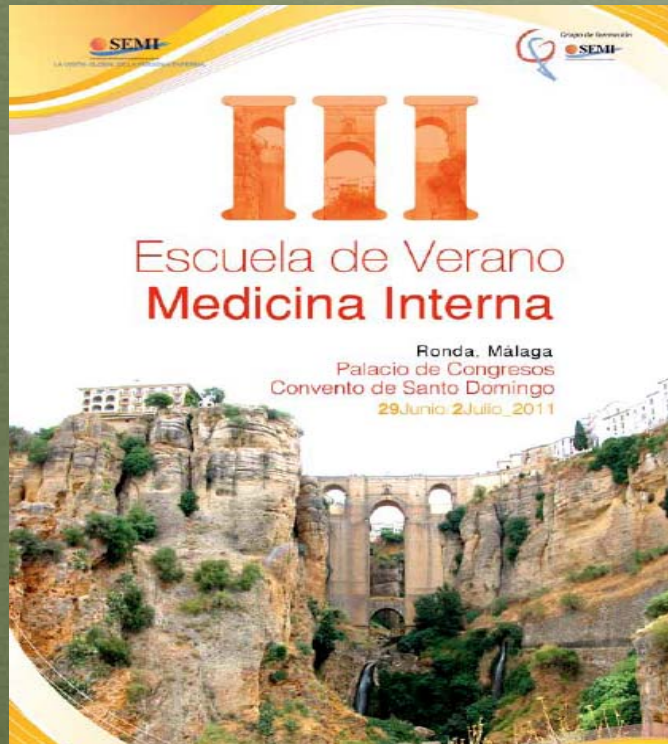


# *¿ Qué hay de nuevo en Insuficiencia Cardiaca ?*



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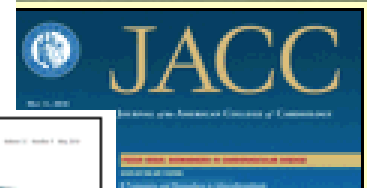
*Lleida*

## Insuficiencia Cardíaca Aguda:

- Diuréticos
- Betabloqueantes

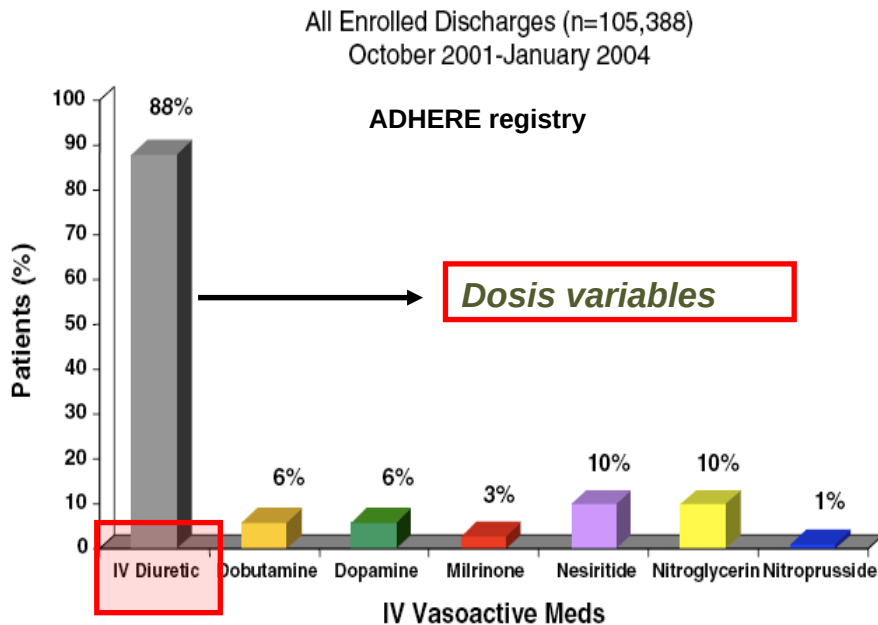
## Insuficiencia Cardíaca Crónica:

- Hierro intravenoso
- Antialdosterónicos
- Ivabradina
- Revascularización



# Tratamiento diurético

# Diuretic Optimization Strategies Evaluation in Acute Heart Failure (DOSE)



[Intervention Review]

