



Sademi
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V Encuentro de Enfermería de Medicina Interna de Andalucía

14, 15 y 16 de Junio 2018
PARANINFO DE LA UNIVERSIDAD DE GRANADA

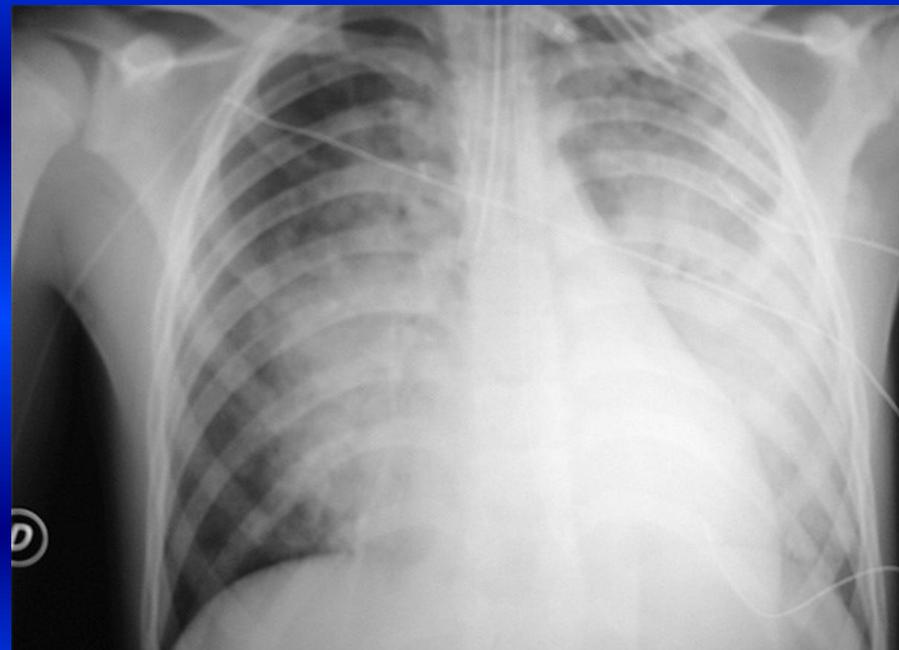
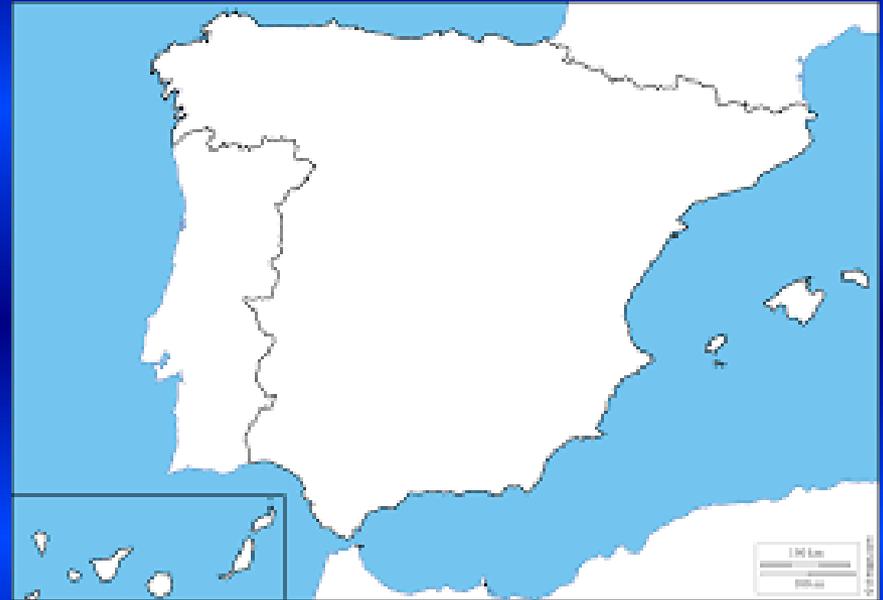
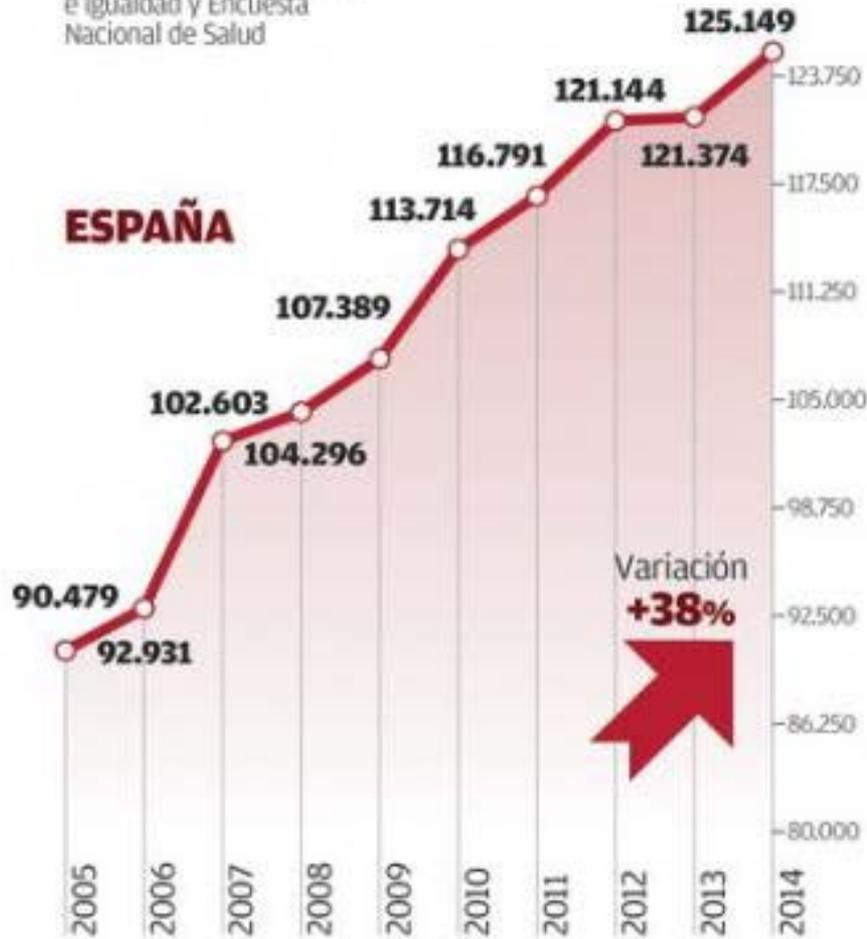
ACTUALIZACIÓN EN I.CARDIACA SACUBITRILO/VALSARTÁN

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Medicina Interna
Hospital Virgen del Camino
Sanlúcar de Bda-Cádiz



Altas por insuficiencia cardíaca

Fuente: Ministerio de Sanidad, Servicios Sociales e Igualdad y Encuesta Nacional de Salud



La hospitalización en el Sistema Nacional de Salud CMBD – Registro de altas

Informe resumen 2010
4.500.000 ALTAS

Gráfico 2: Distribución de altas por grandes grupos de edad y sexo. SNS. Año 2010.

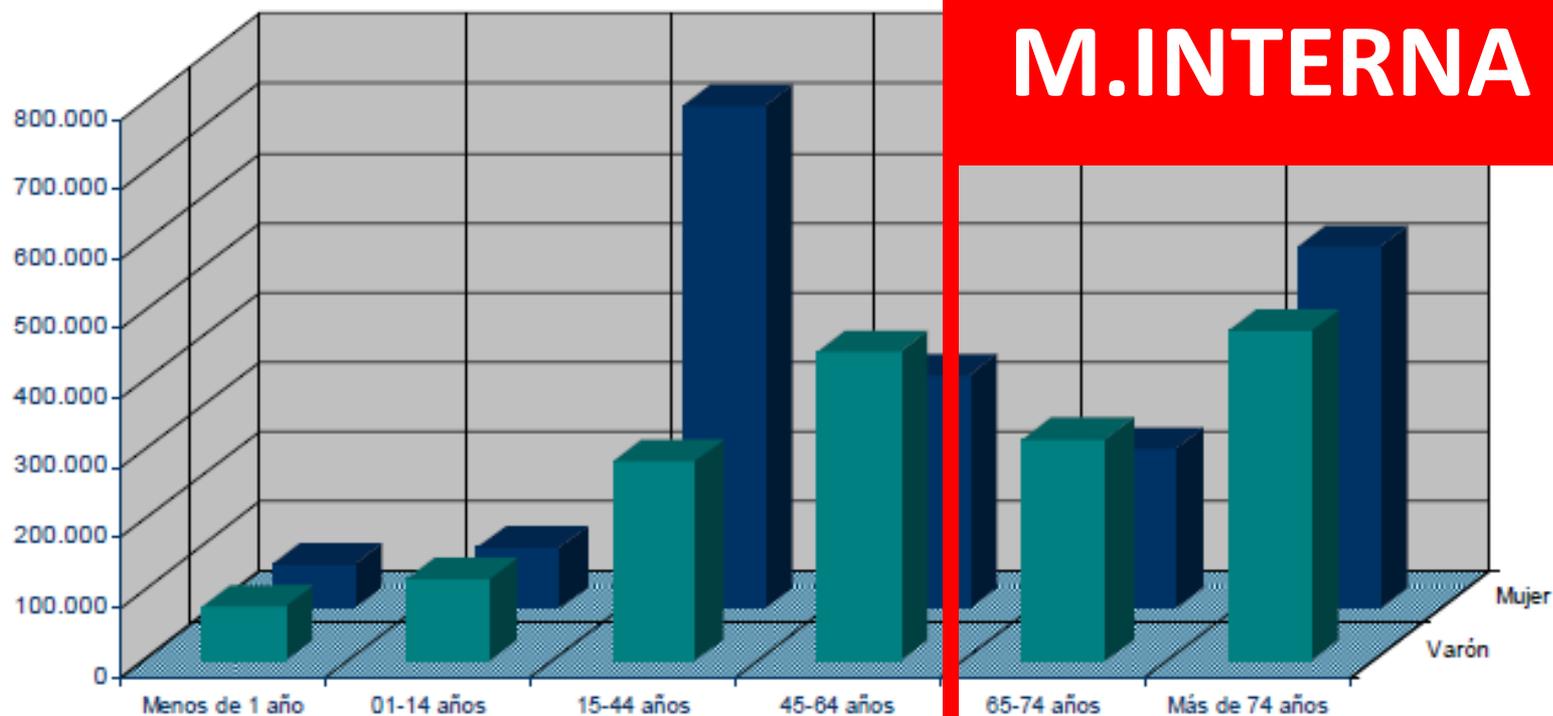
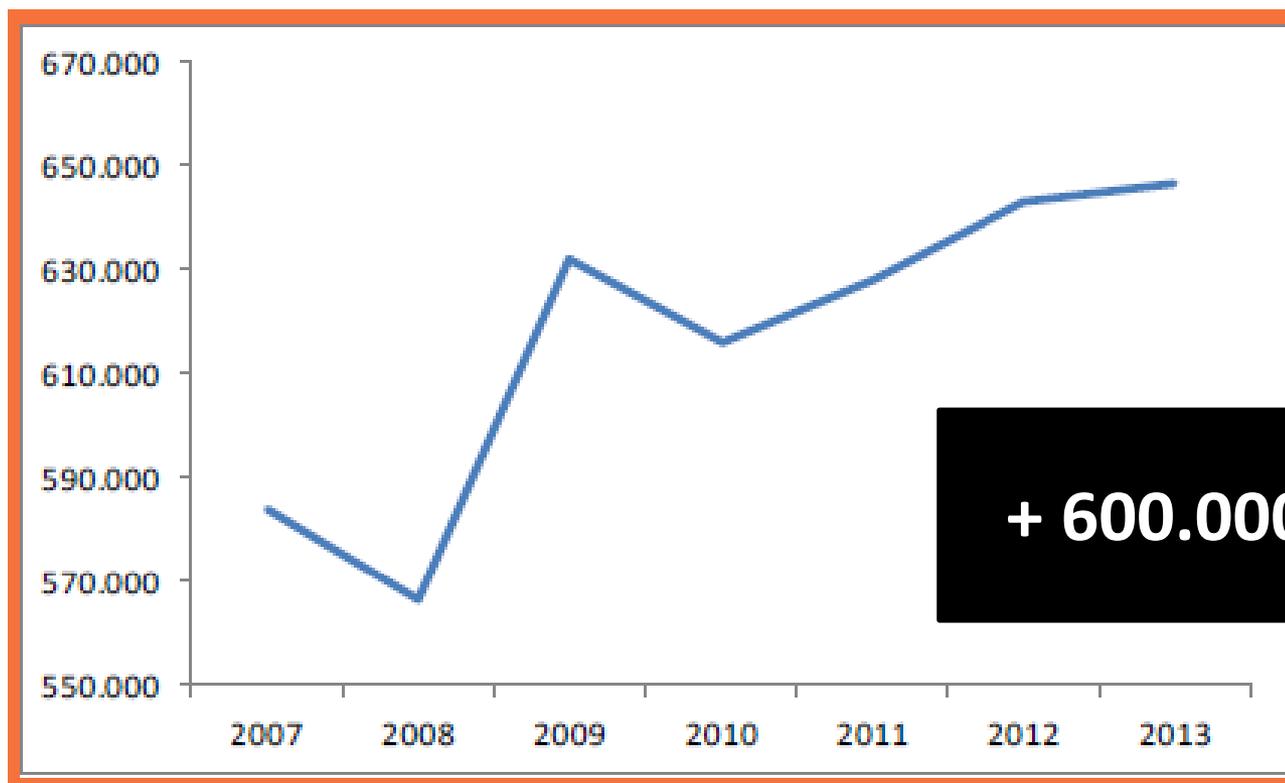


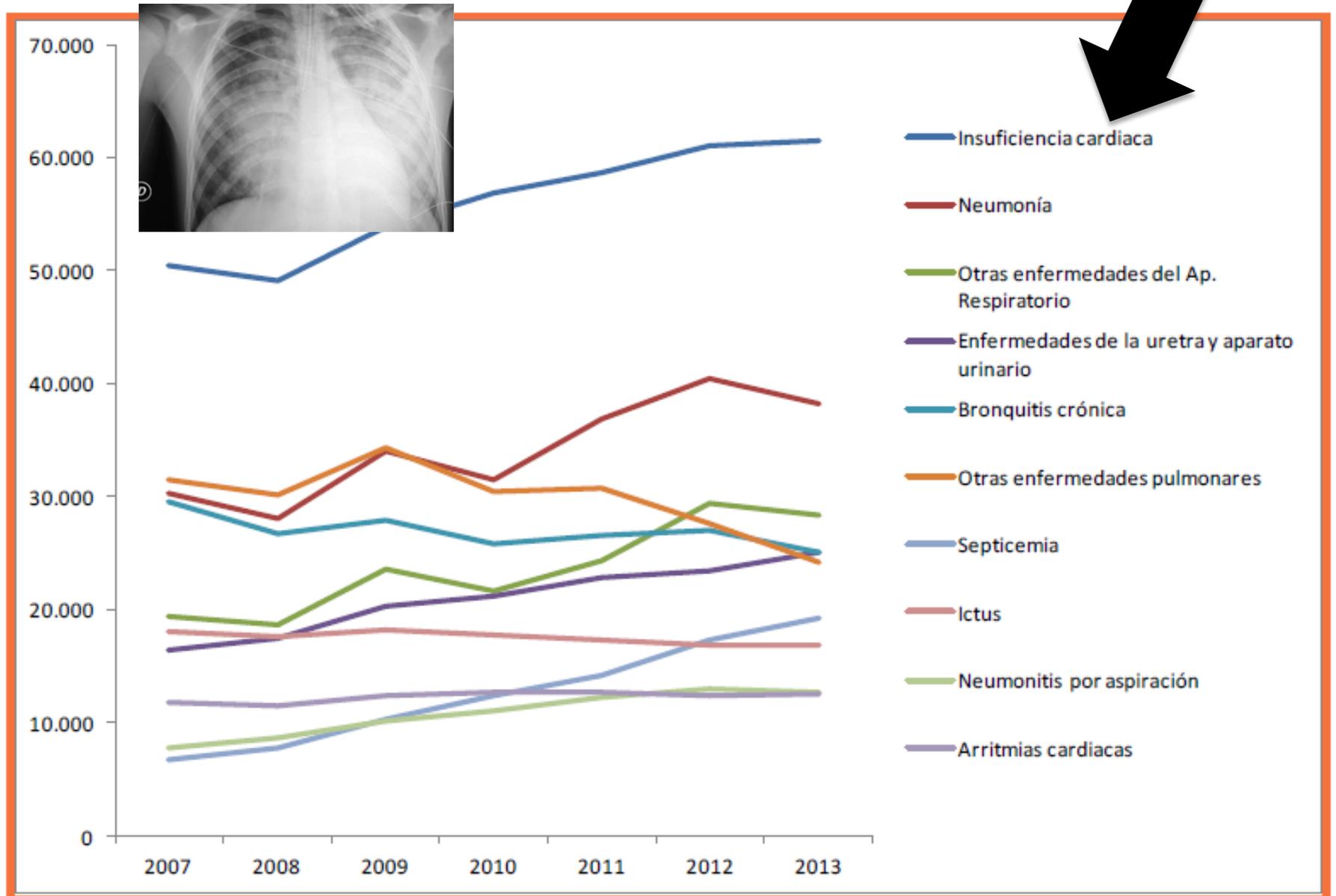
Figura 3.1. Evolución altas MI. 2007-2013

