

ANTICOAGULATION IN ATRIAL FIBRILLATION

Dr Shaba Nabi
Cardiovascular Lead Bristol CCG
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HOW DO WE CHOOSE?

LIFE LINES

50:50

Phone
a Friend

Poll the
Audience

A green arrow-shaped button with a white circle containing the text "50:50".

Random Guess



Ask your colleague



Ask the patient

**RIP
ASPIRIN
18.06.14**

PROS AND CONS OF WARFARIN

ADVANTAGES

- Well established
- Reversible
- GFR < 15
- Significant valve disease
- INR checks compliance
- Long half life means less embolic risk if forget to take
- Once daily

DISADVANTAGES

- Frequent blood tests
- Many drug/alcohol /food interactions
- Overall inferior to NOACS
- Higher bleeding risks
- Poor TTR
- Changing dose so not suitable for blister packs

Meta-analysis

Original research article

NOACs versus warfarin for stroke prevention in patients with AF: a systematic review and meta-analysis

Conclusions NOACs are superior to warfarin for the prevention of the composite of stroke and systemic embolism in patients with AF and an additional risk factor for stroke. There is a significant reduction in intracranial haemorrhage, which drives the finding of significantly lower mortality. During the poststudy switch from NOACs to warfarin there is an excess of the composite of stroke and systemic embolism as well as major bleeding events, which may be of significance in clinical practice.

