

# Co je standardem antikoagulační léčby okolo ablace pro FS ?

Čihák Robert



Symposium Kliniky kardiologie IKEM  
ve spolupráci s MSD

INSTITUT KLINICKÉ A EXPERIMENTÁLNÍ MEDICÍNY  
KLINIKA KARDIOLOGIE



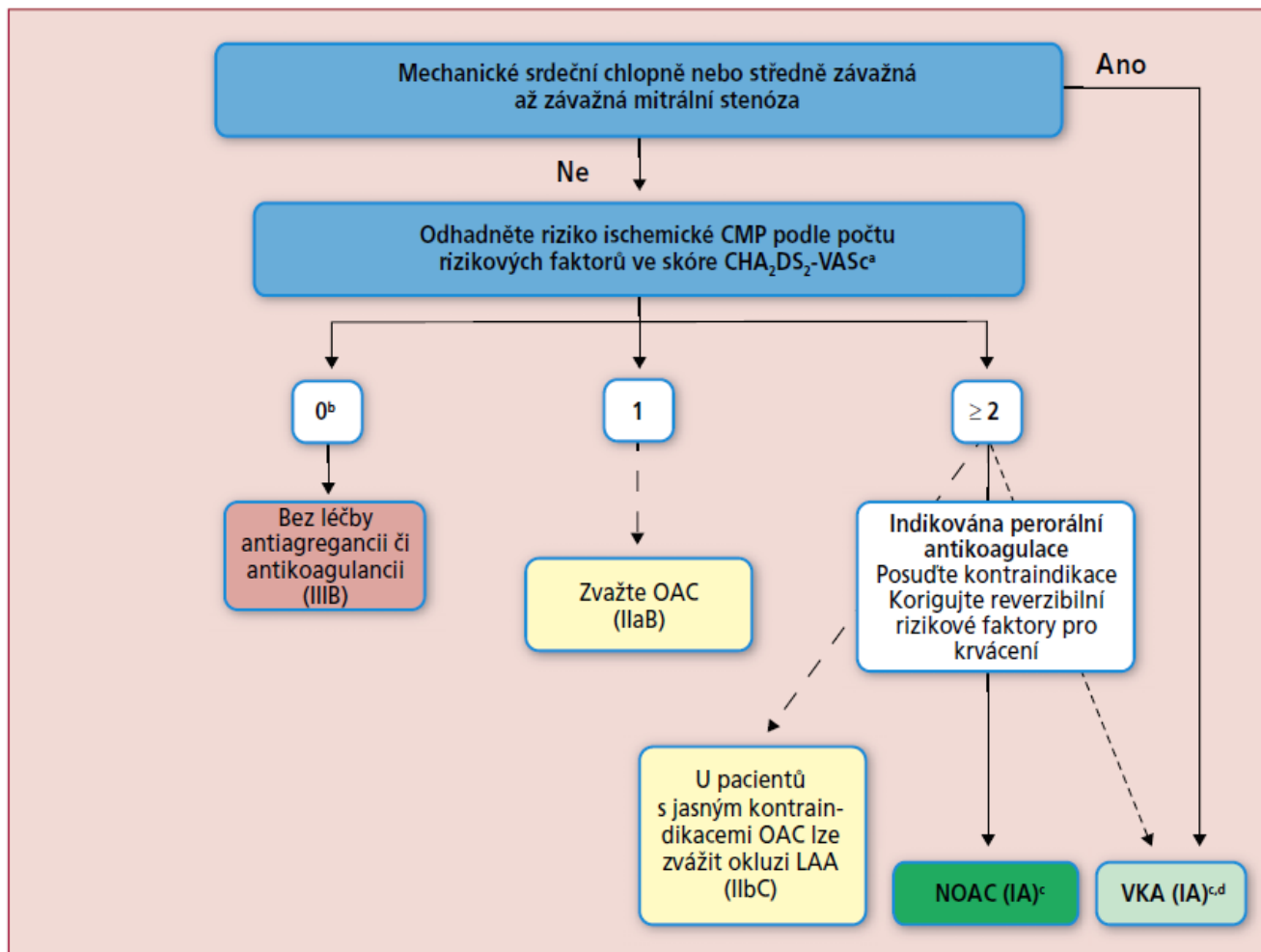
# Agenda

- Prevence TE při zvažování ablace
- Antikoagulační léčba bezprostředně před ablací
- Antikoagulační léčba při ablací
- Antikoagulační léčba po ablací
- Je možné po ablací vysadit antikoagulační léčbu ?

Co je standardem antikoagulační léčby okolo ablace pro FS ?

# Prevence TE při zvažování ablace

## Doporučení pro prevenci TE u pacientů s FS



# Co je standardem antikoagulační léčby okolo ablace pro FS ?

## Doporučení pro prevenci TE u pacientů s FS: skóre 1

### Anticoagulation in Patients With CHA<sub>2</sub>DS<sub>2</sub>VASc 1: *Pro and Contra*

<b>Contra</b>	<b>Pro</b>
Rates of ischaemic stroke are low (further lowered by OMT and life style modification)	Risk of stroke is not "0" (age is the strongest risk factor)
