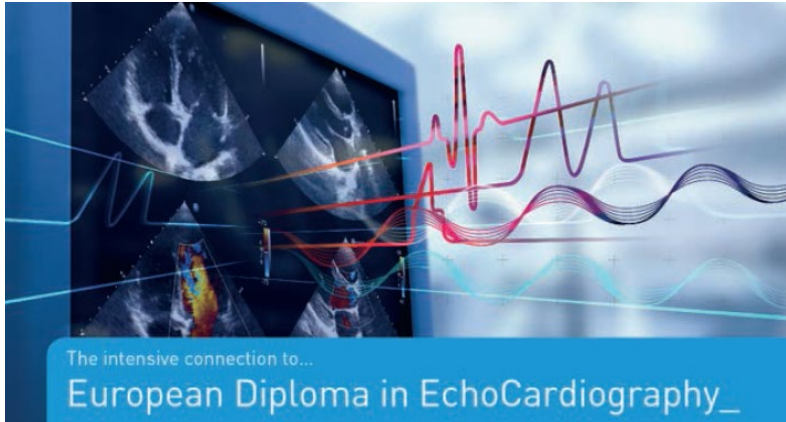




Echo-guided kardioverze

Martin Balík

Komplexní kardiovaskulární centrum, KARIM 1.LF UK a VFN Praha
e-mail: martin.balik@vfn.cz



Conflict of interest

- Research grants: AZV 18-06-00417 (Prospective randomized double-blind study of efficacy and safety of 1c class antiarrhythmic agent (propafenone) for supraventricular arrhythmias in septic shock), AZV NU22-B-147 (Impact of SARS-CoV-2 viral load estimate on therapeutic effects of remdesivir), Gilead Sciences 2021-22 (Immune boosting in severe Covid19)
- Research support: ESICM Stoutenbeek Award (Dutch Society of Critical Care)
- Inventor and patent holder: Lactocitrate[®], EU patent (EP2609915B1), Canadian patent (No.2799624)
- Speaker Fees: FMC, GML-Biomedica, Gilead Sciences, BBraun, AOP Orphan
- Grant to organize educational meetings: None
- Advisory board: AOP Orphan

Incidence, risk factors and outcomes of new-onset atrial fibrillation in patients with sepsis: a systematic review

Critical Care 2014, 18:688 doi:10.1186/s13054-014-0688-5

Clinical review: Treatment of new-onset atrial fibrillation in medical intensive care patients – a clinical framework

Mengalvio E Sleeswijk¹, Trudeke Van Noord², Jaap E Tulleken², Jack JM Ligtenberg², Armand RJ Girbes³ and Jan G Zijlstra²

Critical Care 2007, 11:233 (doi:10.1186/cc6136)

- SV arytmie jako nejčastější spouštěč diastolického srdečního selhání

- Intenzivní péče

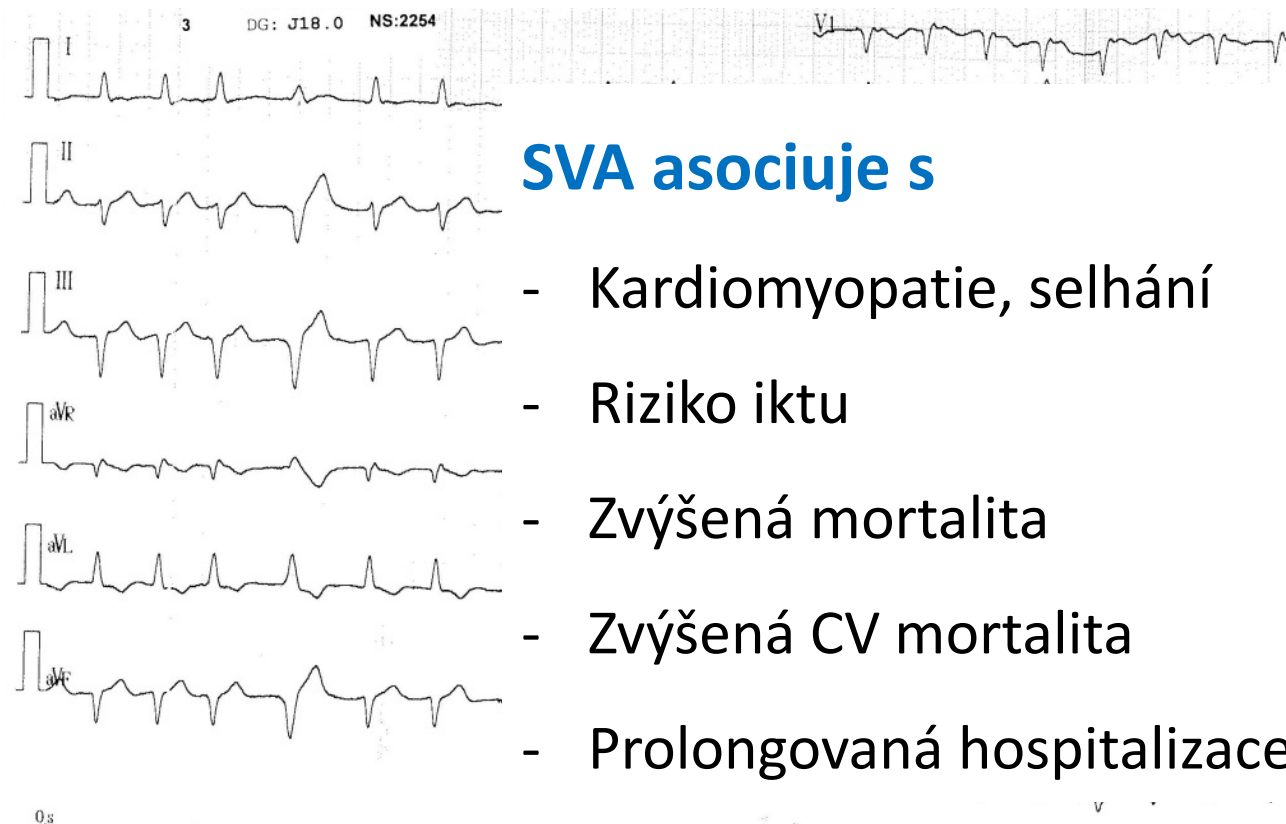
- 4-9-15% (věkově závislé)
- 15-40% perioperační

- Incidence a závažnost stavu

- 10% (4-23%) sepse
- 23% (6-46%) septický šok

- Endemický problém

- 25% populace nad 40 let !



SVA asociuje s

- Kardiomyopatie, selhání
- Riziko iktu
- Zvýšená mortalita
- Zvýšená CV mortalita
- Prolongovaná hospitalizace

SVA a NOAF nezávisle zhoršují prognózu – proč každého nevertujeme na sinus ?

Review Article

Management of Atrial Fibrillation in Critically Ill Patients

Mattia Arrigo, Dominique Bettex, and Alain Rudiger

Ztráta síňové systoly u 6-21% ICU pacientů – asociuje s 2-5x zvýšenou mortalitou !

Received: 7 October 2021 | Revised: 11 November 2021 | Accepted: 30 November 2021

DOI: 10.1111/aas.14007

RESEARCH ARTICLE



Disappointing Success of Electrical Cardioversion for New-Onset Atrial Fibrillation in Cardiosurgical ICU Patients*

Mattia Arrigo, MD^{1,2}; Natalie Jaeger, MD¹; Burkhardt Seifert, PhD³; Donat R. Spahn, MD, FRCA¹; Dominique Bettex, MD¹; Alain Rudiger, MD¹

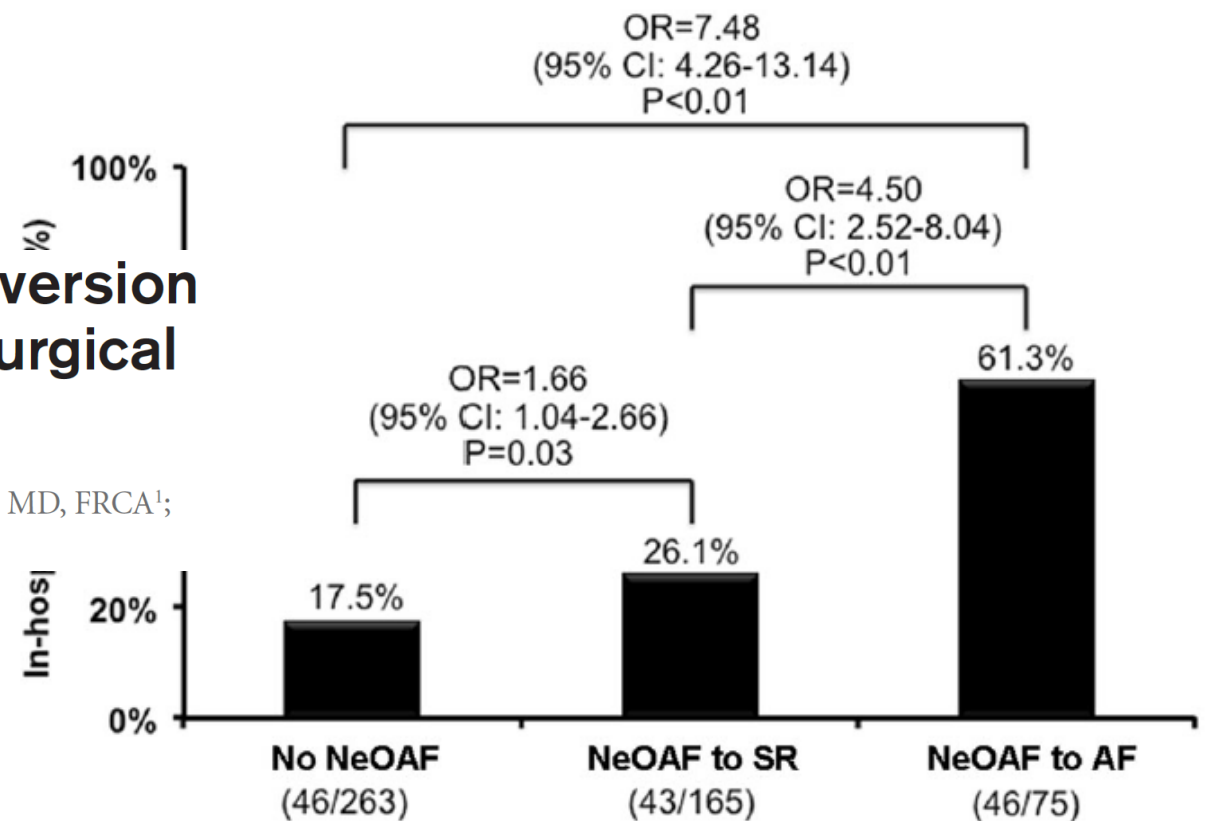
- DC kardioverze – efekt v 24-30%, s antiarytmiky 70-80%
- rate control 4%

Prognostic impact of restored sinus rhythm in patients with sepsis and new-onset atrial fibrillation

Liu et al. *Critical Care* (2016) 20:373

DOI 10.1186/s13054-016-1548-2

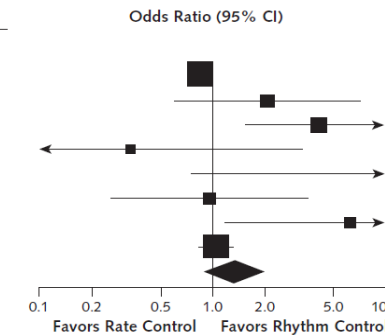
Wen Cheng Liu¹, Wen Yu Lin¹, Chin Sheng Lin¹, Han Bin Huang², Tzu Chiao Lin¹, Shu Meng Cheng¹, Shih Ping Yang¹, Jung Chung Lin³ and Wei Shiang Lin^{1*}



Rhythm nebo rate control ? Potřebujeme EBM+.....

