



Insuficiência Cardíaca

Novos Horizontes no Tratamento

Moderador: Márcia Noya-Rabelo

Debatedores:

Marcus Andrade

Nei Dantas

Fábio Bulhões

Bráulio Pinna

Carlos Vinicius Espírito Santo

Jackson Brandão

Masculino

65 anos

ECG – QRS 150mms

Fe 30%

Cardiopatía Dilatada

Insuficiência Mitral Severa

NYHA III

Enalapril 20mg 2xdia , Carvedilol 25mg 2 xdia, Furosemida
120mg/dia , Espironolactona 25mg/dia

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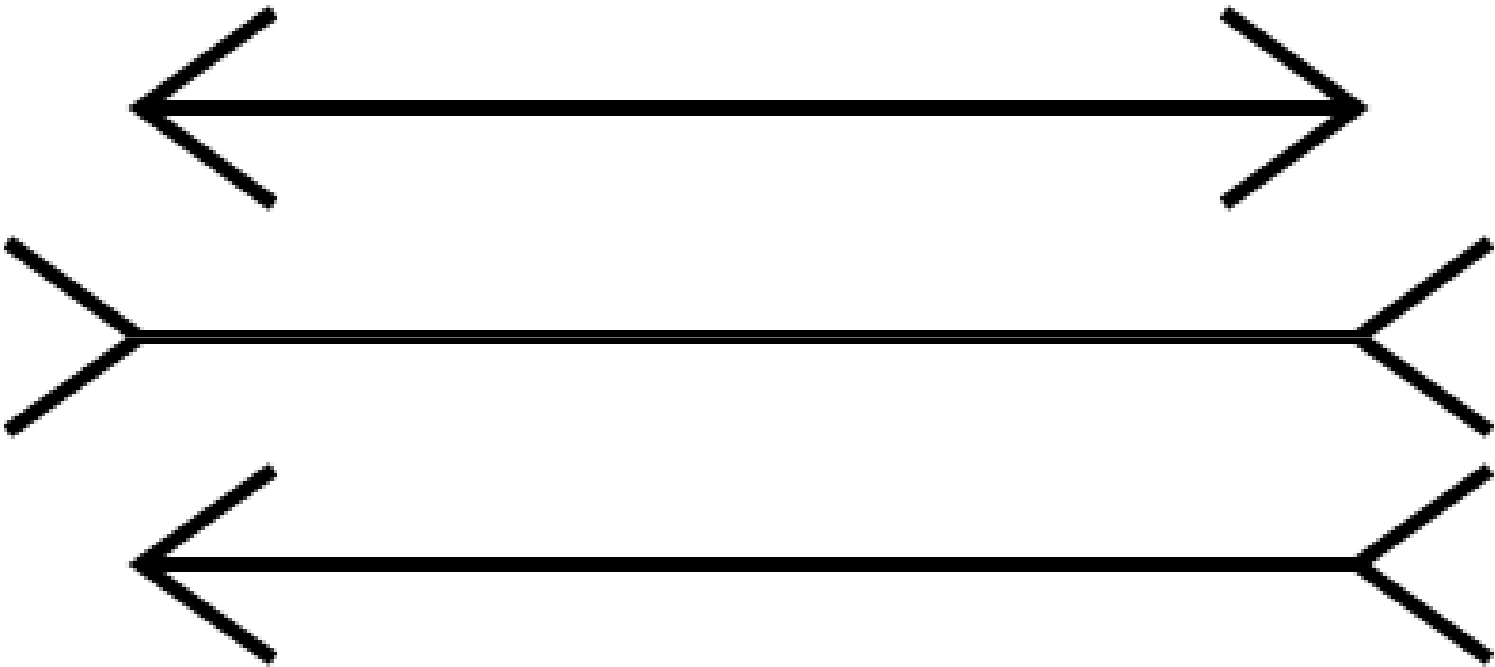
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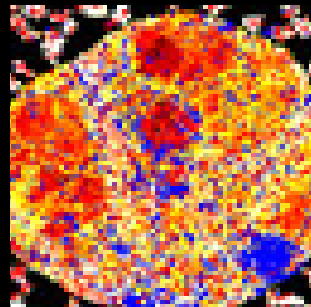
1. Há indicação Valsartana/Sacubutril?
2. Terapia de ressincronização ? Com ou sem CDI?
3. Anticoagulação? Se sim qual anticoagulante?
4. Quando e como abordar a insuficiência mitral secundária?
5. Suporte ventricular mecânico – terapia de destino?
6. Quando e como encaminhar para o Tx?

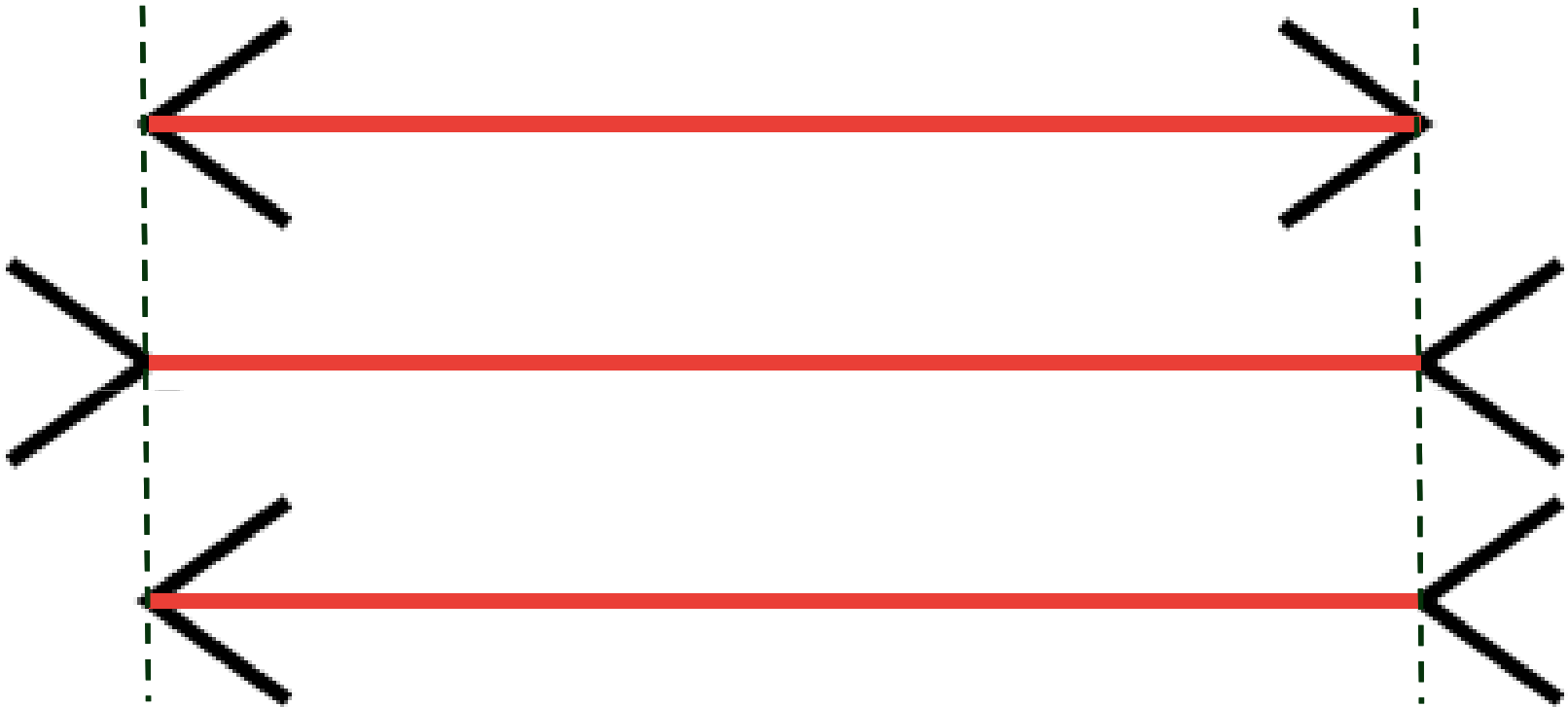


A realidade pode ser tão complexa que as observações feitas de um determinado assunto, vistas de ângulos diferentes, podem parecer bem contraditórias!