Risk of major bleed is not "0"	Risk factors beyond CHA <sub>2</sub> DS <sub>2</sub> VASc need to be considered (CKD, LAE, borderline HTN, ethnicity, genetics)
Net clinical benefit/gain is not as impressive as for high-risk patients	Stroke risk is not "static" and may increase in-between patient review
No adequate RCTs in low-risk patients	The uniform approach is likely be beneficial at the population level (with possible harm in individual patients)
CHA <sub>2</sub> DS <sub>2</sub> VASc 1 patients are not identical (weight of individual components is not equal)	Limited "real-world" experience (but also may serve as <i>contra</i> )
CHA <sub>2</sub> DS <sub>2</sub> -VASc 1 patients are more likely to be treated with rhythm control which may further modify their risk of stroke	? Lower risk of bleeding on NOACs

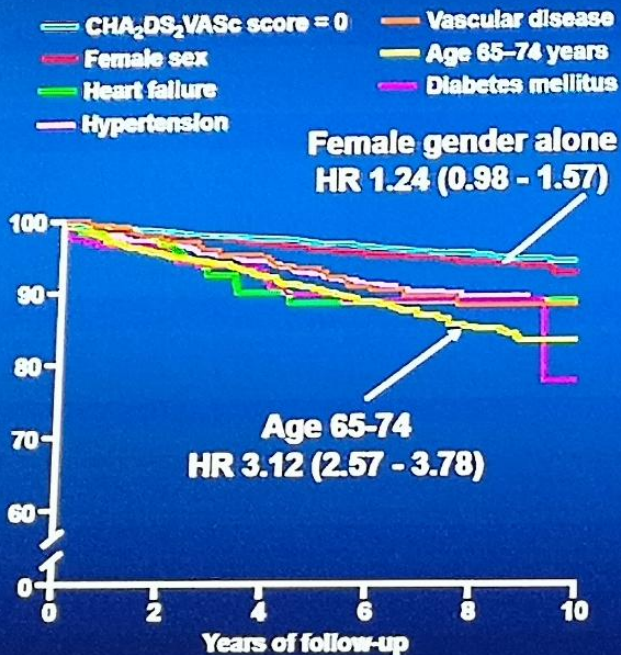
# Co je standardem antikoagulační léčby okolo ablace pro FS ?