- Kardiologie: Rekurence SVA a SE antiarytmik.....důraz na rate control
- PRCT zařazovaly všechny stupně diastolické dysfunkce, bez stratifikace funkce LV a LA
- Hlavní PRCTs neukazují výhody rhythm vs rate control na
 - mortalitu, CV mortalitu
 - srdeční selhání
 - iktus
 - příjmy do nemocnice
- Celková frekvence kardioverze 80%
- AFFIRM Trial (J Am Coll Cardiol 2003)
- RACE Trial (N Engl J Med 2002)
- AF-CHF Trial (included EF_LV<35%, NYHA II-IV (N Engl J Med 2008))

Study, Year (Reference)	Odds Ratio (95% CI)	Deaths/Total, n/N	
		Rate Control	Rhythm Control
Wyse et al, 2002 (27)	0.851 (0.720–1.005)	310/2027	356/2033
Carlsson et al, 2003 (18)	2.087 (0.608–7.167)	8/100	4/100
Okçün et al, 2004 (20)	4.125 (1.562–10.895)	36/84	6/39
Opolski et al, 2004 (21)	0.337 (0.034–3.291)	1/101	3/104
Vora et al, 2004 (26)	14.099 (0.754–263.543)	5/40	0/45
Petrac et al, 2005 (22)	0.957 (0.260–3.532)	5/52	5/50
Yildiz et al, 2008 (28)	6.270 (1.185–33.192)	5/66	2/155
Talajic et al, 2010 (24)	1.048 (0.836–1.314)	228/694	217/682
Overall	1.343 (0.893–2.020)		



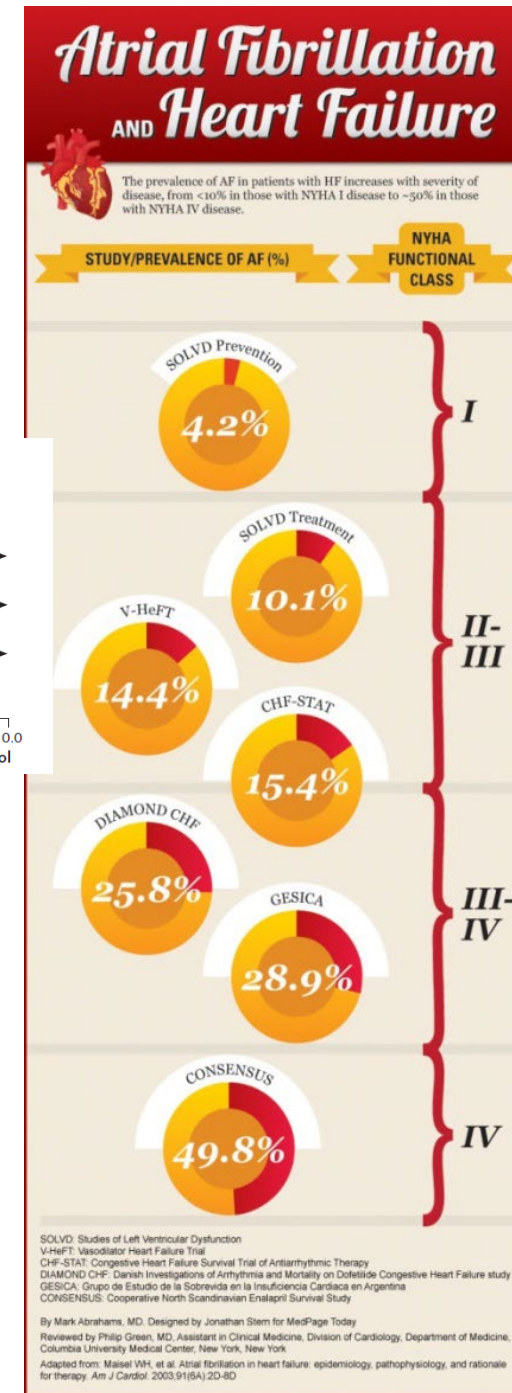
REVIEW

Annals of Internal Medicine

Rate- and Rhythm-Control Therapies in Patients With Atrial Fibrillation

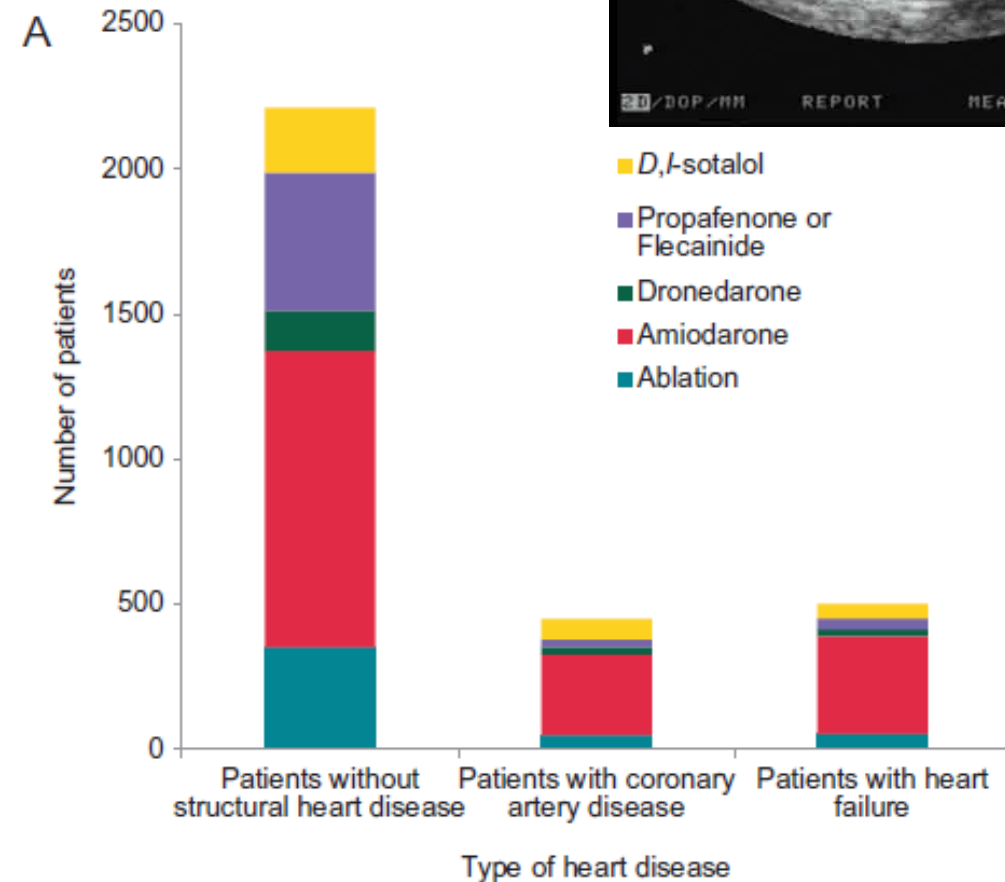
A Systematic Review

Sana M. Al-Khatib, MD, MHS; Nancy M. Allen LaPointe, PharmD; Raneer Chatterjee, MD, MPH; Matthew J. Crowley, MD;



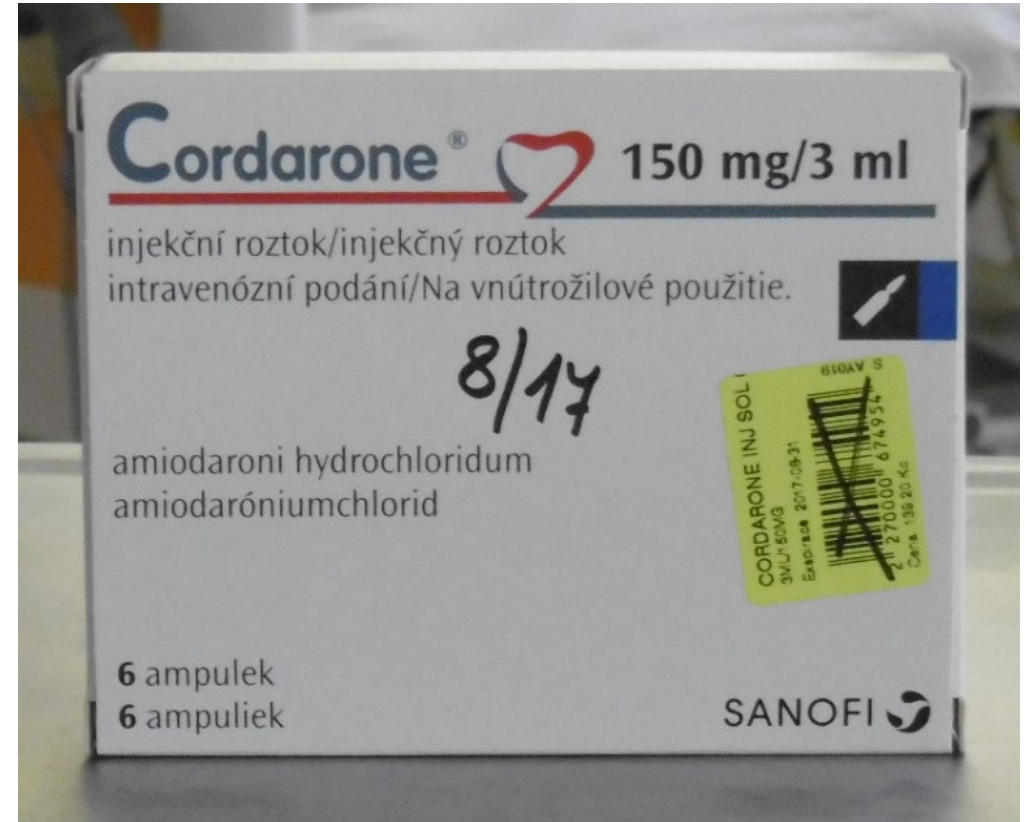
SVA: rozhodovací algoritmy

- Klíčová role echokardiografie:
 - LV velikost, kontraktilita ?
 - LVEDP, preload....katecholaminy ?
 - Chlopně: MS, MR, AS – sig. ?
 - Dilatace LA ?
 - Spont. echokonstrast ?
 - PAPs ?
- Rhythm nebo rate control ?
- Elektrická kardioverze ?
- >48h ? : TEE
- Predictory udržitelnosti SR: 2D a Doppler ?
- Antikoagulace - nastavení
- Předchozí betablokátory
- Medikace při propuštění (betablokátory ?)



Amiodarone

- III class (V-W) antiarytmikum
- Nejméně kardiodepresivní
- Mírně vazodilatační
- Efekt u 80% NOAF – pomalejší než 1c class
- Synergismus s $MgSO_4$ i.v.
- Toxický již za pár dní podávání
 - Prolongovaný QTc
 - Plicní toxicita (3-14 dní...)
 - Hyperthyreóza, dlouhodobě hypothyreóza
 - Hepatotoxicity



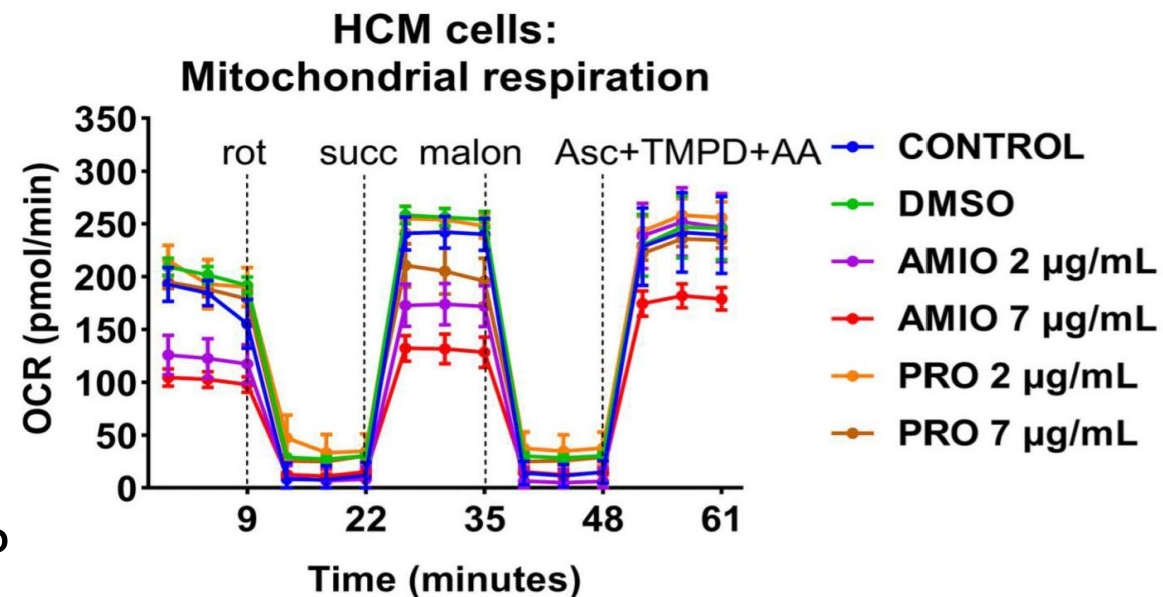
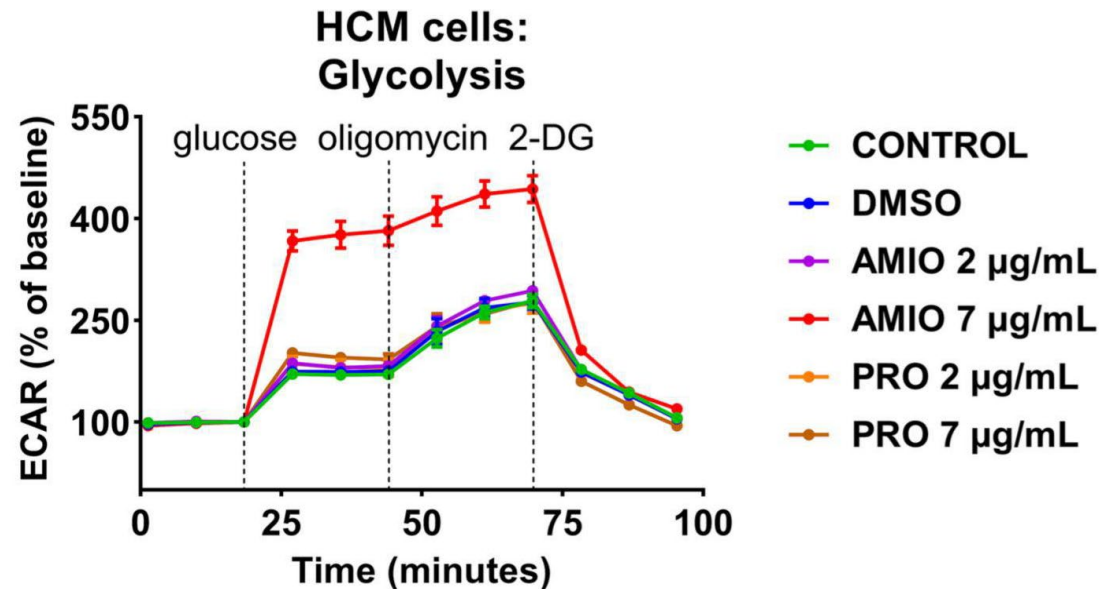
Synonymem antiarytmika v intenzivní péči se stala nejvíce toxická medikace !