Complexidade dos Sistemas Biológicos





Devo trocar IECA por valsartana/sacubitril?

Angiotensin–Neprilysin Inhibition versus Enalapril in Heart Failure

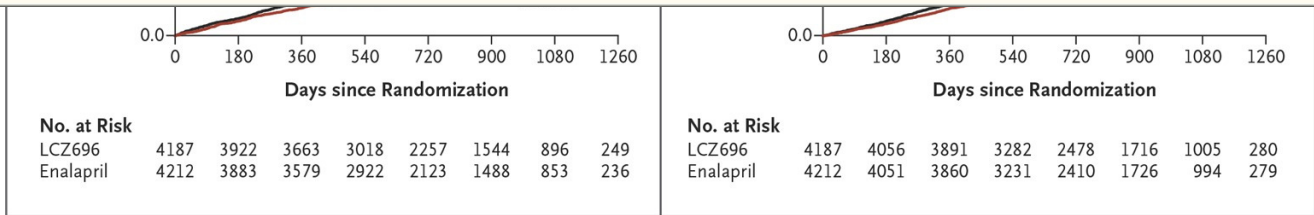
PARADIGM-HF

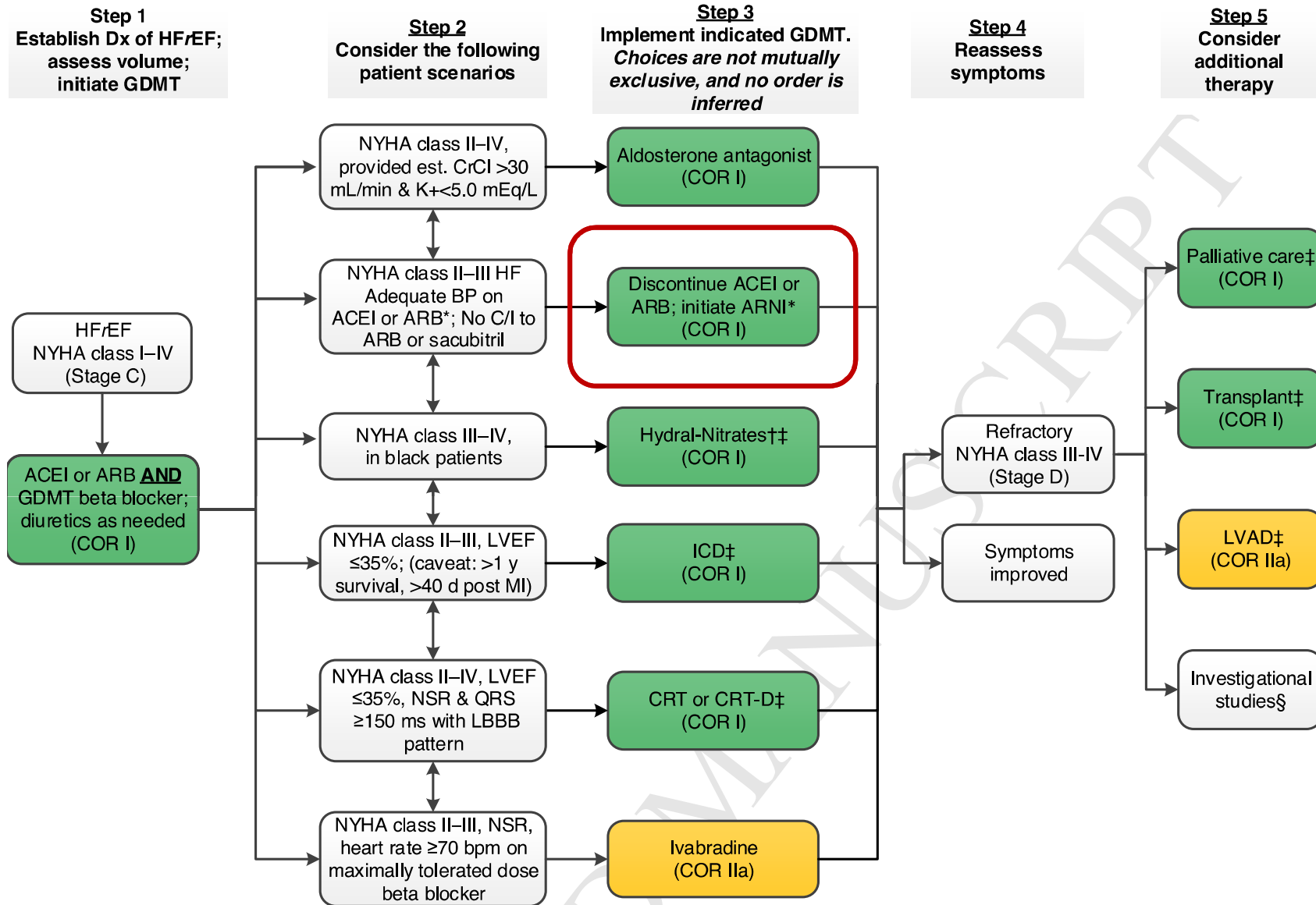


A Primary End Point 1.0	B Death from Cardiovascular Causes 1.0
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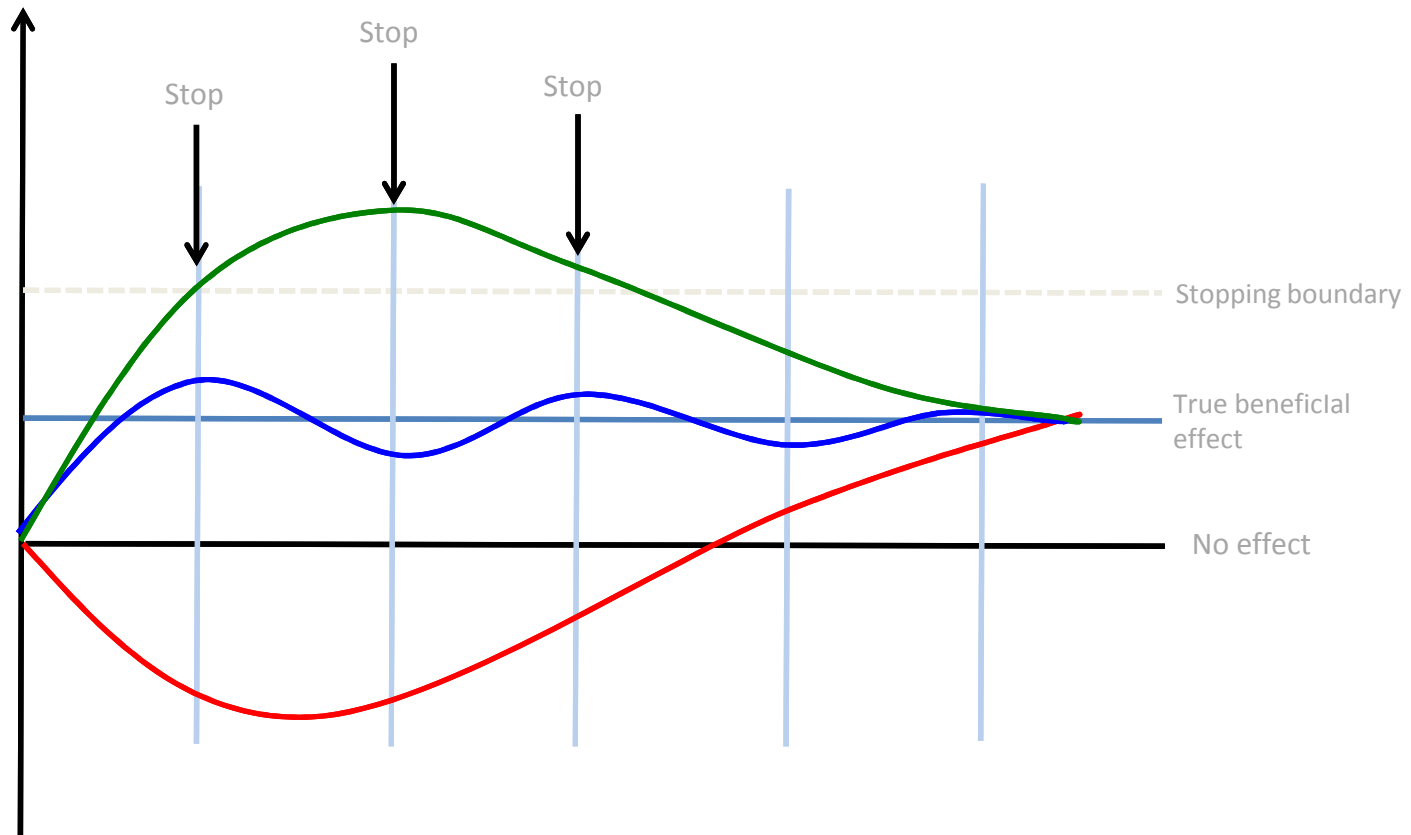
Table 2. Primary and Secondary Outcomes.*

Outcome		LCZ696 (N=4187)	Enalapril (N=4212)	Hazard Ratio or Difference (95% CI)	P Value
Primary composite outcome — no. (%)	NNT: 21				
Death from cardiovascular causes or first hospitalization for worsening heart failure		914 (21.8)	1117 (26.5)	0.80 (0.73–0.87)	<0.001
Death from cardiovascular causes	NNT: 31	558 (13.3)	693 (16.5)	0.80 (0.71–0.89)	<0.001