Doporučení pro prevenci TE u pacientů s FS:  
**všichni si nejsou rovni !! (RF u FS)**

## CHA<sub>2</sub>DS<sub>2</sub>VASc: Contribution of Individual Risk Factors

### Danish cohort

Proportion free from TE, %



Olesen JB, et al. *BMJ* 2011;342:124

### Gender-related differences

- GARFIELD: ↑ 1.3-fold adjusted risk of stroke on OAC (for CHA<sub>2</sub>DS<sub>2</sub>VASc 4 ± 1.4)
- Loire Valley Project: rate of stroke increased from age 70 years or in the presence of CHF, stroke, or vascular disease
- Danish cohort (n = 240,000): female sex is not an independent RF, but a risk modifier in CHA<sub>2</sub>DS<sub>2</sub>VASc ≥ 2 (5-year RR 1.16-1.43)

Camm AJ, et al. *BMJ* 2017;7:e014579

Olesen JB, et al. *Chest* 2012;141:147-53

Nielsen PB, et al. *Circulation* 2016;137:832-40

INSTITUT KLINICKE A EXPERIMENTALNI MEDICINY  
KLINIKA KARDIOLOGIE


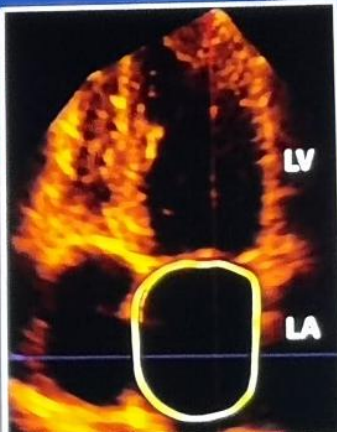



IKE  
M

# Co je standardem antikoagulační léčby okolo ablace pro FS ?

Doporučení pro prevenci TE u pacientů s FS:

## Existují další rizika TE !!

### Beyond CHA<sub>2</sub>DS<sub>2</sub>VASc

	<b>Novel clinical risk factors</b>	<b>Established clinical risk factors (CHA<sub>2</sub>DS<sub>2</sub>VASc)</b>	<b>Echo-cardiogram parameters</b>	
	Chronic kidney disease	Prior stroke/TIA	LA volume	
	Obstructive sleep apnea	Age	LA and LAA function	
	AF burden	Hypertension	Spontaneous contrast	
	<b>Serum biomarkers</b>	Diabetes	<b>Advanced imaging</b>	
	Natriuretic peptides	Heart failure	LA fibrosis	
	Troponin	Female sex	LAA morphology	
	<b>ECG</b>	AF type (paroxysmal vs non-paroxysmal), AHRE		

After Calenda B, et al. Nat Rev Cardiol 2016;13:549-59

Co je standardem antikoagulační léčby okolo ablace pro FS ?

# Antikoagulační léčba bezprostředně před ablací

Antikoagulaci před ablací nevysazujeme ! Žádné překlenování !

2019: RCT evidence on uninterrupted OAC for AF ablation

Study	Treatment arms	Primary outcome	Total follow-up (days)	Main findings
COMPARE <sup>1</sup>	Uninterrupted vs. interrupted Warfarin	Incidence of TE events		Reduced incidence of TE with continued warfarin
VENTURE-AF <sup>2</sup>	Rivaroxaban 20mg od vs. Warfarin	Major bleeding after CA	30	Similarly low event rate, feasible treatment regimen
RE-CIRCUIT <sup>3</sup>	Dabigatran 150mg bid vs. Warfarin	Major bleeding after CA	60	Fewer bleeding events compared to warfarin
AXAFA-AF-NET <sup>4</sup>	Apixaban 5mg bid* vs. Warfarin	Composite: all-cause death, stroke or major bleeding	90	Apixaban safe and effective compared to warfarin
ELIMINATE-AF <sup>5</sup>	Edoxaban 60mg od* vs. VKA	Composite: all-cause death, stroke or major bleeding	120	Low rates of the primary endpoint (1 vs 2 events)

\*Dose reduction as per the respective drug label.

<sup>1</sup>Di Biase L, et al. Circulation. 2014;129:2638-2644; <sup>2</sup>Cappato R, et al. Eur Heart J 2015; 36:1805-1811; <sup>3</sup>Calkins H, et al. N Engl J Med 2017;376:1627-1636; <sup>4</sup>Kirchhof P, et al. Eur Heart J 2018;39:2942-2955; <sup>5</sup>Pending publication.