Amiodarone but not propafenone impairs bioenergetics and autophagy of human myocardial cells

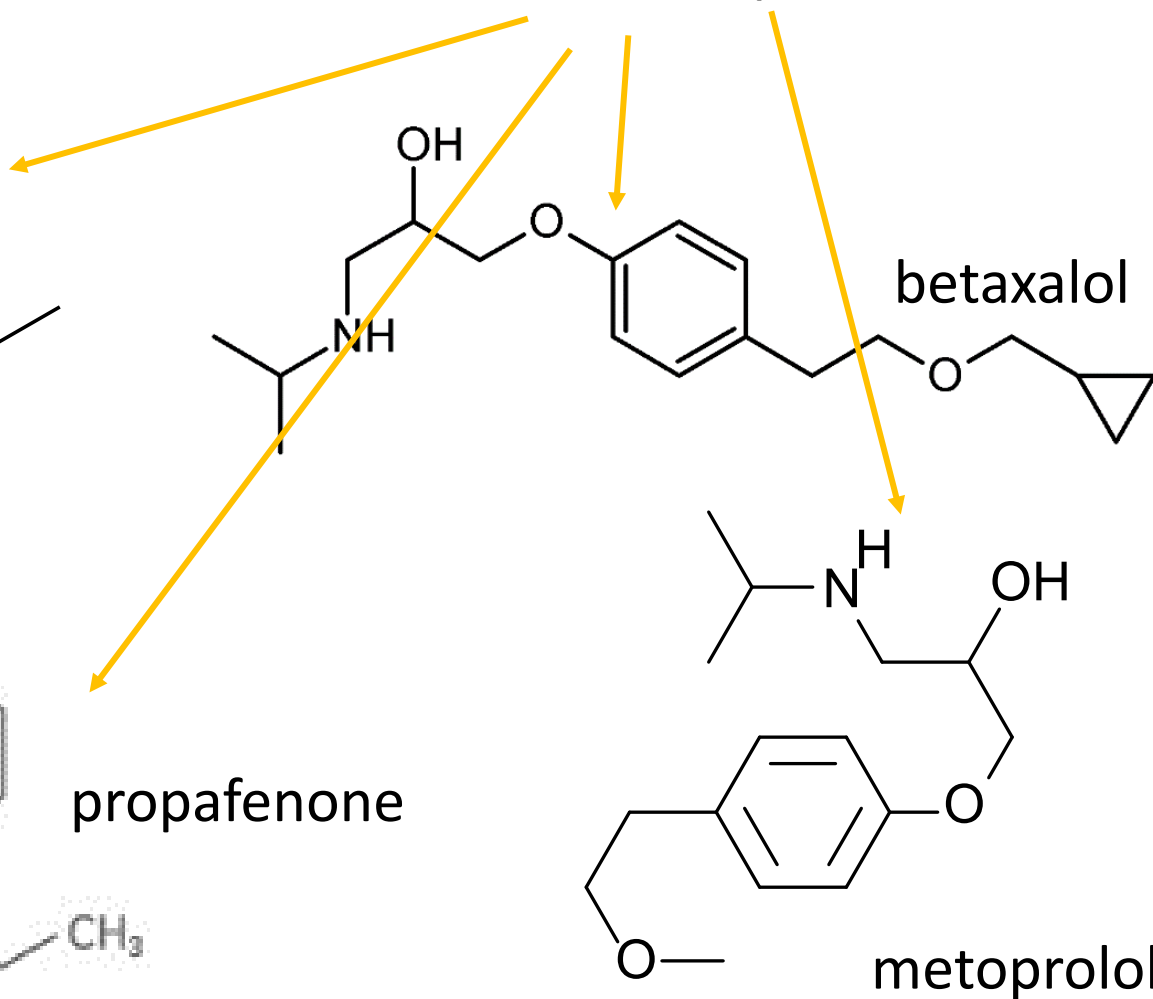
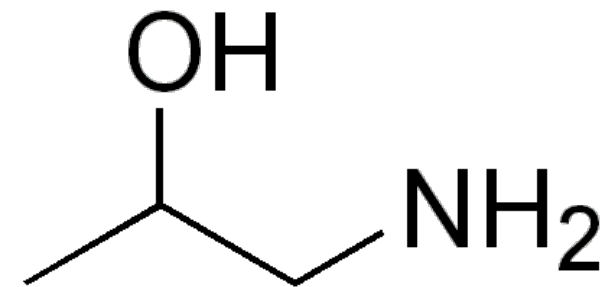
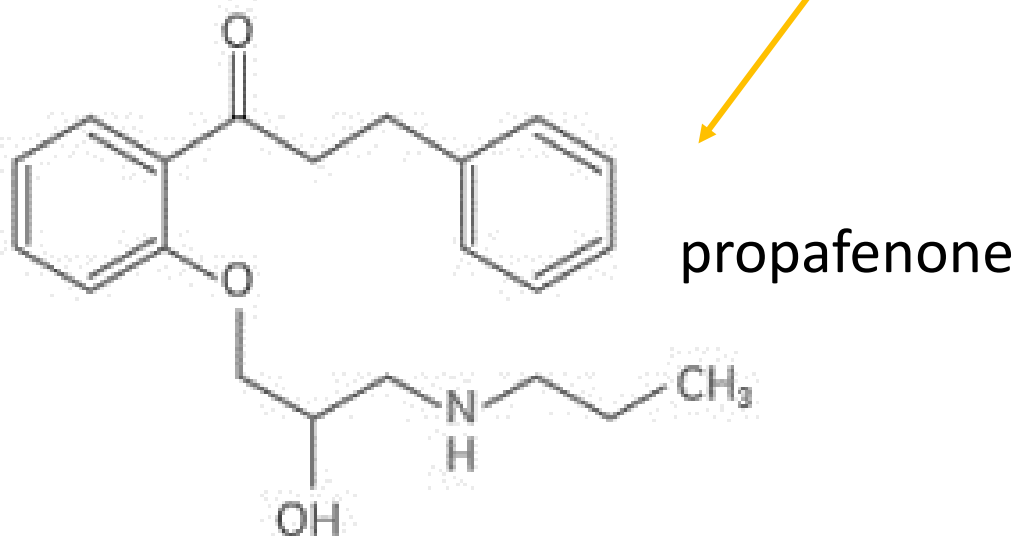
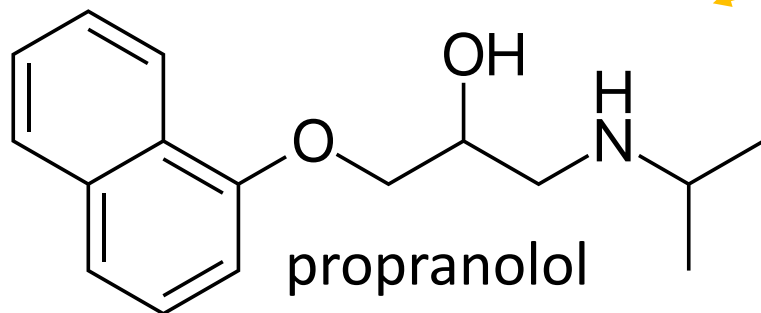
Adéla Krajčová^a, Vlasta Němcová^b, Milada Halačová^{a,c}, Petr Waldauf^a, Martin Balík^d, František Duška^{a,*}

- 24h expozice terapeutickým hladinám amiodaron nebo propafenon: 2 a 7 ug/ml
- Amiodaron s dose-dependentním efektem na mitochondrial uncoupling, přepnutí z oxidativního metabolismu na anaerobní glykolýzu s poklesem ATP
- Autofagie lidských kardiomyocytů
- Propafenone bezpečnější na buněčné úrovni
- Klinická implikace pro vysoké dávky amiodaronu ?



Propanolamine jako chemický základ betablokátorů...a většiny antiarytmik

- Klasifikace antiarytmik: Vaughan-Williams
 - Class II – betablokátoři
 - Class Ic - propafenone

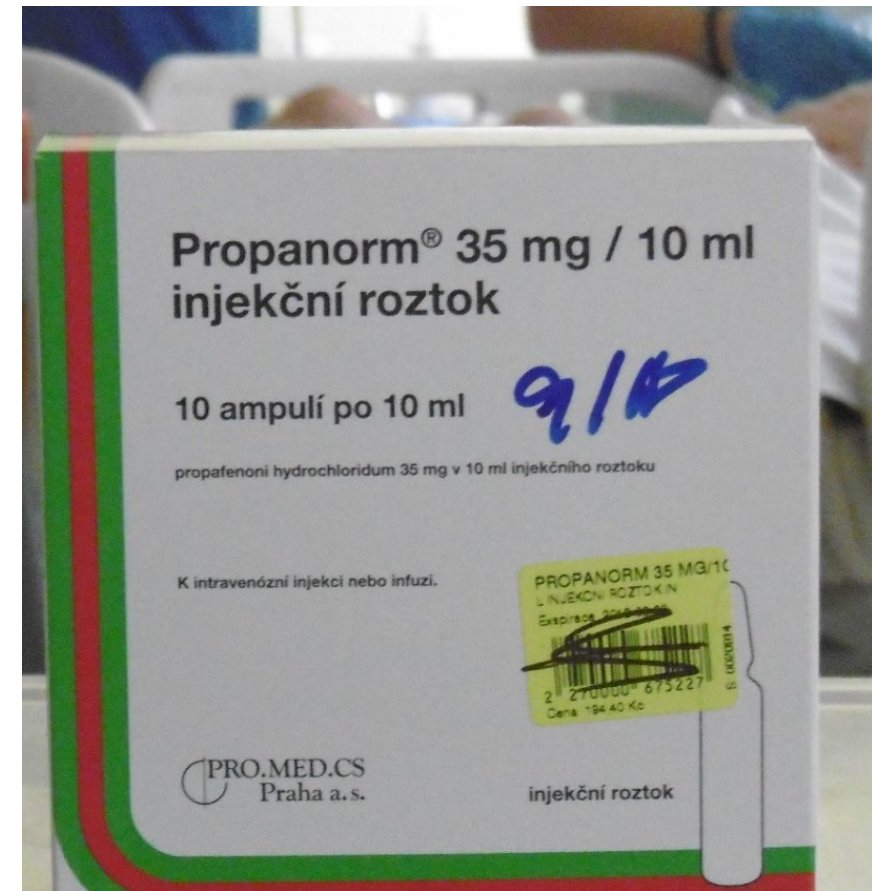


Propafenon

- Účinné a rychlé antiarytmikum (1-2h)
- Kontraindikován u
 - Těžké dysfunkce LK
 - Převodních poruch > Ist AV block
- Monitoring PQ
- Pozor na interakce v ARF a při zhoršení jater
- Flecainide/encainide nepopulární na ICU pro reportované fatální kazuistiky – dose related toxicity



CASE REPORT



Cardiology Journal
2013, Vol. 20, No. 2, pp. 203–205
DOI: 10.5603/CJ.2013.0035
Copyright © 2013 Via Medica
ISSN 1897–5593

Arrhythmogenic effect of flecainide toxicity

Pierre-Yves Courand¹, Franck Sibellas¹, Sylvain Ranc¹, Audrey Mullier²,
Gilbert Kirkorian¹, Eric Bonnefoy¹

Obavy z 1c třídy V-W ?

The New England Journal of Medicine

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Volume 324

MARCH 21, 1991

Number 12

MORTALITY AND MORBIDITY IN PATIENTS RECEIVING ENCAINIDE, FLECAINIDE, OR PLACEBO

The Cardiac Arrhythmia Suppression Trial

- CAST Trial – pacienti s ischemic. chor. srdeční (post AMI) exponováni 1c class vs placebo
- 1c class vyšší mortalita, incl. EF_LV < 30% (!)
- Kardiologie, ambulantni – follow up 10 měsíců
- Nejvíce citovaná studie 1c třídy není aplikovatelná u kriticky nemocných !!!

