









Device detected AF and atrial high rate episodes

Professor Paulus Kirchhof
University of Birmingham Institute of Cardiovascular Sciences
SWBH and UHB NHS trusts, Birmingham, UK
Department of Cardiovascular Medicine, University of Münster, Germany
http://www.birmingham.ac.uk/staff/profiles/cem/CVRS/Kirchhof-Paulus.aspx
email: p.kirchhof@bham.ac.uk

Atrial fibrillation



2-3% of the UK population suffer from AF

AF management requires 1-3% of the entire NHS resources

Premature cardiovascular deaths

Every 4th stroke (or more)

Fequent hospitalizations

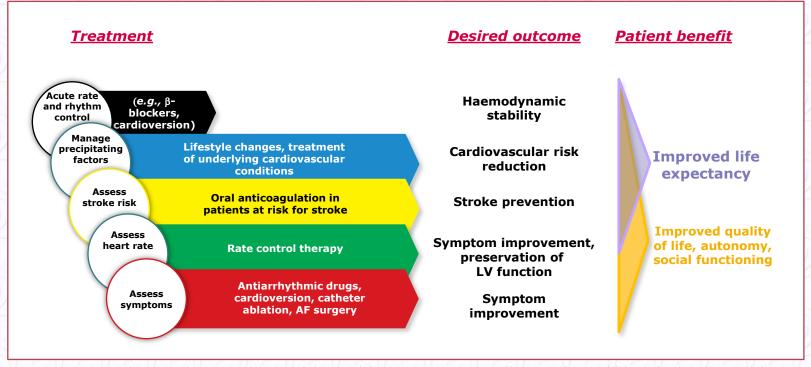
Morbidity, lost autonomy, reduced quality of life

Heart failure and sudden death are common even on optimal management

Timeline of findings from landmark trials in AF

	1995		VKA superior to aspirin for stroke prevention in AF	PVI can suppress	First maze surgery for AF treatment published
	2000	ACEi/ARBs prevent AF in heart failure	VKA reduces stroke in	Rate control not inferior to rhythm con	RF based maze maintains SR after
-)	2005	ARBs prevent AF in hypertension & LVH	AF by 2/3 Ximelagatran as effective as VKA	PVI maintains S better than antiarrhythmic dr Amiodarone not superior to rate	SR The second se
	.0	ARBs do not prevent AF or adverse outcomes in patients without hypertension	Dabigatran at least as effective as VKA in AF	control in heart failure Control in heart outcomes in no permanent AF Lenient rate control acceptable Dronedarone improves Quitcomes in no permanent AF AF ablation improves Quitcomes in no permanent AF	n- :
	20,00	PUFA do not prevent AF MRA prevent AF in HFrEF patients pre- treated with ACEi/beta-blockers	Rixaroxaban and Apixaban at least as effective as VKA in AF Edoxaban at least as effective as VKA in AF	Dronedarone harms in permanent AF First-line PV maintains SI better than antiarrhythm	Bipolar RF more effective than
	2015	ACEi/ARBs prevent AF in hypertension Beta-blockers prevent AF in HFrEF patients pre- treated with ACEi	Meta-analysis and healthcare databases: NOACs safer and slightly more effective compared to VKA	Beta-blockers without prognostic benefit in AF patients with HFrEF drugs PVI alone a effective as complex ablat in persistent A Cryoenergy a effective as RF for	stand-alone AF surgery Concomitant maze surgery maintains SR but increases risk of

The Five Domains of AF Management



To support integrated AF care, the ESC Guidelines task force and the CATCH ME consortium (www.catch-me.info) have developed state-of-the-art interactive tools underpinning integrated AF management. A first version including an overall treatment manager is integrated into the AF section of the ESC pocket guidelines app. Further CATCH ME tools for healthcare professionals and an associated app for AF patients will be released in late 2016 / early 2017. CATCH ME is supported by the European Union grant agreement No 633196 [CATCH ME].