EHRA 2019

LISBON PORTUGAL  
17-19 MARCH



[www.escardio.org/EHRA-Congress](http://www.escardio.org/EHRA-Congress)

19



IKE  
M

# Co je standardem antikoagulační léčby okolo ablace pro FS ?

## Charakteristika populací studií

Variable	VENTURE-AF <sup>1</sup>	RE-CIRCUIT <sup>2</sup>	AXAFA-AF-NET <sup>3</sup>	ELIMINATE-AF <sup>4</sup>
Age (years)	59.5	59	64	60.5
Female sex	29%	27.4%	33%	28.5%
CHA <sub>2</sub> DS <sub>2</sub> -VASc (mean)	1.6	2.0	2.4	NR (50.2% ≥2)

2019: RCT evidence

## Dávkování NOAC kolem ablace

Study	NOAC dose pre-ablation	NOAC dose post-ablation	Additional features
VENTURE-AF <sup>1</sup>	Evening before	≥6h after haemostasis	---
RE-CIRCUIT <sup>2</sup>	Morning on the day of	≥3h after haemostasis	---
AXAFA-AF-NET <sup>3</sup>	Morning on the day of	NA	MRI imaging sub-study QoL and cognitive function assessment
ELIMINATE-AF <sup>4</sup>	Evening before	Mean 6.1h after sheath removal	MRI imaging sub-study



## Úmrtí, ischemický iktus a krvácivé komplikace Kompozitní ukazatel

Incidence of major outcomes in NOAC AF ablation RCTs

Outcome	VENTURE-AF <sup>1</sup>		RE-CIRCUIT <sup>2</sup>		AXAFA-AF-NET <sup>3</sup>		ELIMINATE-AF <sup>4</sup>	
	Riva (124)	VKA (124)	Dabi (317)	VKA (318)	Apix (318)	VKA (315)	Edox (375)	VKA (178)
Death	0.0%	0.8%	0.0%	0.0%	0.3%	0.3%	0.0%	0.0%
Ischemic stroke	0.0%	0.8%	0.0%	0.3%	0.6%	0.0%	0.3%	0.0%
Major bleeding	0.0%	0.8%	1.6%	6.9%	3.1%	4.4%	2.4%	1.7%
Composite	0.0%	2.4%	1.6%	7.2%	4.0%	4.7%	2.7%	1.7%

Summary

## Závěry studií – souhrn:

- Reassuringly low event rates in all 4 trials, but no statistically valid comparisons between NOACs and VKA can be made.

### Take-home messages

- Periprocedural anticoagulation with uninterrupted NOACs appears safe and effective, at least in relatively young and low-to-moderate stroke risk patients.
- However, available evidence is insufficient to justify switching from one to another (or to a particular) OAC drug only for the purpose of AF ablation.

EHRA 2019

LISBON PORTUGAL  
17-19 MARCH



[www.escardio.org/EHRA-Congress](http://www.escardio.org/EHRA-Congress)

2019



INSTITUT KARDIOLOGIE  
KLINIKA KARDIOLOGIE



IKE  
M

## Antikoagulační léčba při ablaci

- Pacient přichází z nepřerušovanou antikoagulační léčbou (NOAC vynecháváme jen dávku ráno před výkonem)
- Při perzistující FS/vysokém skóre - TEE
- Během výkonu heparin iv (první dávka před TS punkcí), cílové ACT 300-350s
- Po ablaci vytažení sheathů na sále, Z steh, heparinizace (částečně) rušena protaminem
- 3-4 hodiny po ablaci TTE k vyloučení separace perikardu a dávka warfarinu/NOAC
- Pokud není možné TTE, při poklesu ACT pod 170 s zahájit infuzi UFH s cílem aPTT ratio 1.5-2.5 (nebo ACT 120-160 s).