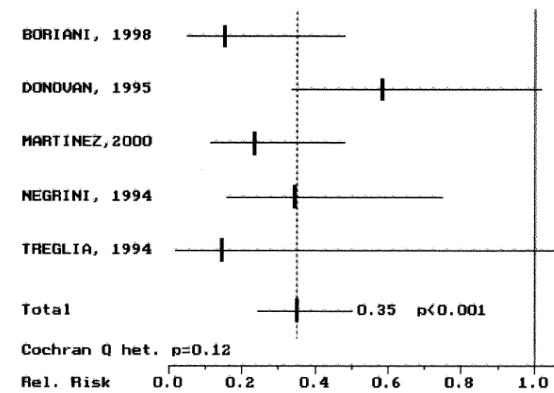
Journal of the American College of Cardiology
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Vol. 41, No. 2, 2003
ISSN 0735-1097/03/\$30.00
PII S0735-1097(02)02705-5

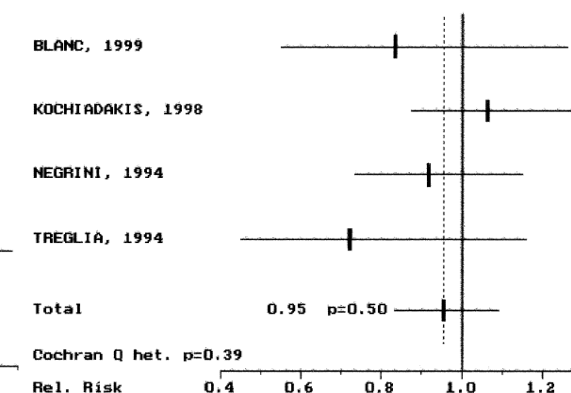
Amiodarone Versus Placebo and Class Ic Drugs for Cardioversion of Recent-Onset Atrial Fibrillation: A Meta-Analysis

Philippe Chevalier, MD, PhD,* Alexis Durand-Dubief, MD,* Haran Burri, MD,* Michel Cucherat, MD,†
Gilbert Kirkorian, MD,* Paul Touboul, MD*

Sinus Rhythm 1-2 Hrs



Sinus Rhythm 24 Hrs



- 6 studií na amiodarone vs placebo a 6 na 1c class vs placebo
- propafenon peak efekt v 1-2h, srovnatelné s amiodaronem ve 24h



CrossMark

234 patients with septic shock and SV arrhythmias

163 (69.7%) atrial fibrillation

34 (14.5%) chronic atrial fibrillation

27 (11.5%) SVT

10 (4.3%) atrial flutter

Propafenone for supraventricular arrhythmias in septic shock—Comparison to amiodarone and metoprolol☆☆☆

M. Balik^{a,*}, I. Kolnikova^a, M. Maly^a, P. Waldauf^b, G. Tavazzi^c, J. Kristof^a**Fáze I: Antiarytmická účinnost (24h)**

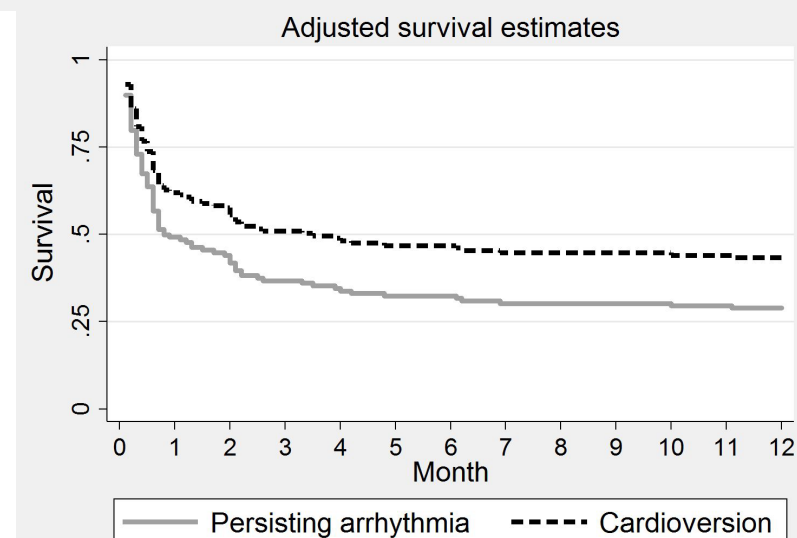
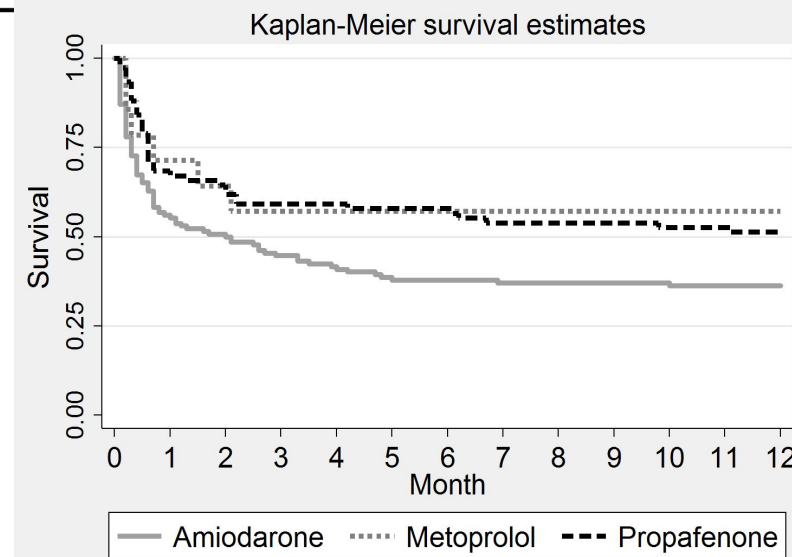
- Současná elektr. KV (23.7% amiodarone, 35.5% propafenone, ns)
- KV: 73.5% amiodarone, 88.9% propafenone, 92.3% metoprolol

Fáze II: Outcome

- ICU mortalita: amio 40.4%, propaf 30.4%, metoprolol 21.4% (all ns)
- 28-d. mortalita: amio 49.6%, propaf 39.5%, metoprolol 21.4% (all ns)

Zdůvodnění PRCT: outcome ve 12 měsících

- Přežití v propafenon jako metoprolol, vyšší než v amiodarone
- Adjusted 12m survival: HR amiod vs propaf 1.58 (1.04; 2.4), p=0.03
- Multivariatní analýza: 12m HR kardioverze versus akutní arytmie: HR 0.67, p=0.113



Propafenone vs Amiodarone in Septic Shock

Journal of Critical Care 41 (2017) 16–23



Contents lists available at ScienceDirect

Journal of Critical Care

journal homepage: www.jccjournal.org



Cardiovascular Drugs and Therapy
<https://doi.org/10.1007/s10557-020-06998-8>

SHORT COMMUNICAITON



Vasopressin in Patients with Septic Shock and Dynamic Left Ventricular Outflow Tract Obstruction

Martin Balik¹ • Adam Novotny¹ • Daniel Suk¹ • Vojtech Matousek¹ • Michal Maly¹ • Tomas Brozek¹ • Guido Tavazzi²

Propafenone for supraventricular arrhythmias in septic shock—Comparison to amiodarone and metoprolol☆☆☆

M. Balik^{a,*}, I. Kolnikova^a, M. Maly^a, P. Waldauf^b, G. Tavazzi^c, J. Kristof^a



Propafenone for supraventricular arrhythmias in septic shock—Comparison to amiodarone and metoprolol



International Journal of Cardiology 266 (2018) 147–148



Contents lists available at ScienceDirect

International Journal of Cardiology

journal homepage: www.elsevier.com/locate/ijcard



Reply



REVIEW



Anaesthesiology Intensive Therapy
2017, vol. 49, no 5, 419–429
ISSN 0209–1712
10.5603/AIT.2017.0061

Editorial

New-onset atrial fibrillation in critically ill patients – Implications for rhythm rather than rate control therapy?☆☆☆☆



M. Balik

Department of Anesthesiology and Intensive Care, 1st Faculty of Medicine, Charles University, General University Hospital, U nemocnice 2, Prague 2 128 08, Czechia

Open access

BMJ Open Efficacy and safety of 1C class antiarrhythmic agent (propafenone) for supraventricular arrhythmias in septic shock compared to amiodarone: protocol of a prospective randomised double-blind study

Management of arrhythmia in sepsis and septic shock

Martin Balik, Vojtech Matousek, Michal Maly, Tomas Brozek

Propafenone vs Amiodarone in Septic Shock, ClinicalTrials.gov ID: NCT03029169

ORIGINAL

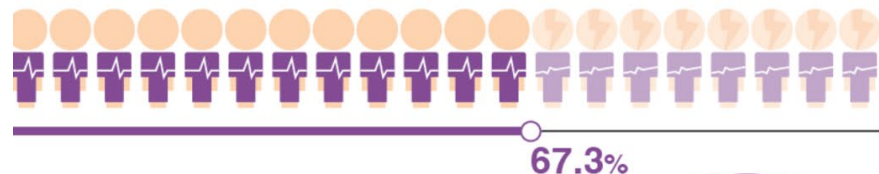
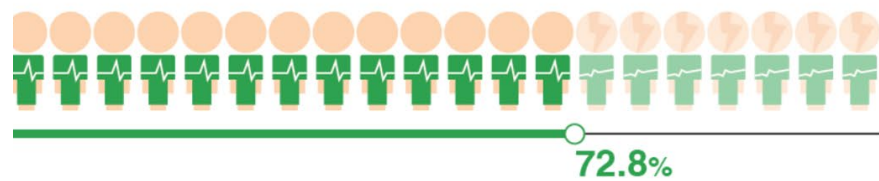
Propafenone versus amiodarone for supraventricular arrhythmias in septic shock: a randomised controlled trial

Martin Balik^{1*}, Michal Maly¹, Tomas Brozek¹, Jan Rulisek¹, Michal Porizka¹, Robert Sachl¹, Michal Otahal¹, Petr Brestovansky¹, Eva Svobodova¹, Marek Flaksa¹, Zdenek Stach¹, Jan Horejssek¹, Lukas Volny¹, Ivana Jurisinova¹, Adam Novotny¹, Pavel Trachta¹, Jan Kunstyr¹, Petr Kopecky¹, Tomas Tencer², Jaroslav Pazout², Jan Belohlavek³, Frantisek Duska², Adela Krajcova² and Petr Waldauf²

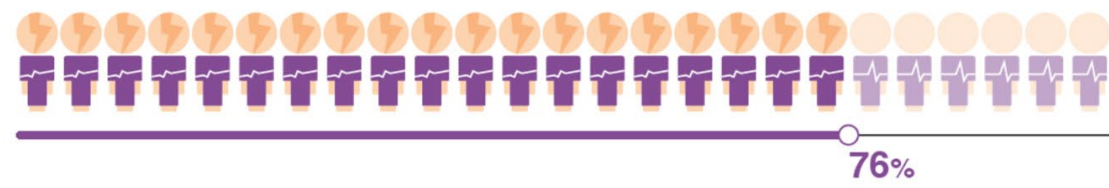
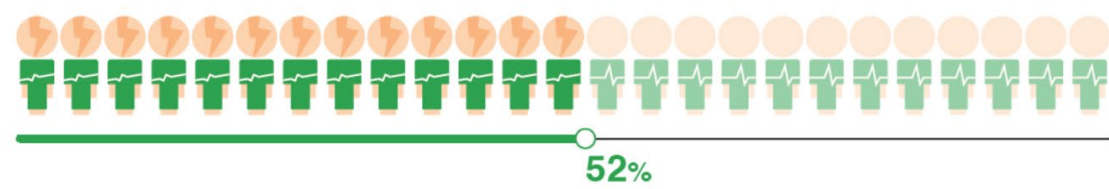


