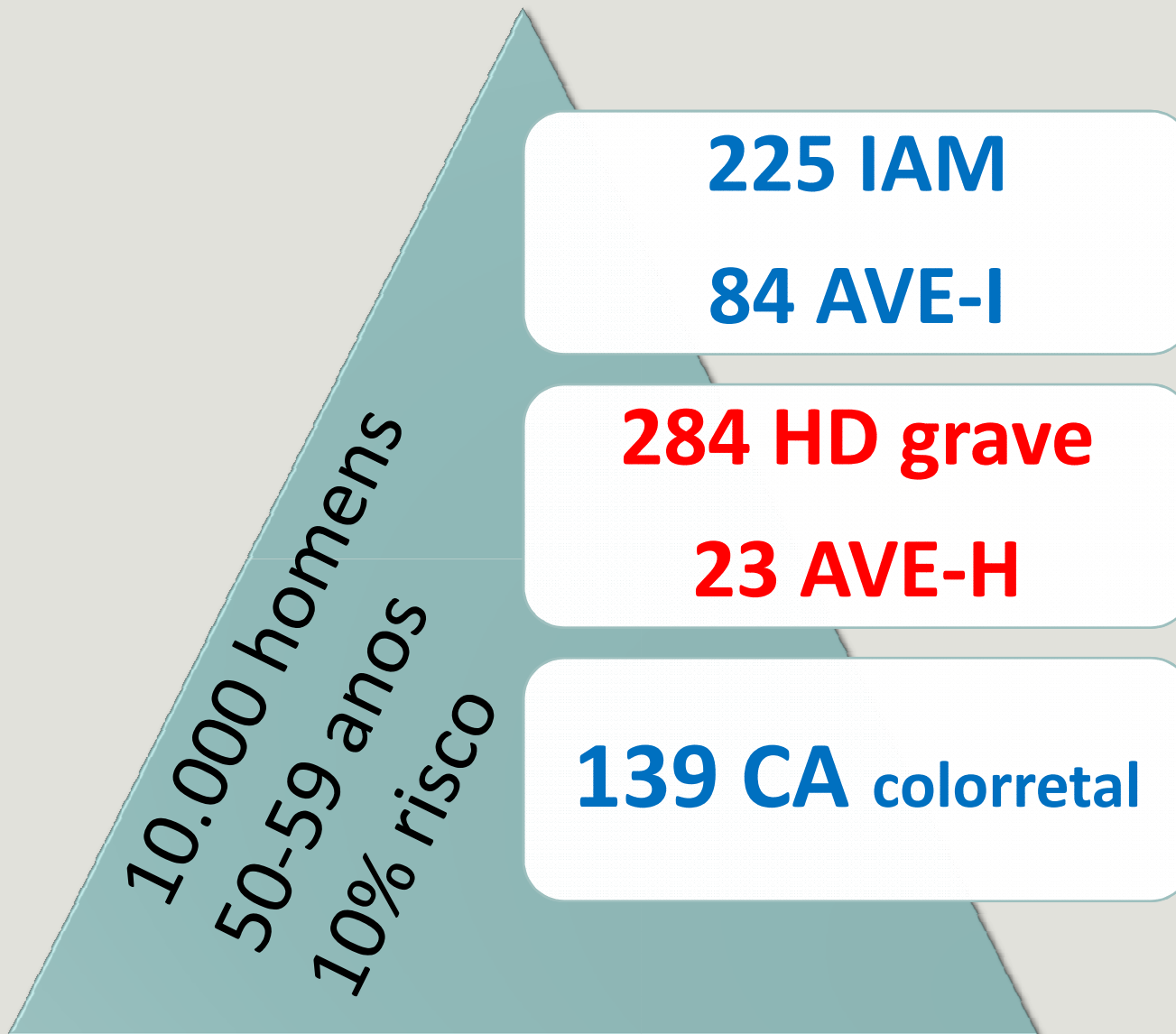
Benefício absoluto depende do risco subjacente