First hospitalization for worsening heart failure		537 (12.8)	658 (15.6)	0.79 (0.71–0.89)	<0.001
Secondary outcomes — no. (%)	NNT: 35				
Death from any cause		711 (17.0)	835 (19.8)	0.84 (0.76–0.93)	<0.001
Change in KCCQ clinical summary score at 8 mo†		-2.99±0.36	-4.63±0.36	1.64 (0.63–2.65)	0.001
New-onset atrial fibrillation‡		84 (3.1)	83 (3.1)	0.97 (0.72–1.31)	0.83
Decline in renal function§		94 (2.2)	108 (2.6)	0.86 (0.65–1.13)	0.28

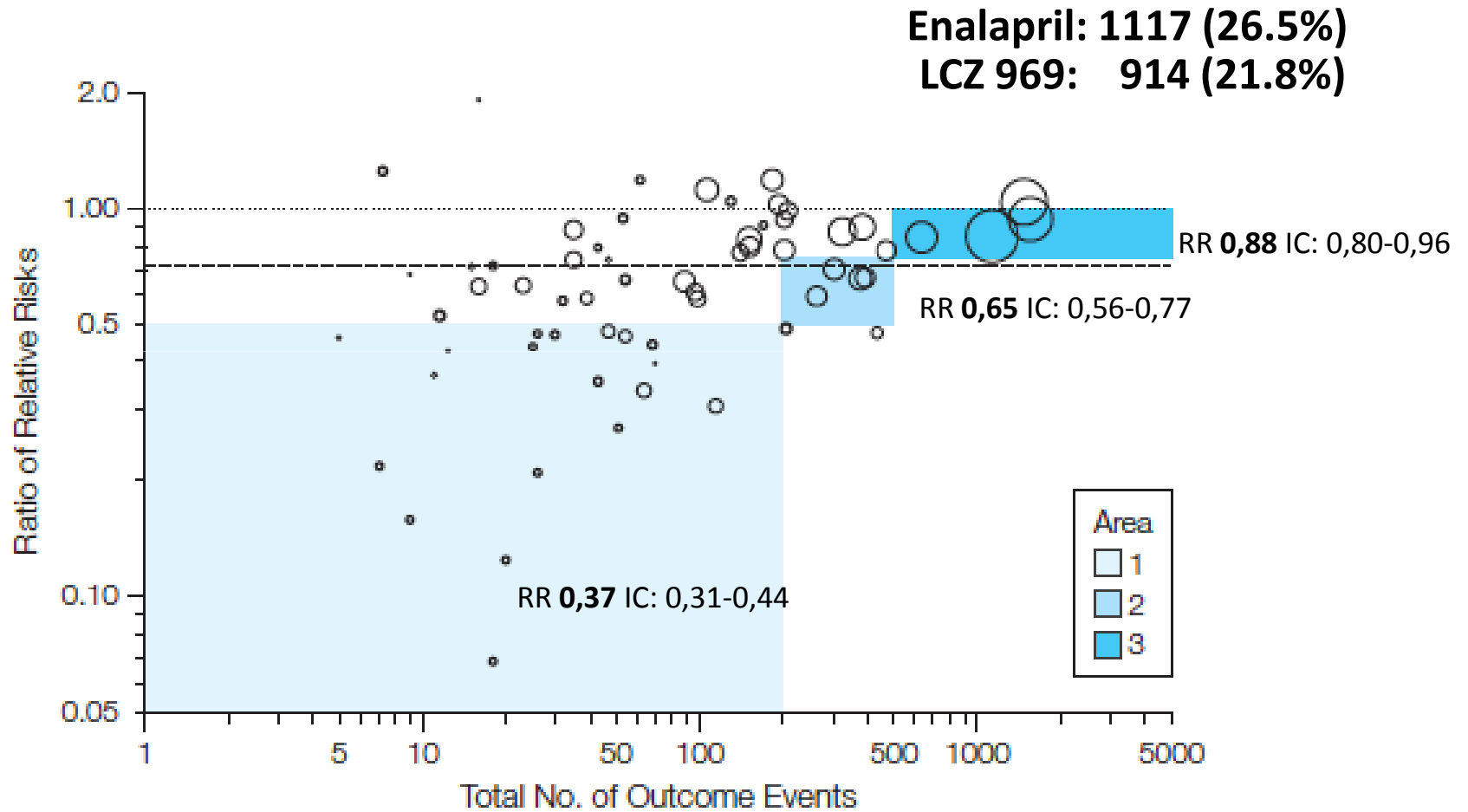




Theoretical Distribution of Randomized Controlled Trial Results as Data Accumulate



Outcomes Events in Truncated RCTs



Entresto Sacubitril / Valsartan

SACUBITRIL; VALSARTAN is a combination of 2 drugs used to reduce the risk of death and hospitalizations in people with long-lasting heart failure. It is usually used with other medicines to treat heart failure. Compare angiotensin receptor neprilysin inhibitors.