240 HOSPITALES



+ 600.000 ALTAS

Figura 3.2. Evolución altas para los 10 primeros diagnósticos. UMI. 2007-2013

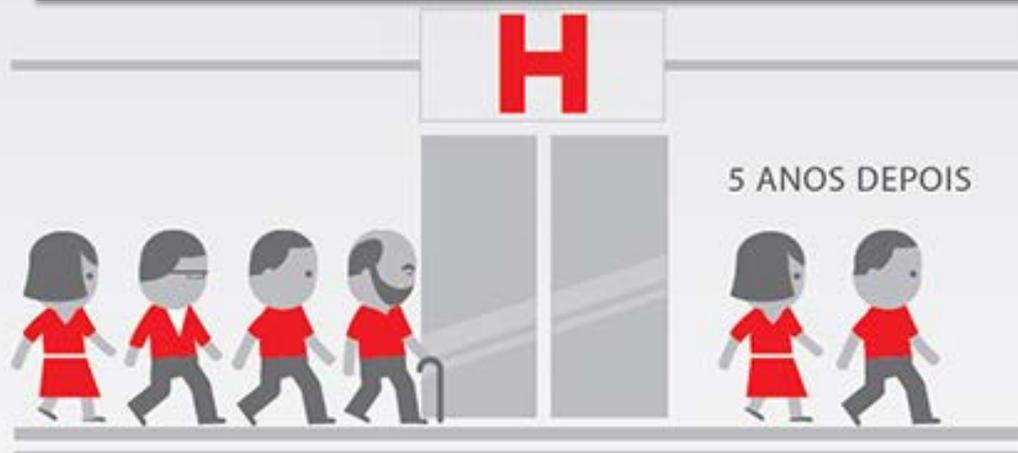


1 de cada 5 personas en el mundo sufrirá I.CARDIACA



**CAUSA + FREC.
HOSPITALIZACIÓN EN
MAYORES DE 65 AÑOS**

**DE LOS QUE INGRESAN, LA MITAD
MUEREN 5 AÑOS DESPUÉS**



#LOESTAMOSCAMBIANDO



Chronic decline

Cardiac function

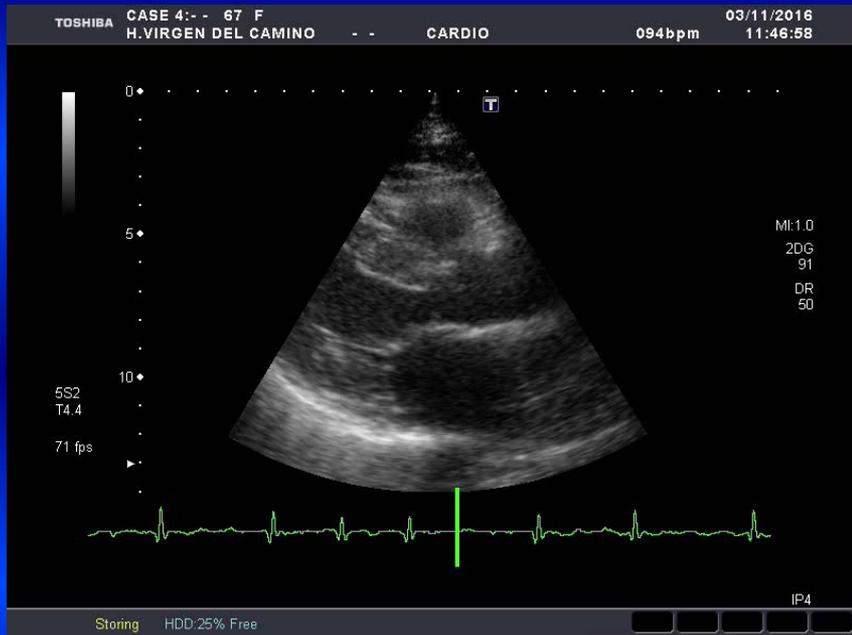


Hospitalizations for acute decompensations

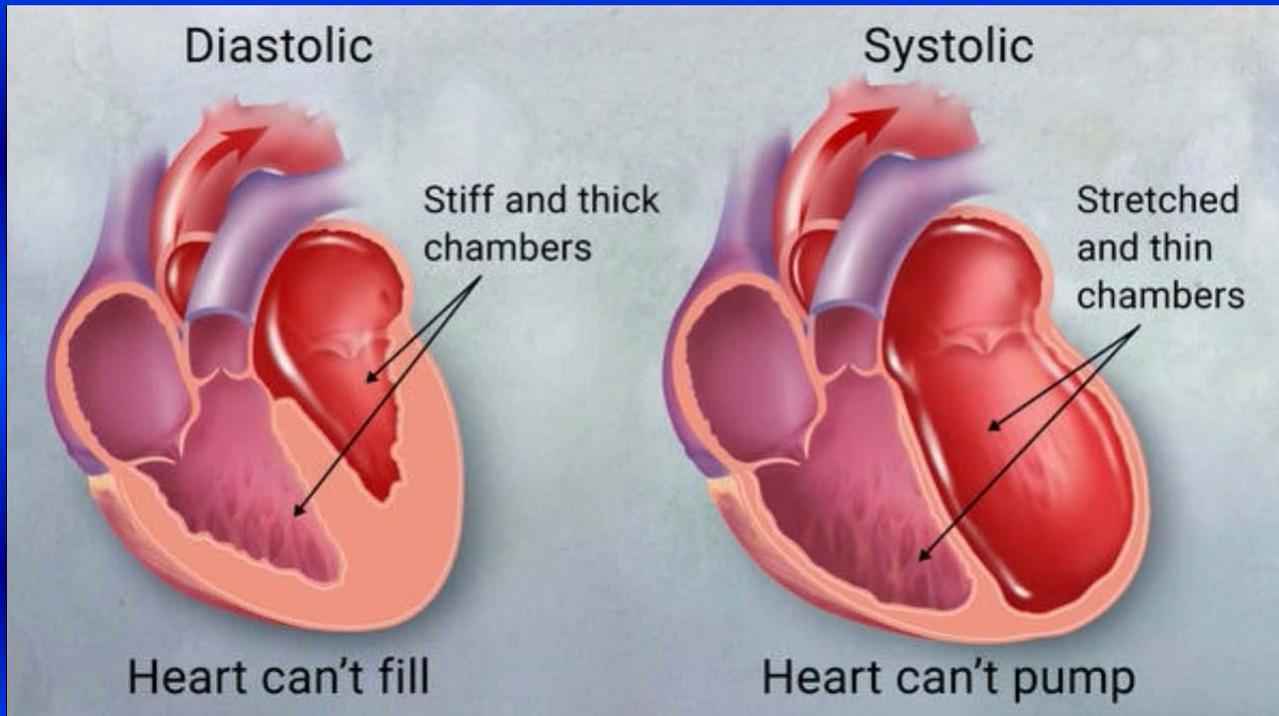
Disease progression

VULNERABILIDAD

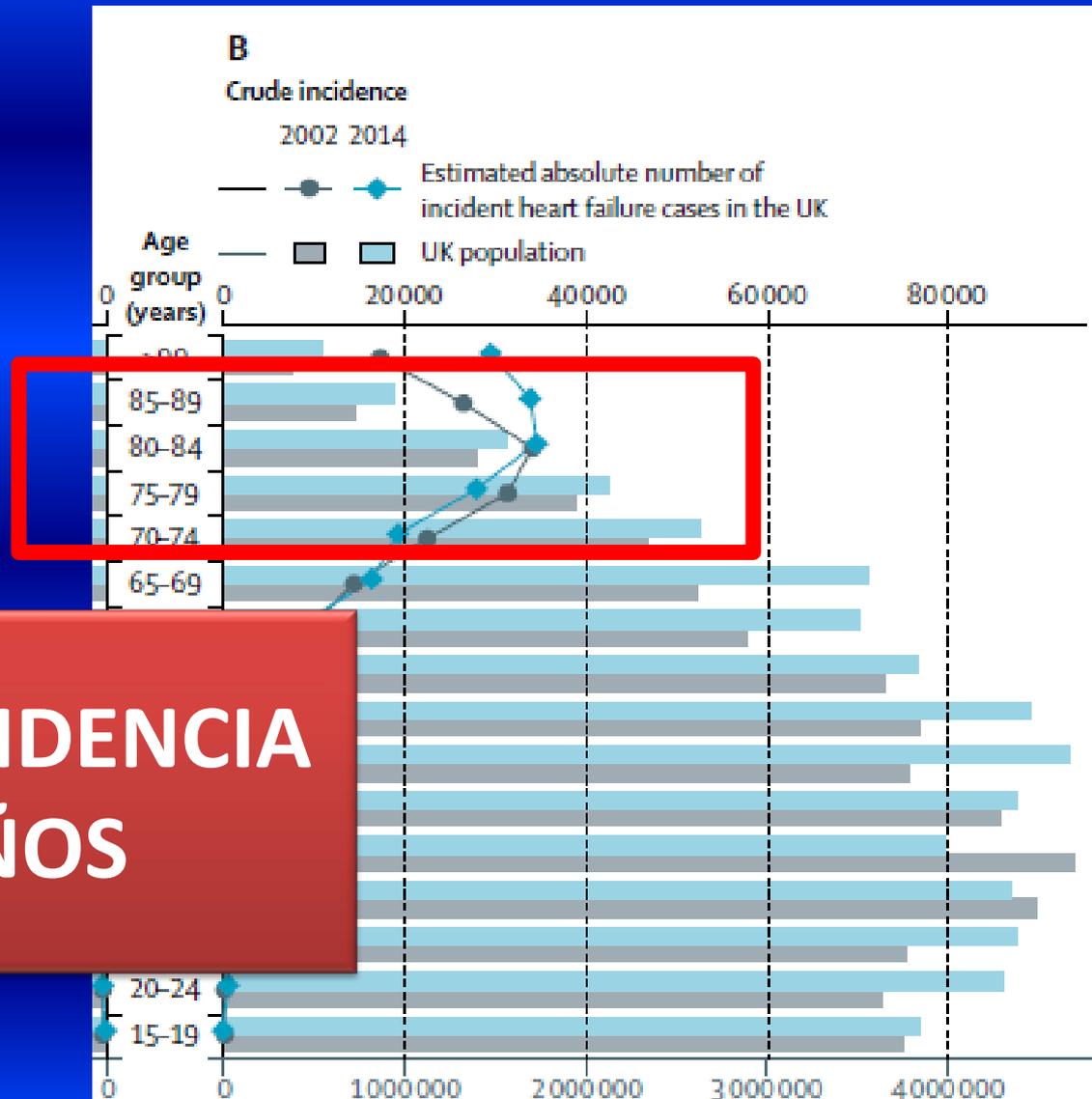




2/3



1/3

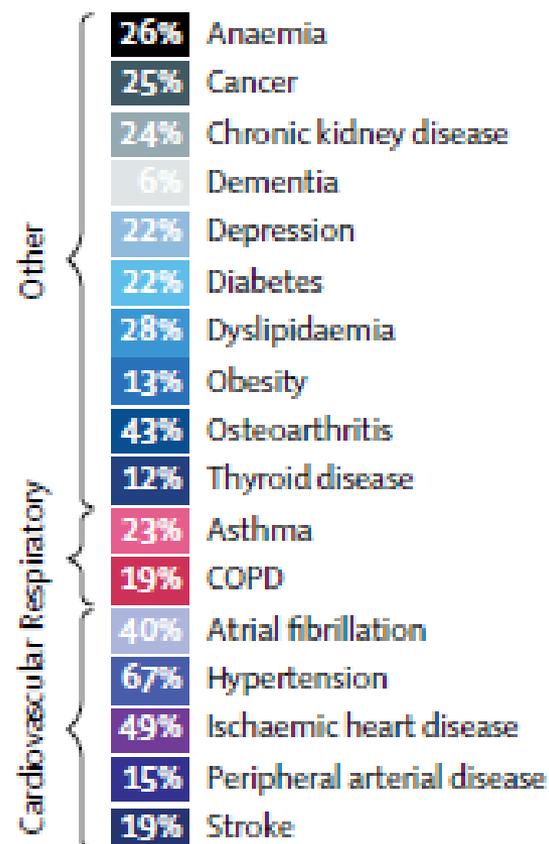
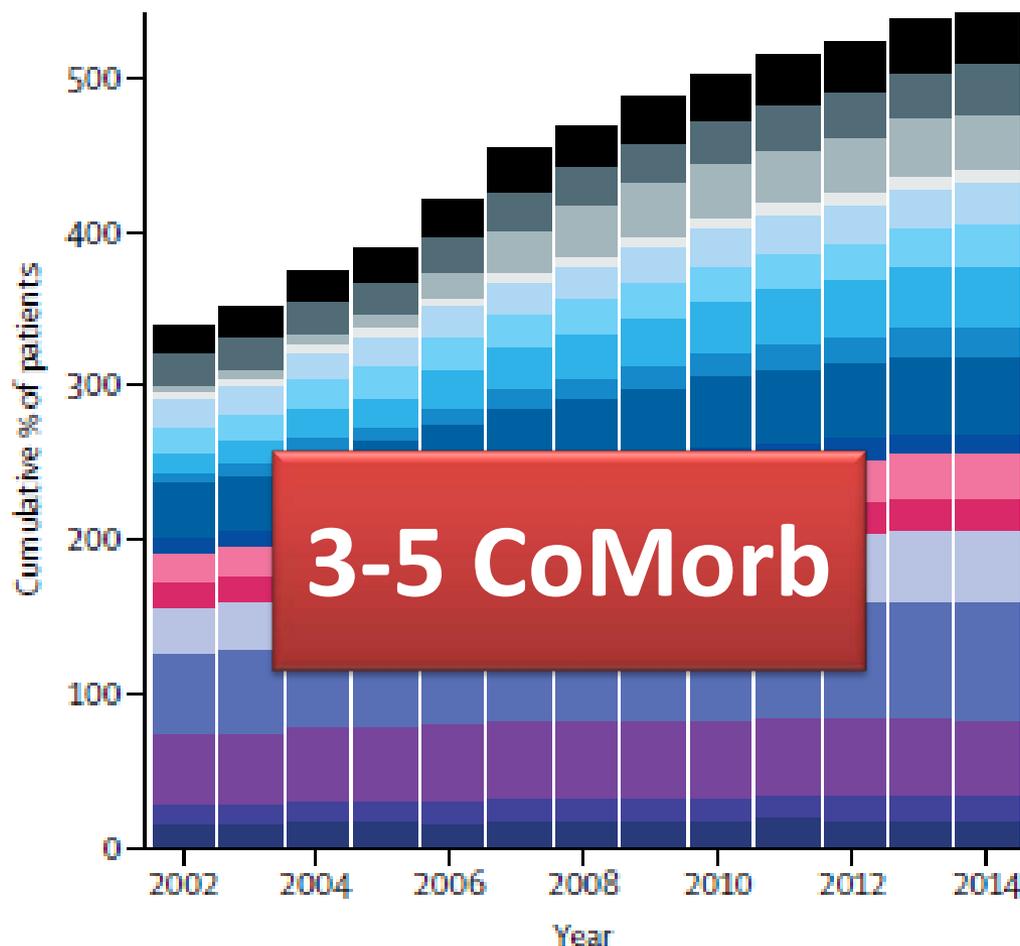


**MÁXIMA INCIDENCIA
70-85 AÑOS**

COMORBILIDADES



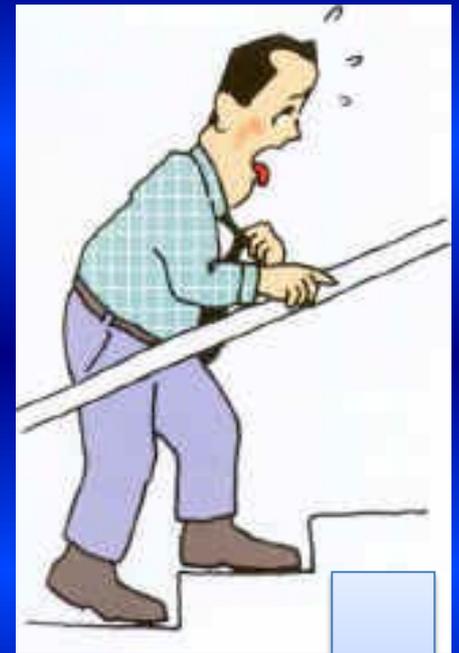
B Individual comorbidities

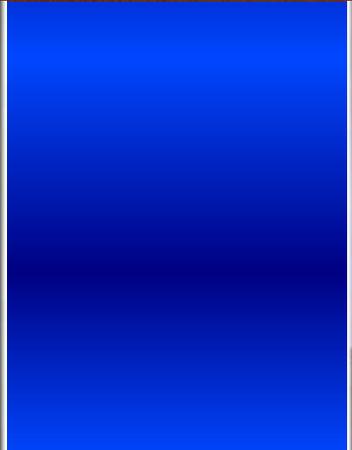
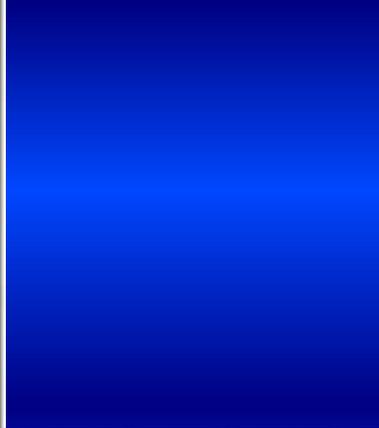




M.INTERNA

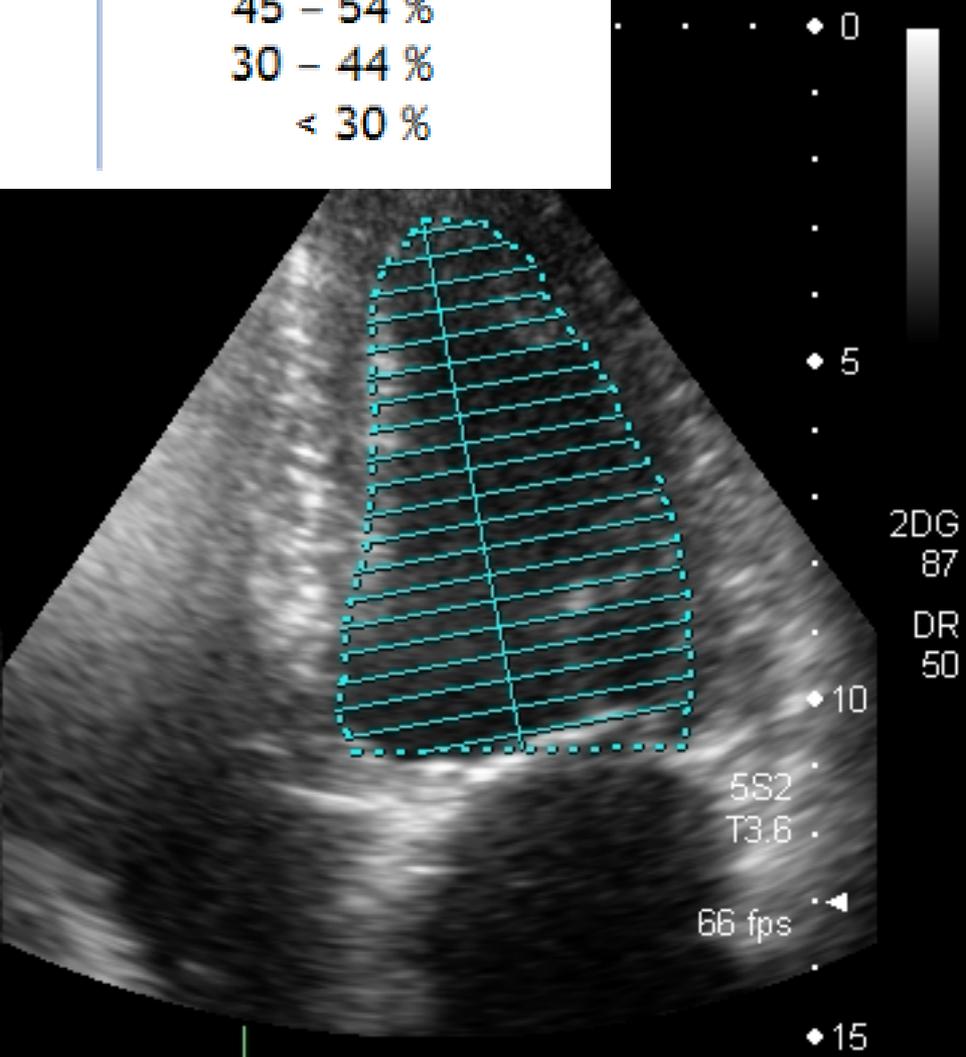
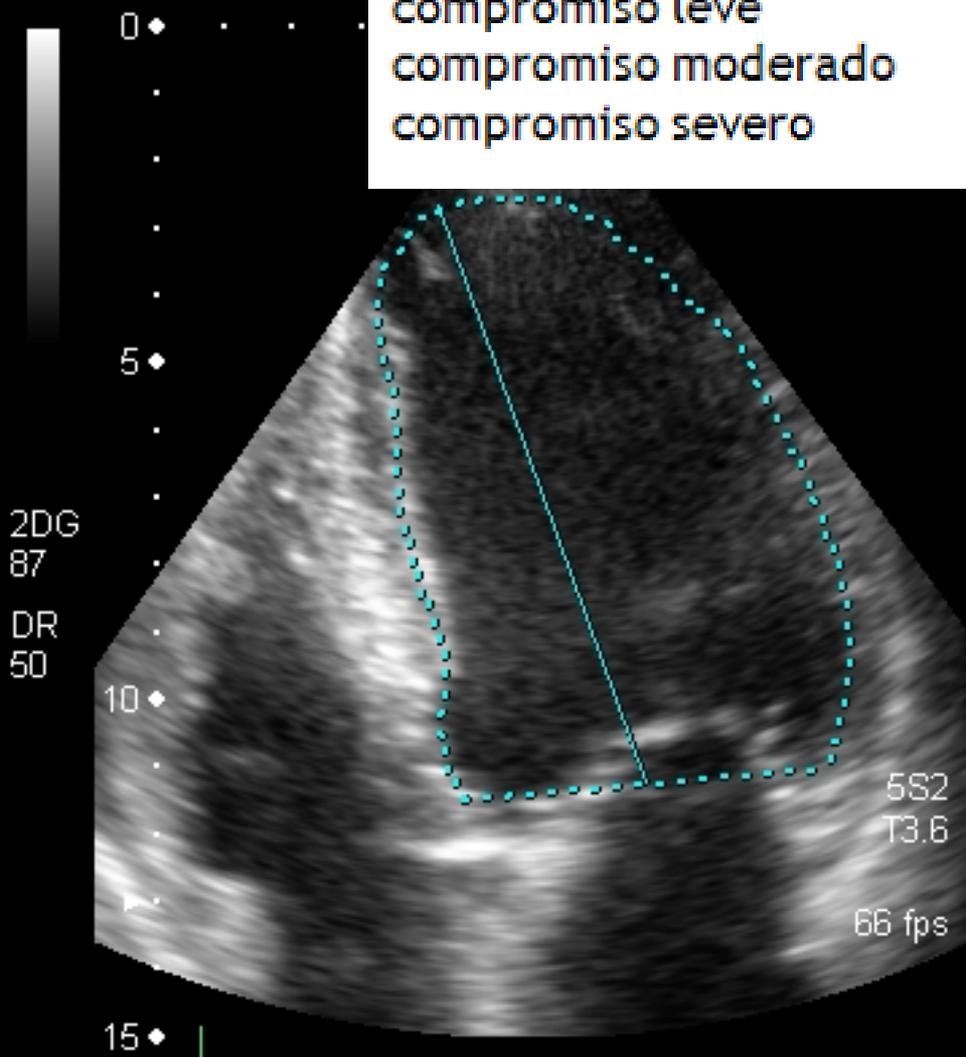
hay un intruso entre nosotros





Descriptiva
normal
compromiso leve
compromiso moderado
compromiso severo

FE in porcentaje*
≥ 55 %
45 - 54 %
30 - 44 %
< 30 %



LV MOD Simpson							
EDV4	193.0 ml	LVLd4	90.8 mm	ESV4	100.1 ml	LVLs4	79.7 mm
EF4	48.1 %	SV4	92.9 ml	CO4	4.924 l/min	HR	53 lpm

2016 ESC Guidelines for the diagnosis and treatment of acute and chronic heart failure

The Task Force for the diagnosis and treatment of acute and chronic heart failure of the European Society of Cardiology (ESC)

Table 3.1 Definition of heart failure with preserved (HFpEF), mid-range (HFmrEF) and reduced ejection fraction (HFrEF)

Type of HF	HFrEF	HFmrEF	HFpEF
CRITERIA	1	Symptoms ± Signs ^a	Symptoms ± Signs ^a
	2	LVEF <40%	LVEF 40–49%
	3	–	1. Elevated levels of natriuretic peptides ^b ; 2. At least one additional criterion: a. relevant structural heart disease (LVH and/or LAE), b. diastolic dysfunction (for details see Section 4.3.2).
			1. Symptoms ± Signs ^a 2. At least one additional criterion: a. relevant structural heart disease (LVH and/or LAE), b. diastolic dysfunction (for details see Section 4.3.2).

SIGNOS + SÍNTOMAS + FE < 40%

Pharmacological treatments indicated in patients with symptomatic (NYHA Class II-IV) heart failure with reduced ejection fraction

Recommendations	Class ^a	Level ^b	Ref ^c
An ACE-I ^d is recommended, in addition to a beta-blocker, for symptomatic patients with HFrEF to reduce the risk of HF hospitalization and death.	I	A	2, 163-165
A beta-blocker is recommended, in addition an ACE-I ^d , for patients with stable, symptomatic HFrEF to reduce the risk of HF hospitalization and death.	I	A	167-173
An MRA is recommended for patients with HFrEF, who remain symptomatic despite treatment with an ACE-I ^d and a beta-blocker, to reduce the risk of HF hospitalization and death.	I	A	174, 175

IECA

B-BLOQ

ARM



**HOSPITALIZACIÓN
MUERTE**



7.3 Other treatments recommended in selected symptomatic patients with heart failure with reduced ejection fraction

7.3.1 Diuretics

Diuretics are recommended to reduce the signs and symptoms of congestion in patients with HFrEF, but their effects on



- IECA
- BB
- ARM
- DIURÉTICOS

Patient with symptomatic^a HFrEF^b

■ Class I
■ Class IIa

Therapy with ACE-I^c and beta-blocker
(Up-titrate to maximum tolerated evidence-based doses)

Still symptomatic
and LVEF $\leq 35\%$

No

Yes

Add MR antagonist^{d,e}
(up-titrate to maximum tolerated evidence-based dose)

Still symptomatic
and LVEF $\leq 35\%$

No

Yes



ARNI to replace
ACE-I

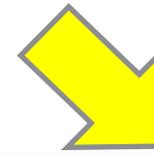
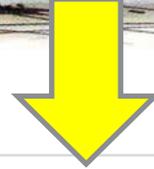
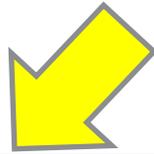
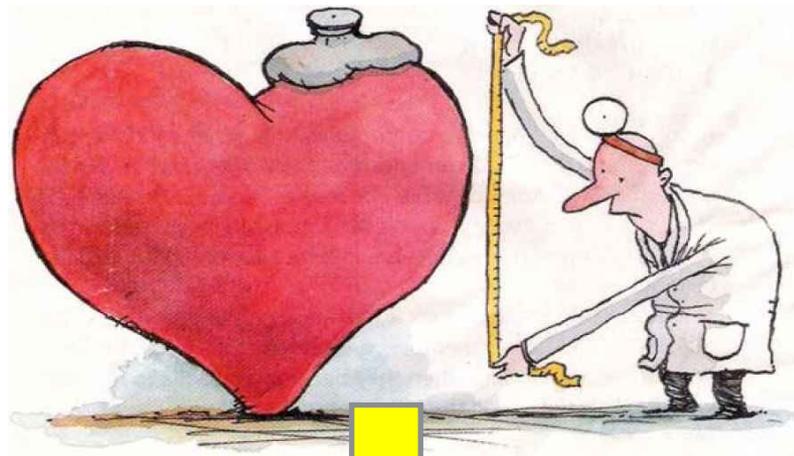


INHIBIDORES DE LA NEPRILISINA Y RECEPTORES DE ANGIOTENSINA

LCZ696

SACUBITRIL

VALSARTAN

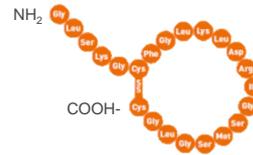


Péptido natriurético atrial (ANP)



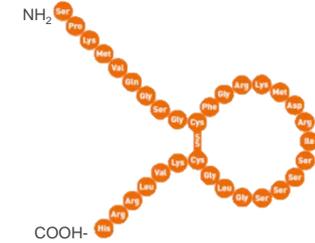
- Expresado en la **aurícula**
- Medible en plasma

Péptido natriurético tipo C (CNP)