Revolutionise Your Clinical Decision - Making for AFib Patients



Download the ESC Pocket Guidelines App to access:

2016 ESC Clinical Practice Guidelines on AFib

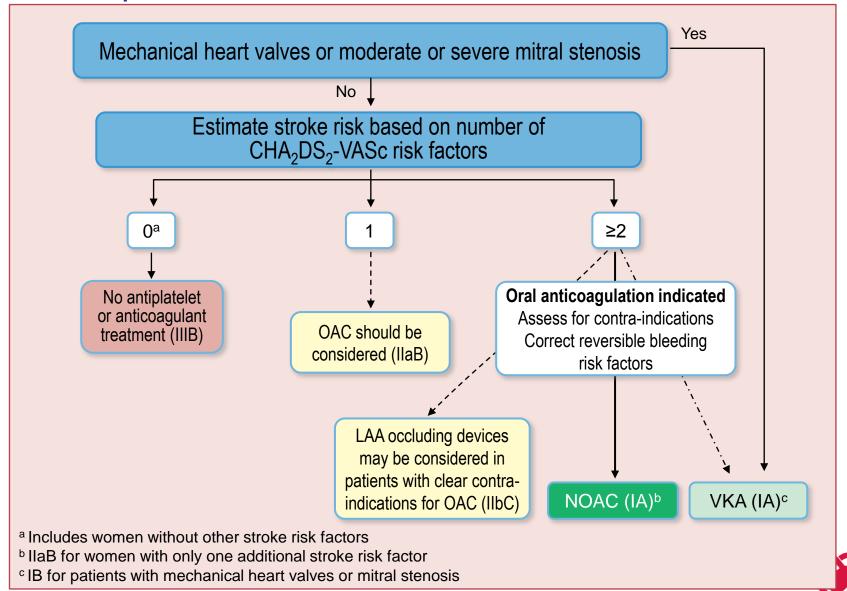
Exciting new tools from **CATCH ME*** to personalise prevention and management of your AFib patients







Stroke prevention in atrial fibrillation



The 2016 ESC AF guidelines in 17 bullet points

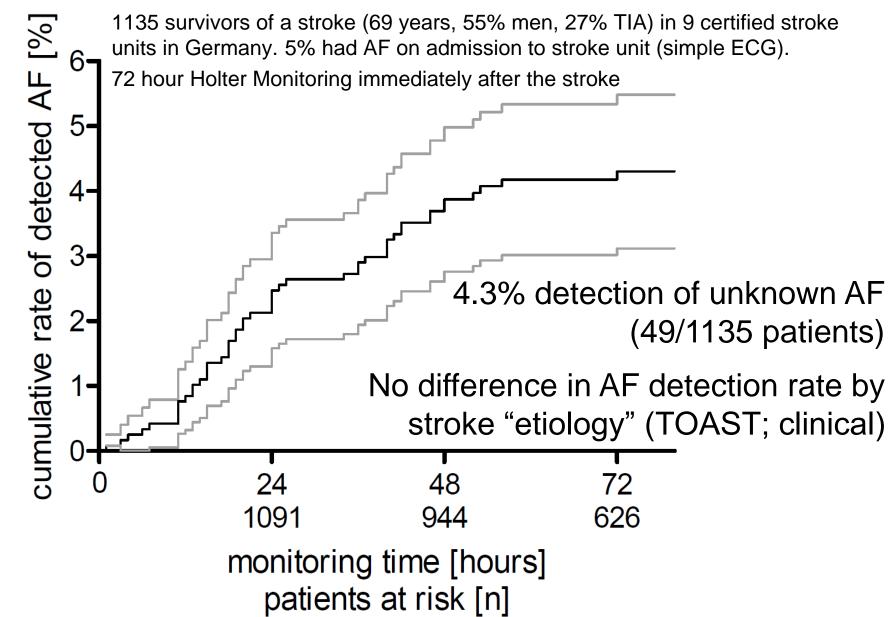
Here, we provide 17 simple rules to guide diagnosis and management of AF patients according to the 2016 ESC/EACTS/ESO Guidelines for the management of atrial fibrillation

- 1. Use ECG screening in at risk populations for atrial fibrillation, especially stroke survivors and the Elderly.
- 2. Document AF by ECG before starting treatment.



Silent AF in stroke survivors: IDEAS





Grond M, et al. Stroke 44: 3357-3364. (2013)

Silent, undiagnosed AF is a common cause of ischemic stroke

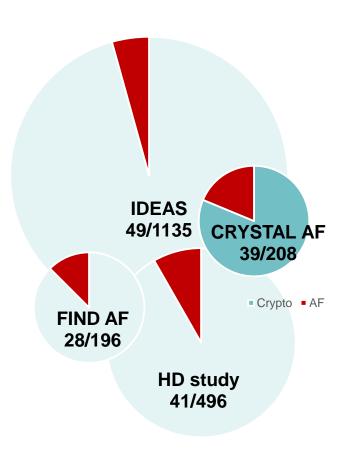


5% of patients presenting with an acute stroke have previously undiagnosed AF on admission (detected by ECG).