Prescription Settings brand tablet 97mg/103mg 60 tablets SAVE SHARE


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Prices and coupons for 60 tablets of Entresto 97mg/103mg

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Target (CVS)	\$422.29 with free coupon	GET FREE COUPON
Membership warehouse <small>Name cannot be shown. Why not?</small>	\$423.14 with free coupon	GET FREE COUPON
Walmart	\$423.91 with free discount	GET FREE DISCOUNT
Safeway	\$426.83 with free coupon	GET FREE COUPON
SAVINGS TIP Manufacturer Coupon Pay as little as \$10 per month. See Tips		
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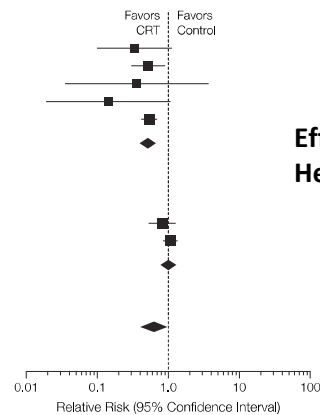
Save even more on your family's prescriptions at Kroger

Terapia de Ressincronização ?
Com ou sem CDI?

Cardiac Resynchronization Therapy for Patients With Left Ventricular Systolic Dysfunction

A Systematic Review

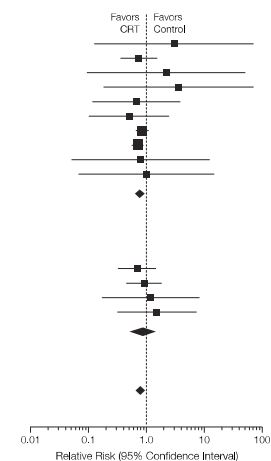
Source	Hospitalization for Heart Failure, No./Total		Relative Risk (95% Confidence Interval)
	CRT	Control	
CRT Alone vs Medical Therapy			
MUSTIC-SR, ⁹ 2001	3/29	9/29	0.33 (0.10-1.11)
MIRACLE, ⁸ 2002	18/228	34/225	0.52 (0.30-0.90)
MUSTIC-AF, ¹¹ 2002	1/25	2/18	0.36 (0.04-3.67)
RD-CHF, ¹⁴ 2003	1/22	7/22	0.14 (0.02-1.07)
CARE-HF, ¹⁹ 2005	72/409	133/404	0.53 (0.42-0.69)
Subtotal	95/713	185/698	0.51 (0.41-0.64)
Test for Heterogeneity: $\chi^2_4 = 2.27$; $P = .68$; $I^2 = 0\%$ Test for Overall Effect: $Z = 5.88$; $P < .001$			
CRT + ICD vs ICD Alone			
CONTAK-CD, ¹³ 2003	32/245	39/245	0.82 (0.53-1.26)
MIRACLE-ICD, ¹² 2003	85/187	78/182	1.06 (0.84-1.33)
Subtotal	117/432	117/427	1.00 (0.80-1.24)
Test for Heterogeneity: $\chi^2_1 = 1.10$; $P = .29$; $I^2 = 8.6\%$ Test for Overall Effect: $Z = 0.04$; $P = .87$			
Total	212/1145	302/1125	0.63 (0.43-0.93)
Test for Heterogeneity: $\chi^2_{14} = 23.08$; $P < .001$; $I^2 = 74.0\%$ Test for Overall Effect: $Z = 2.34$; $P = .02$			



Effect of CRT on Proportion of Patients Hospitalized for Heart Failure

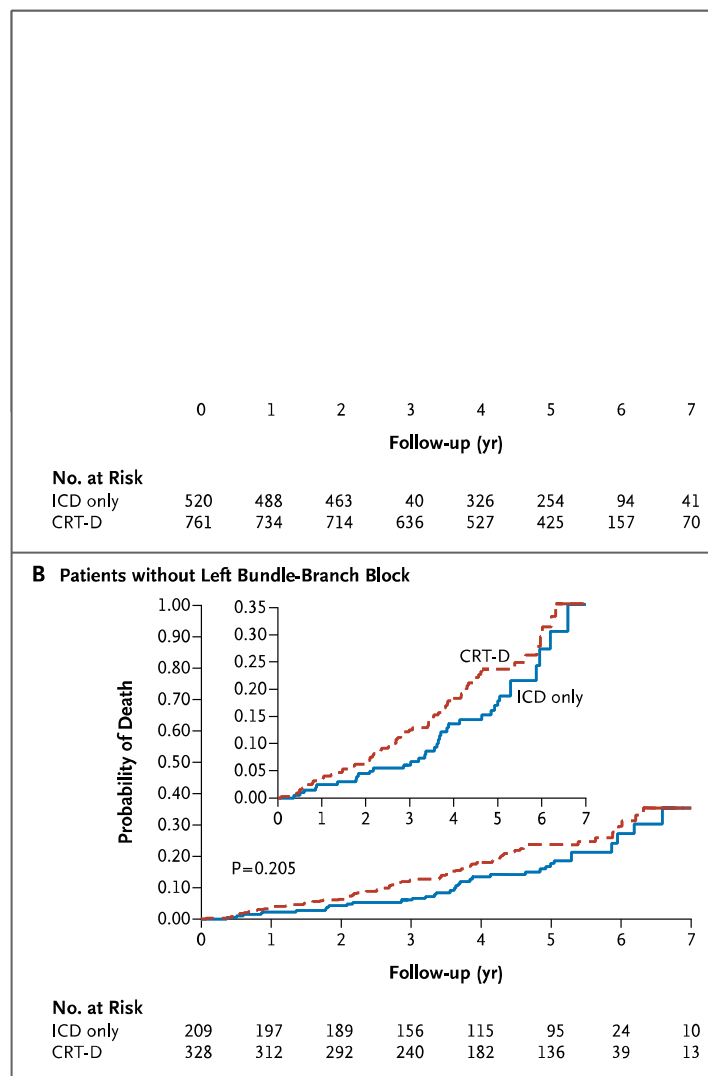
Effect of CRT on All-Cause Mortality

Source	All-Cause Mortality, No./Total		Relative Risk (95% Confidence Interval)
	CRT	Control	
CRT Alone vs Medical Therapy			
MUSTIC-SR, ⁹ 2001	1/29	0/29	3.00 (0.13-70.74)
MIRACLE, ⁸ 2002	12/228	16/225	0.74 (0.36-1.53)
MUSTIC-AF, ¹¹ 2002	1/25	0/18	2.19 (0.09-50.93)
PATH-CHF, ¹⁵ 2002	2/24	0/17	3.60 (0.16-70.54)
PATH-CHF II, ¹⁶ 2003	2/43	3/43	0.67 (0.12-3.79)
RD-CHF, ¹⁴ 2003	2/22	4/22	0.50 (0.10-2.45)
COMPANION, ¹⁸ 2004	131/617	77/308	0.85 (0.66-1.09)
CARE-HF, ¹⁹ 2005	92/409	129/404	0.70 (0.56-0.89)
VECTOR, ²⁰ 2005	1/69	1/47	0.80 (0.05-12.40)
HOBIPACE, ¹⁷ 2006	1/16	1/16	1.00 (0.07-14.64)
Subtotal	245/1472	231/1129	0.77 (0.66-0.91)
Test for Heterogeneity: $\chi^2_5 = 3.72$; $P = .59$; $I^2 = 0\%$ Test for Overall Effect: $Z = 3.16$; $P = .002$			
CRT + ICD vs ICD Alone			
CONTAK-CD, ¹³ 2003	11/245	16/245	0.69 (0.33-1.45)
MIRACLE-ICD, ¹² 2003	14/187	15/182	0.91 (0.45-1.83)
MIRACLE-ICD II, ¹³ 2004	2/85	2/101	1.19 (0.17-8.76)
RHYTHM-ICD, ²¹ 2005	6/119	2/60	1.51 (0.31-7.27)
Subtotal	33/636	35/588	0.86 (0.54-1.39)
Test for Heterogeneity: $\chi^2_3 = 0.97$; $P = .81$; $I^2 = 0\%$ Test for Overall Effect: $Z = 0.60$; $P = .55$			
Total	279/2108	266/1717	0.78 (0.67-0.91)
Test for Heterogeneity: $\chi^2_{17} = 4.90$; $P = .38$; $I^2 = 0\%$ Test for Overall Effect: $Z = 3.18$; $P = .001$			