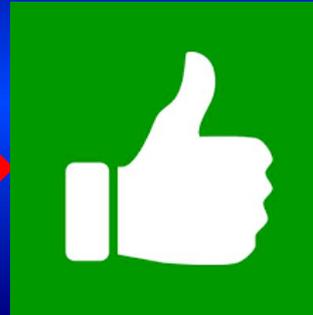
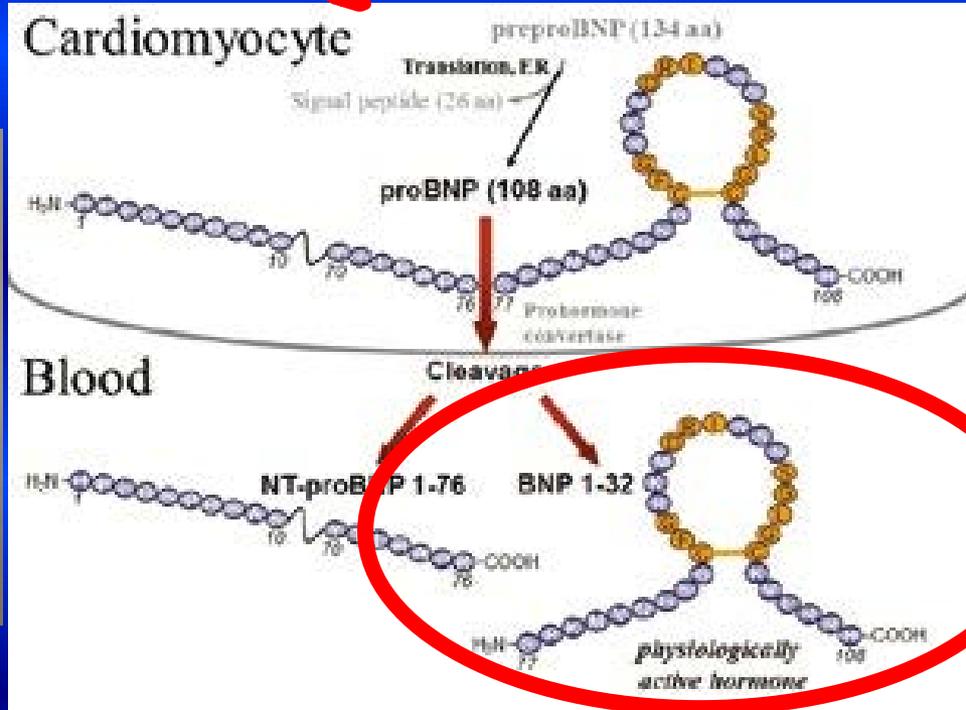
- Expresado en **células endoteliales vasculares** y **sistema nervioso central**
- No detectable en plasma – actúa localmente en tejidos

Péptido natriurético tipo B (BNP)



- Expresado en **aurícula** y **tejido ventricular**
- Medible en plasma

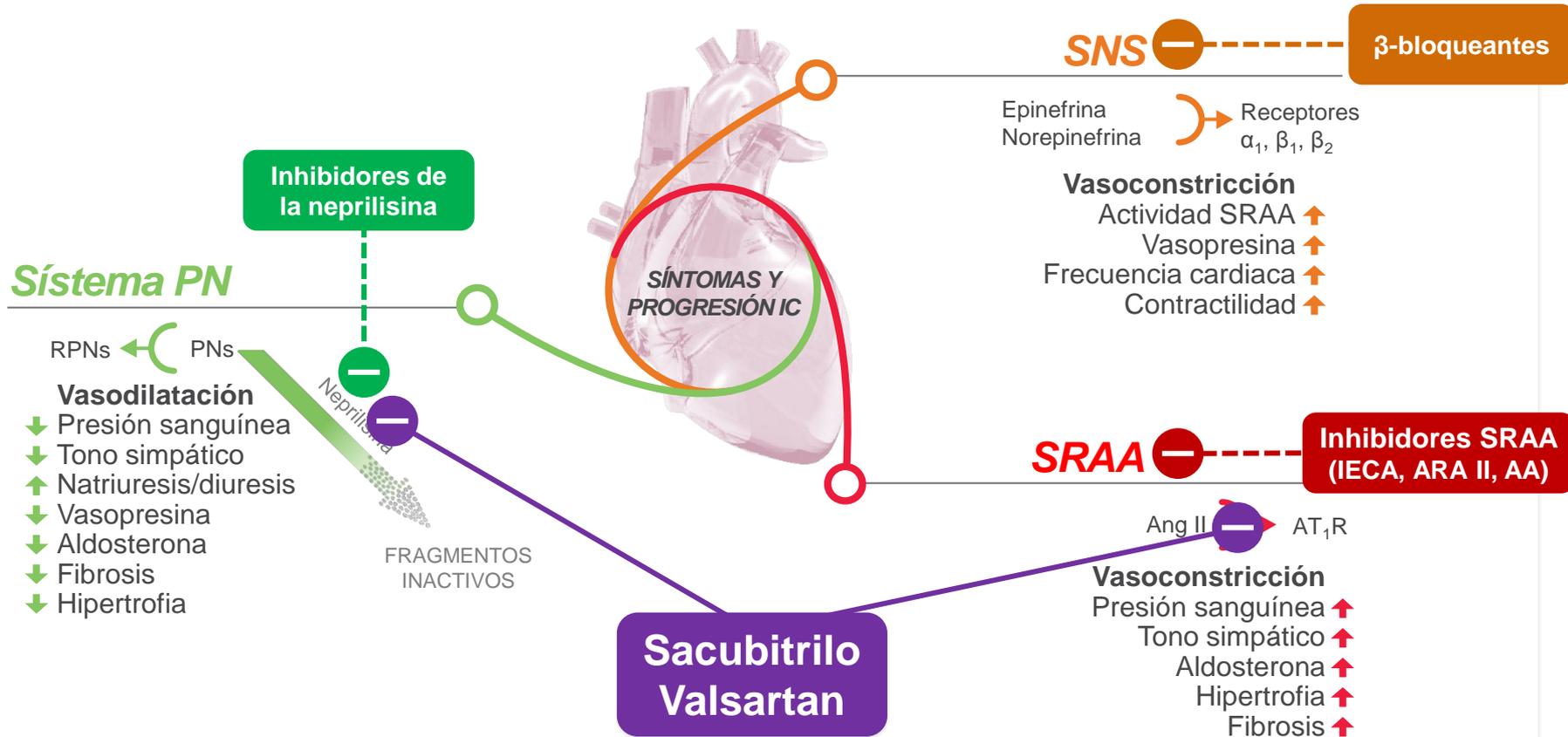
EFFECTOS BENEFICIOSOS



SACUBITRILO

NEPRILISINA

SACNAL nueva alternativa a los IECAs en pacientes con IC crónica con disfunción sistólica



- LCZ696: aumento de péptidos natriuréticos (inhibiendo Neprilisina) con supresión simultánea del SRAA

IECA

Omapatrilato

ECA



Aminopectidasa



Angioedema

Adrenomedulina

Sustancia P

Bradiquinina

Endotelina



NEP



DPP IV

Valsartan

Sacubitril

**SIN AUMENTO DE
ANGIOEDEMA**



The **NEW ENGLAND**
JOURNAL *of* **MEDICINE**

ESTABLISHED IN 1812

SEPTEMBER 11, 2014

VOL. 371 NO. 11

Angiotensin–Neprilysin Inhibition versus Enalapril in Heart Failure

8442 Pacientes
IC NYHA II-IV
FE DEPRIMIDA
FGe >30 ml/min/1,73m²

ENALAPRIL
10 mg/12h

LCZ696
200 mg/12h

MUERTE CV Y HOSPITALIZACIÓN POR I.CARDIACA



PARADIGM-HF: Resumen características basales pacientes

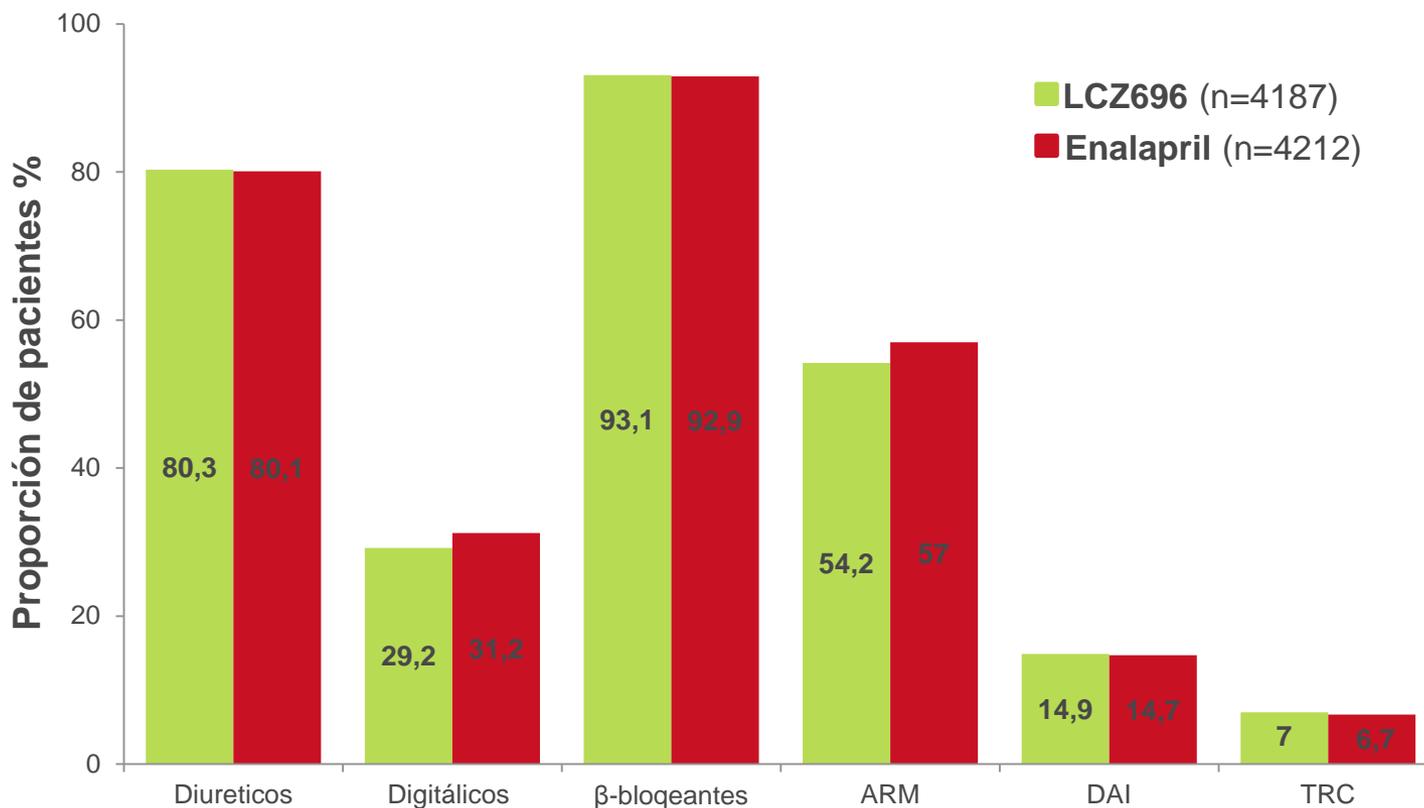
Características*	LCZ696 (n=4187)	Enalapril (n=4212)	
Edad, años	63.8 ± 11.5	63.8 ± 11.3	
Mujeres, n (%)	879 (21.0)	953 (22.6)	
Cardiopatía isquémica, n (%)	2506 (59.9)	2530 (60.1)	
Clase funcional NYHA, n (%)			
II	2998 (71.6)	2921 (69.3)	
III	969 (23.1)	1049 (24.9)	
Diabetes, n (%)	1451 (34.7)	1456 (34.6)	
FEVI, %	29.6 ± 6.1	29.4 ± 6.3	
PAS, mmHg	122 ± 15	121 ± 15	
FC, latidos/min	72 ± 12	73 ± 12	
NT pro-BNP, pg/mL (IQR)	1631 (885–3154)	1594 (886–3305)	
BNP, pg/mL (IQR)	255 (155–474)	251 (153–465)	

■ LCZ696
■ Enalapril

VARÓN
63 AÑOS
FE 29%
NYHA II
TAS:122 mmHg
Fc: 72 lpm
NT pro-BNP:1600 pg/ml

PARADIGM-HF: Resumen características basales pacientes

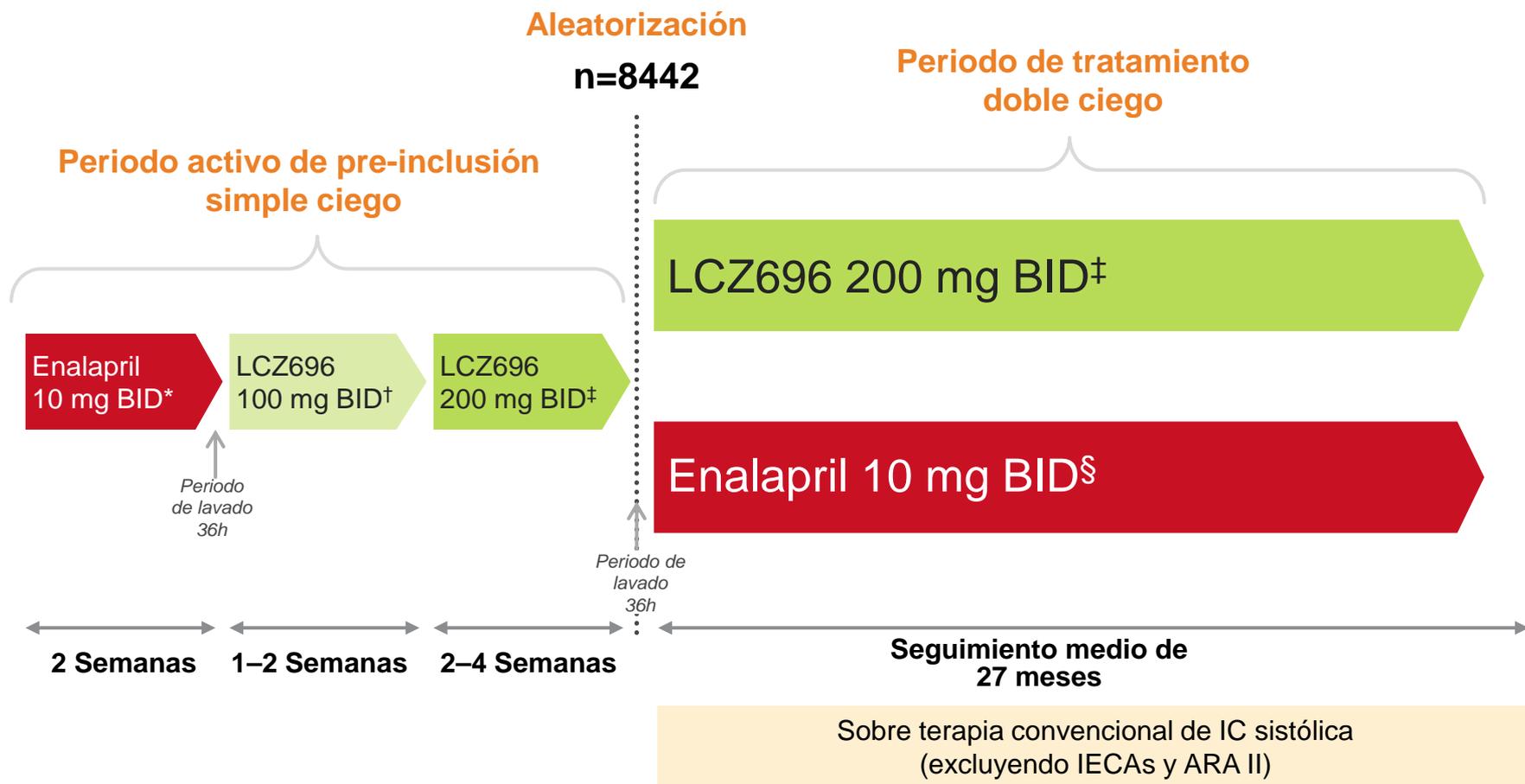
Tratamientos en el momento de la randomización



*media ± standard deviation, unless stated

Recreated from Novartis internal data, CLCZ696B2314
[Table 1](#) of McMurray, et al. *N Engl J Med* 2014

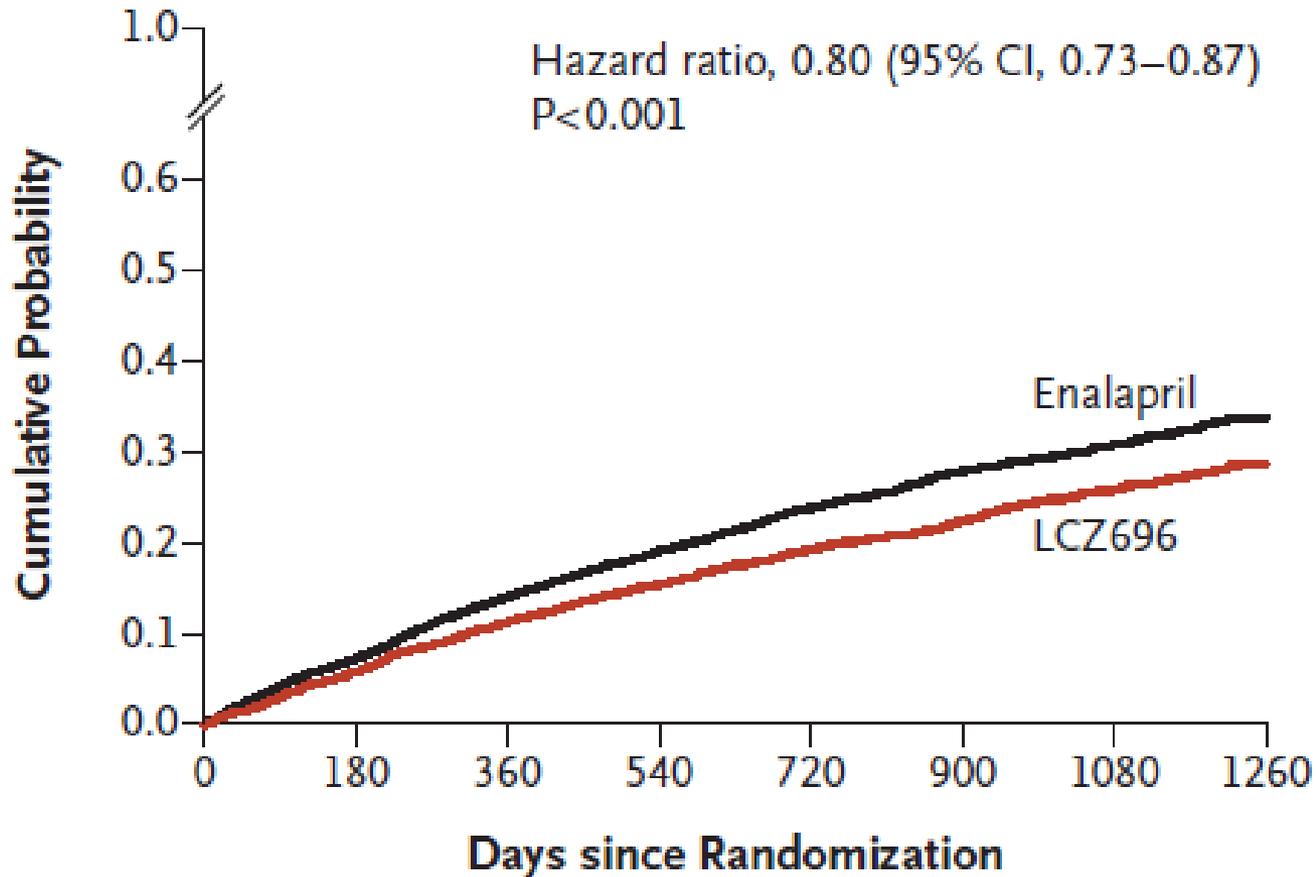
PARADIGM-HF: *Diseño del estudio*





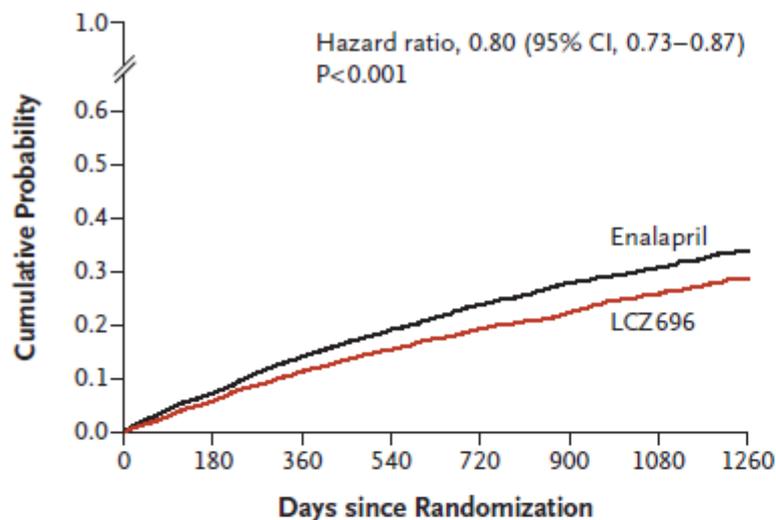
27 meses

A Primary End Point

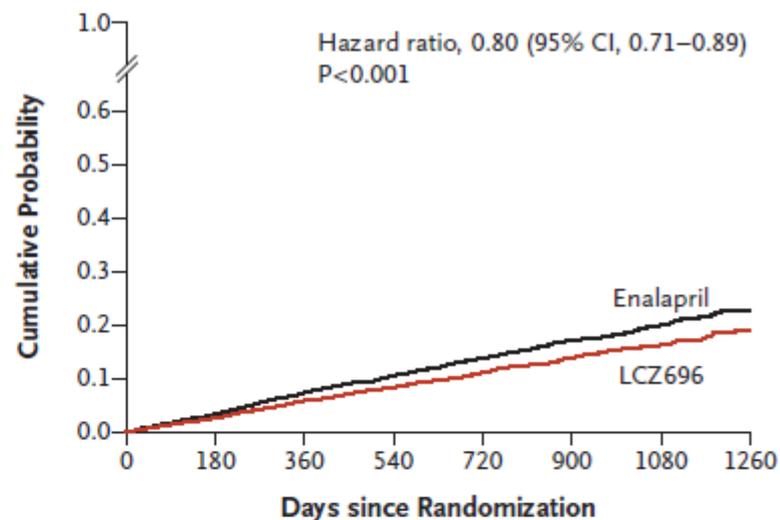


No. at Risk

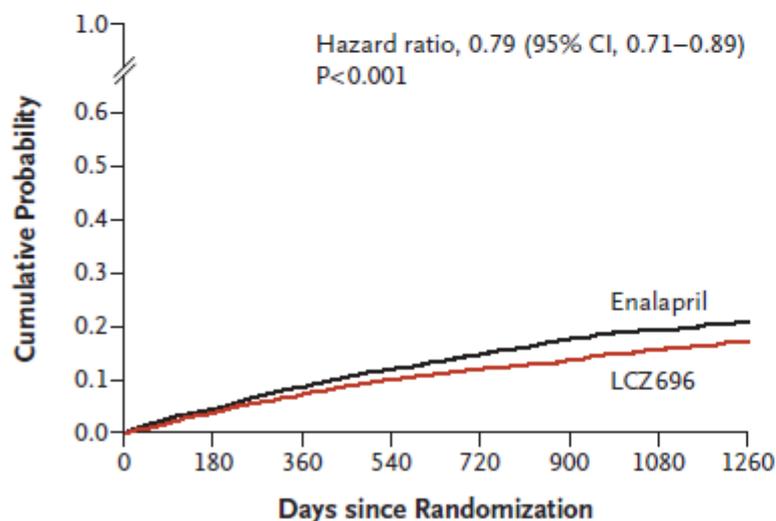
LCZ696	4187	3922	3663	3018	2257	1544	896	249
Enalapril	4212	3883	3579	2922	2123	1488	853	236