RESULTS

PATIENTS IN SINUS RHYTHM AFTER 24H



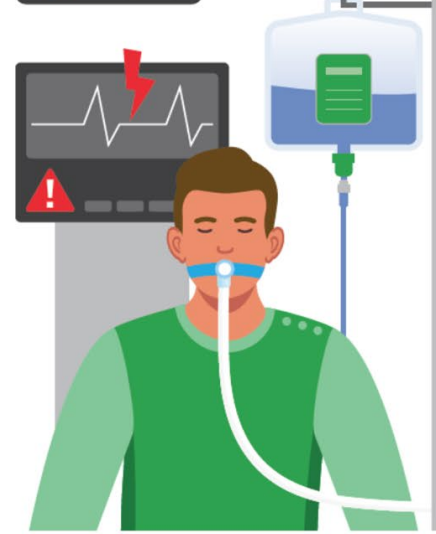
ARRHYTHMIA RECURRENCES



104 PATIENTS with supraventricular arrhythmia and a left ventricular ejection fraction above 35%.

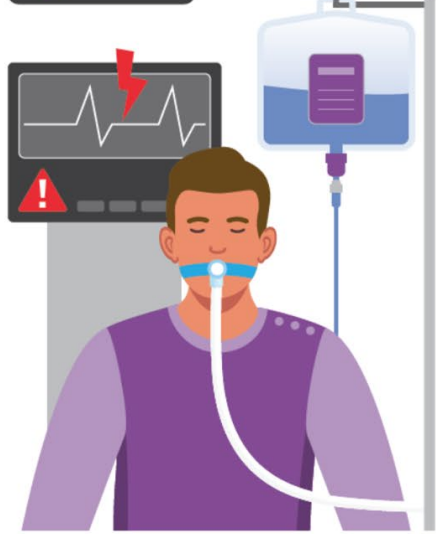
PROPAFENONE GROUP

104 PATIENTS



AMIODARONE GROUP

105 PATIENTS

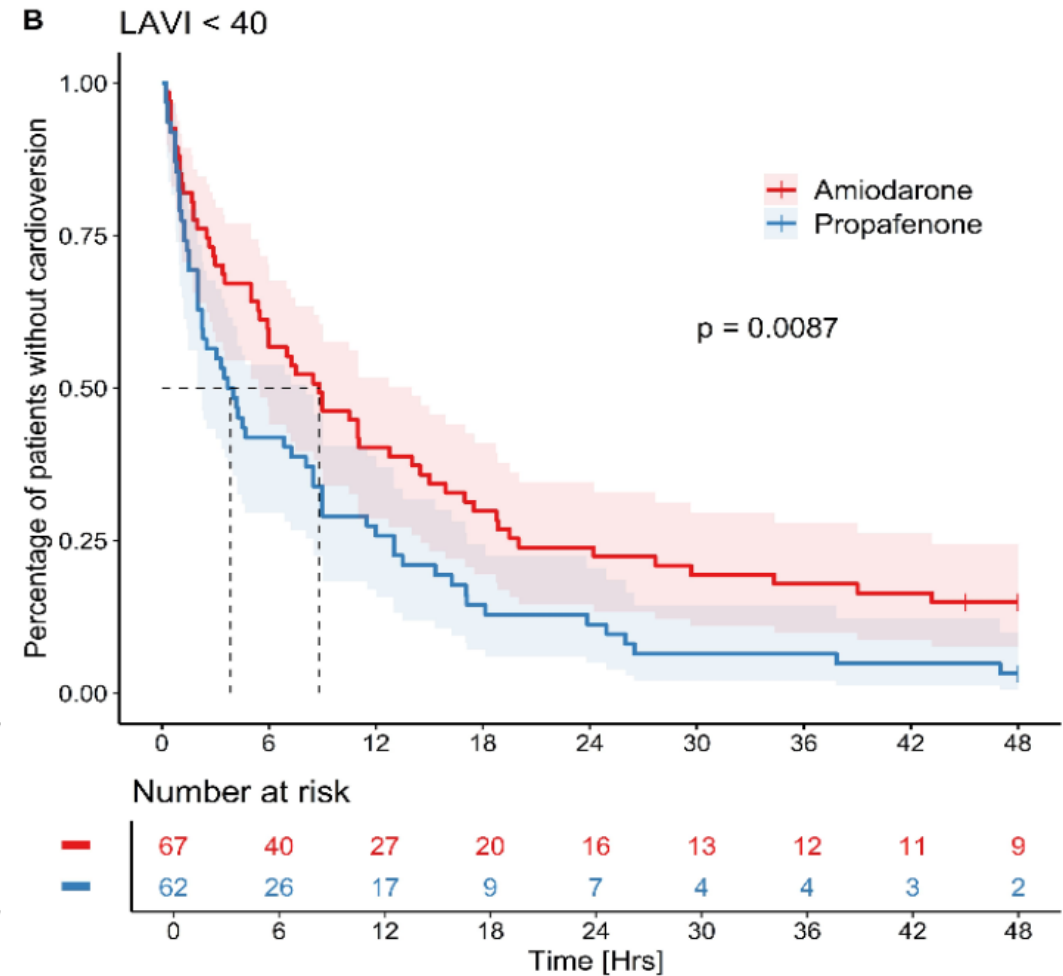
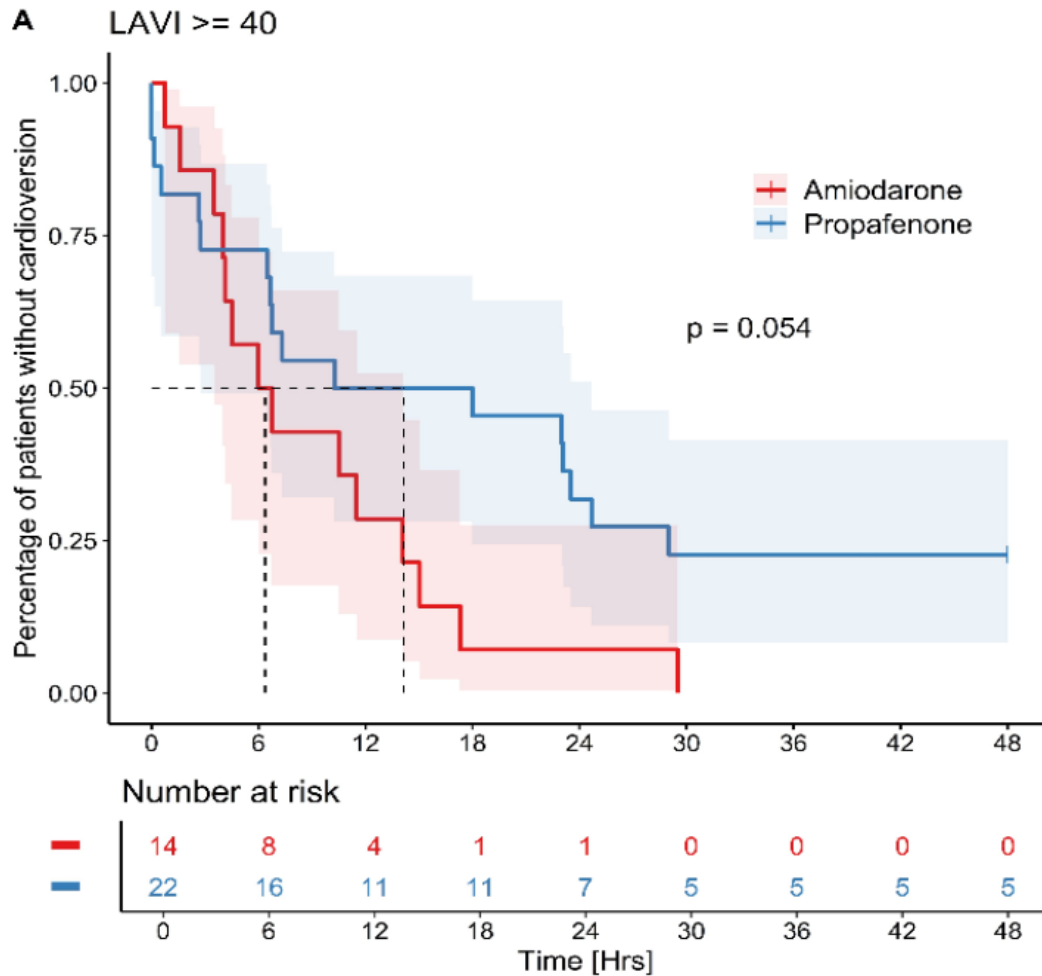


Nedilatovaná LA: p=0.04 pro primary outcome SR ve 24h (propafenon vs amio)

24h: LAVI < 40 ml/m² sinus v 81.3% propafenon vs 65.7% amiodaron (p=0.04)

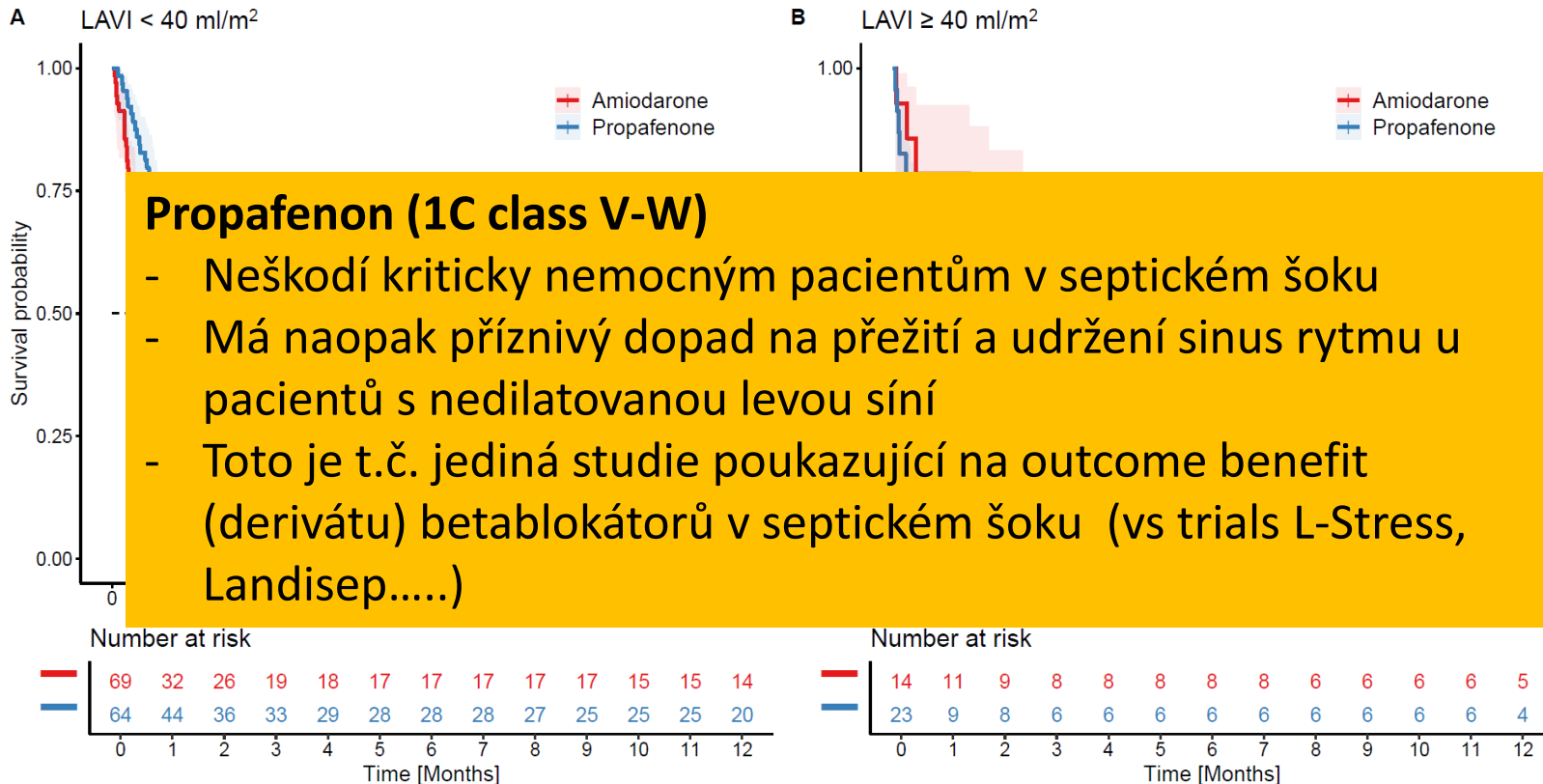
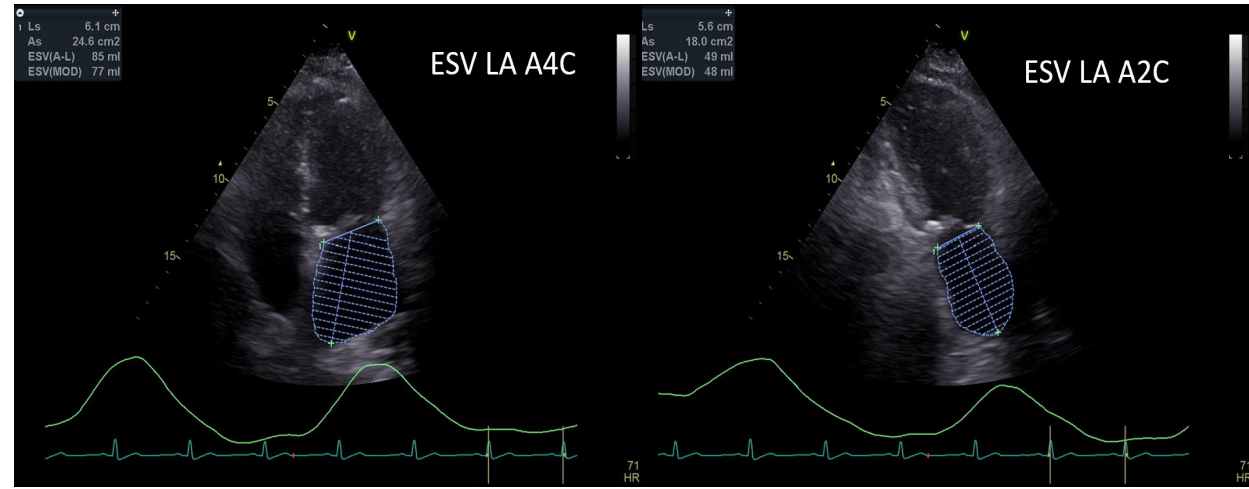
LAVI < 40 ml/m² lépe s propafenon (p=0.009): 3.6 (2.0; 7.25)h vs 8.5 (5.0;12.8)h v amiodaron