ECG monitoring identifies AF in stroke survivors:

- ~ 10% of unselected stroke patients
- ~ 30% of "cryptogenic stroke" patients

ECG detection of AF prior to the first stroke is highly desirable.



Stroke AF

Rizos T, et al. *Stroke.* 43:2689-2694 (2012)

Grond M, et al. *Stroke*. 44:3357-3364 (2013)

Sanna T, et al. N Engl J Med. 370:2478-2486 (2014)

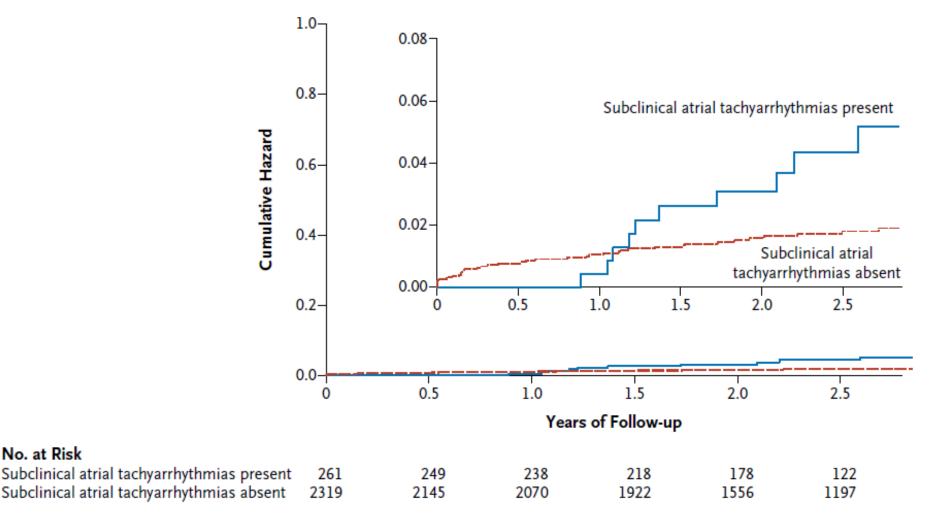
Gladstone DJ, et al. *N Engl J Med.* 370:2467-2477 (2014)

AHRE and stroke

No. at Risk



2580 pacemaker patients over 65 years with hypertension (CHADS₂VA₂Sc ≥ 2) without known AF; 3 months monitoring for AHRE (subclinical AT) by pacemaker



Healey JS, et al. N Engl J Med 366: 120-129. (2012)

Stroke risk in patients with AHRE



Trial	Study type and duration	Study population	Criteria for the diagnosis of AHRE	
MOST ²⁹	Subgroup analysis of RCT, 6 years	n = 312, median age 74 years,55% female, and 60% had a history of SND	Atrial rate >220 bpm for 10 consecutive beats	HR 2.8 for death or stroke
TRENDS ³⁰	Prospective observational study, mean follow-up 1.4 years	$n = 2486$ with ≥ 1 risk factor for stroke	AT/AF burden = longest total AT/AF duration on any given day during the prior 30-day period and classified as subsets: zero, low (<5.5 h [median duration]), and high (≥ 5.5 h)	HR 0.98 (<5.5 AHRE hours/30 days) HR 2.2 (>5.5 AHRE hours/30 days)
ASSERT ³¹	Prospective observational study, mean follow-up 2.5 years	$n=2580$, age ≥ 65 years, with hypertension and no history of AF	Atrial rate >190 bpm for >6 min	HR 2.49 (so far no "burden" analysis)
Carelink/VA ³⁴	Case crossover study, analysis of data 30 days preceding a stroke	n = 9850, median age 68 years,99% male, and 98% had a defibrillator	≥5.5 h of AF on ≥1 day in the preceding 30 days	HR 4 for times with AHRE vs times without
Belgrade Atrial Fibrillation Study ³⁵	Single-centre registry study and mean follow-up 9.9 ± 6.1 years	$n=$ 1100, mean age 52.7 \pm 12.2 years, 13.3%) had asymptomatic AF	Asymptomatic presentation of first diagnosed AF	
SOS AF project ³⁶	Pooled analysis of individual patient data from five prospective studies	 n = 10 016, median age 70 years. Pts without permanent AF with ICDs were included if they had at least 3 months of follow-up 	Device-detected AF. Cutoff points of AF burden defined as: 5 min, 1, 6, 12, and 23 h	No risk increase for short AHRE HR 2.1 for >1 ARHE hour