A Primary End Point**No. at Risk**

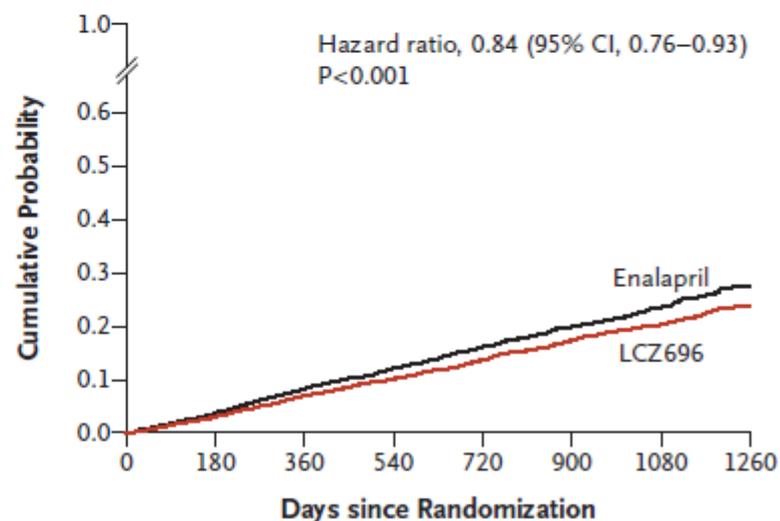
LCZ696	4187	3922	3663	3018	2257	1544	896	249
Enalapril	4212	3883	3579	2922	2123	1488	853	236

B Death from Cardiovascular Causes**No. at Risk**

LCZ696	4187	4056	3891	3282	2478	1716	1005	280
Enalapril	4212	4051	3860	3231	2410	1726	994	279

C Hospitalization for Heart Failure**No. at Risk**

LCZ696	4187	3922	3663	3018	2257	1544	896	249
Enalapril	4212	3883	3579	2922	2123	1488	853	236

D Death from Any Cause**No. at Risk**

LCZ696	4187	4056	3891	3282	2478	1716	1005	280
Enalapril	4212	4051	3860	3231	2410	1726	994	279

Table 3. Adverse Events during Randomized Treatment.*

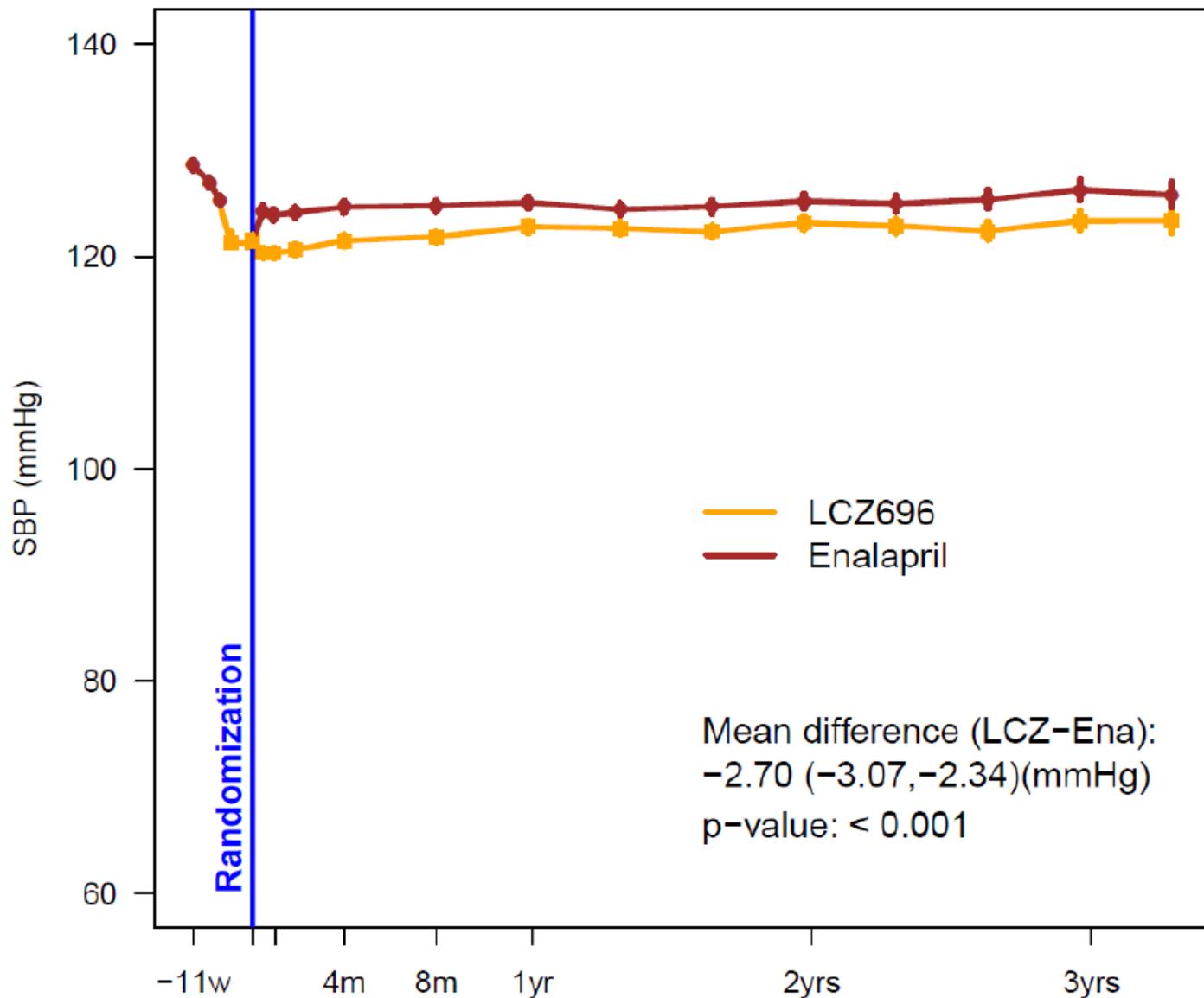
Event	LCZ696 (N=4187)	Enalapril (N=4212)	P Value
	<i>no. (%)</i>		
Hypotension			
Symptomatic	588 (14.0)	388 (9.2)	<0.001
Symptomatic with systolic blood pressure <90 mm Hg	112 (2.7)	59 (1.4)	<0.001
Elevated serum creatinine			
≥2.5 mg/dl	139 (3.3)	188 (4.5)	0.007
≥3.0 mg/dl	63 (1.5)	83 (2.0)	0.10
Elevated serum potassium			
>5.5 mmol/liter	674 (16.1)	727 (17.3)	0.15
>6.0 mmol/liter	181 (4.3)	236 (5.6)	0.007
Cough	474 (11.3)	601 (14.3)	<0.001
Angioedema†			
No treatment or use of antihistamines only	10 (0.2)	5 (0.1)	0.19
Use of catecholamines or glucocorticoids without hospitalization	6 (0.1)	4 (0.1)	0.52
Hospitalization without airway compromise	3 (0.1)	1 (<0.1)	0.31
Airway compromise	0	0	—

+ hipotensión

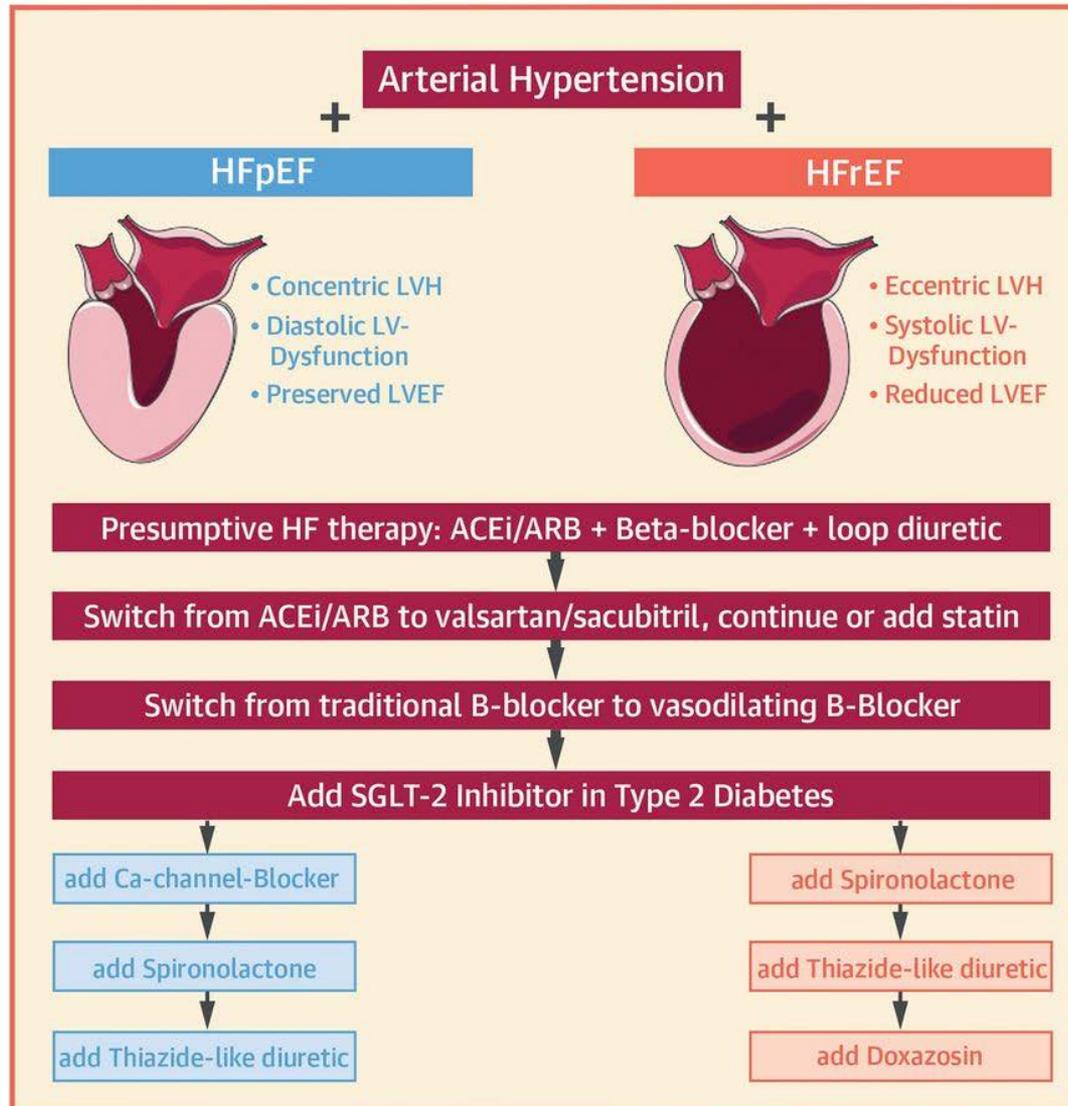
- Deterioro renal

- Hiper K

Sin aumentar angioedema



CENTRAL ILLUSTRATION: Suggested Empirical Antihypertensive Strategy in HF Patients With Persisting Hypertension



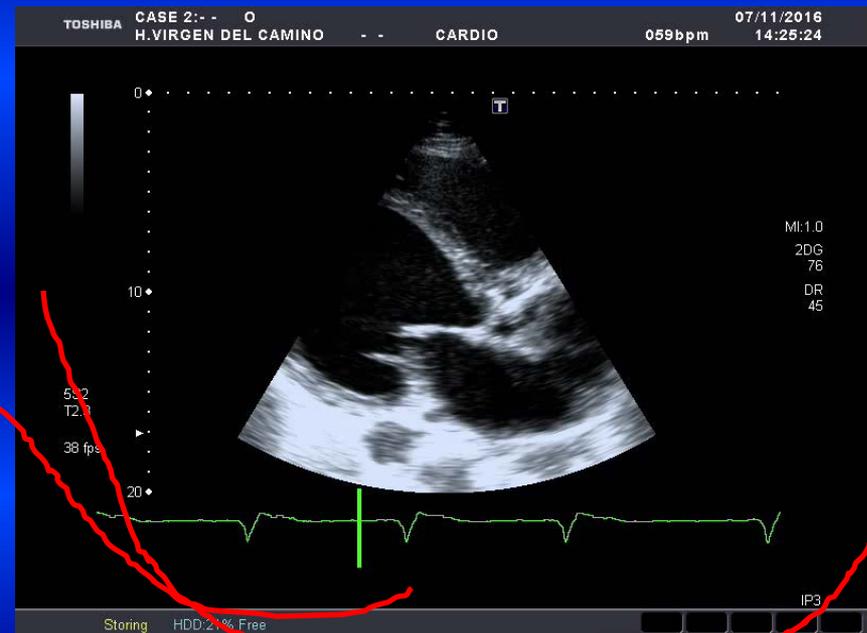
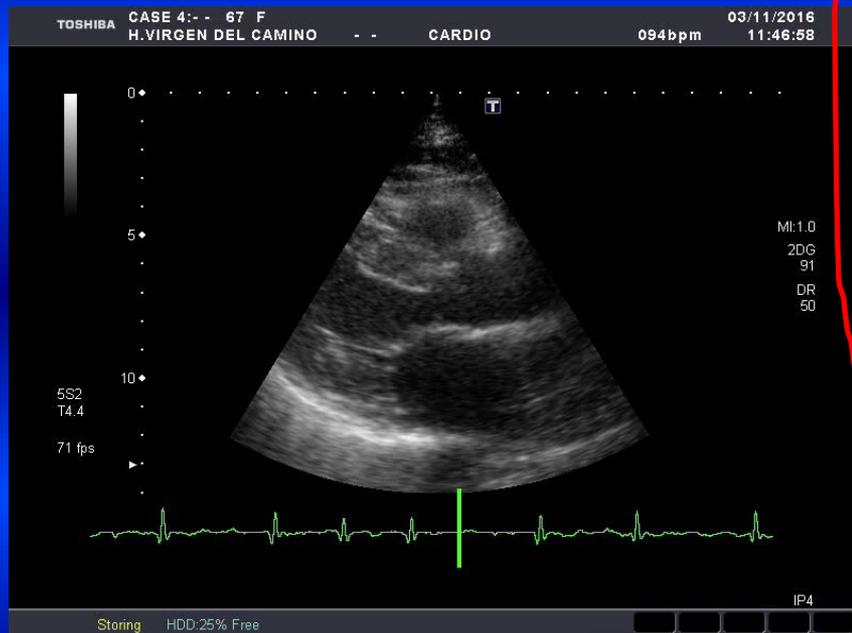
MANEJO PRÁCTICO

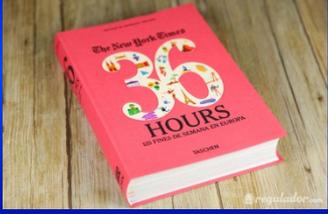


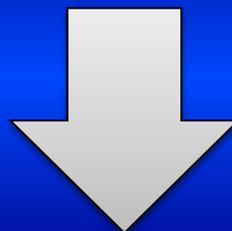
¿Cuál es el paciente?

FE PRESERVADA

FE DEPRIMIDA









1

2

3



European Journal of Heart Failure (2017)

Rationale for and design of the TRUE-AHF trial: the effects of ularitide on the short-term clinical course and long-term mortality of patients with acute heart failure

Serelaxin in addition to standard therapy in acute heart failure: rationale and design of the RELAX-AHF-2 study

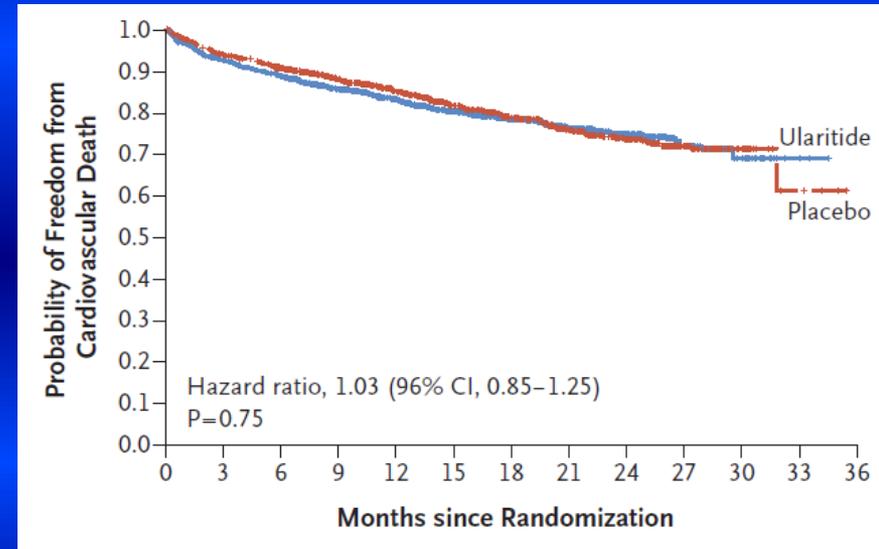
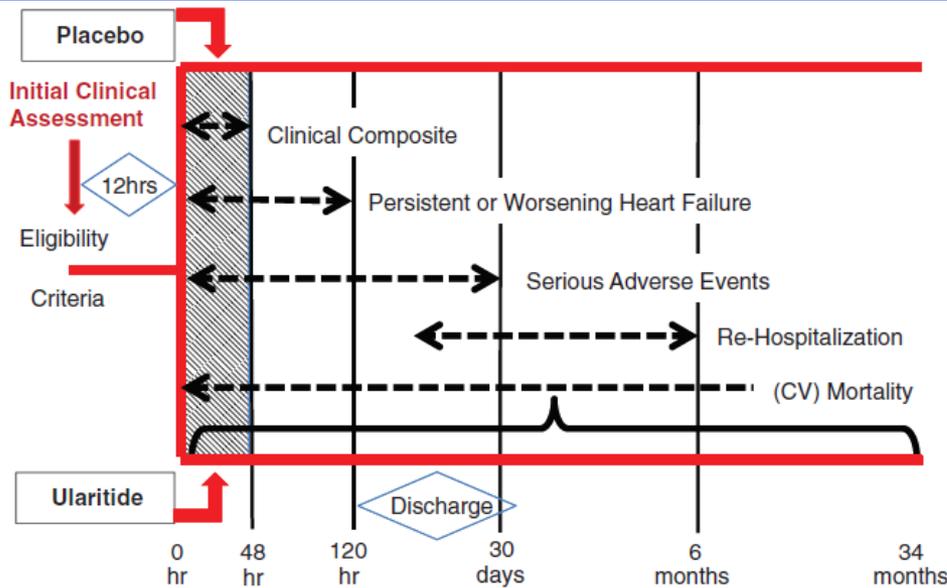
Eur Heart J. 2017 Aug 7;38(30):2364-2373. doi: 10.1093/eurheartj/ehx196.
Biased ligand of the angiotensin II type 1 receptor in patients with acute heart failure: a randomized, double-blind, placebo-controlled, phase IIB, dose ranging trial (BLAST-AHF).
Pang PS¹, Butler J², Collins SP³, Cotter G⁴, Davison BA⁴, Ezekowitz JA⁵, Filippatos G⁶, Levy PD⁷, Metra M⁸, Ponikowski P⁹, Teerlink JR¹⁰, Voors AA¹¹, Bharucha D¹², Goin K¹³, Soergel DG¹³, Felker GM¹⁴.

Effect of Ularitide on Cardiovascular Mortality in Acute Heart Failure

MAY 18, 2017

M. Packer, C. O'Connor, J.J.V. McMurray, J. Wittes, W.T. Abraham, S.D. Anker,

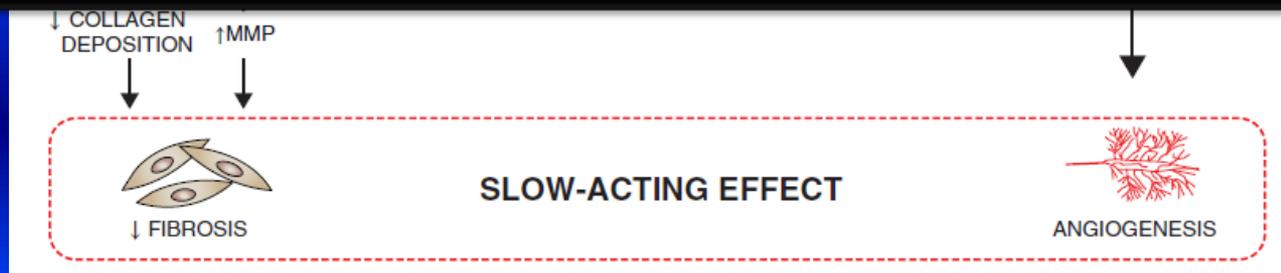
65% FEVI REDUCIDA



CONCLUSIONS

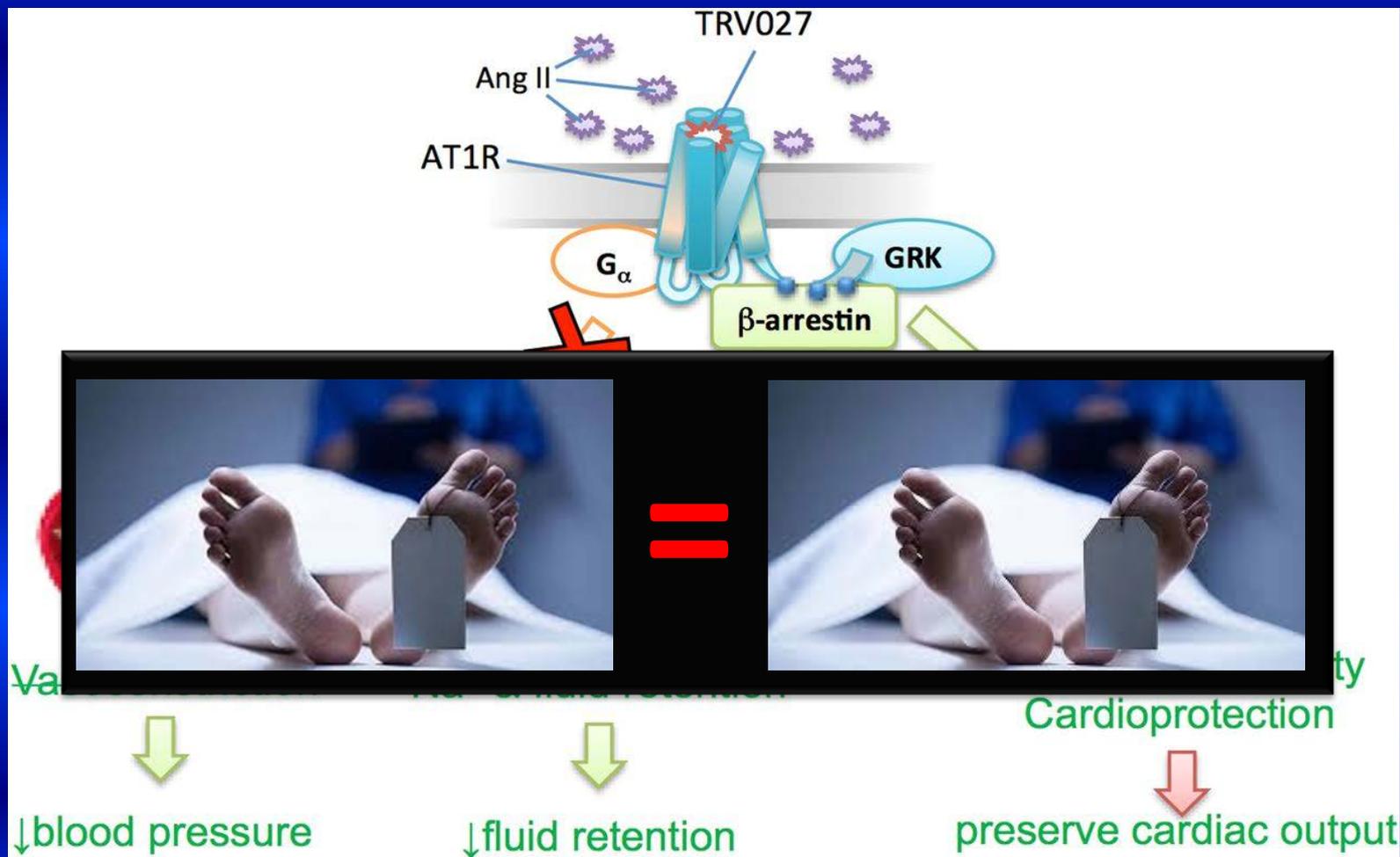
In patients with acute heart failure, ularitide exerted favorable physiological effects (without affecting cardiac troponin levels), but short-term treatment did not affect a clinical composite end point or reduce long-term cardiovascular mortality.