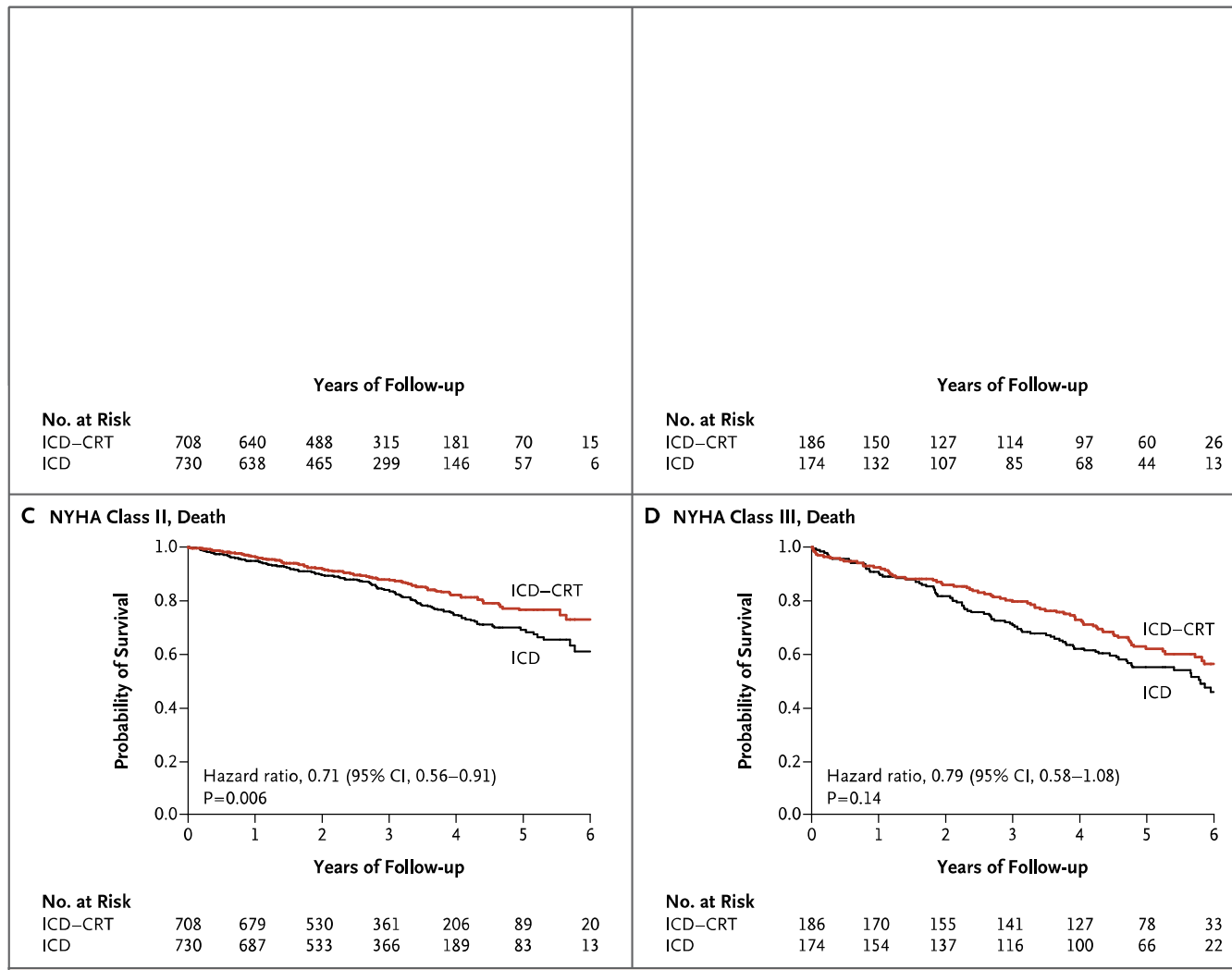


Survival with Cardiac-Resynchronization Therapy in Mild Heart Failure

MADIT-CRT – 7 year follow-up



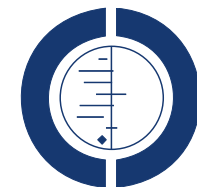
Cardiac-Resynchronization Therapy for Mild-to-Moderate Heart Failure RAFT Trial



Anticoagulação?

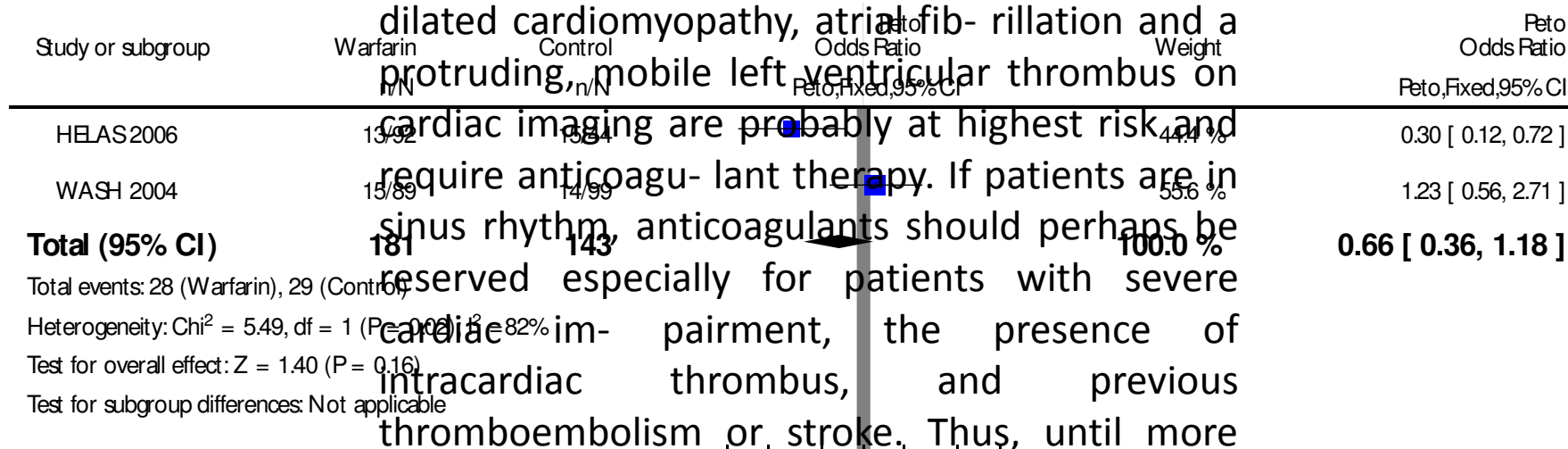
Se sim qual anticoagulante?