LAVI ≥ 40 ml/m² v amiodaronu (p=0.05): 6.4 (3.5; 14.1)h do KV vs 18 (2.8; 24.7)h v propafenonu



The outcomes of patients with septic shock treated with propafenone compared to amiodarone for supraventricular arrhythmias are related to end-systolic left atrial volume

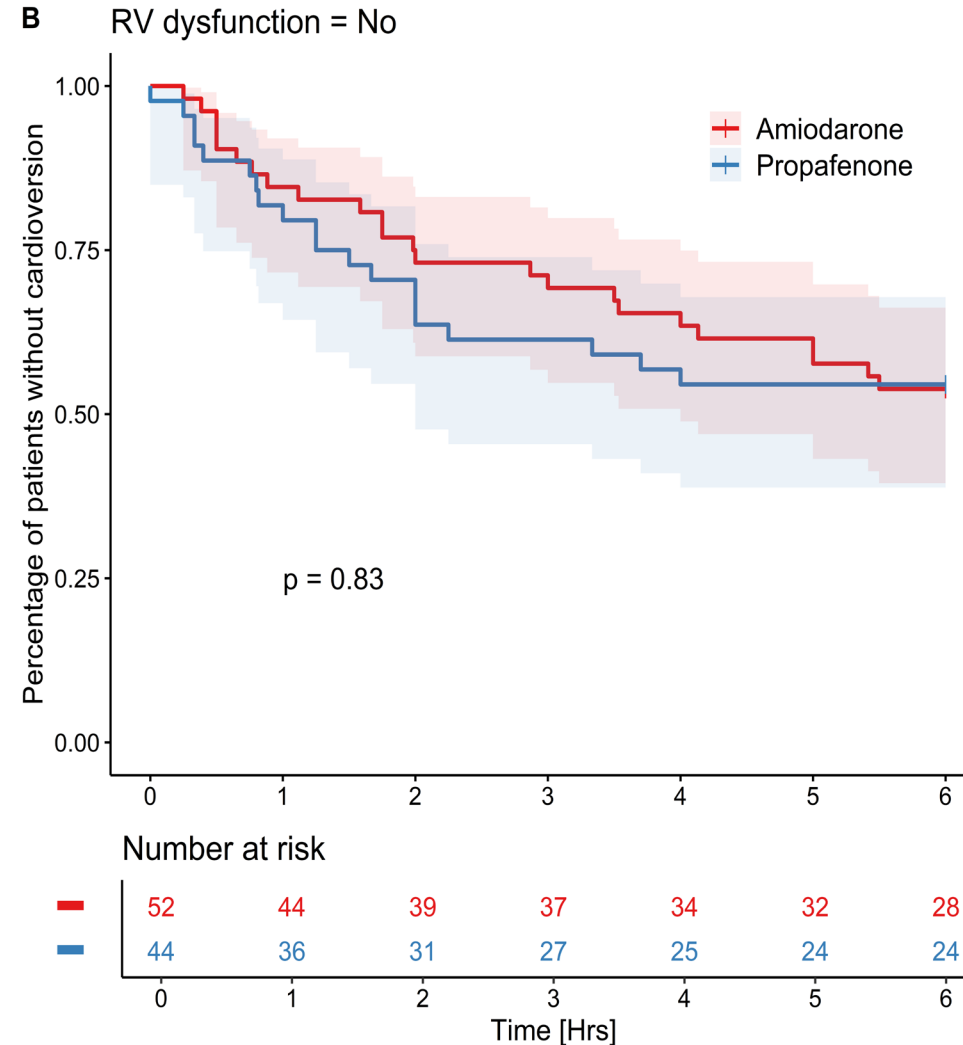
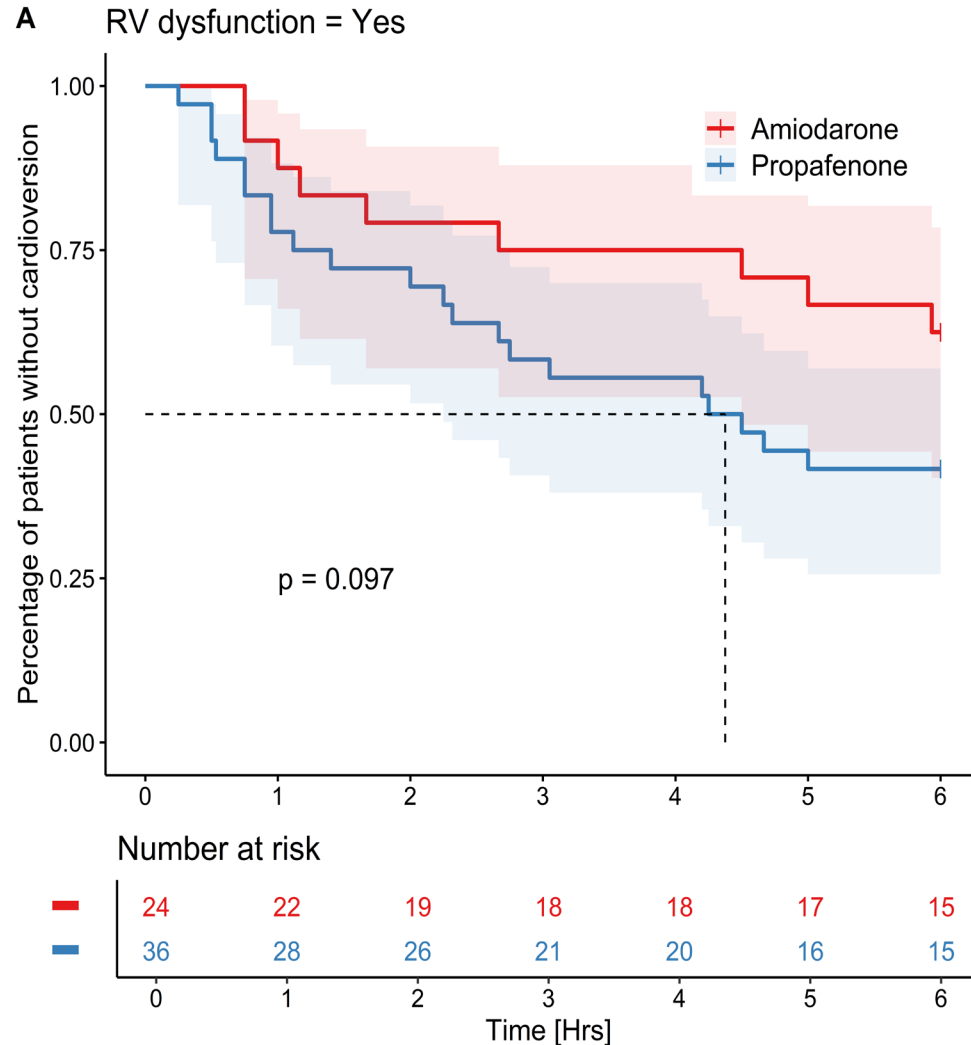
Petr Waldauf¹, Michal Porizka², Jan Horejsek², Michal Otahal², Eva Svobodova², Ivana Jurisinova², Michal Maly², Tomas Brozek², Jan Rulisek², Pavel Trachta², Tomas Tencer¹, Adela Krajcova¹, Frantisek Duska¹, and Martin Balik^{2*}



- LAVI < 40 ml/m²
 - mortality benefit propafenonu 1-měsíc (p=0.007), 1-rok (p=0.013)
 - HR 0.60 pro mortalitu 1-rok (p=0.014)
- LAVI ≥ 40 ml/m²
 - Mortalita horší s propafenone 1-měsíc (p=0.03) a 6-m (p=0.052)
 - HR 3.6 pro mortalitu 1-měsíc (p=0.045)

Dysfunkční PK a SVA – propafenon nesignifikantně lepší

- 61(37.9%) pac. se středně-těžce dilatovanou PK, TAPSE<15 mm, PAPs>40 mmHg versus 100(62.1%) pac. s normální až mírně dysfunkční PK.

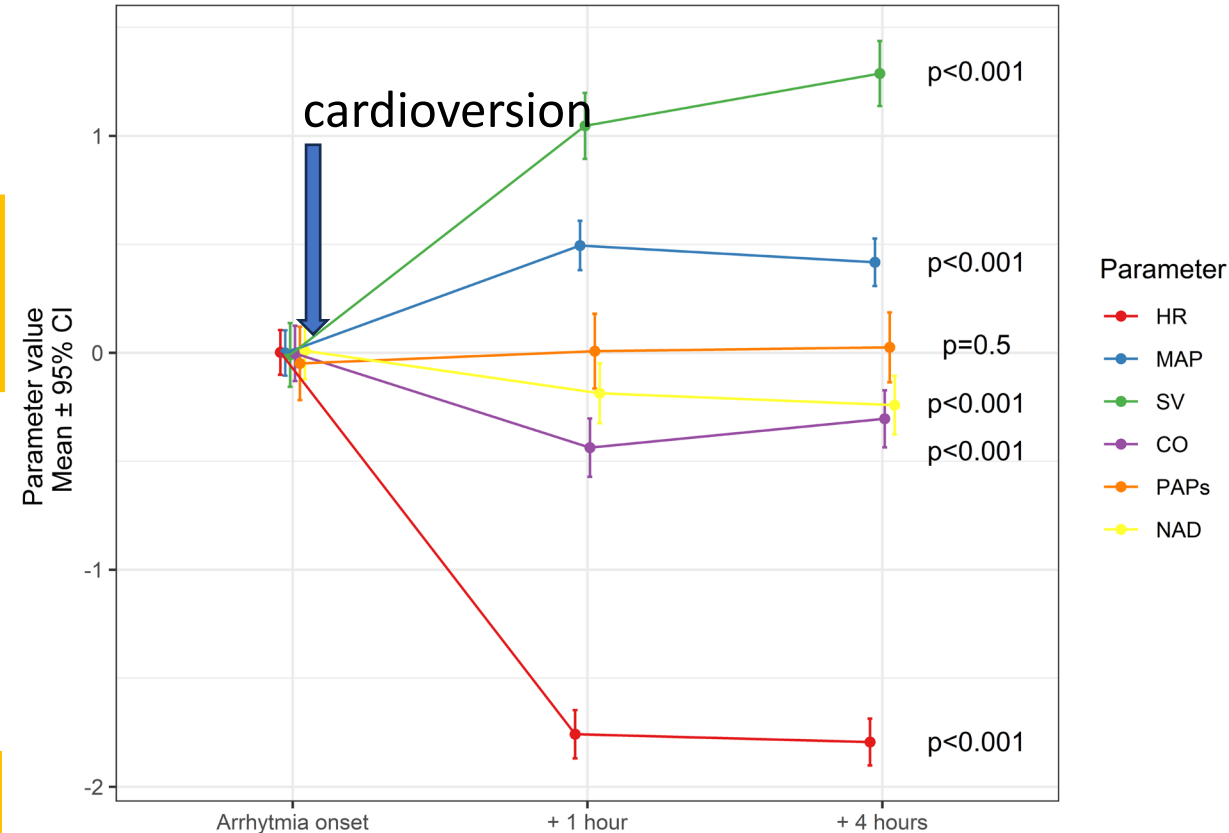


Co se změní s kardioverzí v septickém šoku ? Vliv pulzatilního toku....