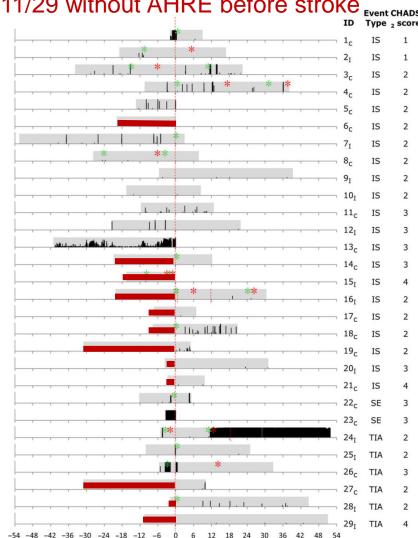
modified from Camm AJ et al Europace, published on line 4 Oct16

Timing of AHRE and ischemic stroke / TIA



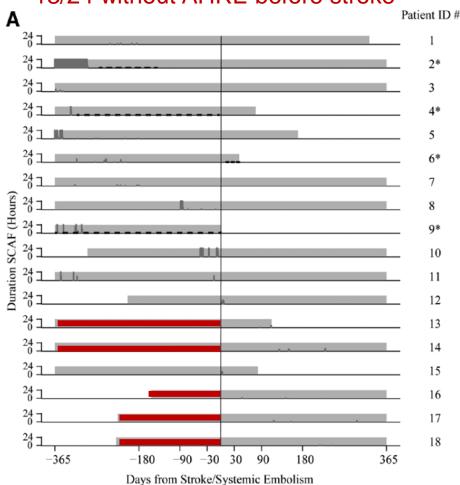


11/29 without AHRE before stroke



ASSERT

13/24 without AHRE before stroke

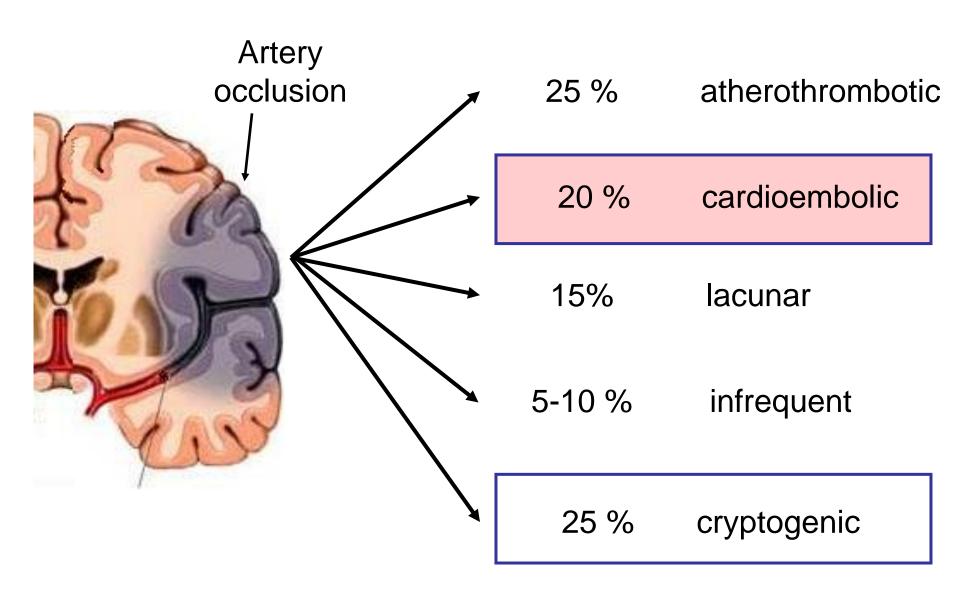


A further 8 patients did not have AHRE either one year before or after the event

Brambatti M, et al. *Circulation*. 27;129(21):2094-9. (2014) Martin, D. T., et al. et al. Eur Heart J 36: 1660-1668. (2015)