Anticoagulation versus placebo for heart failure in sinus rhythm



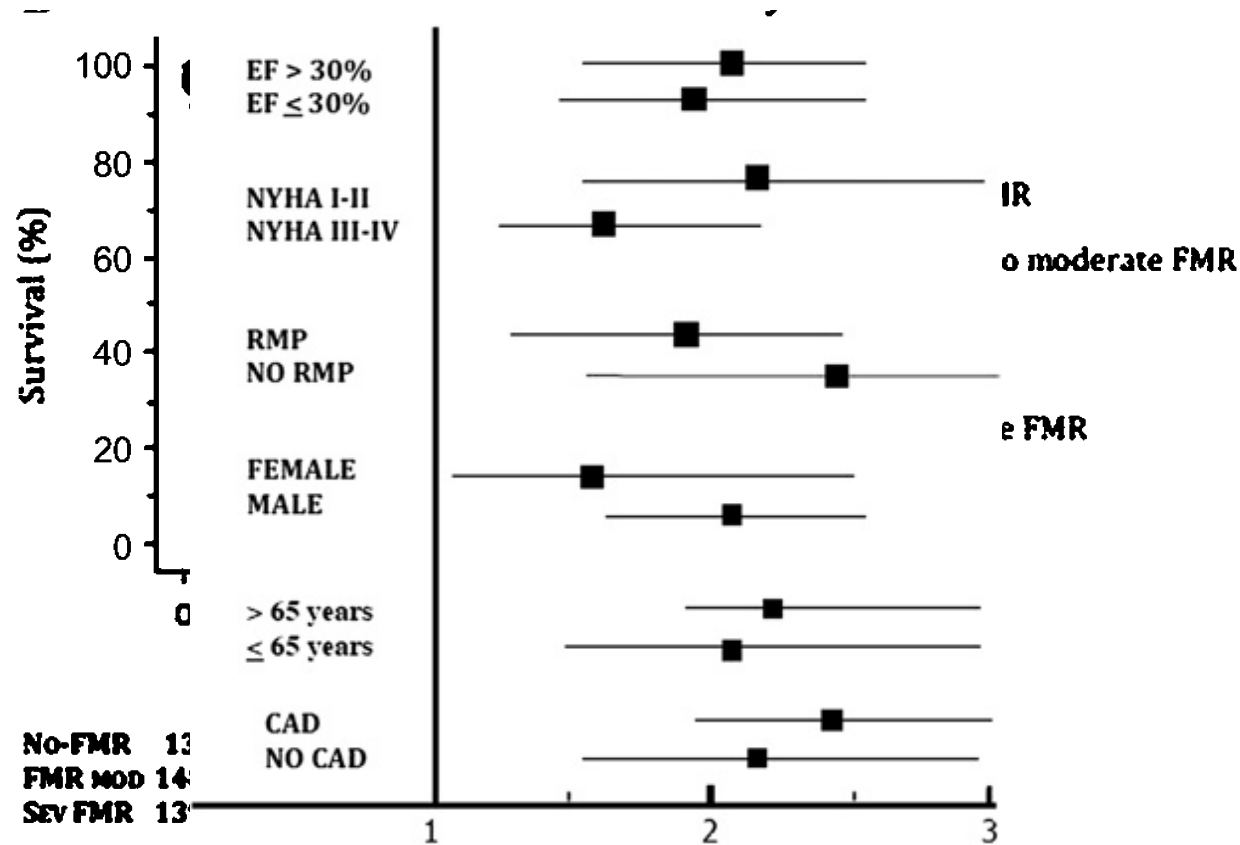
THE COCHRANE COLLABORATION®

Based on current evidence, patients with heart failure with poor cardiac function or idiopathic dilated cardiomyopathy, atrial fibrillation and a protruding mobile left ventricular thrombus on cardiac imaging are probably at highest risk and require anticoagulant therapy. If patients are in sinus rhythm, anticoagulants should perhaps be reserved especially for patients with severe cardiac impairment, the presence of intracardiac thrombus, and previous thromboembolism or stroke. Thus, until more evidence becomes available clinical decisions to treat patients with heart failure with anticoagulants must be made on an individual basis, based upon individual benefits and risks.



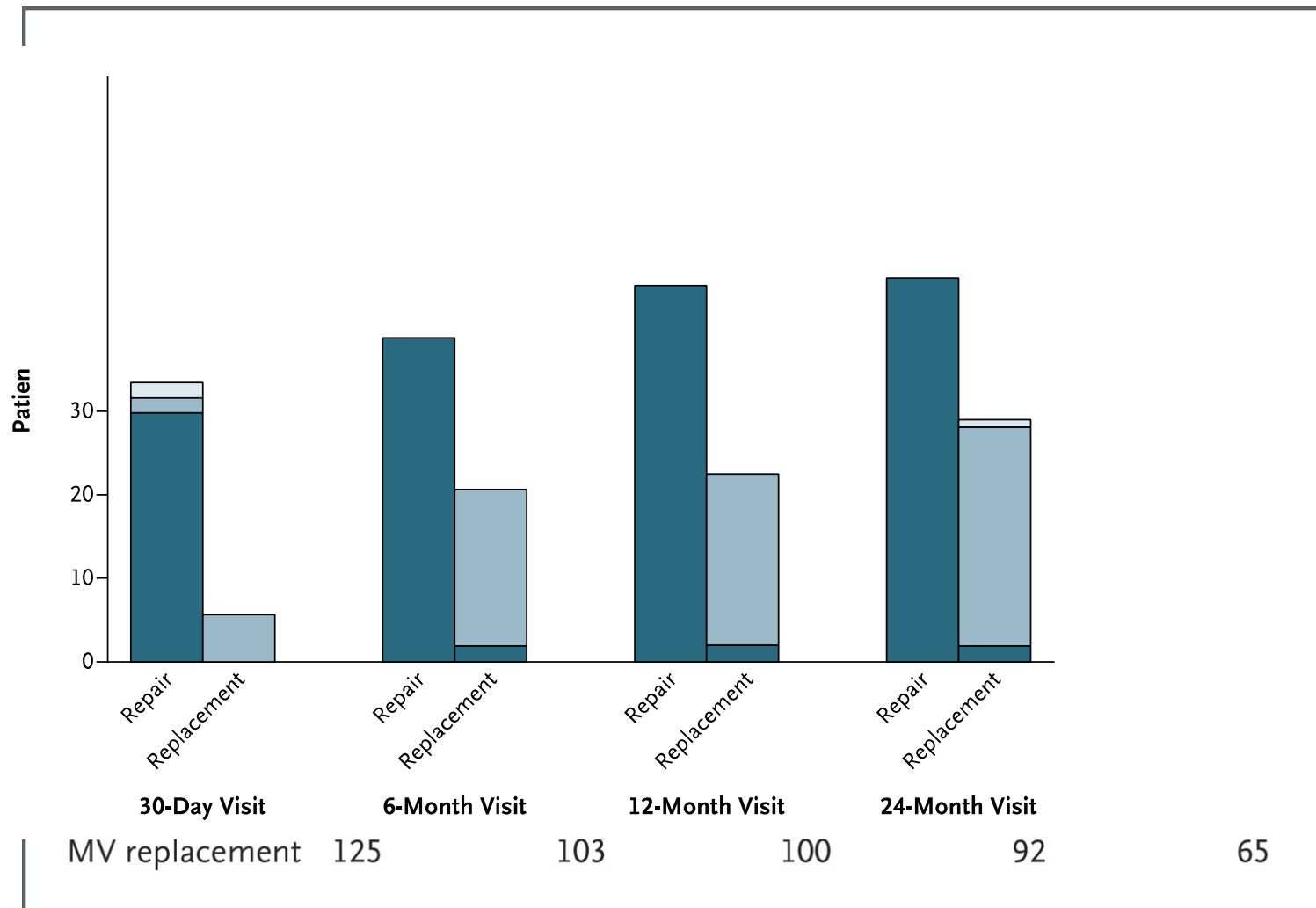
Quando e como abordar a insuficiência mitral secundária?