Serelaxin in addition to standard therapy in acute heart failure: rationale and design of the RELAX-AHF-2 study



Biased ligand of the angiotensin II type 1 receptor in patients with acute heart failure: a randomized, double-blind, placebo-controlled, phase IIB, dose ranging trial (BLAST-AHF).

Pang PS¹, Butler J², Collins SP³, Cotter G⁴, Davison BA⁴, Ezekowitz JA⁵, Filippatos G⁶, Levy PD⁷, Metra M⁸, Ponikowski P⁹, Teerlink JR¹⁰, Voors AA¹¹, Bharucha D¹², Goin K¹³, Soergel DG¹³, Felker GM¹⁴.

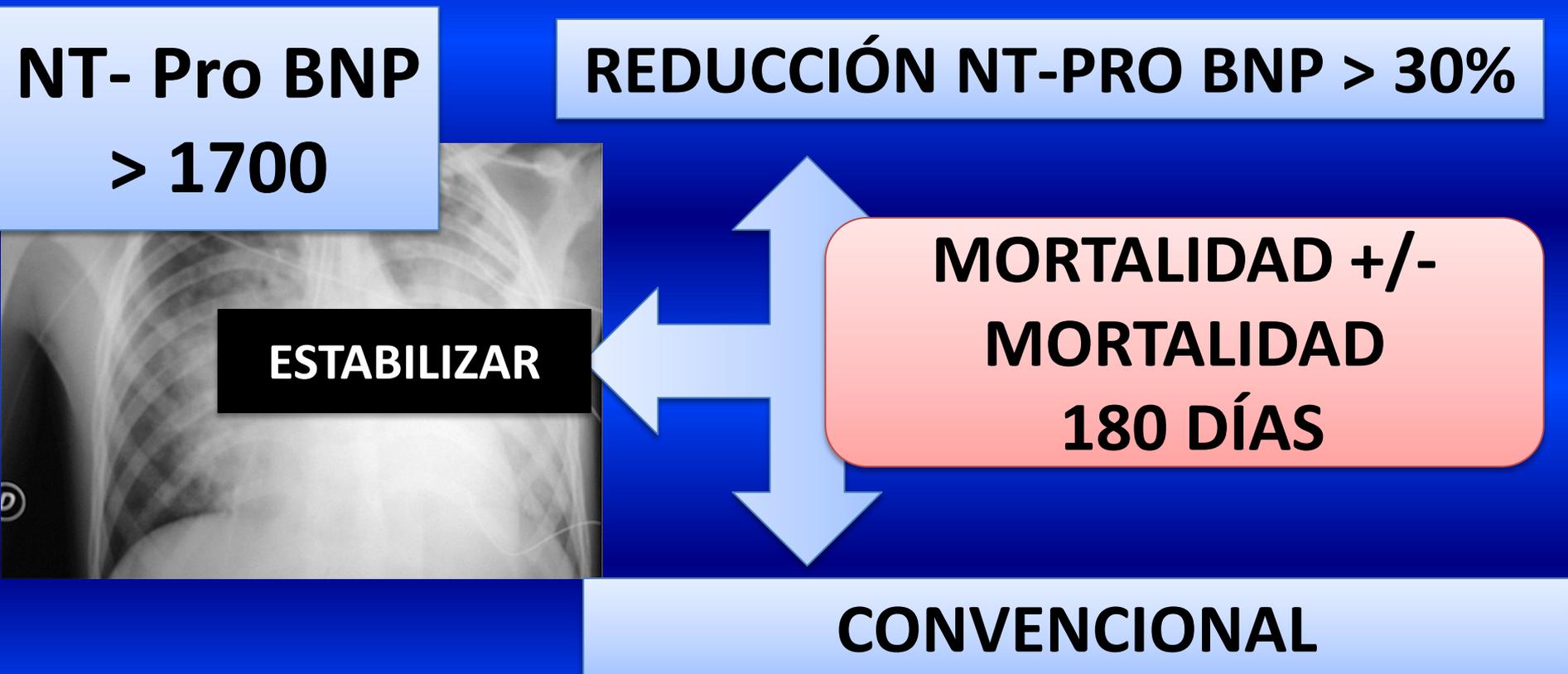


ORIGINAL RESEARCH ARTICLE

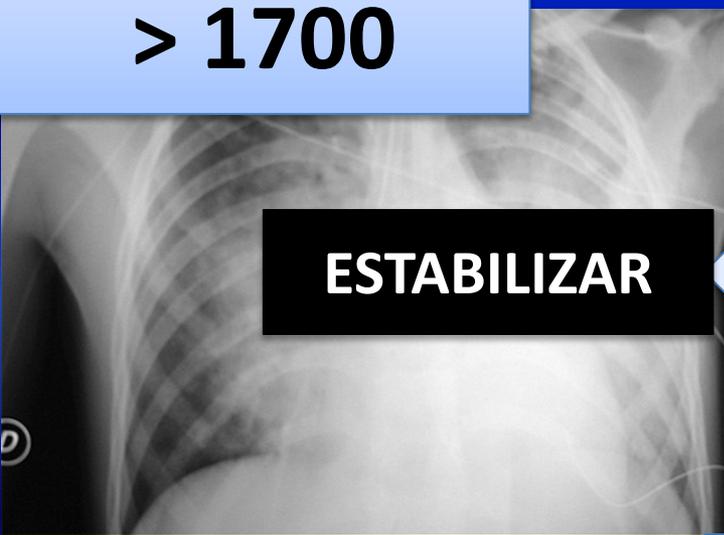


NT-proBNP (N-Terminal pro-B-Type Natriuretic Peptide)-Guided Therapy in Acute Decompensated Heart Failure

PRIMA II Randomized Controlled Trial (Can NT-ProBNP-Guided Therapy During Hospital Admission for Acute Decompensated Heart Failure Reduce Mortality and Readmissions?)



**NT- Pro BNP
> 1700**



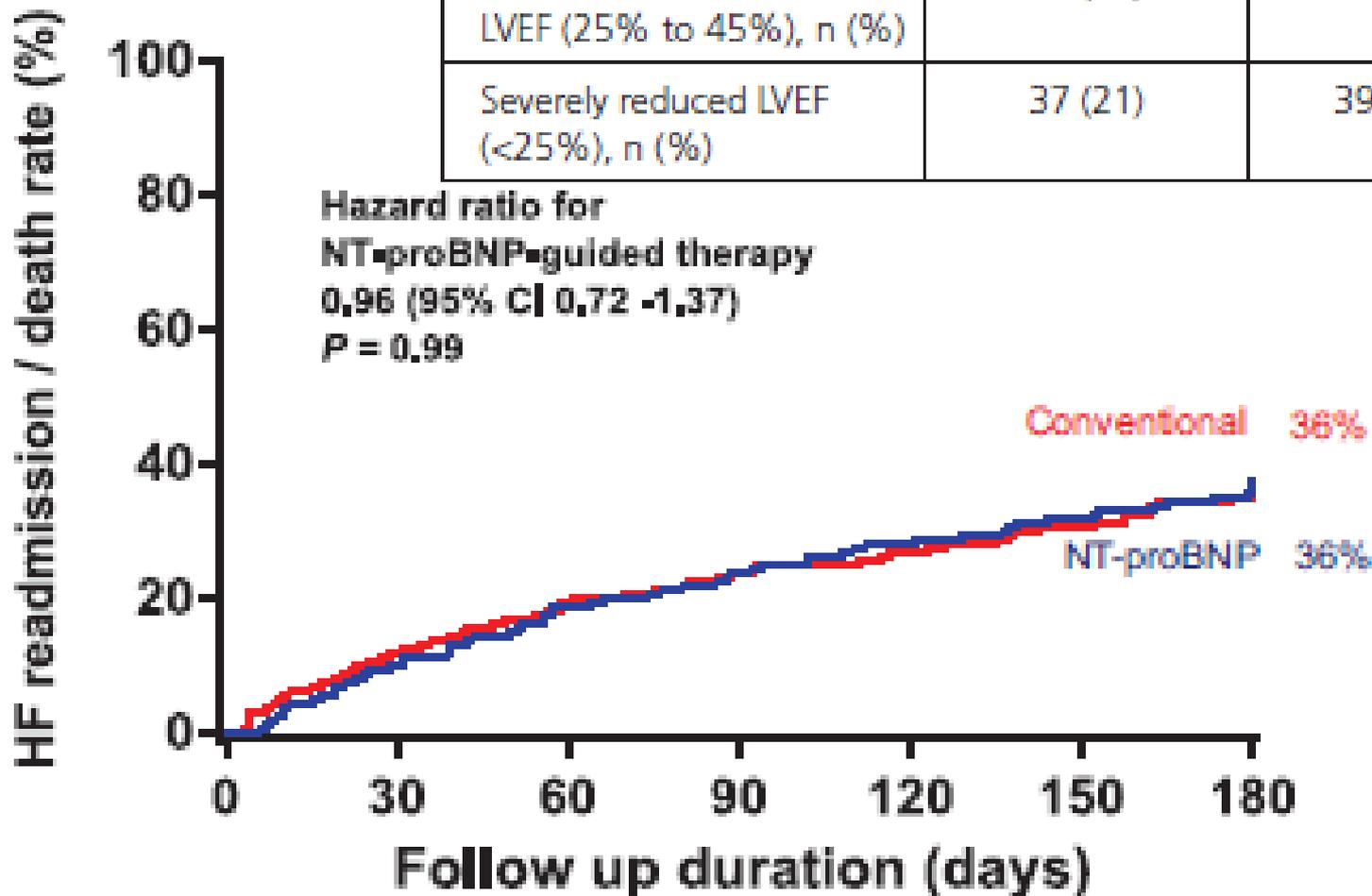
ESTABILIZAR

REDUCCIÓN NT-PRO BNP > 30%

**MORTALIDAD +/-
MORTALIDAD
180 DÍAS**

CONVENCIONAL

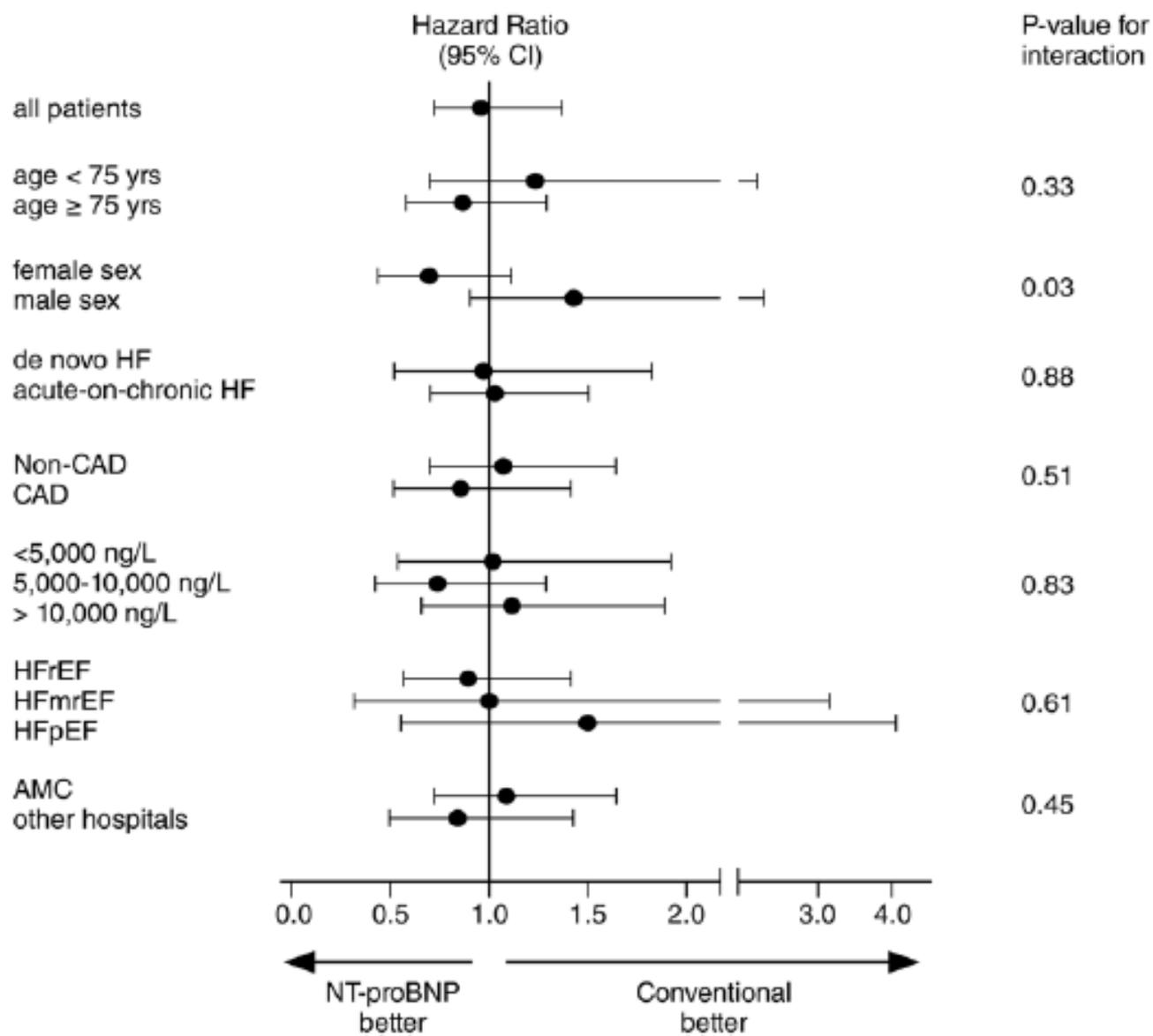
Preserved LVEF (>45%), n (%)	47 (27)	49 (30)
Mild-moderate reduced LVEF (25% to 45%), n (%)	89 (51)	78 (47)
Severely reduced LVEF (<25%), n (%)	37 (21)	39 (24)



No. at Risk

NT-proBNP	201	182	166	156	145	138	130
Conventional	203	181	166	156	149	143	132

All-cause mortality / HF readmissions

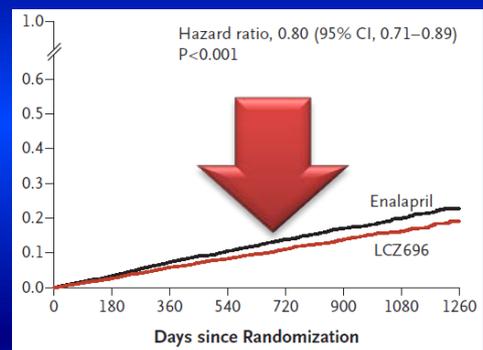
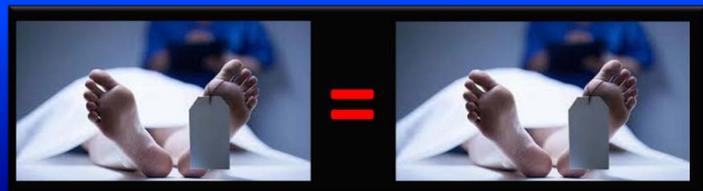




1

2

3



Rationale for and design of the TRUE-AHF trial: the effects of ularitide on the short-term clinical course and long-term mortality of patients with acute heart failure

Serelaxin in addition to standard therapy in acute heart failure: rationale and design of the RELAX-AHF-2 study

Eur Heart J, 2017 Aug 7;38(30):2364-2373. doi: 10.1093/eurheartj/ehx196.

Biased ligand of the angiotensin II type 1 receptor in patients with acute heart failure: a randomized, double-blind, placebo-controlled, phase IIB, dose ranging trial (BLAST-AHF).

Pana PS¹, Butler J², Collins SP³, Cotter G⁴, Davison BA⁴, Ezekowitz JA⁵, Filippatos G⁶, Levy PD⁷, Metra M⁸, Ponikowski P⁹, Teerlink JR¹⁰, Voors AA¹¹, Bharucha D¹², Golin K¹³, Soergel DG¹³, Felker GM¹⁴.

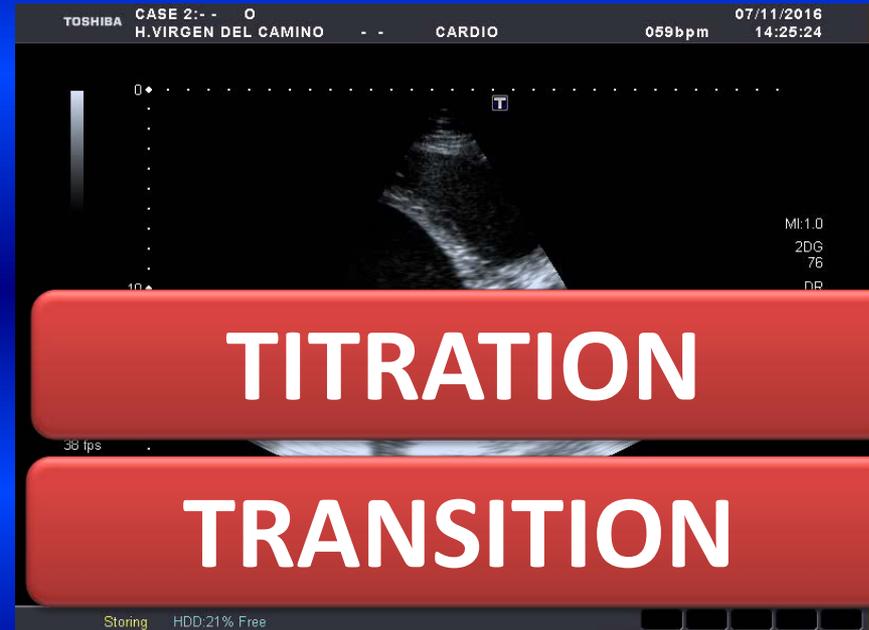
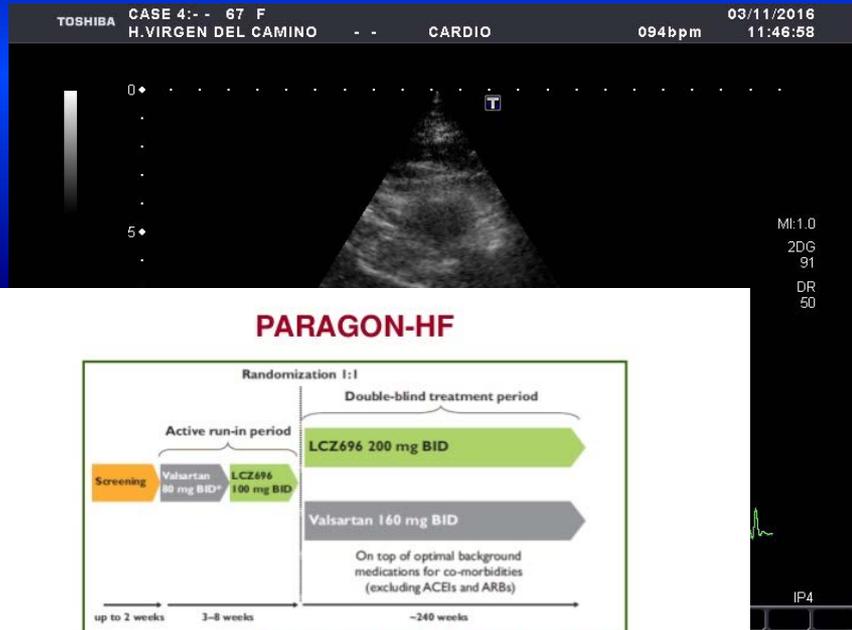
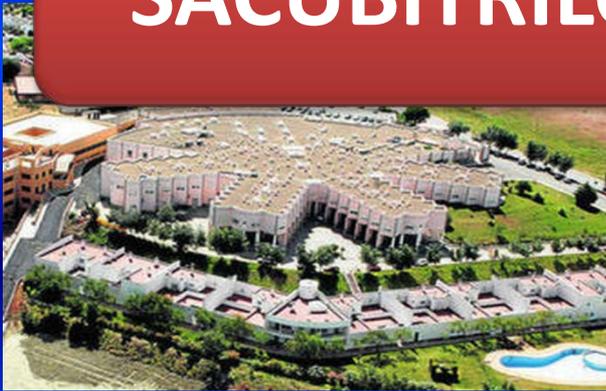


14 internistas

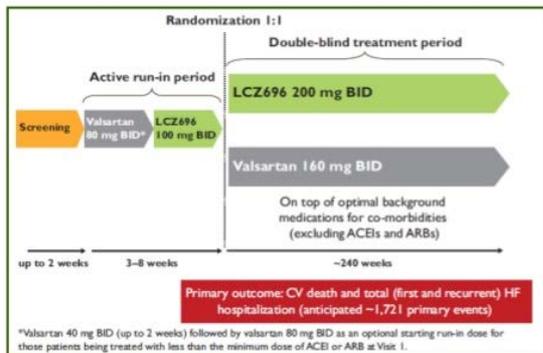


120 camas MI

SACUBITRILO/VALSARTAN



PARAGON-HF



Primary objective

- The primary objective of this trial is to compare LCZ696 to valsartan in reducing the rate of the composite endpoint of CV death and total (first and recurrent) HF hospitalizations, in HF patients (NYHA Class II-IV) with preserved EF (left ventricular ejection fraction [LVEF] $\geq 45\%$).