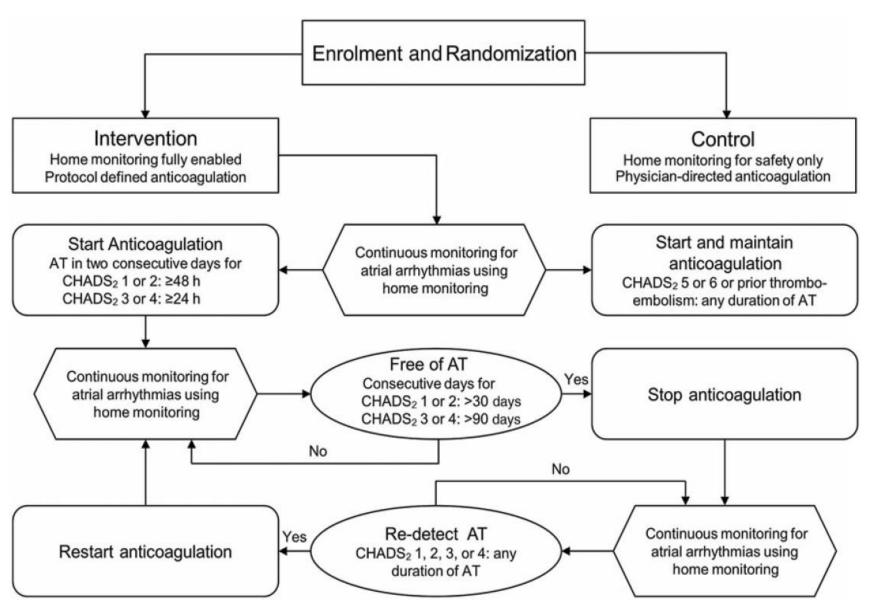
Etiology of ischemic stroke





Anticoagulation in patients with AHRE: IMPACT



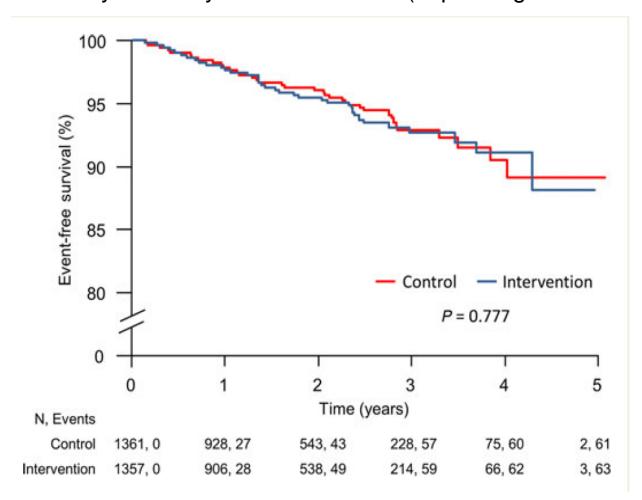


Martin DT, et al. et al. *Eur Heart J* 36: 1660-1668. (2015)

Anticoagulation in patients with AHRE: IMPACT



2718 pacemaker patients wit AHRE (median CHADSVASC score 4) randomized to no anticoagulation (usual care) or anticoagulation initiated at times of AHRE and continued for 30 days / 90 days after last AHRE (depending on CHADS).



Outcomes:

22 strokes, 41 major bleeds

OAC vs usual care

Stroke 0.7 vs 1.3%

Bleed 1.6 vs 1.2%

Death 5.4 vs 5.1 %

AHRE and stroke prevention:



ASSERT

Stroke risk in patients with AHRE is lower than in patients with "overt" AF.

Half of strokes occurred **before** the first detected AHRE.

ASSERT, IMPACT, observational data sets

Strokes in device patients rarely happen in the days after AHRE events.

EXPECT-AF

20% of AHRE episodes are artefacts or other atrial arrhythmias

IMPACT

Anticoagulation around the time of AHRE episodes does not modify stroke rate compared to no anticoagulation in patients with AHRE.

Glotzer TV, et al. *Circulation*. 107:1614-1619 (2003) Glotzer TV, et al. *Circ Arrhythm Electrophysiol*. 2:474-480 (2009) Hindricks G, et al. *Circ Arrhythm Electrophysiol*. 3:141-147 (2010) Healey JS, et al. *N Engl J Med*. 366:120-129 (2012) Al-Khatib SM, et al. *Eur Heart J* 34:2464-71 (2013) Brambatti M, et al. *Circulation*. 27;129(21):2094-9. (2014) Boriani G, et al. *Eur Heart J*. 35:508-516 (2014)

Martin, D. T., et al. et al. *Eur Heart J* 36: 1660-1668. (2015)

AHRE and stroke prevention: We need more data.