## Co je standardem antikoagulační léčby okolo ablace pro FS ?

# Antikoagulační léčba po ablaci

Calkins et al Catheter and Surgical Ablation of Atrial Fibrillation

**Table 4** Anticoagulation strategies: pre-, during, and postcatheter ablation of AF

	Recommendation	Class	LOE
Postablation	In patients who are not therapeutically anticoagulated prior to reverse heparin is reasonable.	I	C-E0
	Adherence to AF anticoagulation guidelines is recommended for patients who have undergone an AF ablation procedure, regardless of the apparent success or failure of the procedure.	I	C-E0
	Decisions regarding continuation of systemic anticoagulation more than 2 months post ablation should be based on the patient's stroke risk profile and not on the perceived success or failure of the ablation procedure.	I	C-E0
	In patients who have not been anticoagulated prior to catheter ablation of AF or in whom anticoagulation with a NOAC or warfarin has been interrupted prior to ablation, administration of a NOAC 3 to 5 hours after achievement of hemostasis is reasonable postablation.	IIa	C-E0
	Patients in whom discontinuation of anticoagulation is being considered based on patient values and preferences should consider undergoing continuous or frequent ECG monitoring to screen for AF recurrence.	IIb	C-E0

AF = atrial fibrillation; LOE = Level of Evidence; NOAC = novel oral anticoagulant; TEE = transesophageal electrocardiography  
\*Time in therapeutic range (TTR) should be > 65% – 70% on warfarin.

- Antikoagulační léčbu podáváme alespoň dva měsíce po výkonu
- Vysazování po ablaci lze podle rizikové stratifikace, ne (jen) podle průběhu po výkonu

Co je standardem antikoagulační léčby okolo ablace pro FS ?

## Je možné po ablaci vysadit antikoagulační léčbu ?

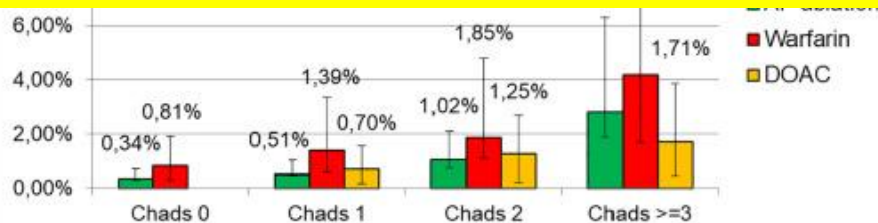
Incidence of thromboembolic events following **atrial fibrillation catheter ablation and rate control strategies** according to the kind of oral **anticoagulation**: A systematic review and meta-analysis.

- 27 studies were selected, including 50,973 patients in the AF catheter ablation group; 281,595 patients in the VKA group; 54,811 patients in the NOAC group

Stroke/year

**AF catheter ablation significantly reduces the incidence of long-term thromboembolic events compared to OAT through both VKA and DOAC associated to rate control strategies.**

**This lower thromboembolic risk is strengthened by the reduction in hemorrhagic complications provided by AF ablation, as a large proportion of patients can safely interrupt OAT and avoid related bleedings**



ricks G<sup>5</sup>,

Jared Bunch T<sup>6</sup>, Saliba W<sup>7</sup>. *Int J Cardiol.* 2018 Nov 1;