Targets of New Oral Anticoagulants

Initiation Phase

Propagation Phase

Thrombin Activity

Contact

TF

VIIa

IX
Platelet Surface

Warfarin

Apixaban
Rivaroxaban
Edoxaban

Dabigatran
etexilate

Fibrinogen → Fibrin



Xa

Common Pathway

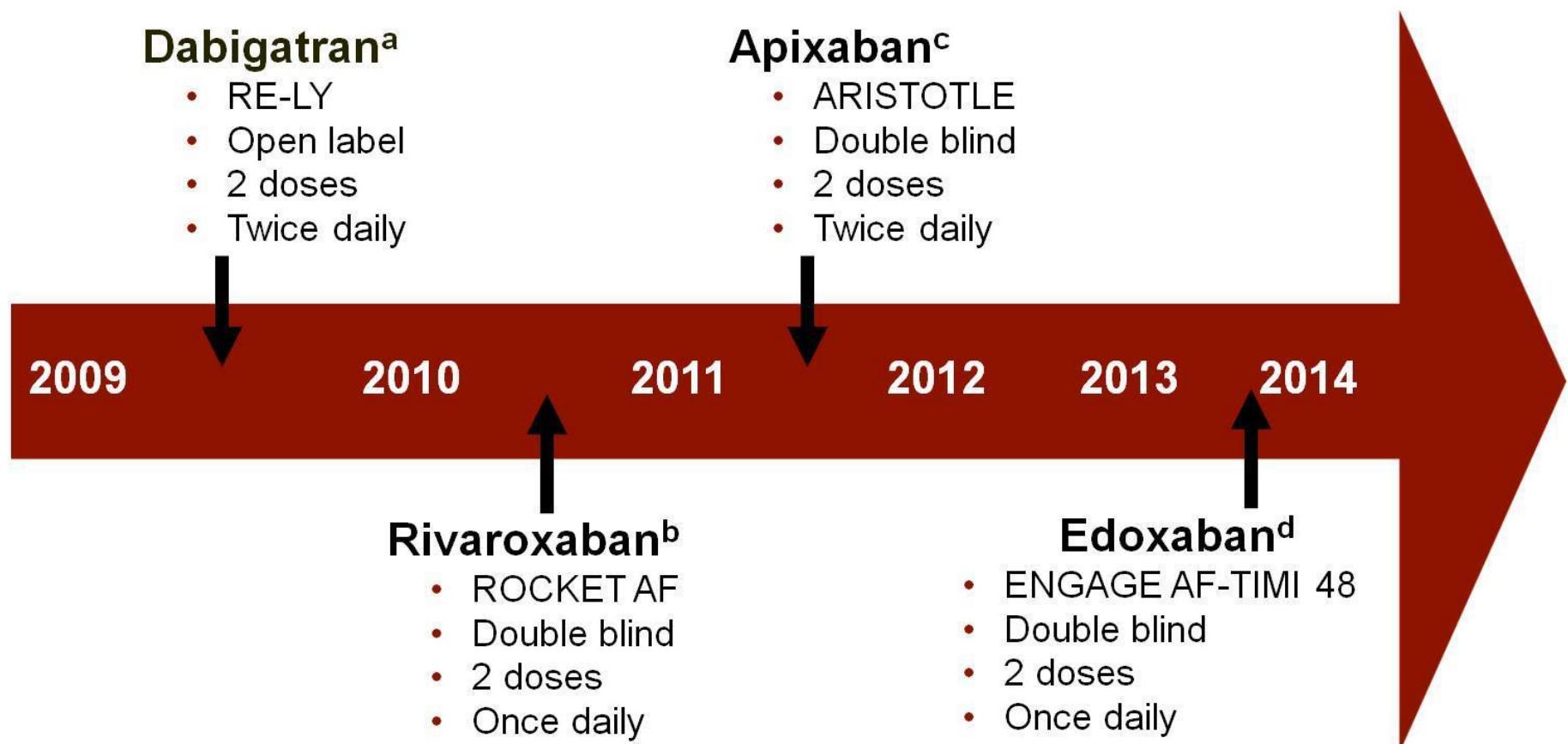
Thrombin

VIII

IX

VIIa

NOACs for Stroke Prevention in AF



a. Connolly SJ, et al. *N Engl J Med.* 2009;361:1139-1151^[4]; b. Patel MR, et al. *N Engl J Med.* 2011;365:883-891^[5]; c. Granger CB, et al. *N Engl J Med.* 2011;365:981-992^[6]; d. Giuliano RP, et al. *N Engl J Med.* 2013;369:2093-2104.^[7]

TRIAL DATA – ALL AGAINST WARFARIN

	RE-LY^a (Dabigatran)	ROCKET-AF^b (Rivaroxaban)	ARISTOTLE^c (Apixaban)	ENGAGE AF^d (Edoxaban)
Randomized, N	18,113	14,264	18,201	21,105
Age, y	72 ± 9	73 [65-78]	70 [63-76]	72 [64-78]
Female, %	37	40	35	38
Paroxysmal AF, %	32	18	15	25
VKA naive, %	50	38	43	41
Aspirin use, %	40	36	31	29
CHADS₂				
■ 0-1	32		34	
■ 2	35	13	36	47
■ 3-6	33	87	30	53

a. Connolly SJ, et al. *N Engl J Med.* 2009;361:1139-1151^[3]; b. Patel MR, et al. *N Engl J Med.* 2011;365:883-891^[4]; c. Granger CB, et al. *N Engl J Med.* 2011;365:981-992^[5]; d. Giuliano RP, et al. *N Engl J Med.* 2013;369:2093-2104.^[6]



DABIGATRAN

RIVAROXABAN

APIXABAN

EDOXABAN

WARFARIN

	DABIGATRAN 150/110 mg	RIVAROXABAN 20/15 mg	APIXABAN 5/2.5 mg	EDOXABAN 60/30 mg
STROKE RISK	↓↓↓ (D150) ↓ (D110)	↓	↓↓	↓
INTRACRANIAL HAEMORRHAGE	↓↓	↓	↓	↓
MAJOR BLEEDING	= (D150) ↓(D110)	=	↓↓	↓
GI BLEEDING	↑	↑	=	=
DYSPEPSIA	↑	-	-	-

	DABIGATRAN	RIVAROXABAN	APIXABAN	EDOXABAN
DOSING	150 MG BD	20 MG OD	5 MG BD	60 MG OD
BLISTER PACK	N	Y	Y	Y
INTAKE WITH FOOD	N	Y	N	N
AGE DOSE ADJUSTMENTS	➢ 80 years ➢ 110 mg bd	NIL	2.5MG DOSE IF 2 OUT OF 3	NIL
WEIGHT DOSE ADJUSTMENTS	NIL	NIL	SEE ABOVE	30MG DOSE IF WT < 60KG
RENAL DOSE ADJUSTMENTS	NOT FOR GFR < 30	NOT FOR GFR < 15 GFR 15 – 49: 15MG DOSE	NOT FOR GFR < 15 GFR 15 – 29: 2.5MG DOSE	NOT FOR GFR < 15 GFR 15 – 49: 30MG DOSE
CONVERSION FROM WARFARIN	Start when INR < 2	Start when INR ≤ 3	Start when INR < 2	Start when INR ≤ 2.5