## Continuous infusion versus bolus injection of loop diuretics in congestive heart failure

Existen 8 ensayos con 254 pacientes

### *Infusión continua:*

- mayor efecto diurético
- menos efectos secundarios
- menor estancia hospitalaria
- reducción de mortalidad global pero no cardiovascular



No hay recomendaciones definitivas

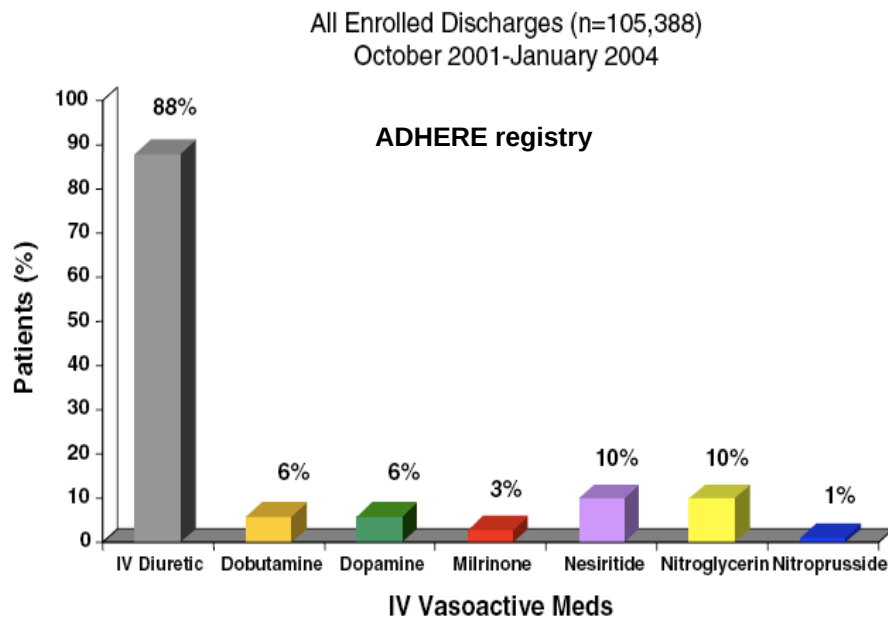
Heart Fail Rev (2007) 12:125–130

Cochrane Database Syst Rev 2009  
1:CD003178

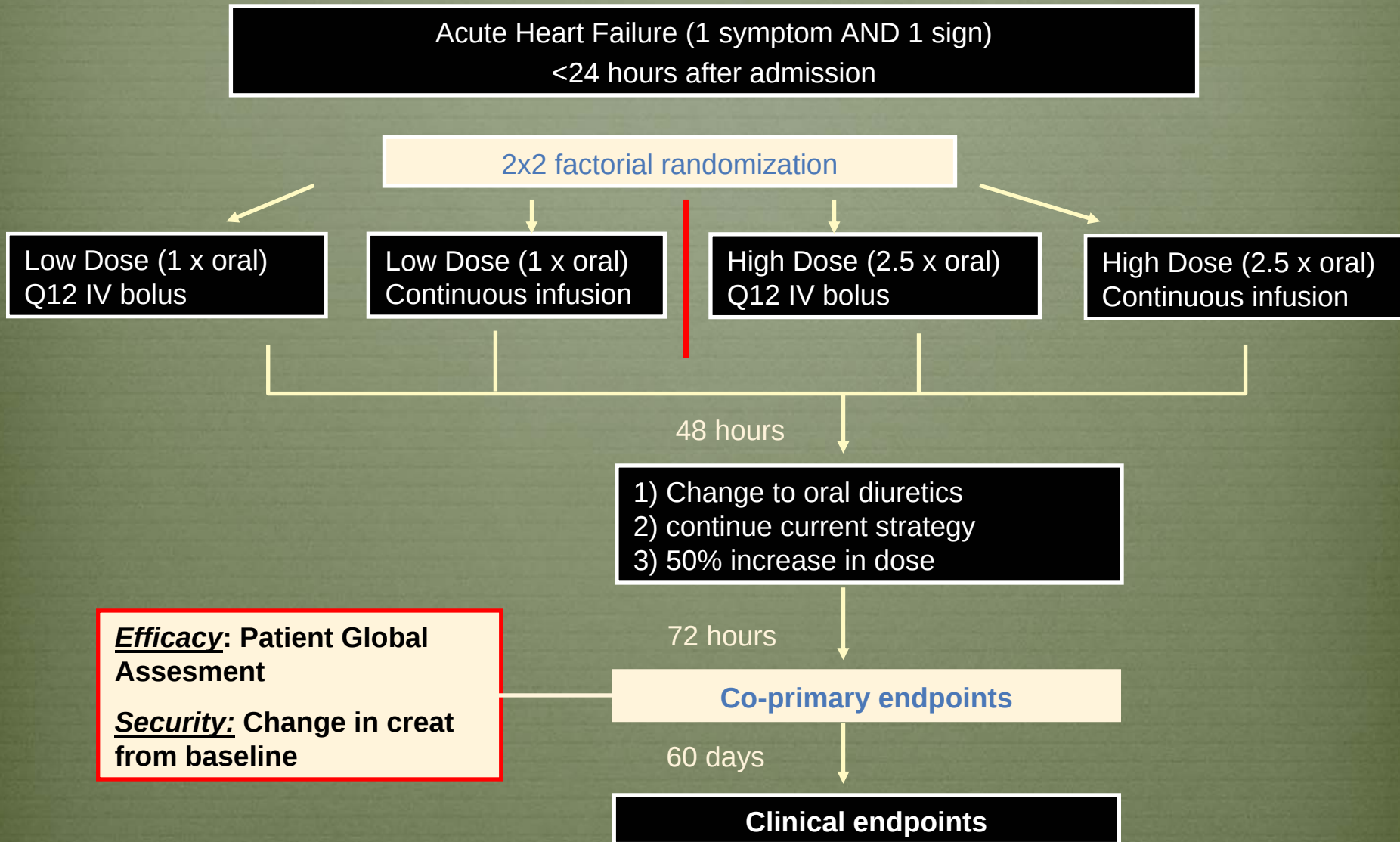
# Diuretic Strategies in Patients with Acute Decompensated Heart Failure