INSTITUT KLINICKÉ A EXPERIMENTÁLNÍ MEDICÍNY  
KLINIKA KARDIOLOGIE

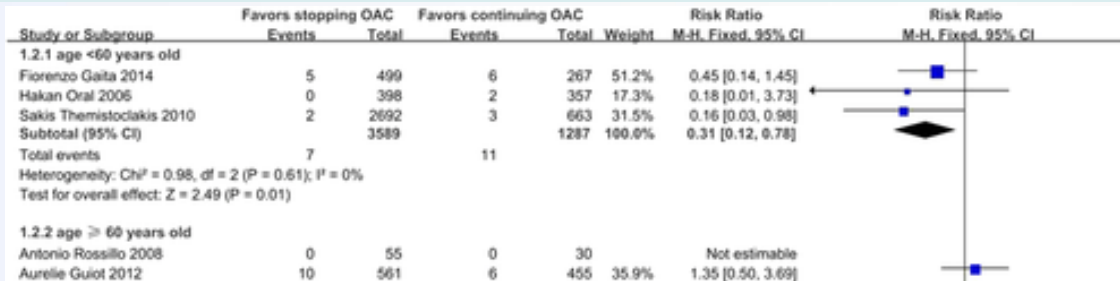


IKEM

Co je standardem antikoagulační léčby okolo ablace pro FS ?

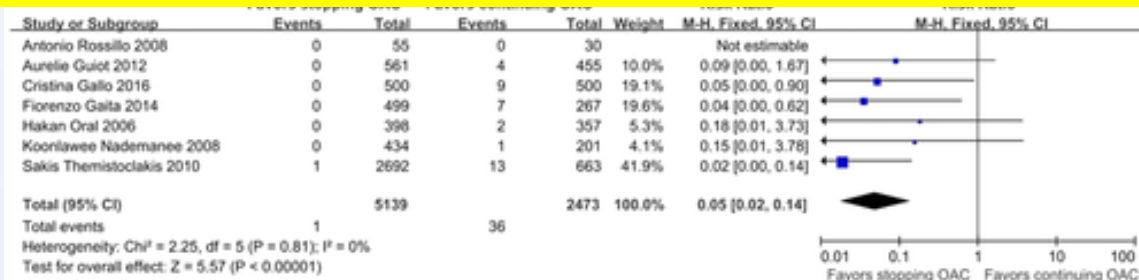
# Je možné po ablaci vysadit antikoagulační léčbu ?

Withdrawal of oral anticoagulants 3 months after successful radiofrequency catheter ablation in patients with atrial fibrillation: A meta-analysis (7 studies).



TE risk

**The withdrawal of oral anticoagulants 3 months after successful radiofrequency catheter ablation for patients with AF may be safe and feasible**

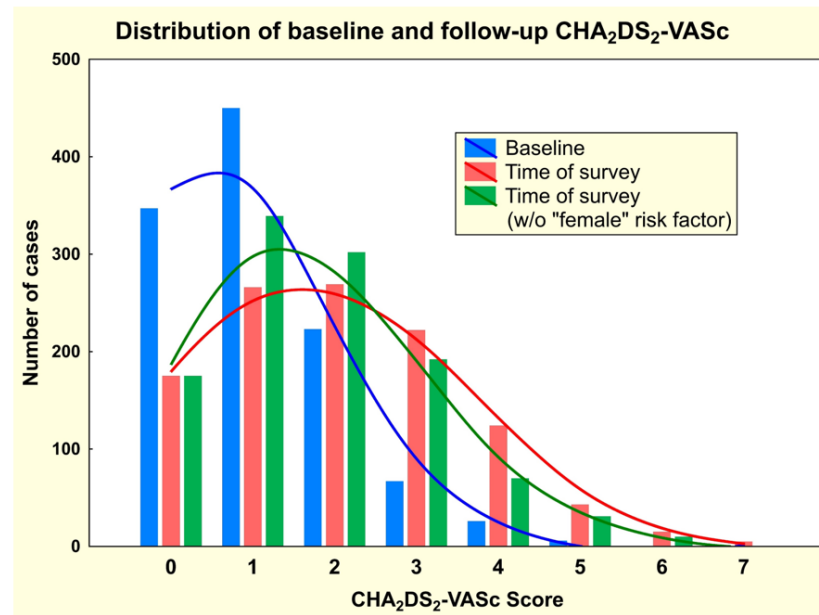
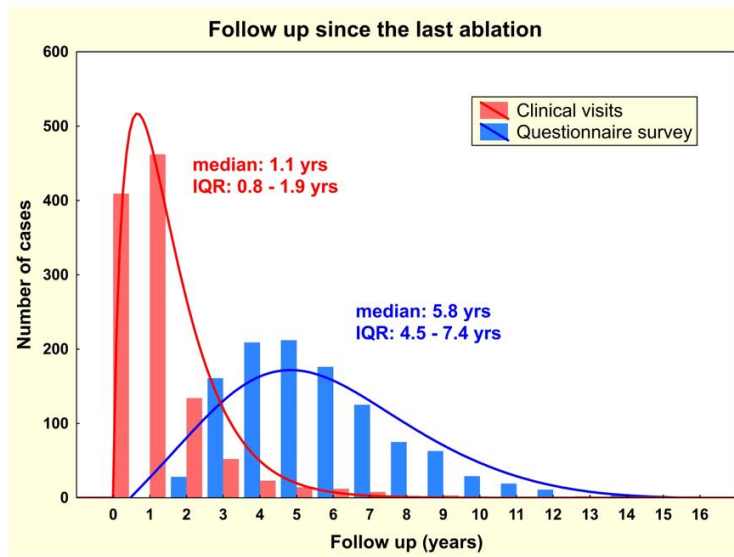