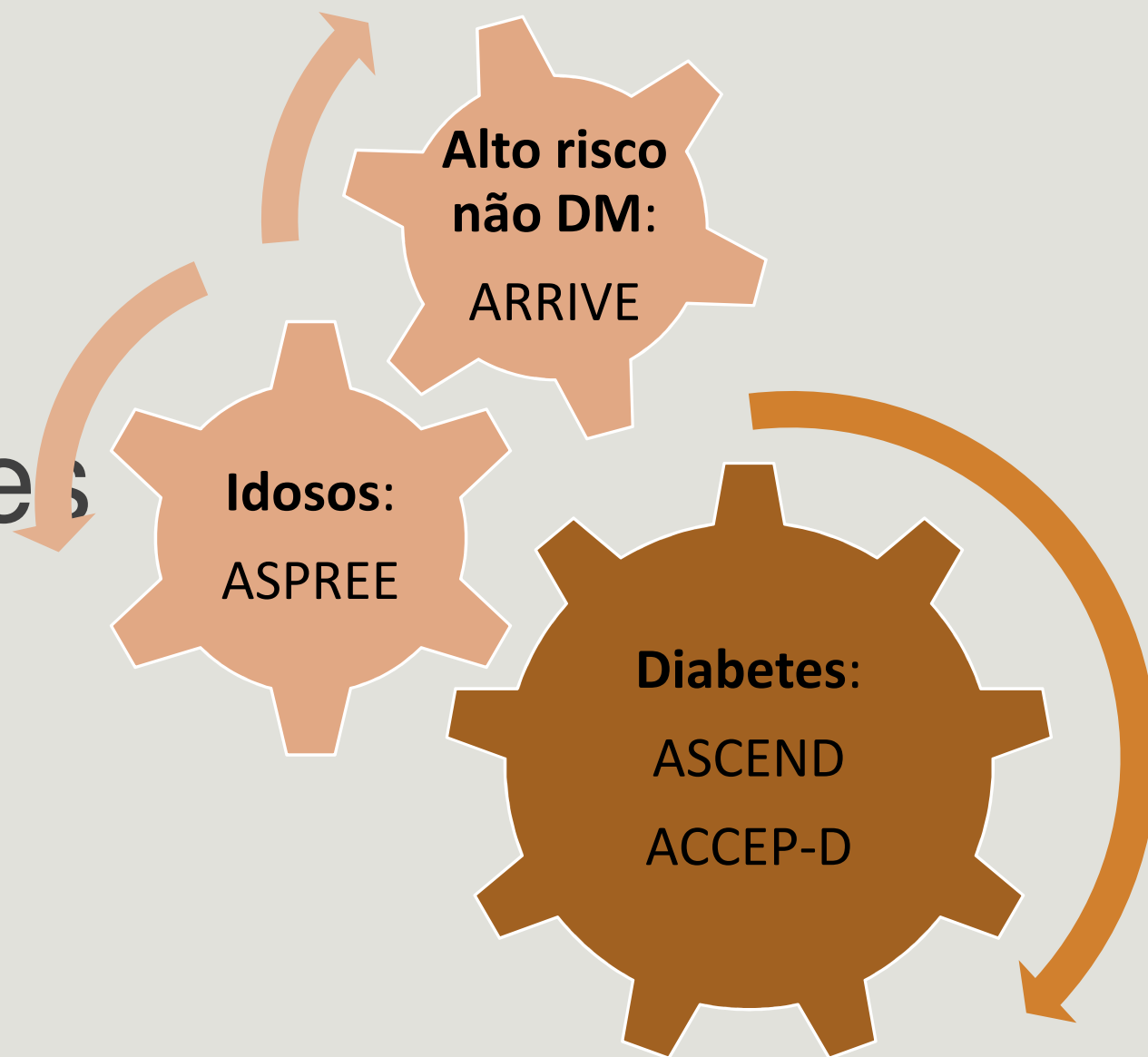
AAS aumenta sangramento em 50%

AAS reduz risco em ...





40.000 pacientes





The Aspirin-Guide app from researchers at Brigham and Women's Hospital, Harvard Medical School, helps clinicians decide which patients are candidates for the use of low-dose aspirin (75 to 81 mg/d) in the primary prevention of atherosclerotic cardiovascular disease (ASCVD) by balancing the ASCVD benefits against the risk of harm due to gastrointestinal (GI) or other bleeding.