## BACKGROUND

Loop diuretics are an essential component of therapy for patients with acute decompensated heart failure, but there are few prospective data to guide their use.



- 308 pacientes con FEVI media 35% en tratamiento con furosemida ( entre 80 y 240 mg/día como dosis basal ) que ingresan por agudización IC
- Se excluyen tto inotrope, vasodilatador o ultrafiltración, creat > 3 mg/dl.
- **Objetivos:** Comparar de dosis de furosemida y via de administración en términos de eficacia y seguridad

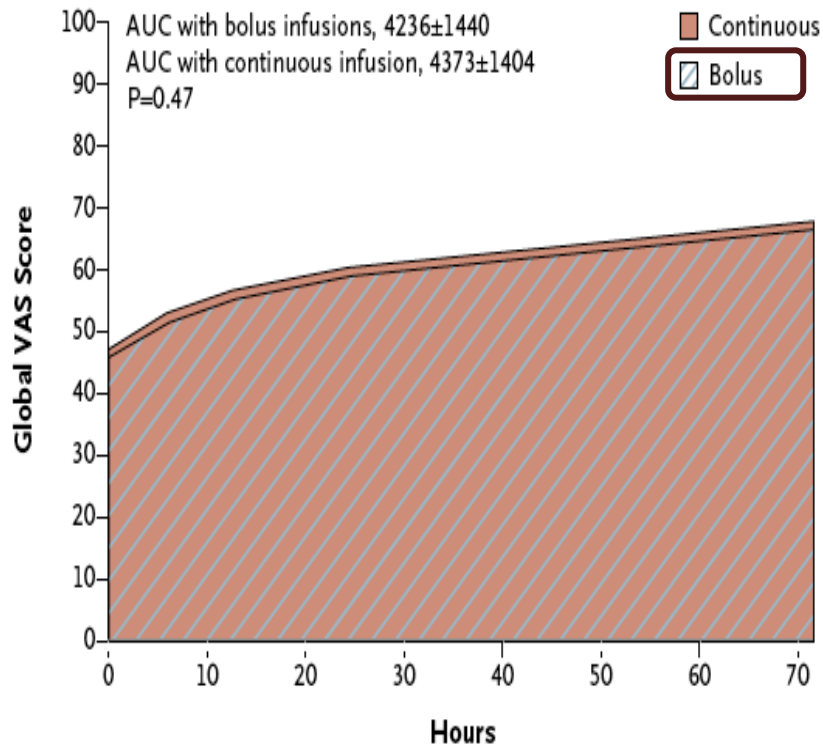


## Diuretic Strategies in Patients with Acute Decompensated Heart Failure

| Characteristic                                               | Bolus Every 12 Hr<br>(N = 156) | Continuous Infusion<br>(N = 152) | Low Dose<br>(N = 151) | High Dose<br>(N = 157) |
|--------------------------------------------------------------|--------------------------------|----------------------------------|-----------------------|------------------------|
| Age — yr                                                     | 66.2±13.2                      | 65.8±14.1                        | 65.9±13.3             | 66.2±13.9              |
| Male sex — no. (%)                                           | 115 (74)                       | 111 (73)                         | 110 (73)              | 116 (74)               |
| Dose of oral furosemide or furosemide equivalent<br>— mg/day | 134±53                         | 127±50                           | 131±52                | 131±51                 |