Independent Prognostic Value of Functional Mitral Regurgitation in Patients with Heart Failure



Two-Year Outcomes of Surgical Treatment of Severe Ischemic Mitral Regurgitation

CTSN



Percutaneous Repair or Surgery for Mitral Regurgitation EVEREST II Trial

Table 2. Primary Efficacy End Point at 12 Months and Major Adverse Events at 30 Days in the Intention-to-Treat Population.*

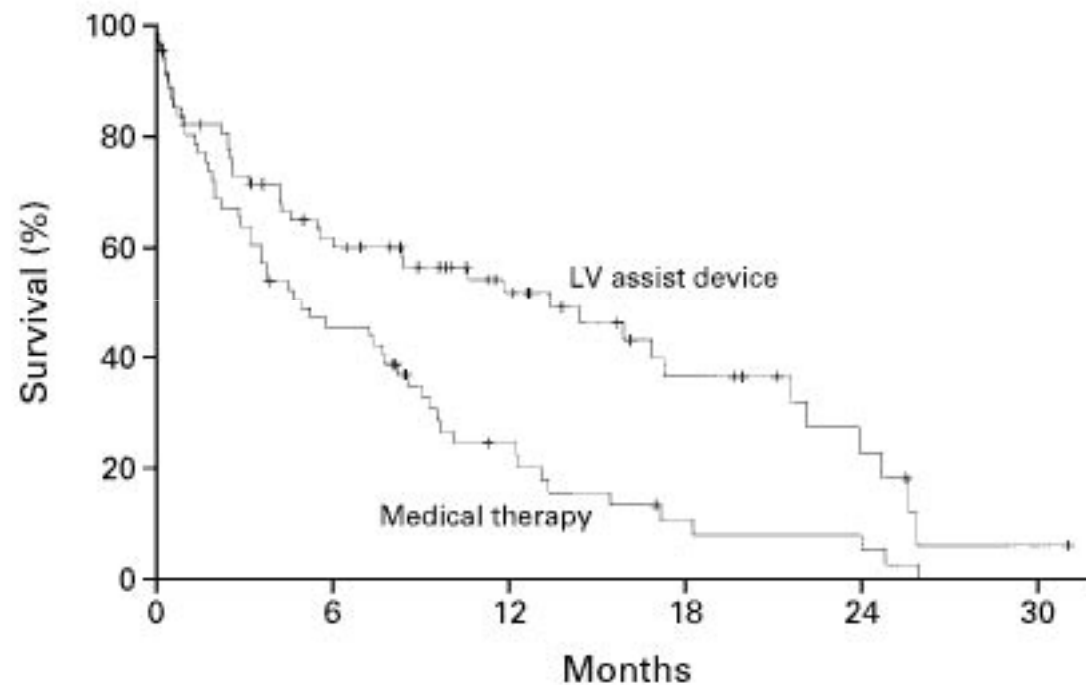
Event	Percutaneous Repair no. (%)	Surgery	P Value
Freedom from death, from surgery for mitral-valve dysfunction, and from grade 3+ or 4+ mitral regurgitation†	100 (55)	65 (73)	0.007
Surgery for mitral-valve dysfunction‡	37 (20)	2 (2)	<0.001
Major adverse event at 30 days§			
Any major adverse event	27 (15)	45 (48)	<0.001¶
Any major adverse event excluding transfusion	9 (5)	9 (10)	0.23
Death	2 (1)	2 (2)	0.89
Myocardial infarction	0	0	NA
Reoperation for failed surgical repair or replacement	0	1 (1)	0.74
Urgent or emergency cardiovascular surgery for adverse event	4 (2)	4 (4)	0.57
Major stroke	2 (1)	2 (2)	0.89
Renal failure	1 (<1)	0	1.00
Deep wound infection	0	0	NA
Mechanical ventilation for >48 hr	0	4 (4)	0.02
Gastrointestinal complication requiring surgery	2 (1)	0	0.78
New onset of permanent atrial fibrillation	2 (1)	0	0.78
Septicemia	0	0	NA
Transfusion of ≥2 units of blood	24 (13)	42 (45)	<0.001

Suporte ventricular mecânico – terapia de destino?

Quando e como encaminhar para o Tx?

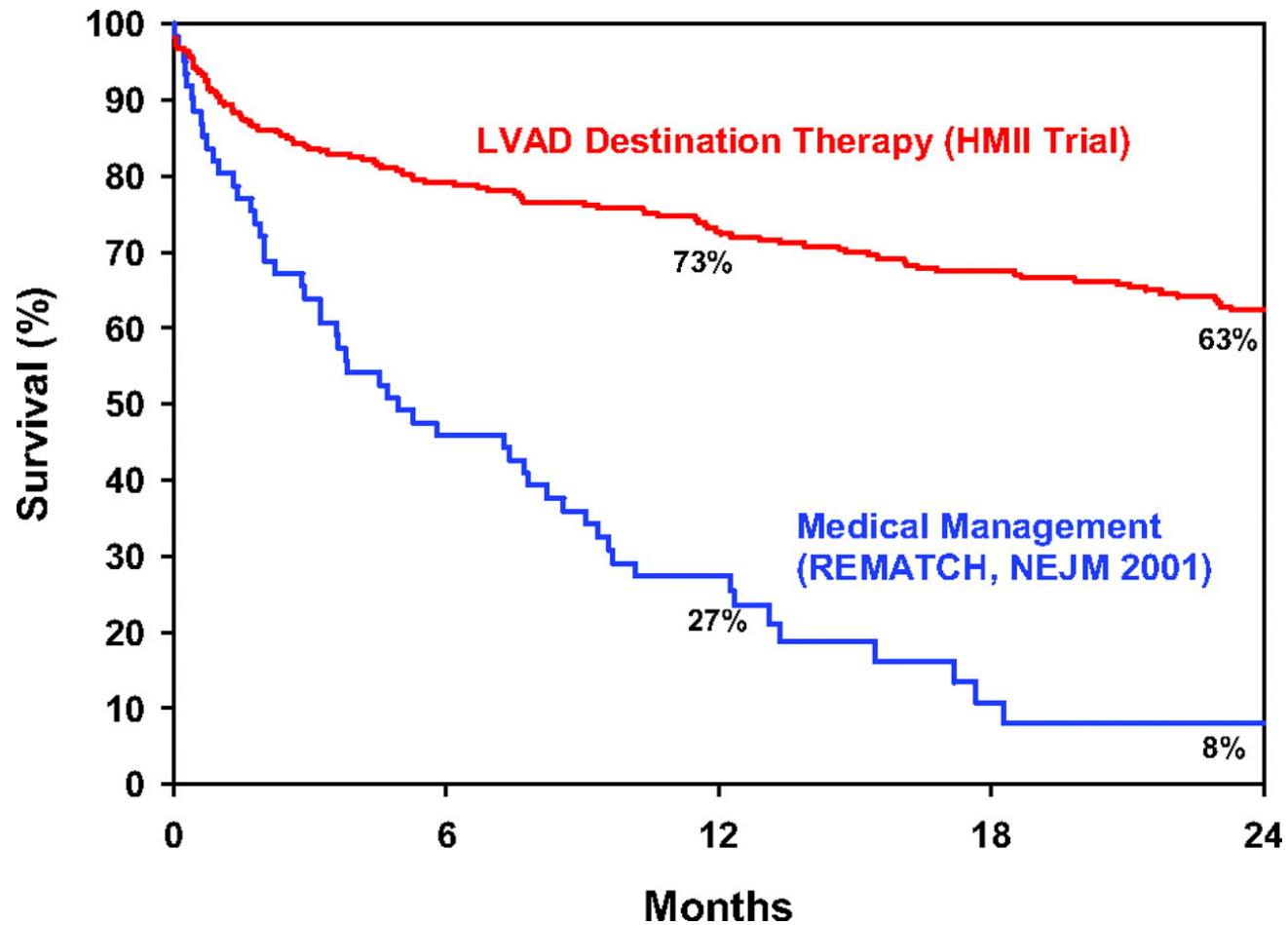
Long Term of A Left Ventricular Assist Device for End-Stage Heart failure