Age: 50
Sex: Male
Race: Black
Smoker: No
Hypertension Medication: Yes
Systolic BP: 125 mmHg
Diabetes: No
Cholesterol Medication: Yes
Total Cholesterol: 180 mg/dL
LDL Cholesterol: 40 mg/dL
Atrial Fibrillation: No
Peptic Ulcer: No
Dyspepsia/GI Pain: Yes
Regular NSAID use: No
Prednisone/Systemic Corticosteroids: No

Guidance: Do not advise aspirin

ASCVD Risk Score: 0.70% over 10 years without aspirin

Bleeding Risk Score: 2.40% over 10 years without aspirin
2 risk factors for increased risk of bleeding

- Male
- Upper gastrointestinal pain or dyspepsia

Number Needed to Treat (NNT) with aspirin over 10 years to prevent one ASCVD event: 77

Number Needed to Harm (NNH): number needed to treat with aspirin over 10 years to result in one aspirin-related bleeding event: 72

50
Male
Race: Black
Smoker: Yes
Hypertension Medication: Yes
Systolic BP: 125 mmHg
Diabetes: Yes
Lipid Medication: Yes
Total Cholesterol: 180 mg/dL
LDL Cholesterol: 40 mg/dL
Atrial Fibrillation: No
Gastric Ulcer: No
Dyspepsia/GI Pain: No
Regular NSAID use: No
Prednisone/Systemic Corticosteroids: No

Guidance: Advise low-dose aspirin (75-81 mg/d)

ASCVD Risk Score: >20% over 10 years without aspirin use

Bleeding Risk Score: 1.20% over 10 years without aspirin use

1 risk factor for increased risk of bleeding

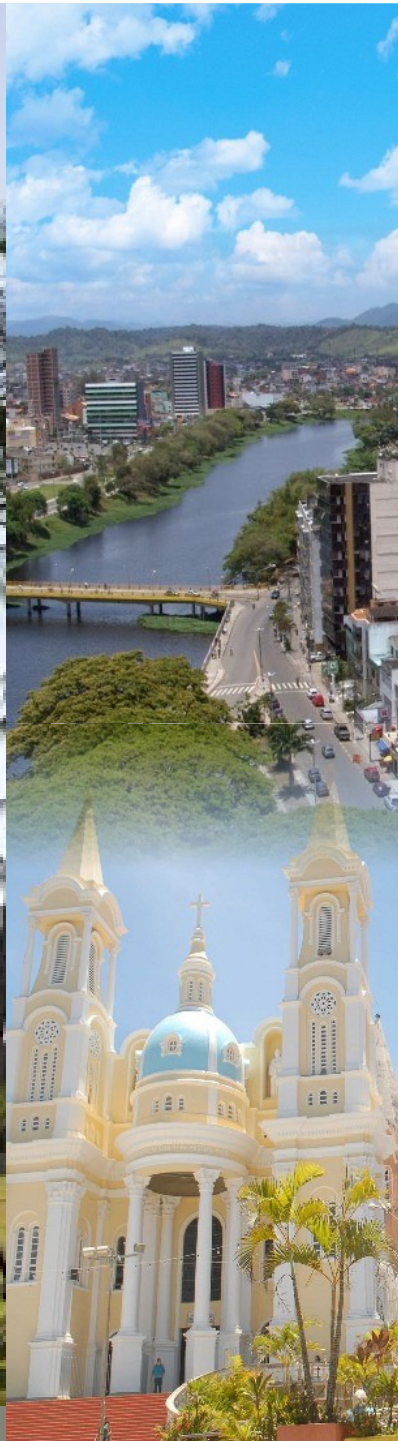
• Male

Number Needed to Treat (NNT) with aspirin over 10 years to prevent one ASCVD event: 26

Number Needed to Harm (NNH): number needed to treat with aspirin over 10 years to result in one aspirin-related bleeding event: 144

Goal: NNT less than NNH (i.e., benefit is greater than harm)

[\[more\]](#)



XV JORNADA DE CARDIOLOGIA DO SUL DA BAHIA

29 E 30 DE SETEMBRO DE 2017
ITABUNA | ILHÉUS