bleeding risk

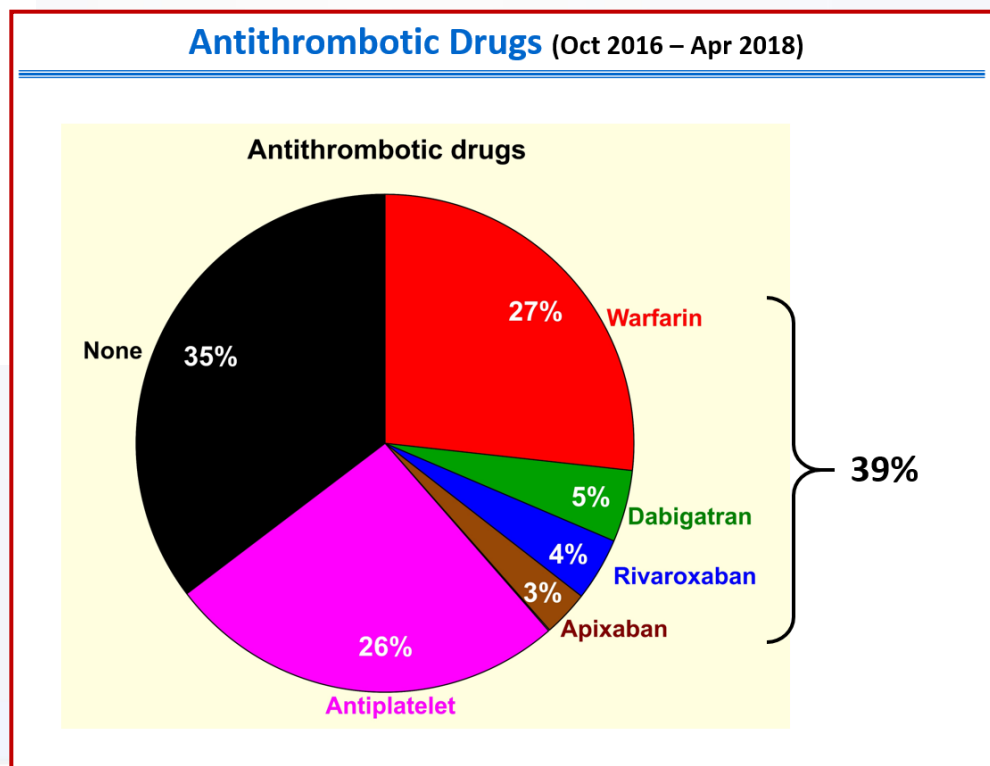
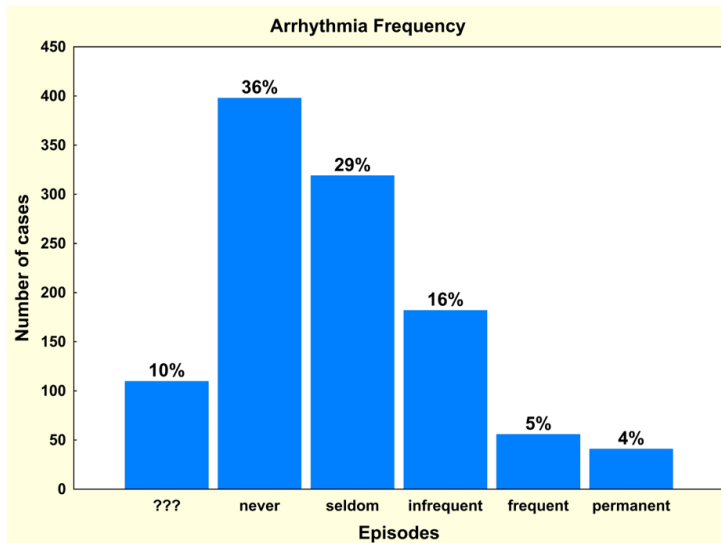
# Co je standardem antikoagulační léčby okolo ablace pro FS ?