REMATCH Study Group



NO. AT RISK							
LV assist device	68	38	22	11	5	1	
Medical therapy	61	27	11	4	3	0	

Outcomes in Advanced Heart Failure Patients with Ventricular Assist Devices for destination Therapy

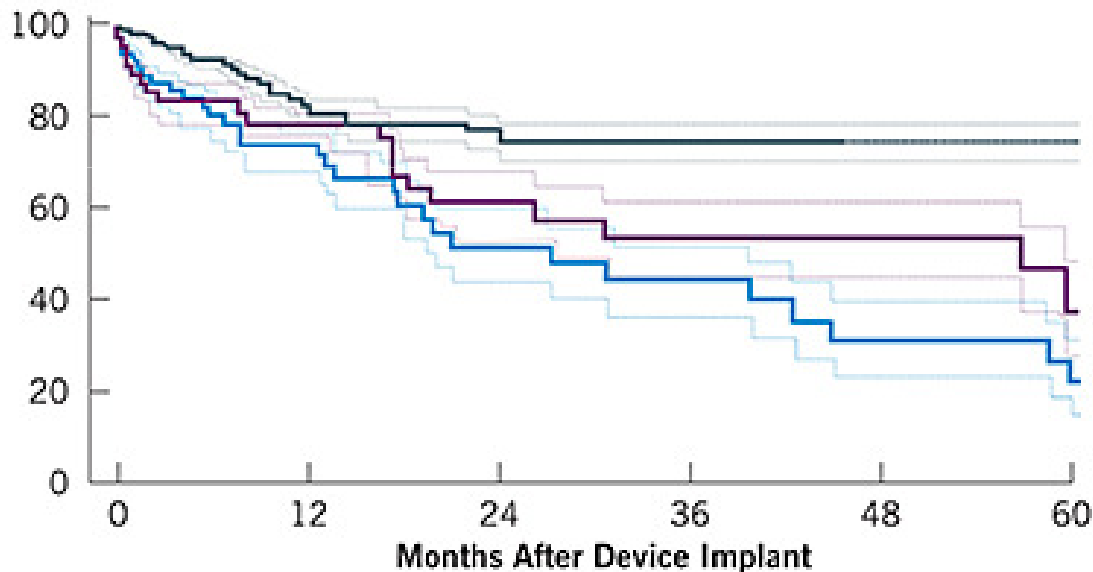


Survival Rate – Left Ventricular Assist Device Cleveland Clinic



Left Ventricular Assist Device Implantation, Survival Rate

Survival (%)



— Prior to 2010 (N = 71, deaths = 32)
— 2010 - 2011 (N = 54, deaths = 21)
— 2012 - 2015 (N = 164, deaths = 29)
 Log-rank $P = 0.002$

Percent Survival (Number at Risk)

	0	12	24	36	48	60
164	164	81	38	10		
54	54	27	18	13	11	3
71	71	30	15	11	7	5

Diretriz Brasileira Transplante Cardíaco

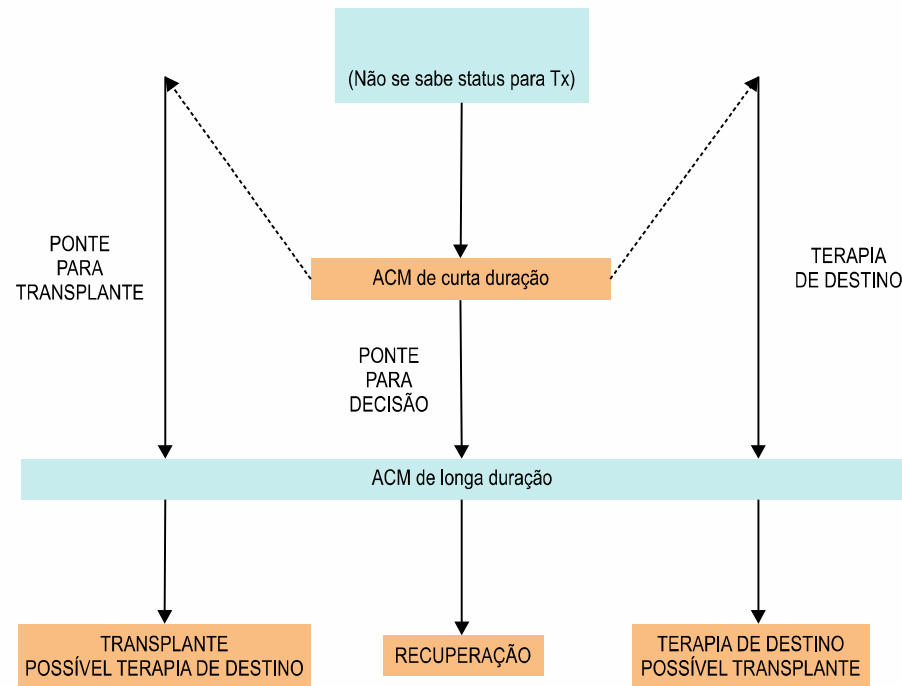
Indicações



Classe de Recomendação	Indicações	Nível de Evidência
Classe I	IC refratária na dependência de drogas inotrópicas e/ou de suporte circulatório e/ou ventilação mecânica;	C
	VO ₂ pico ≤ 10 ml/Kg/min;	C
	Doença isquêmica com angina refratária sem possibilidade de revascularização;	C
	Arritmia ventricular refratária;	C
	Classe funcional III/IV persistente.	C
Classe IIa	Teste da caminhada dos 6 minutos < 300 metros;	C
	Uso de BB com VO ₂ pico ≤ 12 ml/Kg/min;	C
	Sem uso de BB com VO ₂ pico ≤ 14 ml/Kg/min;	C
	Teste cardiopulmonar com relação VE/VCO ₂ > 35 e VO ₂ pico ≤ 14 ml/Kg/min.	C
Classe III	Presença de disfunção sistólica isolada;	C
	Classe funcional III ou IV sem otimização terapêutica.	C

VO₂ - consumo de oxigênio; BB - betabloqueador.

Diretriz de Assistência Circulatoria Mecânica



Demonstrated benefits of guideline-recommended heart failure therapies

Guideline-recommended therapy	Relative risk reductions in pivotal randomized clinical trial (s) (%)	Number needed to treat for mortality benefit (standardized to 12 m)	Relative risk reduction in meta-analysis
Angiotensin converting enzyme inhibitor OR angiotensin II receptor blocker	17	77	20%
Beta-blocker therapy (carvedilol, bisoprolol, extended release metoprolol succinate)	34	28	31%
Aldosterone antagonist	30	18	25%
Hydralazine plus nitrate	43	21	Not available
Cardiac resynchronization therapy	36	24	29/22%
Implantable cardioverter defibrillator	23	70	26%