Parameter	Arrhythmia onset (n=209)	Cardioversion +1h (n=173)	Cardioversion +4h (n=187)	p-value
HR (/min)	130 (111; 147)	85 (78; 96)	85 (77; 94)	<0.001
MAP (mmHg)	75 (70; 82)	80 (75; 85)	80 (75; 85)	<0.001
CVP (mmHg)	8.6 (8; 9.2)	8 (7.4; 8.6)	8.1 (7.5; 8.7)	0.01
SV (ml)	50 (42; 60)	66 (58; 78)	69 (60; 79)	<0.001
CO (l/min)	6.315 (5.332; 7.654)	5.496 (4.849; 6.656)	5.787 (4.862; 6.901)	<0.001
CI (l/min.m ²)	3.165 (2.618; 3.792)	2.839 (2.401; 3.304)	2.932 (2.450; 3.371)	<0.001

- MAP (↑) = CO (↓) x SVR(...NAD↓)..?
- arytmie vs sinus po kardioverzi

PAPs (mmHg)	44 (37; 54)	47 (38; 54)	46 (37; 54)	0.5
SVR (dyn/s.cm ⁻⁵)	867 (825; 909)	1064 (1020; 1107)	1003 (961; 1046)	<0.001
NAD (µg/kg.min)	0.30 (0.15; 0.45)	0.25 (0.12; 0.40)	0.21 (0.10; 0.40)	<0.001
change of dosage		-0.05 (-0.09; -0.02)	-0.07 (-0.1; -0.04)	
AVP (IU/h)	2.00 (2.00; 3.00)	2.00 (0.00; 2.00)	2.00 (0.00; 2.00)	<0.001
change of dosage		-0.76 (-1.09; -0.42)	-0.84 (-1.17; -0.52)	
N of patients	N=55	N=42	N=49	
Dobutamine (µg/kg.min)	3.00 (2.75; 3.00)	3.00 (2.50; 4.00)	4.00 (3.00; 5.00)	0.6
N of patients	N=11	N=9	N=7	



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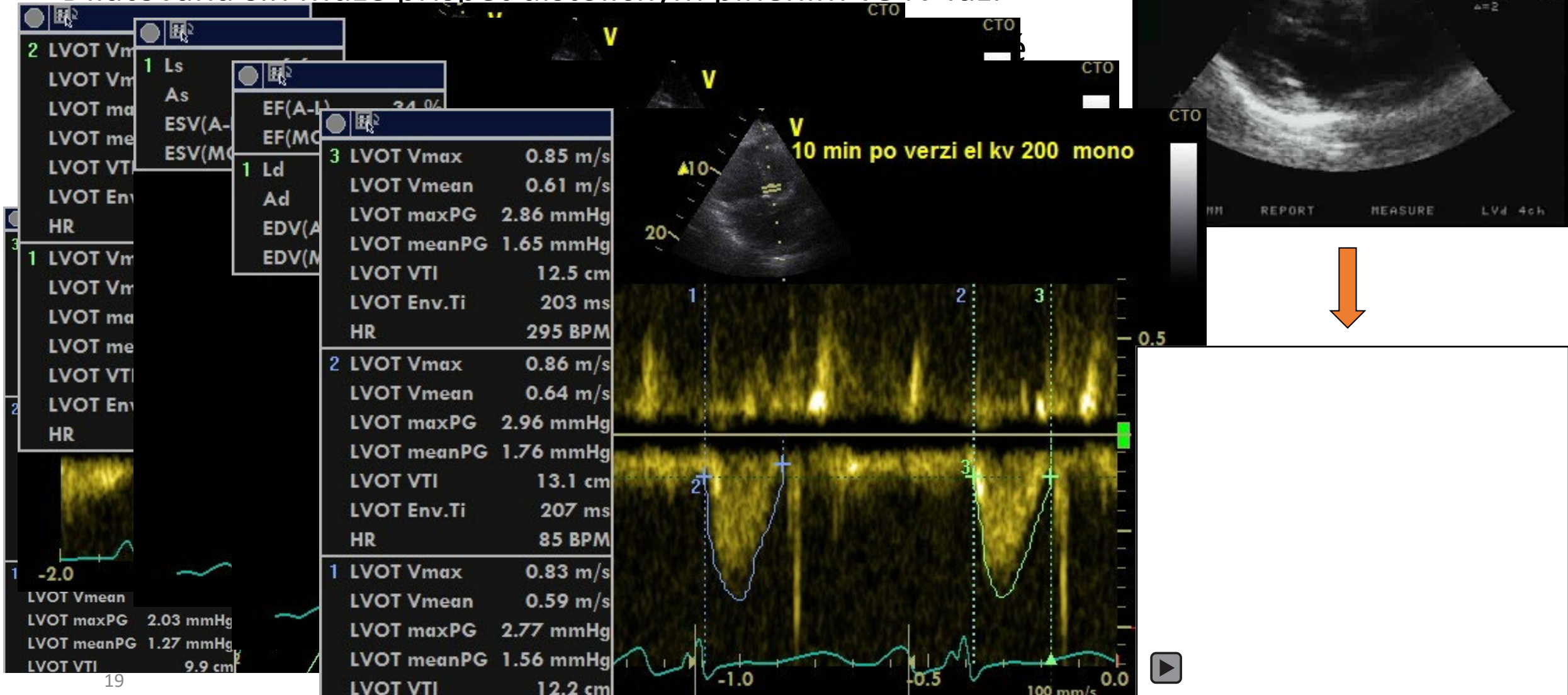


Echocardiography predictors of sustained sinus rhythm after cardioversion of supraventricular arrhythmia in patients with septic shock

M. Balik^{a,*}, P. Waldauf^{b,1}, M. Maly^a, T. Brozek^a, J. Rulisek^a, M. Porizka^a, R. Sachl^a,

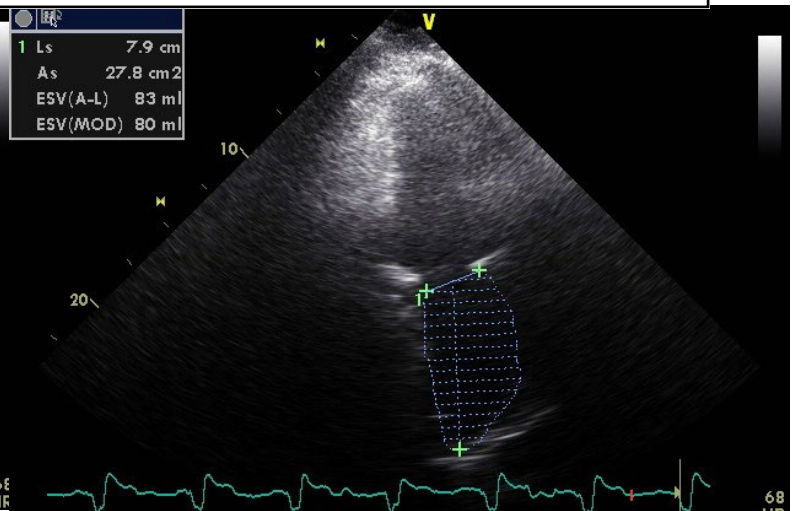
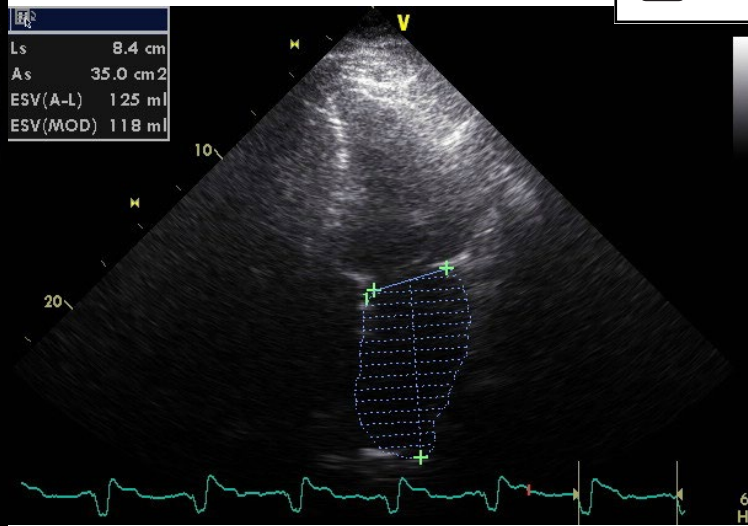
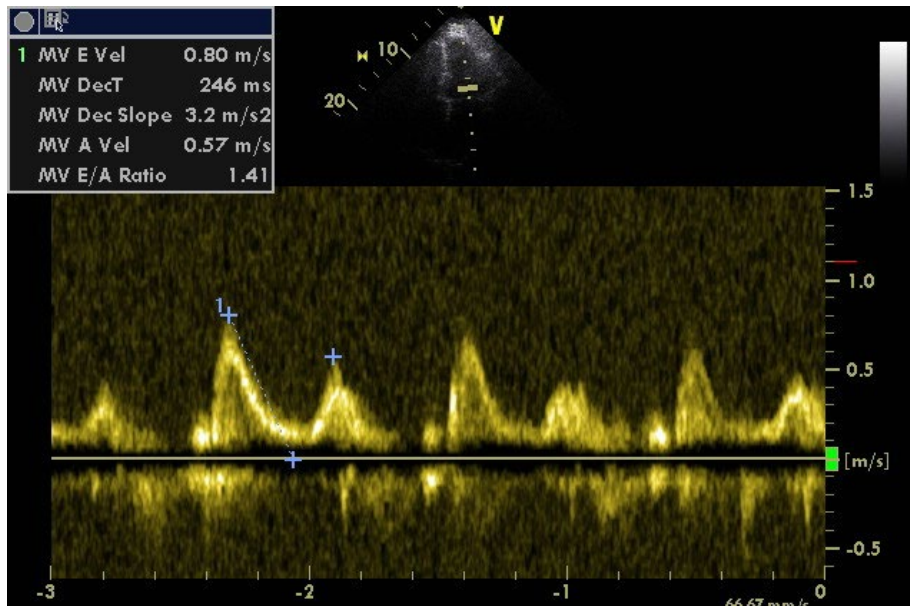
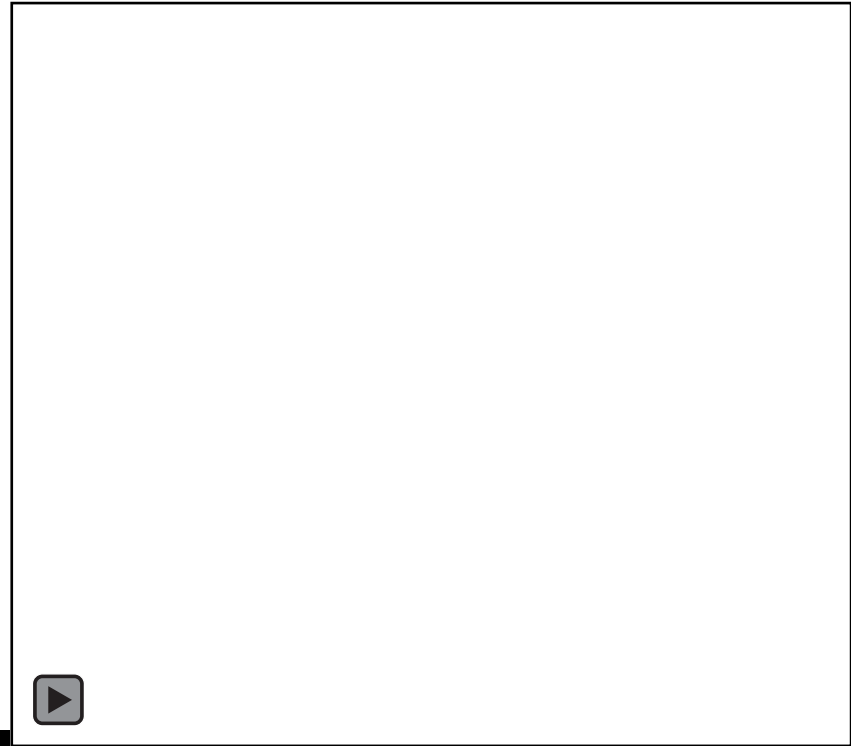
2D a Doppler monitoring atriální funkce – může zdůvodnit rhythm control strategy