TITRATION

TRANSITION

PARADISE-MI

Initiating sacubitril/valsartan (LCZ696) in heart failure: results of TITRATION, a double-blind, randomized comparison of two uptitration regimens

NIH U.S. National Library of Medicine

ClinicalTrials.gov

Find Studies About Studies Submit Studies Resources

TRANSITION -- IC

Save this study

Comparison of Pre- and Post-discharge Initiation of LCZ696 Therapy in HFrEF Patients
Decompensation Event (TRANSITION)

NIH U.S. National Library of Medicine

ClinicalTrials.gov

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PARADISE--MI

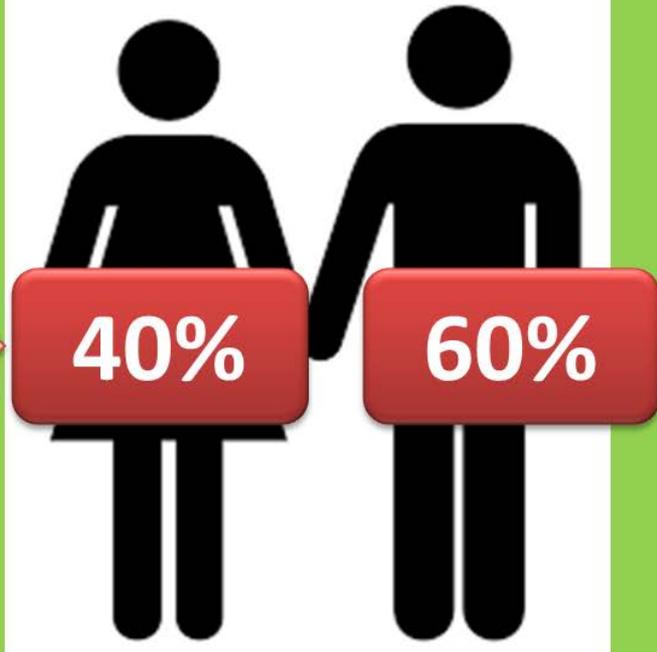
Save this study Saved Studies (0)

Prospective ARNI vs ACE Inhibitor Trial to Determine Superiority in Reducing Heart Failure Events After
MI (PARADISE-MI)





80

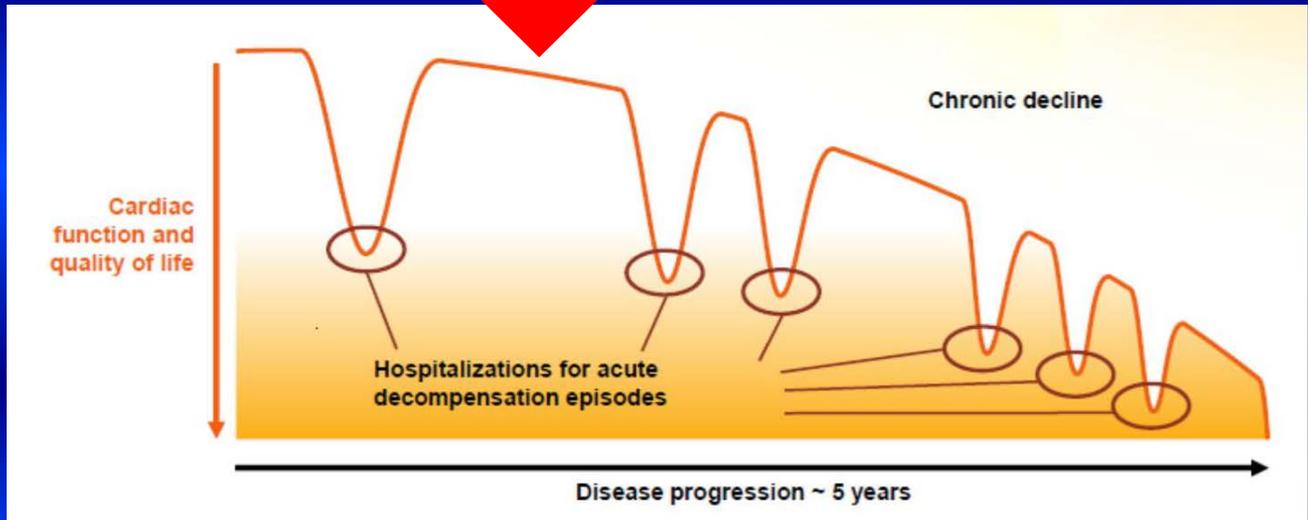
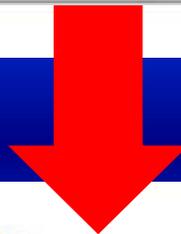


Angiotensin–Neprilysin Inhibition versus Enalapril
in Heart Failure



VARÓN
63 AÑOS
FE 29%
NYHA II
TAS:122 mmHg
Fc: 72 lpm
NT pro-BNP:1600
pg/ml

78 años

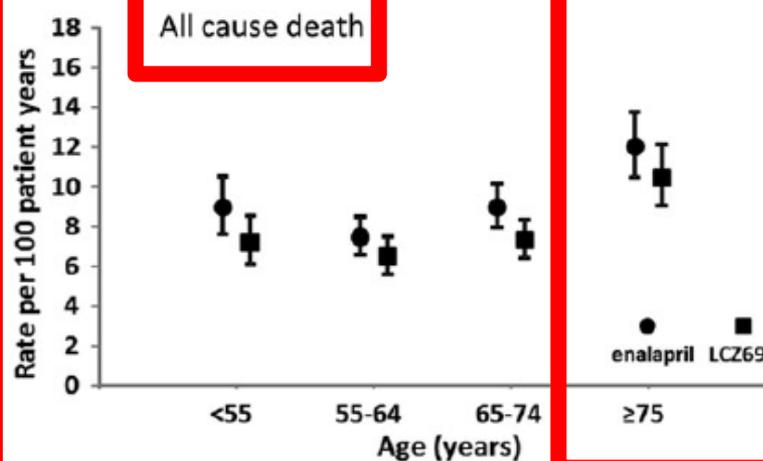
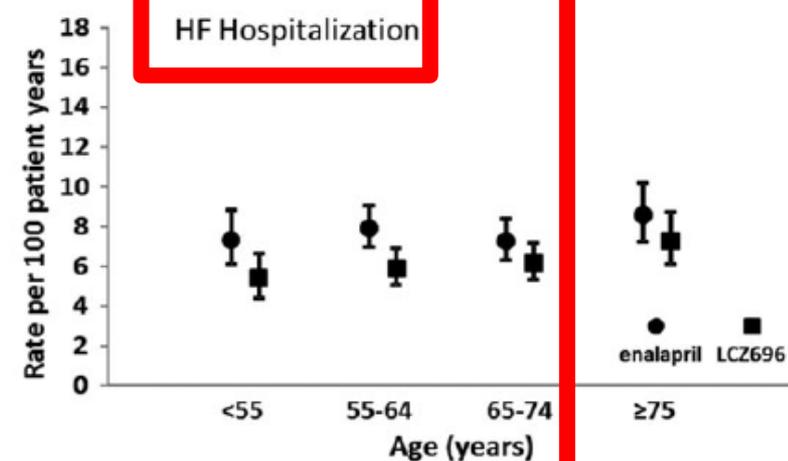
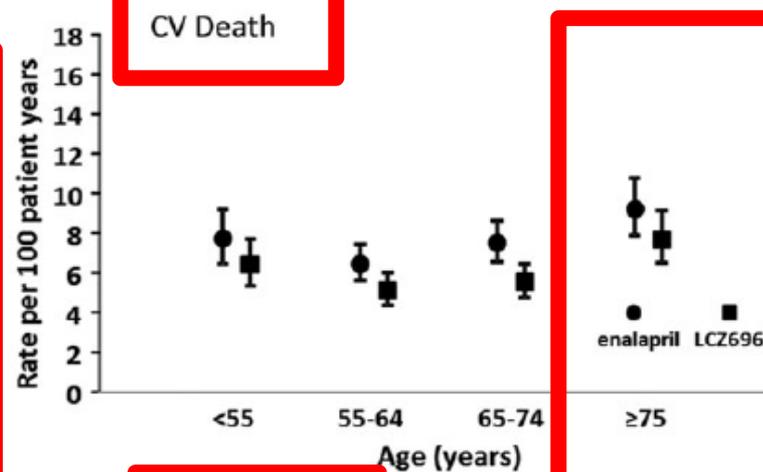
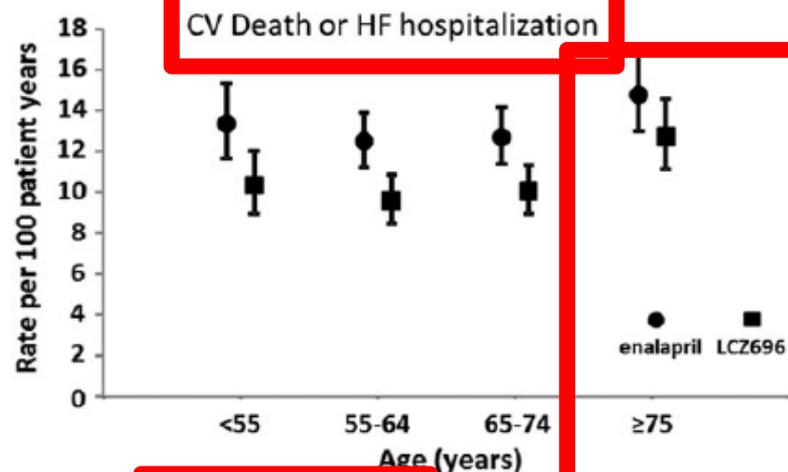


20%

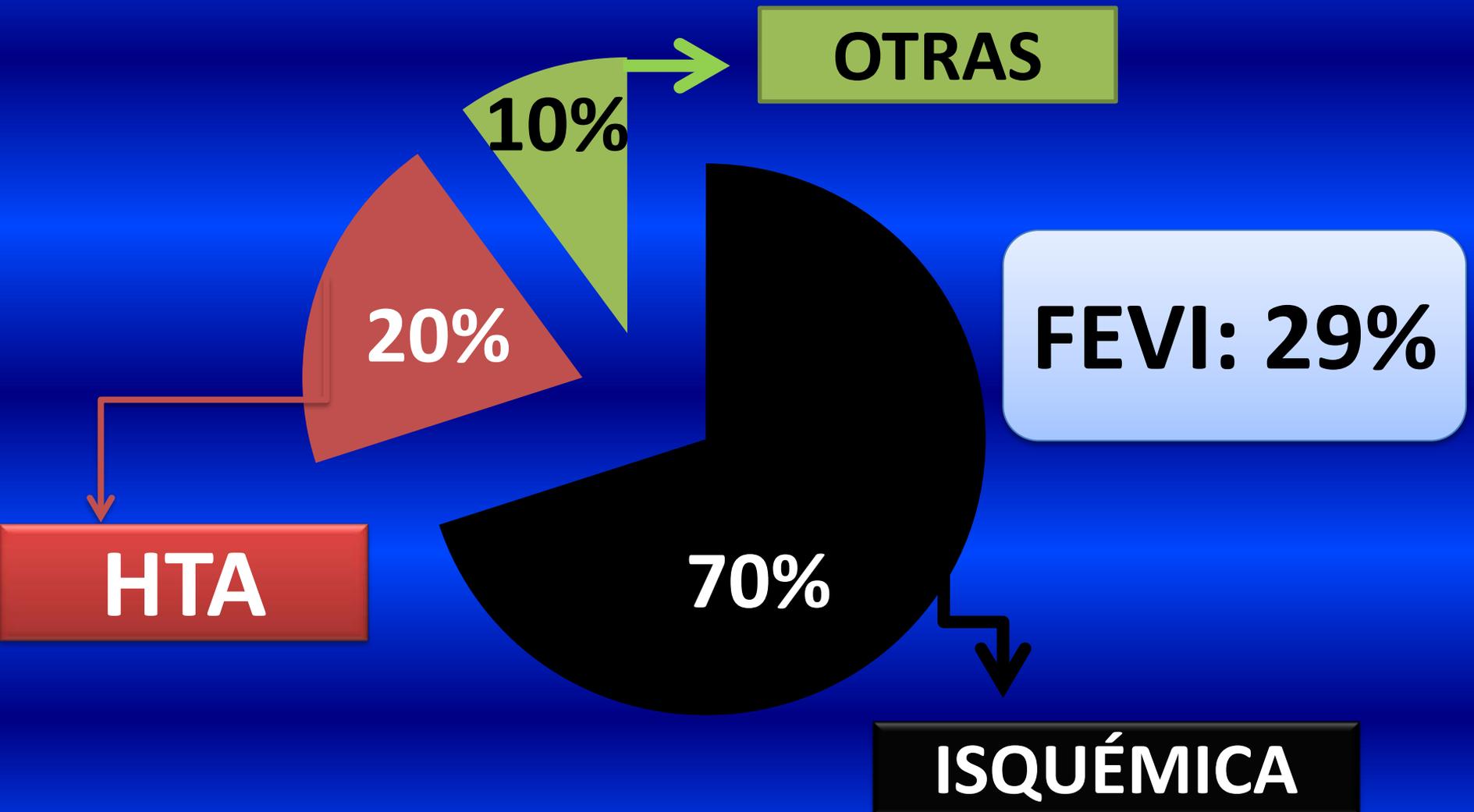
≥ 75 years
(n = 1563)

65–74 years
(n = 2557)

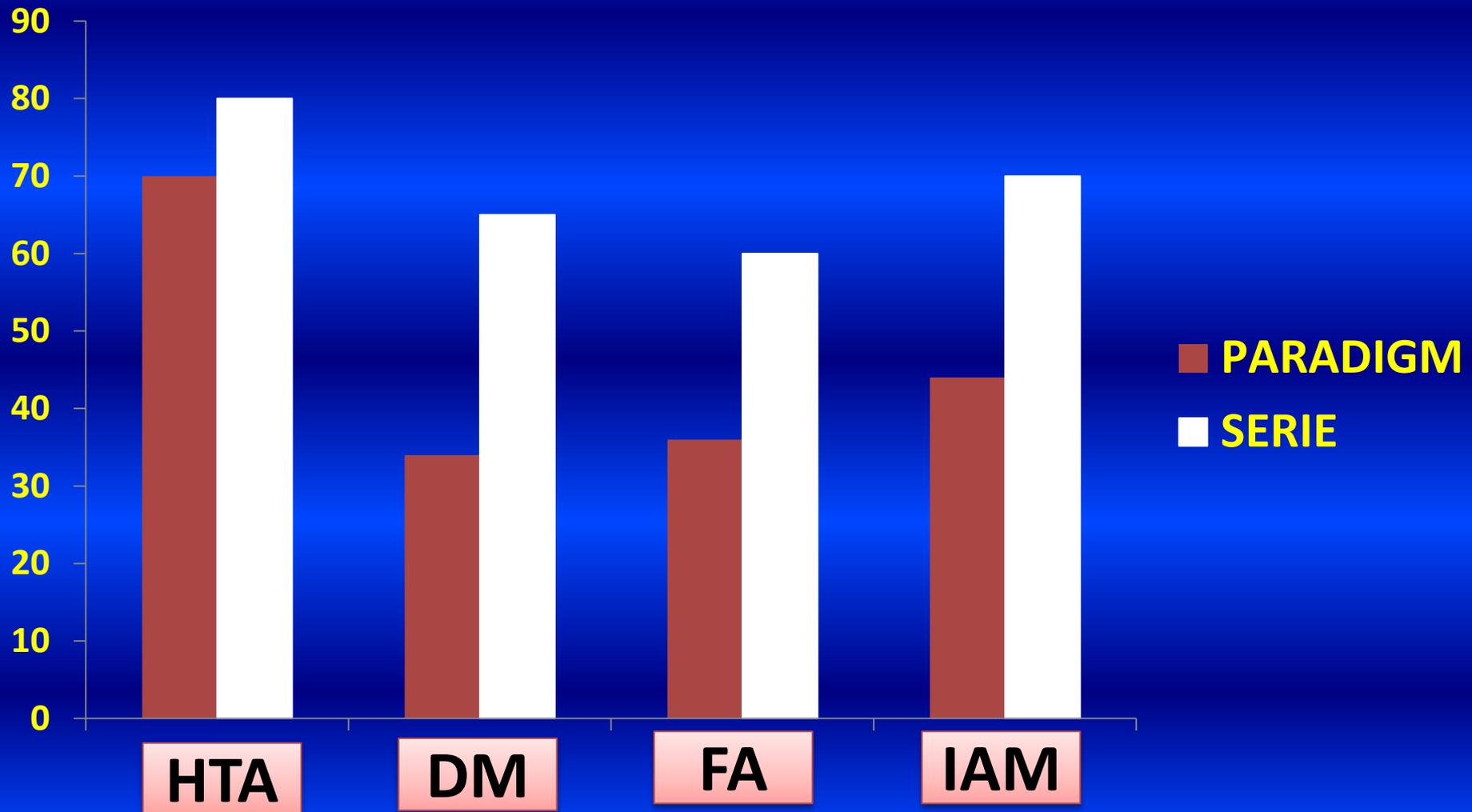
Efficacy and safety of LCZ696 (sacubitril-valsartan) according to age: insights from PARADIGM-HF



ETIOLOGÍA



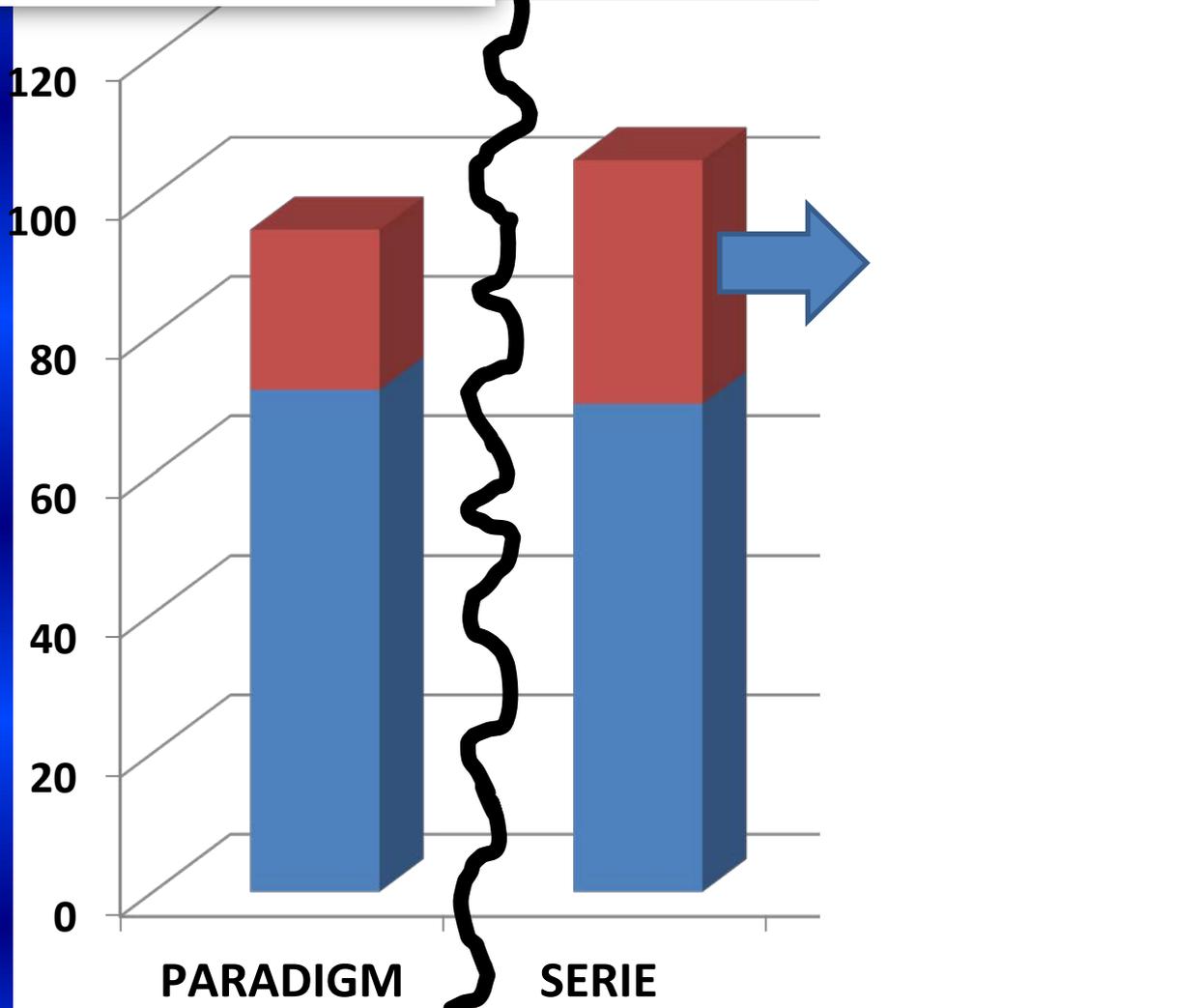
COMORBILIDADES



Angiotensin–Nepriylsin Inhibition versus Enalapril in Heart Failure

John J.V. McMurray, M.D., Milton Packer, M.D., Akshay S. Desai, M.D., M.P.H., Jianjian Gong, Ph.D., Martin P. Lefkowitz, M.D., Adel R. Rizkala, Pharm.D., Jean L. Rouleau, M.D., Victor C. Shi, M.D., Scott D. Solomon, M.D., Karl Swedberg, M.D., Ph.D., and Michael R. Zile, M.D., for the PARADIGM-HF Investigators and Committees*

CLASE FUNCIONAL



10% PASA A CLASE II

- NYHA III
- NYHA II

DOSIS DE FUROSEMIDA



60 mg



80 mg



80 mg

Sac/Val
24/26

80 mg

3 días
iv

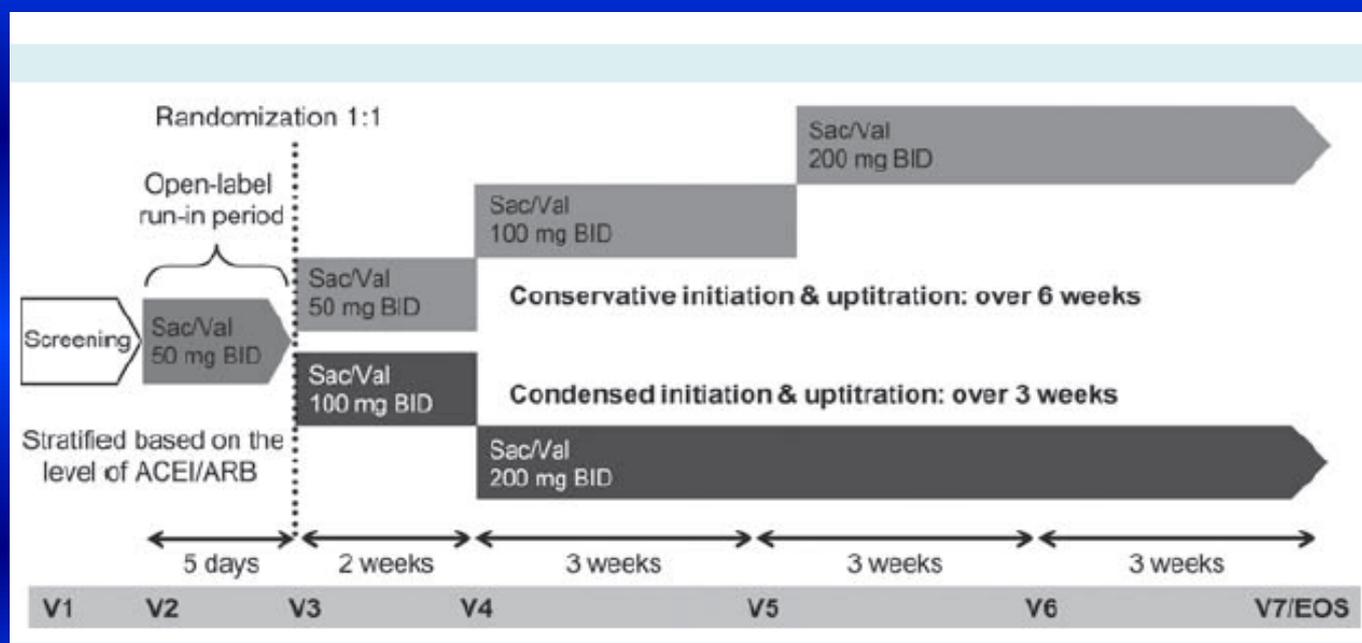
2 días
vo

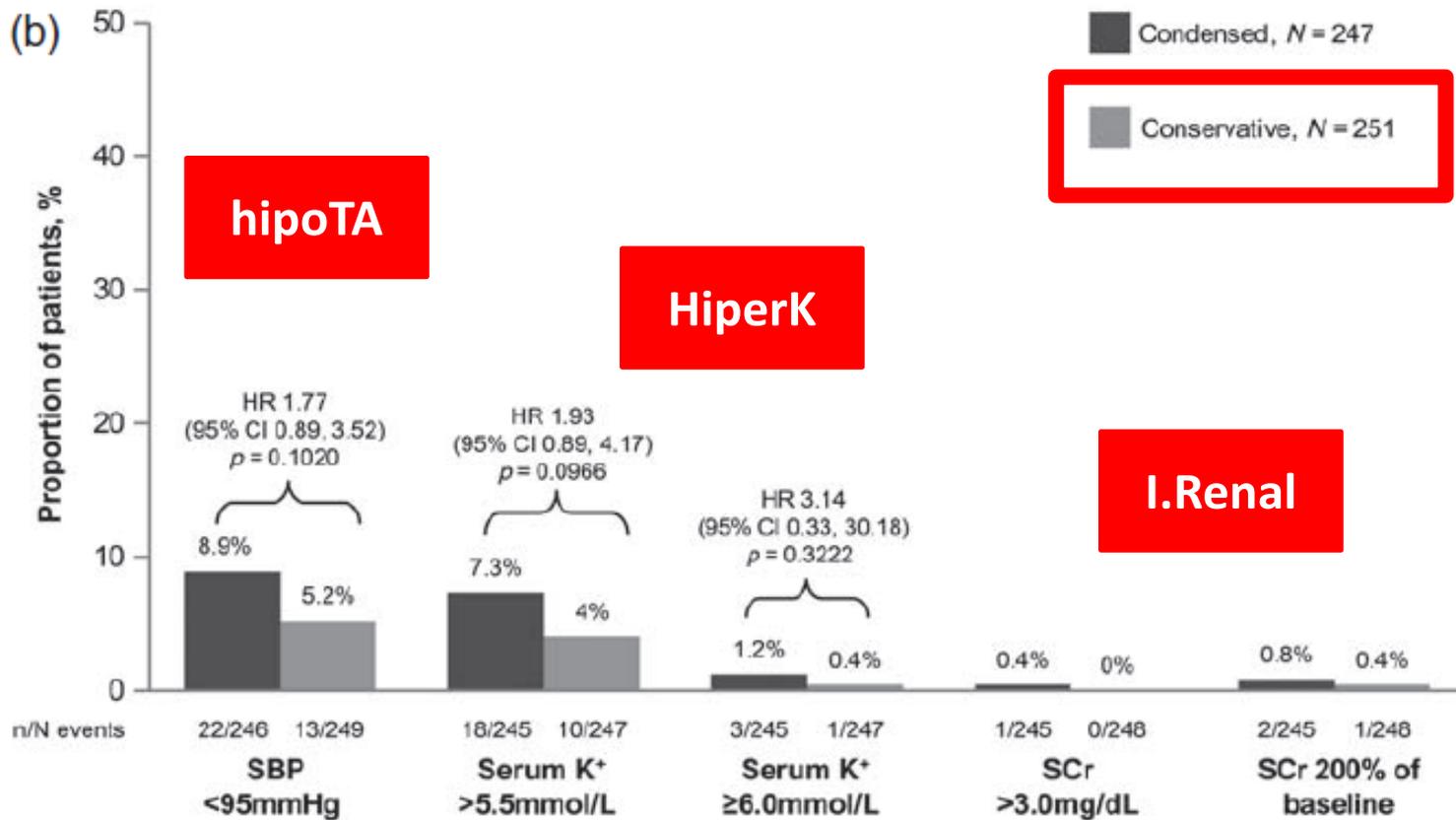
2 días
vo

7 DÍAS



Initiating sacubitril/valsartan (LCZ696) in heart failure: results of TITRATION, a double-blind, randomized comparison of two uptitration regimens