The pathological and prognostic significance of AHRE has not been fully established. There is a need to identify and validate further markers of risk in patients with AHRE. Finally, the use of oral anticoagulation for stroke prevention in patients with AHRE must be evaluated. Given the risks and inconvenience of OAC therapy, there are currently insufficient data to support their routine use in patients with AHRE, but no clinically detected AF.

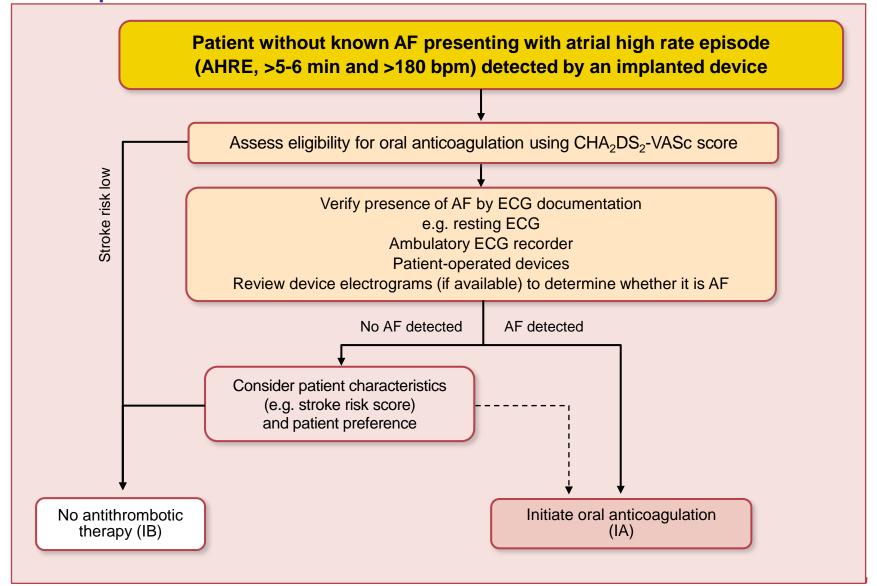
Current uncertainties relating to the detection of AHRE and to the management of patients with AHRE



- Patients with AHRE (but without diagnosed AF) have an increased risk of future stroke compared to patients without AHRE.
- 2. Short AHRE episodes (e.g. shorter than 5 minutes) are prone to be artefacts.
- 3. The prognostic impact (e.g. increase in stroke risk) of rare and short bouts of atrial fibrillation is probably lower than that of ECG-diagnosed atrial fibrillation.
- 4. Some "atrial high rate episodes" reflect other atrial arrhythmias not necessarily requiring stroke prevention therapy.
- 5. The timing of AHRE is not related to stroke in patients with AHRE.
- 6. One study did not find an effect of intermittent anticoagulation at the time of AHRE and a few weeks afterwards compared to no anticoagulation.

There is equipoise for oral anticoagulation in AHRE patients.

Management of atrial high rate episodes detected by an implanted device



Design of the NOAH – AFNET 6 trial



1st patient enrolled in June 2016, 15 European countries, 3400 patients

Patients at risk for cardiovascular events

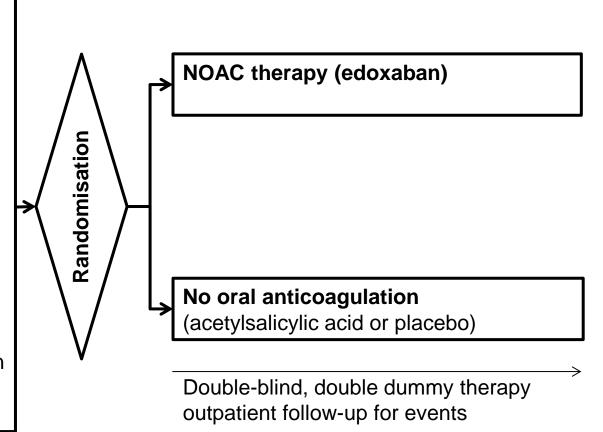
(Age 65 years or more and one additional CHA₂DS₂VASc factor)

and

documented atrial high rate episode by implanted device

Main exclusion criteria

conventionally diagnosed AF indication for oral anticoagulation contraindication for NOAC therapy













Document AF before initiating therapy in patients with device detected atrial high rate episodes

Thank you