## Jak pokračují pacienti v antikoagulační léčbě po ablaci v IKEM ? (dotazníková data)

### Follow Up: Clinical versus Survey



# Co je standardem antikoagulační léčby okolo ablace pro FS ? Jak pokračují pacienti v antikoagulační léčbě po ablaci v IKEM ?





Co je standardem antikoagulační léčby okolo ablace pro FS ?

## Je možné po ablaci vysadit antikoagulační léčbu ?

**Pill (NOAC)-in-the pocket**

**“As Needed”** nonvitamin K antagonist oral anticoagulants for infrequent atrial fibrillation episodes following atrial fibrillation ablation guided by diligent pulse monitoring: A feasibility study

- 99 patients (age  $64 \pm 8$  years), CHA<sub>2</sub>DS<sub>2</sub>-VASc score  $\geq 1$  in men and  $\geq 2$  in women, capable of pulse assessment twice daily
- All patients were instructed to start NOAC if AF  $>1$  hour or recurrent shorter episodes. Duration of NOAC use after restart was typically 2 to 4 weeks.

**The use of as needed NOACs when AF is suspected with pulse monitoring is effective and safe**

Co je standardem antikoagulační léčby okolo ablace pro FS ?  
**Je možné po ablaci vysadit antikoagulační léčbu ?**

Anticoagulation after successful AF ablation:

**Brushing you teeth may keep you off blood thinners**





- Prevence TE při zvažování ablace  
**podle zásad rizikové stratifikace**
- Antikoagulační léčba bezprostředně před ablací  
**nevysazujeme warfarin ani NOAC**
- Antikoagulační léčba při ablaci  
**heparin s cílovým ACT 300-350**
- Antikoagulační léčba po ablaci  
**pokračujeme vždy alespoň dva měsíce**
- Je možné po ablaci vysadit antikoagulační léčbu ?  
**ponecháváme dle rizikové stratifikace, ale při středním riziku lze v individuálních případech možné vysazení zvážit**

# Co je standardem antikoagulační léčby okolo ablace pro FS ?



*Botanická zahrada Praha, 11.5.2019*

## **Děkuji za pozornost !**

Co je standardem antikoagulační léčby okolo ablace pro FS ?

## Je možné po ablaci vysadit antikoagulační léčbu ?

Anticoagulation after catheter ablation of atrial fibrillation: An unnecessary evil? A systematic review and meta-analysis (16 studies).

bleeding risk

The results of our pooled analysis seem to suggest that the risk-benefit ratio favors the discontinuation of OAT after successful AF ablation even in patients at moderate-high risk of TE.