MENOS PROBABILIDAD DE PROBLEMAS

DOSIS DE FUROSEMIDA



60 mg



80 mg



60 mg



40-60 mg

24/26 mg



49/51 mg

60%

97/103 mg

40%

49/51 mg

DOSIS DE SAC/VALSARTAN

DOSIS DE FUROSEMIDA



60 mg



80 mg



60 mg



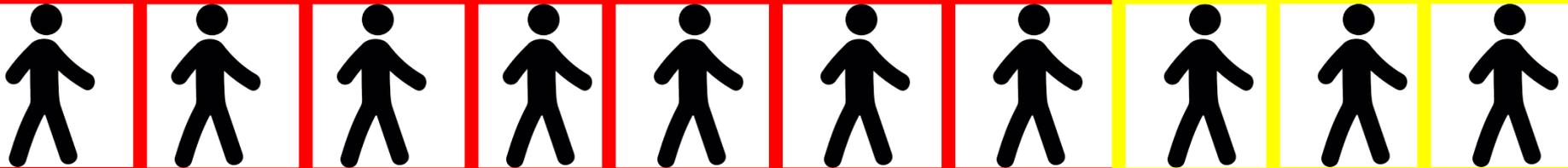
40-60 mg

24/26 mg

49/51 mg

60%

97/103 mg

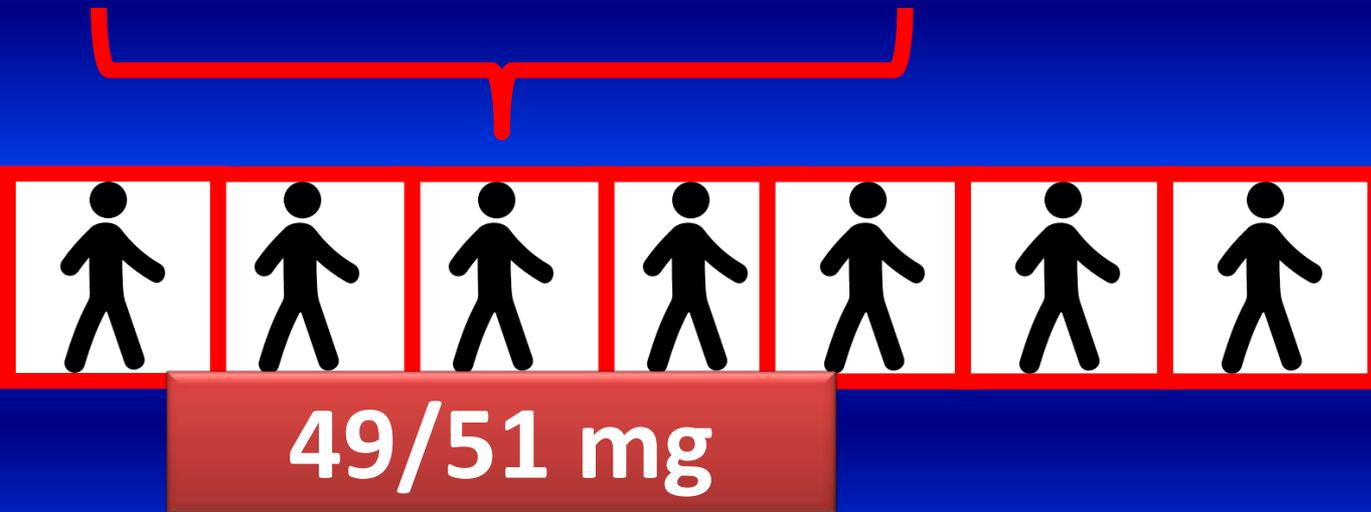


TA NO LO PERMITE

FG BAJA 10-20%

PACIENTES QUE NO ALCANZAN DOSIS MÁXIMAS HIPOTENSIÓN

97/103 mg

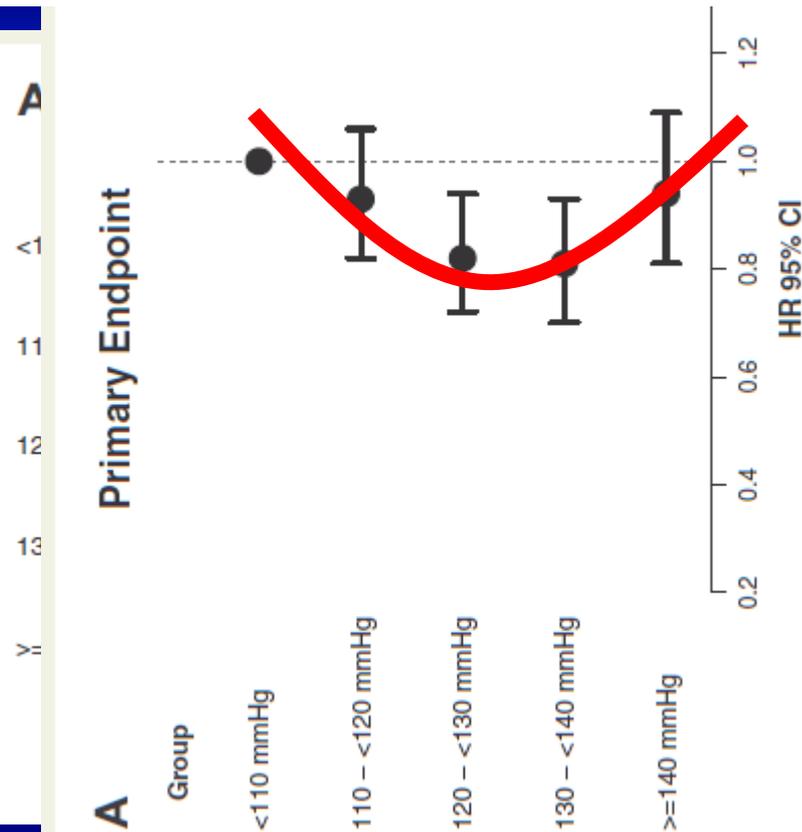


Systolic blood pressure, cardiovascular outcomes and efficacy and safety of sacubitril/valsartan (LCZ696) in patients with chronic heart failure and reduced ejection fraction: results from PARADIGM-HF

European Heart Journal (2017) 0, 1–12



PAS BASAL 110 –
140 mmHg
MAYOR BENEFICIO



Efficacy of sacubitril/valsartan vs. enalapril at lower than target doses in heart failure with reduced ejection fraction: the PARADIGM-HF trial

**LOS PACIENTES
CON DOSIS
MEDIA, OBTIENEN
BENEFICIO**

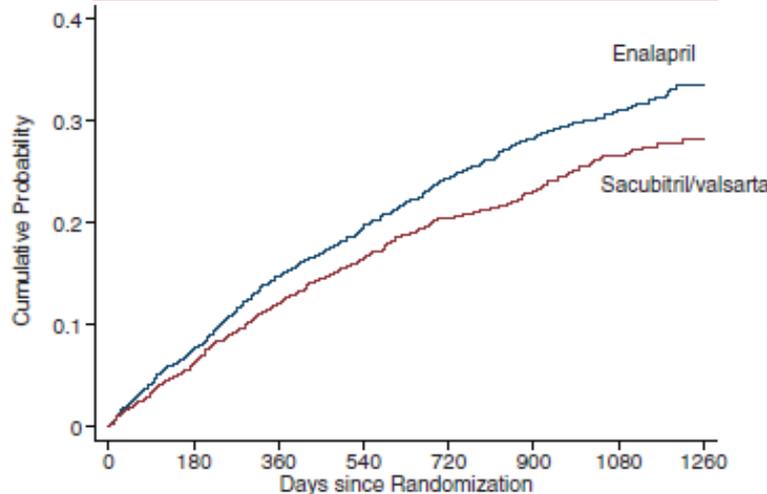


Effects of Sacubitril/Valsartan in the PARADIGM-HF Trial (Prospective Comparison of ARNI with ACEI to Determine Impact on Global Mortality and Morbidity in Heart Failure) According to Background Therapy

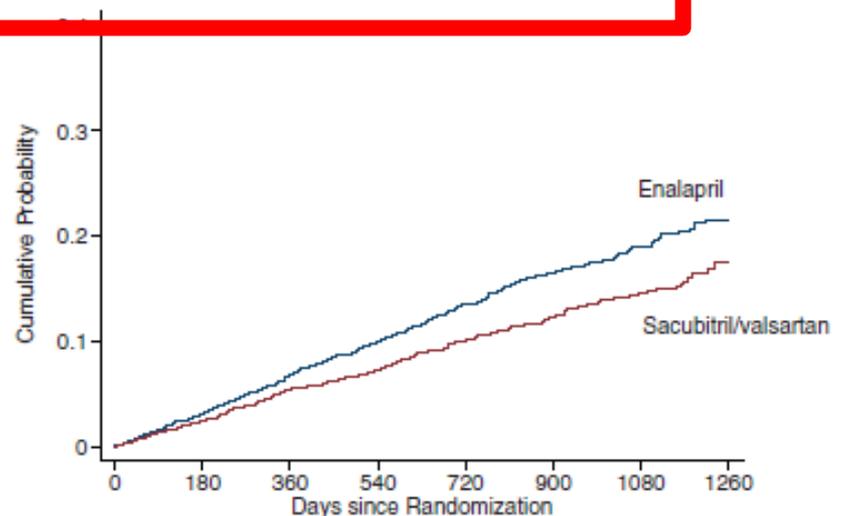
Naoki Okumura, Pardeep S. Jhund, Jianjian Gong, Martin P. Lefkowitz, Adel R. Rizkala, Jean L. Rouleau, Victor C. Shi, Karl Swedberg, Michael R. Zile, Scott D. Solomon, Milton Packer and John J.V. McMurray

on behalf of the PARADIGM-HF Investigators and Committees*

Beta-blocker dose <50% target dose



No mineralocorticoid receptor antagonist



PACIENTES QUE NO ALCANZAN DOSIS MÁXIMAS POR DISFUNCIÓN RENAL



FGe: **45-60**
ml/min/1,73m²



FGe: **30-45**
ml/min/1,73m²



FGe: **45-60**
ml/min/1,73m²

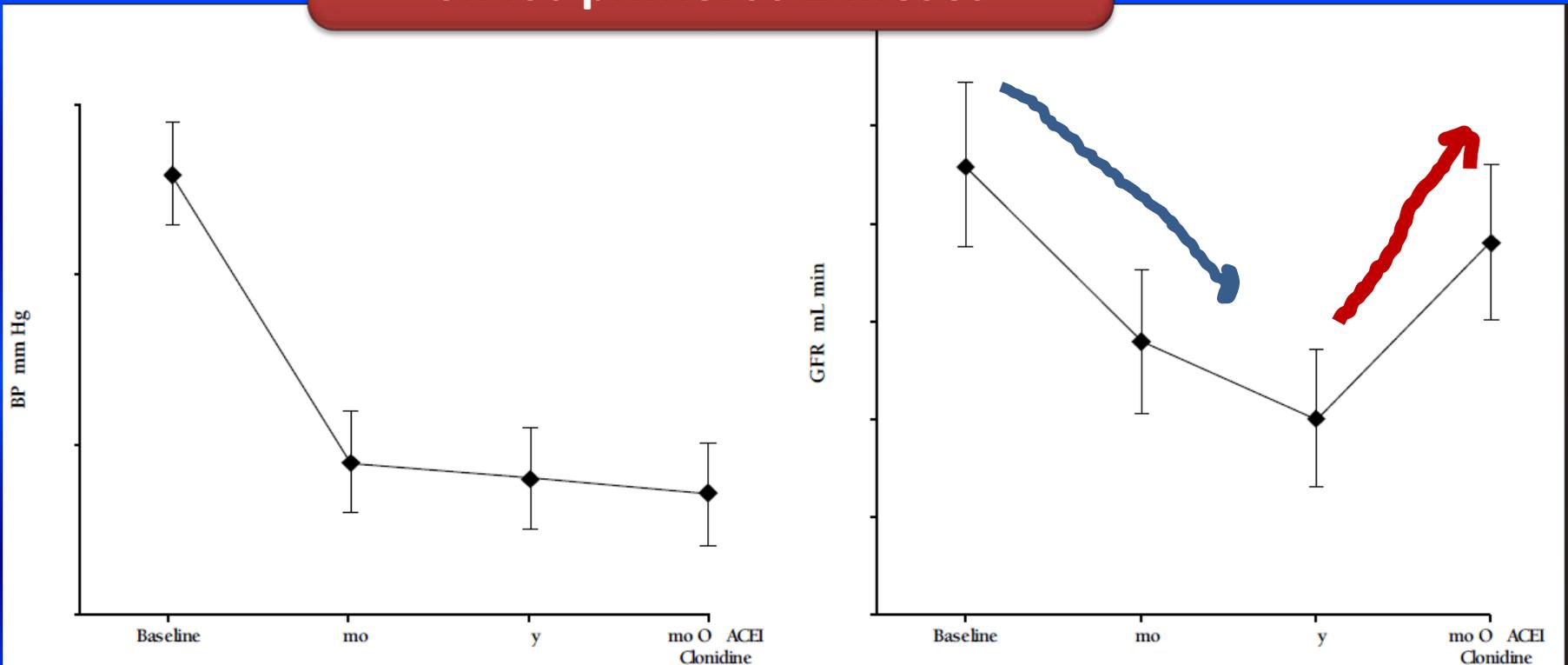
Angiotensin-Converting Enzyme Inhibitor–Associated Elevations in Serum Creatinine

Is This a Cause for Concern?

Arch Intern Med. 2000;160:685-693

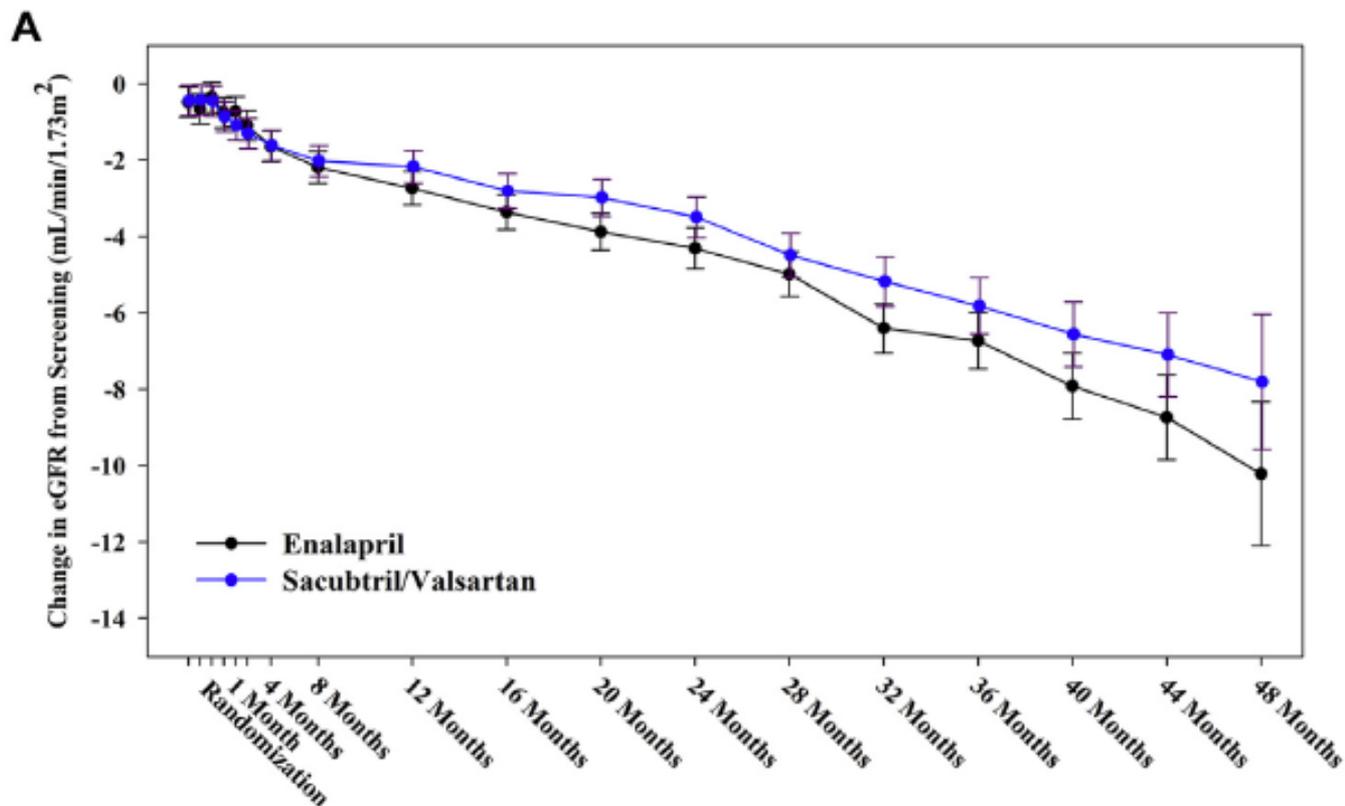
George L. Bakris, MD; Matthew R. Weir, MD

**Frecuente bajada < 30% de FG
en los primeros 2 meses**



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Renal Effects and Associated Outcomes During Angiotensin-Nepriylsin Inhibition in Heart Failure



RETIRADA DE TRATAMIENTO

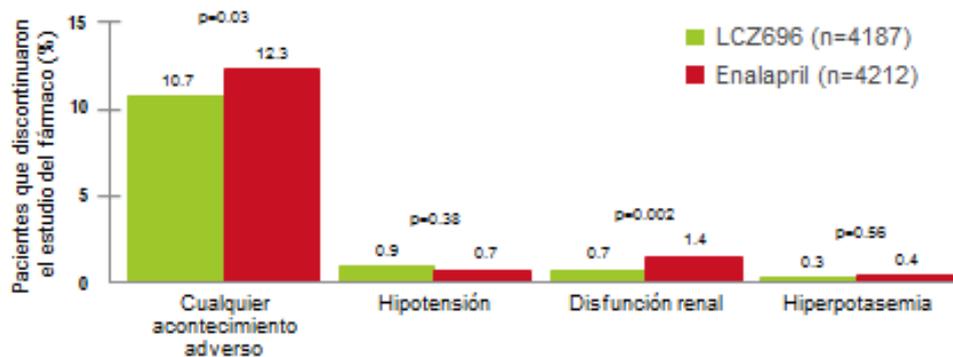
4/80 (5%)

3 HIPOTENSIÓN

1 DISPEPSIA



Acontecimientos adversos que provocaron una discontinuación permanente del estudio

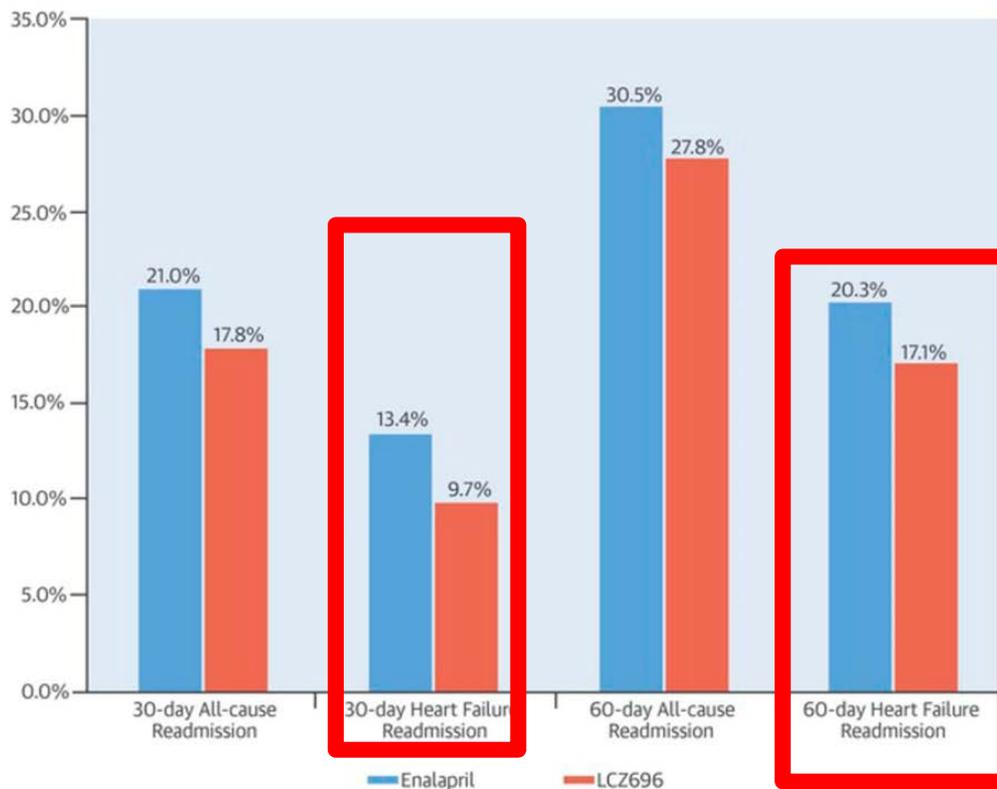


REINGRESO POR I.CARDIACA



1m: 4/80 (5%)
3m: 7/80 (9%)