www.mdcalc.com/creatinine-clearance-cockcroft-gault-equation

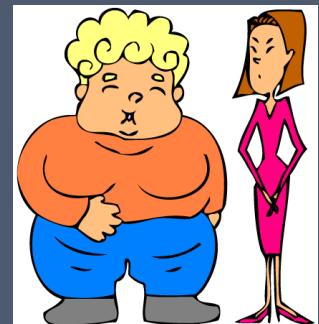
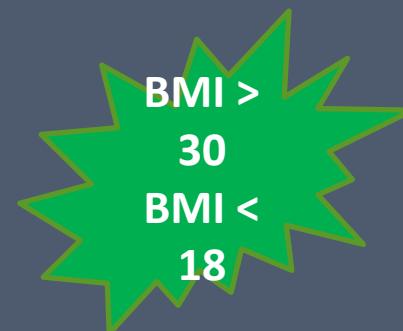
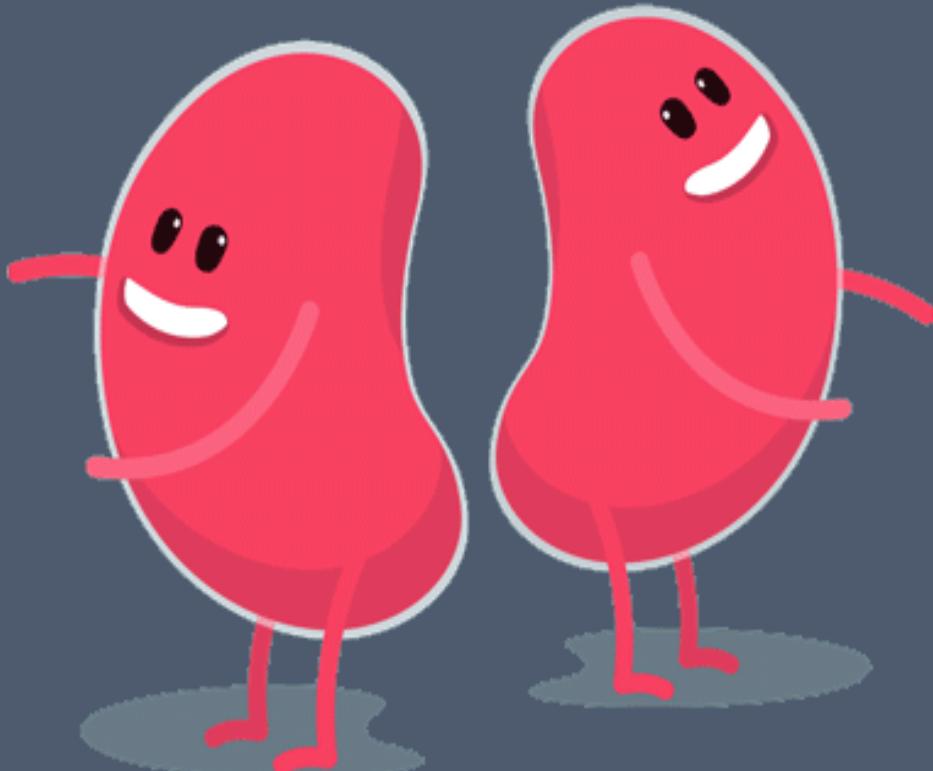
eGFR vs Cr Clearance

Labs

MDRD formula

Drug Trials

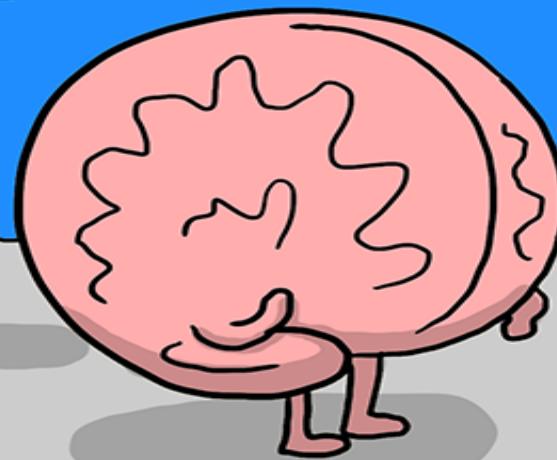
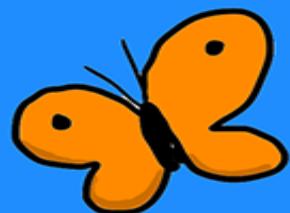
Cockcroft and Gault formula



Low GFR < 60

We've got a problem,
Heart, what do you
think we should do?

Let's COMPLAIN
about it until it
MAGICALLY
DISAPPEARS!!!



NOACS AND ANTI-PLATELETS

- Stable CAD and AF managed on OACs alone
- Prasugrel and Ticagrelor not recommended with NOACS
- Dual anti-plts + OAC as short time as possible
- Single anti-plt + OAC for 1 year only
- Reduce dose NOAC if combined with anti-plt
- Consider PPI for all combination therapies

AF patient with CHADS2 Score ≥2

Assess bleeding risk, renal function, patient's preference & comorbid conditions

High risk of stroke, low bleeding risk
(HAS-BLED) score <3

HAS-BLED score ≥3,
high risk of bleeding

Renal impairment.
h/o g.i. bleed &
dyspepsia

Preference for once/
daily regimen

Consider NOAC with best efficacy, most experience post FDA approval

Consider dose-adjustment and NOAC with lowest incidence of bleeding

Consider NOAC with less predominant renal excretion

Consider NOAC with longer half-life

Dabigatran 150 bid

Apixaban, reduced dose dabigatran

Apixaban, rivaroxaban

Rivaroxaban

BNSSG FORMULARY DECISION AID

- <http://www.bnssgformulary.nhs.uk/includes/documents/BNSSG%20NOAC%20Decision%20guideV5%20update%20March16.pdf>



"Oh waiter! Will you pass me
the anticoagulant please?"