- Dilatovaná síň může přispět distolickým plněním ve IV fázi



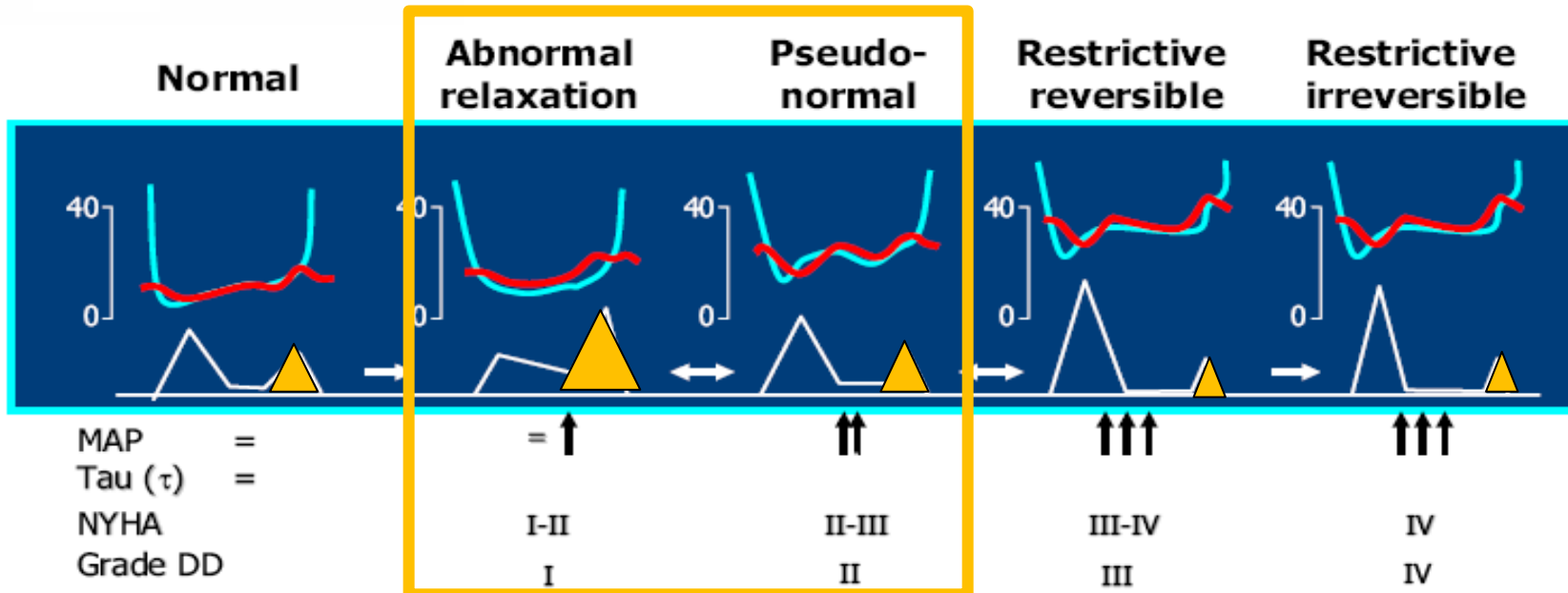
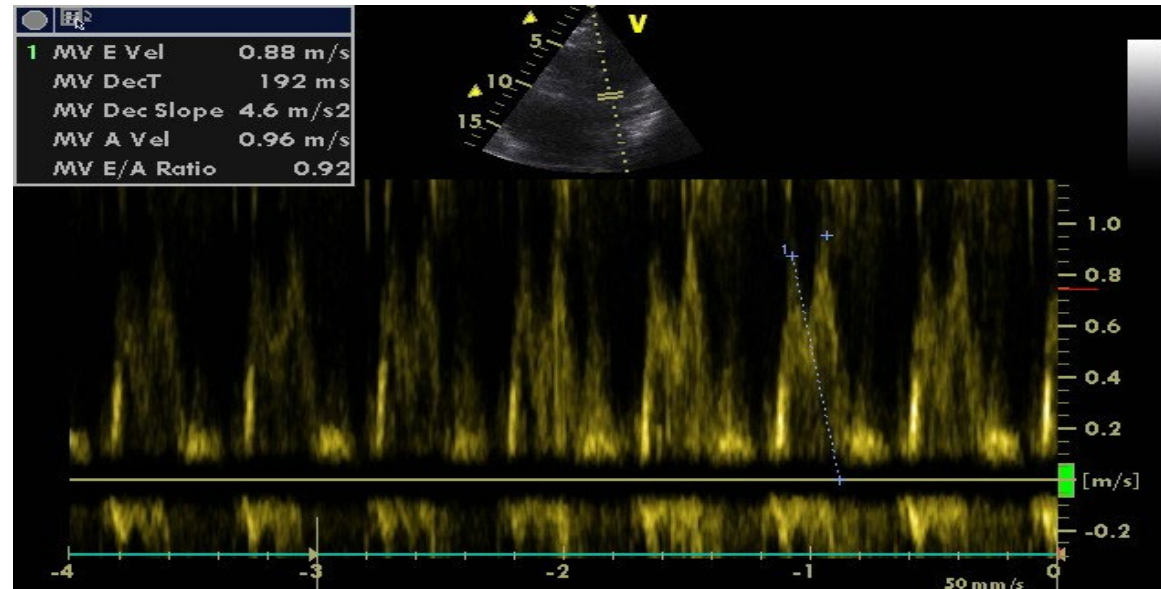
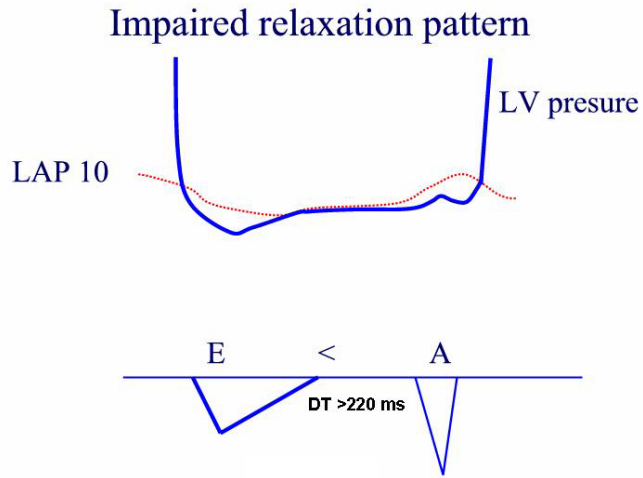
LA_EF a návrat systoly síní jako prediktor udržení sinusového rytmu

- Cut off 40 ml/m² LAVi modifikován
- Chronicky zvýšeným LAP
- Preload
- Indikátor diastolic. dysfunkce – ne kontraindikace ke KV a rhythm control
- A wave a Avti 1-4h po kardioverzi mohou zdůvodnit rhythm control strategii



LA emptying fraction 32%

Stupeň závislosti komorového plnění na systole síní (= sinus rytmus)



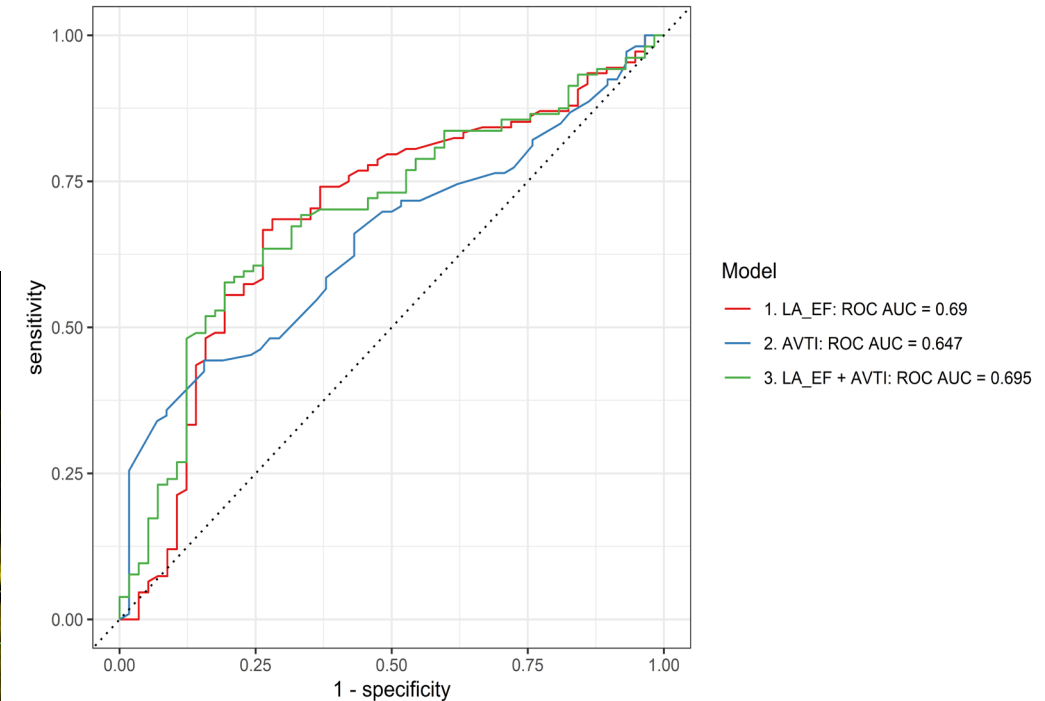
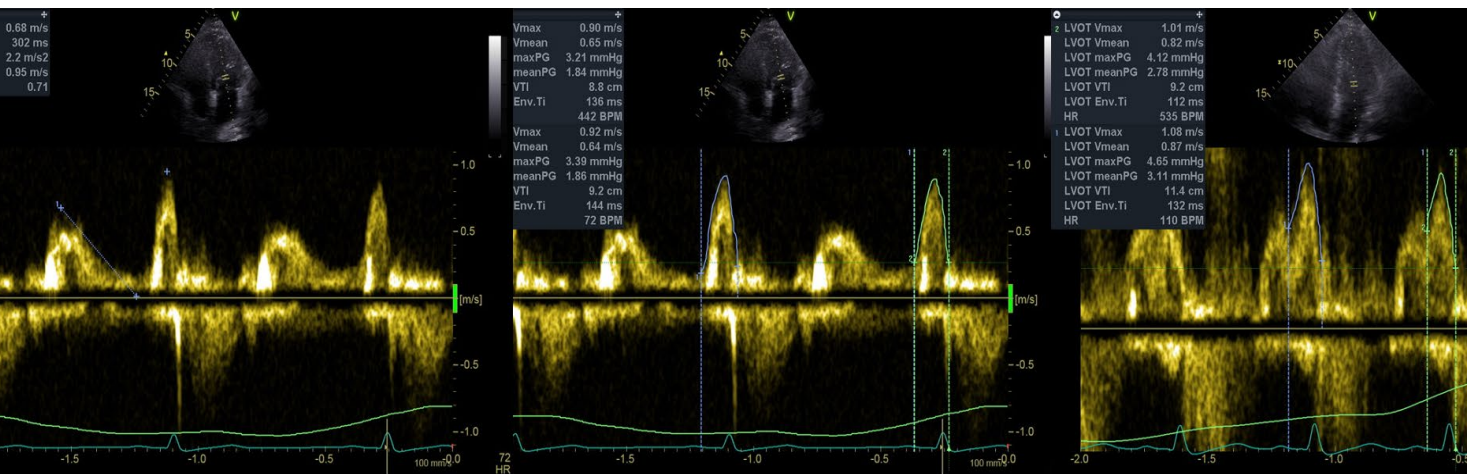


- LA_ESD a LAVI nestačí k predikci udržení SR (cut off ?)

• Echo guided kardioverze !

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Parameter at CV +4h N of patients	0 recurrence N=65	1 recurrence N=41	2 recurrences N=28	3 recurrences N=16	>3 recurrences N=37	p-value
LA_EF (%)	44 (36, 49)	37 (28, 43)	31 (28, 38)	35 (31, 42)	30 (25, 36)	<0.001
Avti (cm)	8.65 (7.13, 9.50)	8.00 (6.25, 9.85)	7.85 (5.88, 8.85)	6.80 (6.00, 8.10)	6.50 (5.65, 8.30)	0.007

Echo a nastavení antikoagulace

Roční riziko centrální embolizace ve vztahu k velikosti LA a funkci LV

	<i>risk(%)</i>
• AF + normal echocardiogram	1·5
• AF + LA > 25 mm/m ²	8·8
• AF + global LV dysfunction	12·6
• AF + LA > 25 mm/m ² + moderate/severe LV dysf.	20·0
• Stroke Prevention in Atrial Fibrillation Study Group Investigators. Ann Intern Med 1992	
• Aiyrala S, et al: JASE 2011; 24(5): 499-505	
	
• Moderate to severe MR may reduce risk of thrombus formation in dilated LA (Nakagami H: Am Heart J 1998; 136: 528-32)	

Echokardiografie u SV arytmií

- Pravidelný RR (incl. pacing) a $HR < 100/\text{min}$Doppler jako u sinus rhythm (spec +/-80%)
- LVEDP assessment vč. dynamic manuevres
- Valvular. etiology
- NOAF and chronic paroxysmal or persistent: Echo guided cardioversion
- Rate control in chronic permanent AF or on high vasopressors
- LV dysfunkce asociuje s prevalencí AF (až 50% u těžké LV dysfunkce na ICU)
- LA_ESD (PLAX), $\leq 45 \text{ mm}$, biplanar LAVi, $\leq 40 \text{ ml/m}^2$ jako stratifikátory šance na rhythm control
- LA_EF, $\leq 37\%$, A wave vč. Avti $\leq 8 \text{ cm}$ po KV jako prediktor udržení SR
- Echo guided priority pro antikoagulaci, vyloučení rizika embolizace (TEE) SVA>48h

Echokardiografie esenciální pro farmakoterapii + ev. elektrickou KV

- Ic class (propafenone) u normální až moderate LV dysf., Mg^{2+} , K^+
- III class (amiodarone) u kontraindikace propafenonu
- II class (betablockers) u hyperkinetické cirkulace a hypertenze



Děkuji za pozornost !

Komplexní kardiovaskulární centrum
1. lékařská fakulta, Karlova Univerzita
Všeobecná fakultní nemocnice

U nemocnice 2; 128 08, Praha, EU

T: +420 224 962 243

F: +420 224 962 118

E: martin.balik@vfn.cz

www.karim-vfn.cz