CENTRAL ILLUSTRATION: Influence of LCZ696 on Readmission: Rates After Investigator-Reported HF Hospitalization According to Treatment Assignment





EXITUS-3 MESES

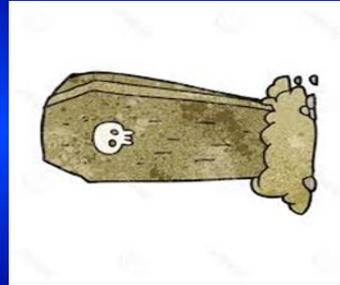
9/80 (11%)



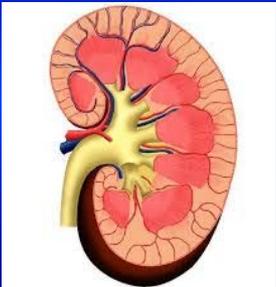
3 ICC



1 IAM



1 MS



1 IRENAL



1 TEP



2 SEPSIS

NO ESTAMOS SOLOS



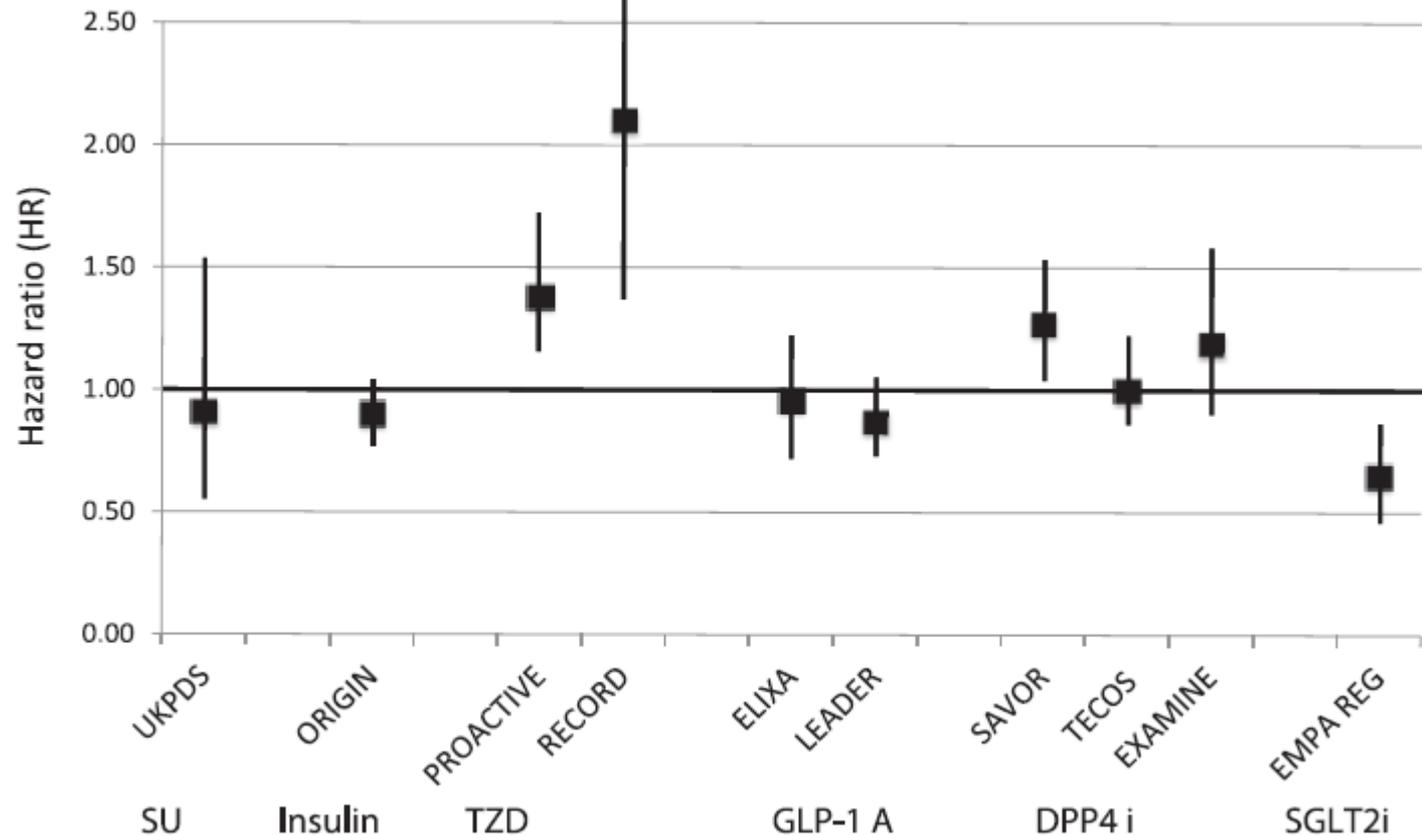
Heart failure outcomes in clinical trials of glucose-lowering agents in patients with diabetes

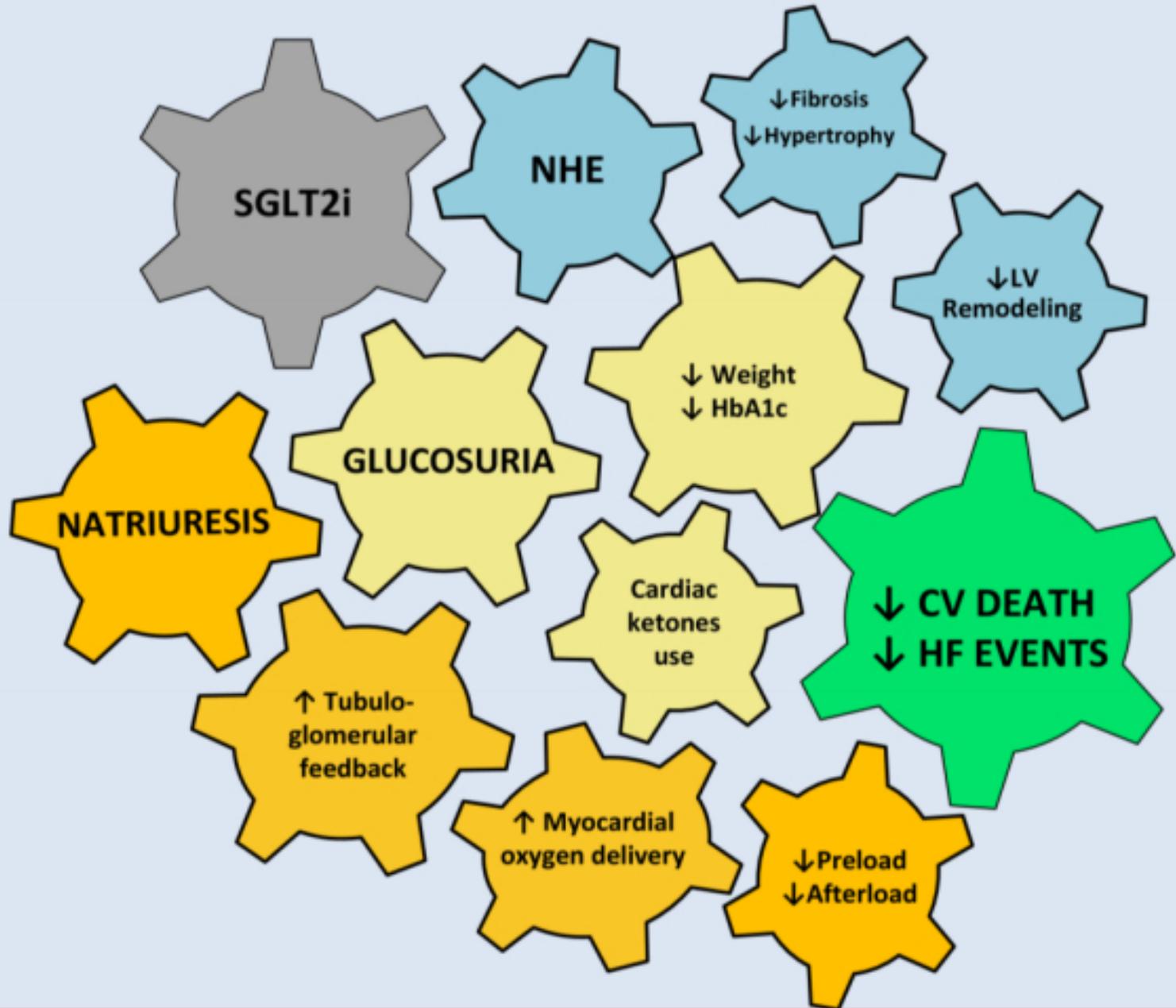
European Journal of Heart Failure (2016)



David H. Fitchett^{1*}, Jacob A. Udell², and Silvio E. Inzucchi³

HOSPITALIZACIÓN POR I.CARDIACA

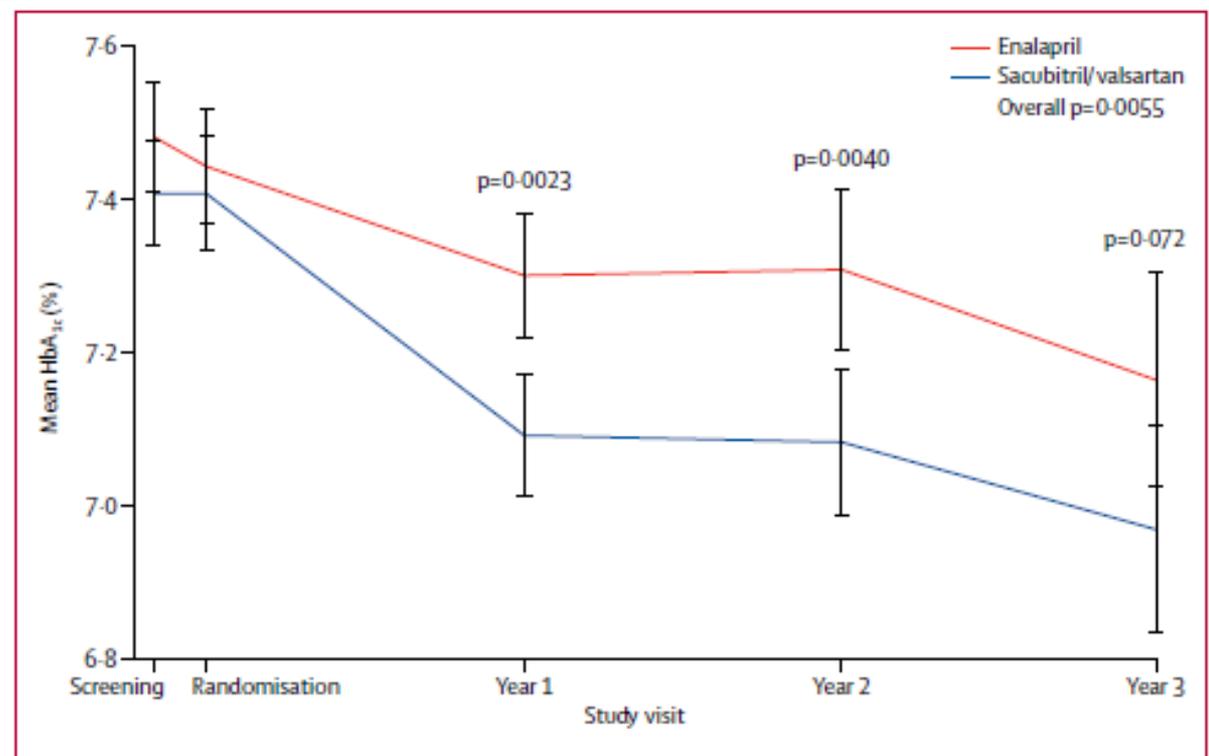




Effect of sacubitril/valsartan versus enalapril on glycaemic control in patients with heart failure and diabetes: a post-hoc analysis from the PARADIGM-HF trial

Jelena P Seferovic, Brian Claggett, Sara B Seidemann, Ellen W Seely, Milton Packer, Michael R Zile, Jean L Rouleau, Karl Swedberg, Martin Lefkowitz, Victor C Shi, Akshay S Desai, John J V McMurray, Scott D Solomon

Lancet Diabetes Endocrinol 2017
Published Online
March 18, 2017
[http://dx.doi.org/10.1016/S2213-8587\(17\)30087-6](http://dx.doi.org/10.1016/S2213-8587(17)30087-6)



(n=3778)

Figure 1: Changes in mean HbA_{1c} and confidence intervals by treatment group at screening, randomisation, 1-year, 2-year, and 3-year visits

INTRODUCCIÓN DE INSULINA

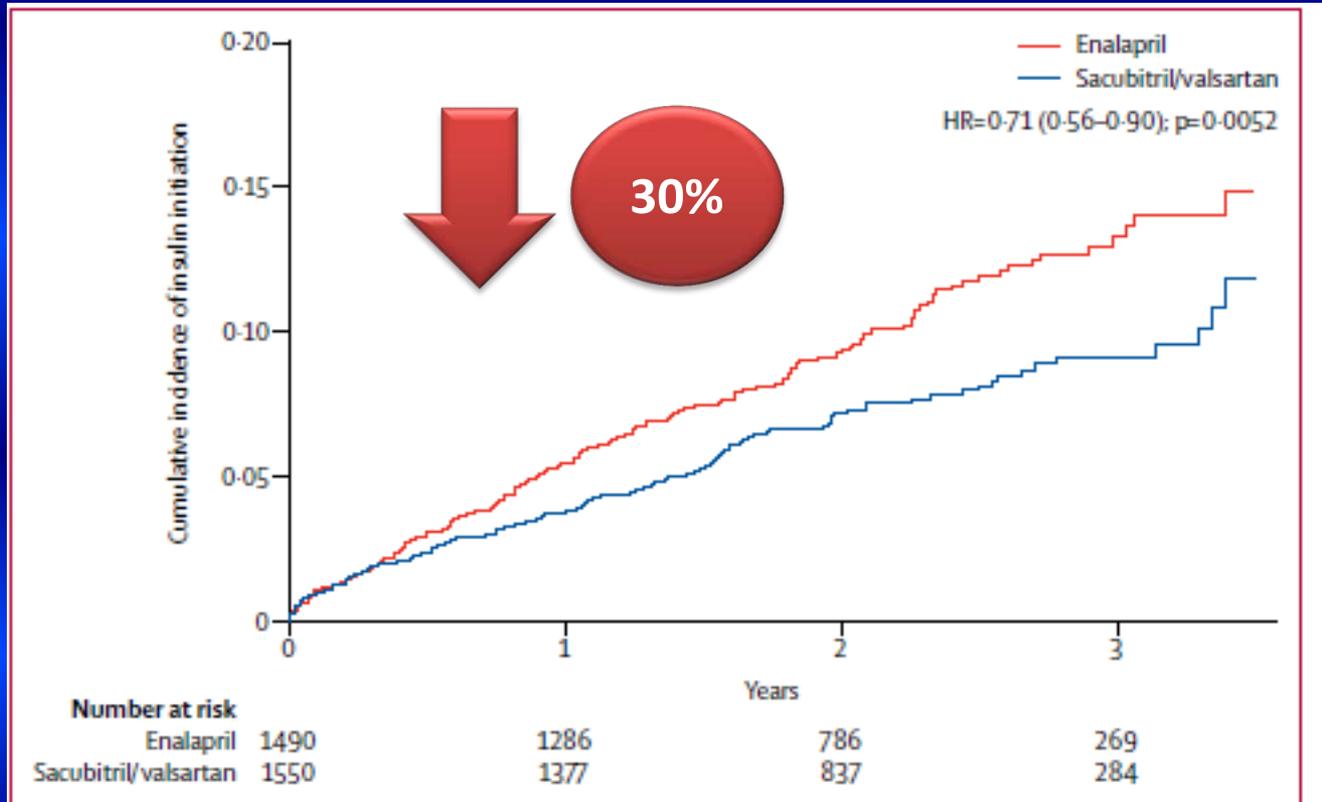


Figure 2: Kaplan-Meier curve showing time to insulin initiation in the sacubitril/valsartan and enalapril groups, in patients previously not treated with insulin

Effect of the angiotensin-receptor-neprilysin inhibitor **LCZ696** compared with enalapril on mode of death in heart failure patients

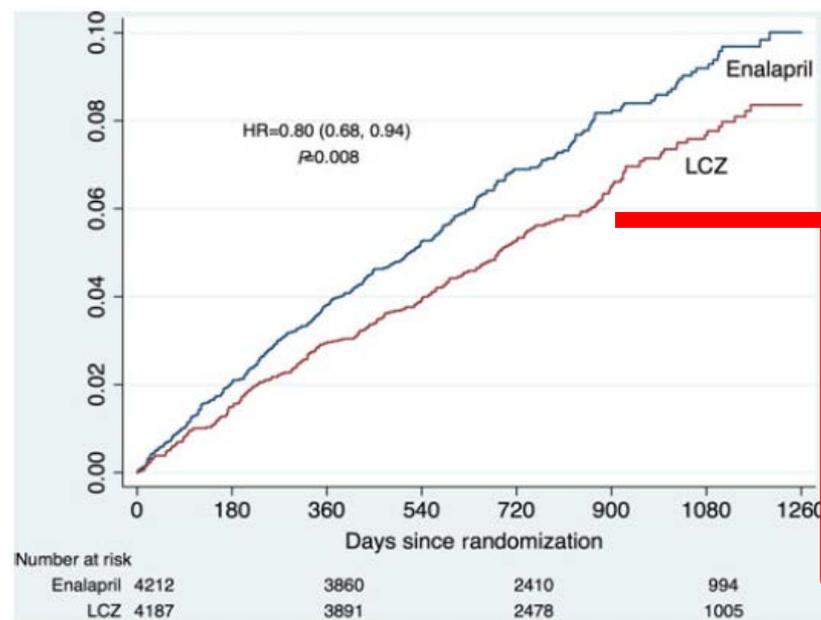
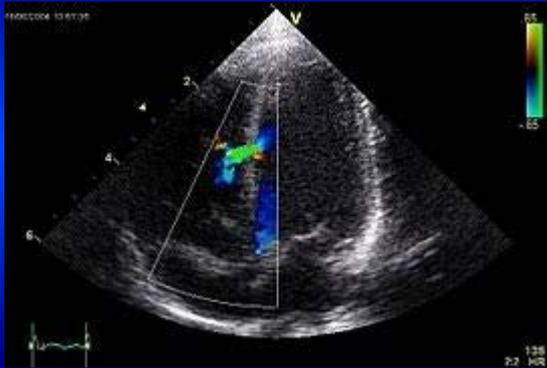


Figure 1 Kaplan–Meier survival curve for sudden death, by treatment. HR, hazard ratio.





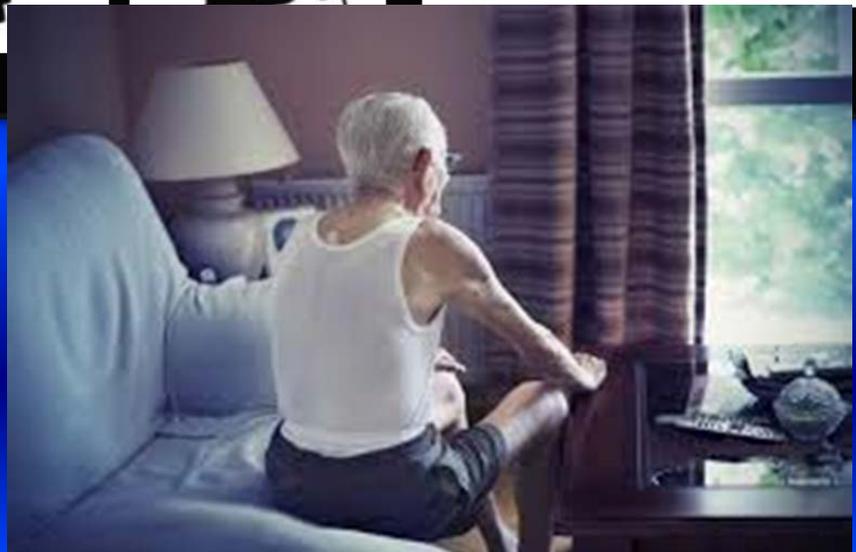
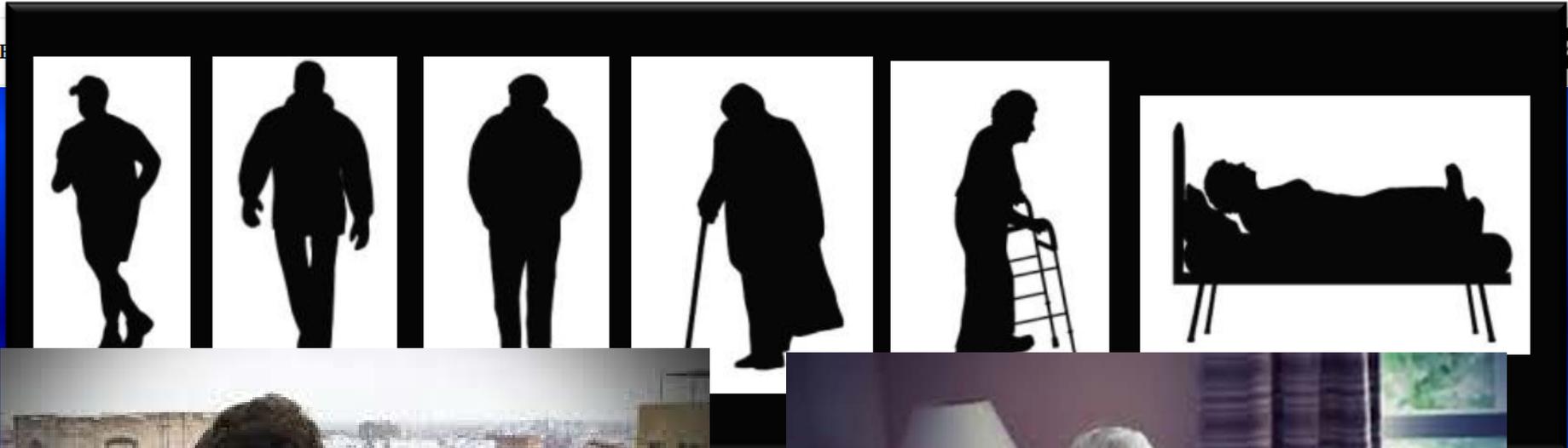
VALORACIÓN CLÍNICA

Sociedad

1 DE CADA 3 MAYORES DE 65 a

ANTROPOLOGÍA • Las investigacion

La epidemia de la soledad ya supera a la obesidad como amenaza para la salud



NUESTRA EXPERIENCIA “AL ALTA”

- Inicio de tratamiento cuando llevamos 2 días de diuréticos orales.
- Normalmente empezamos por dosis bajas.
- Revisamos al mes y 3 meses como mínimo.
- Subimos a dosis media.
- No corremos para subir a dosis altas.
- Tasa de reingresos, retirada, mortalidad → algo menor que PARADIGM.

CADA MAESTRILLO.....

- Es raro que se retire por fallo renal.
- A veces hay que bajar diuréticos, betabloq. o retirar ARM.
- Escribimos en todos los informes de alta que no se nos ha olvidado el IECA.

MUCHAS GRACIAS