| Ejection fraction (%)                                        | 35±18                          | 35±18                            | 33±17                 | 36±18                  |
| <b>Creatinine — mg/dl</b>                                    | <b>1.5±0.5</b>                 | <b>1.5±0.5</b>                   | <b>1.5±0.5</b>        | <b>1.5±0.5</b>         |
| Systolic blood pressure — mm Hg                              | 118±19                         | 121±22                           | 120±19                | 119±21                 |
| Heart rate — beats/min                                       | 76±14                          | 80±17                            | 78±15                 | 79±17                  |
| ACE inhibitor or ARB — no. (%)                               | 104 (67)                       | 93 (61)                          | 94 (62)               | 103 (66)               |
| Beta-blocker — no. (%)                                       | 133 (85)                       | 123 (81)                         | 125 (83)              | 131 (83)               |
| Aldosterone antagonist — no. (%)                             | 42 (27)                        | 44 (29)                          | 43 (28)               | 43 (27)                |

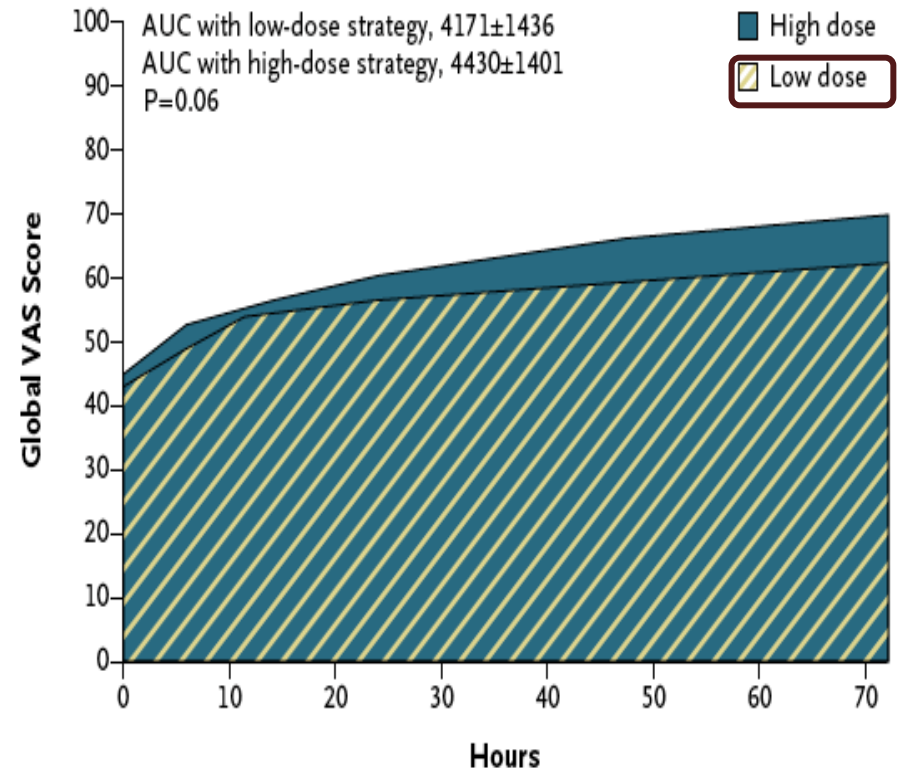
## Efficacy: Patient Global Assessment

**A Bolus vs. Continuous Infusion**



A las 48 horas el grupo Bolus precisó mayor incremento de dosis ( 21 vs 11% -  $p= 0.01$  )

**B Low-Dose vs. High-Dose Strategy**

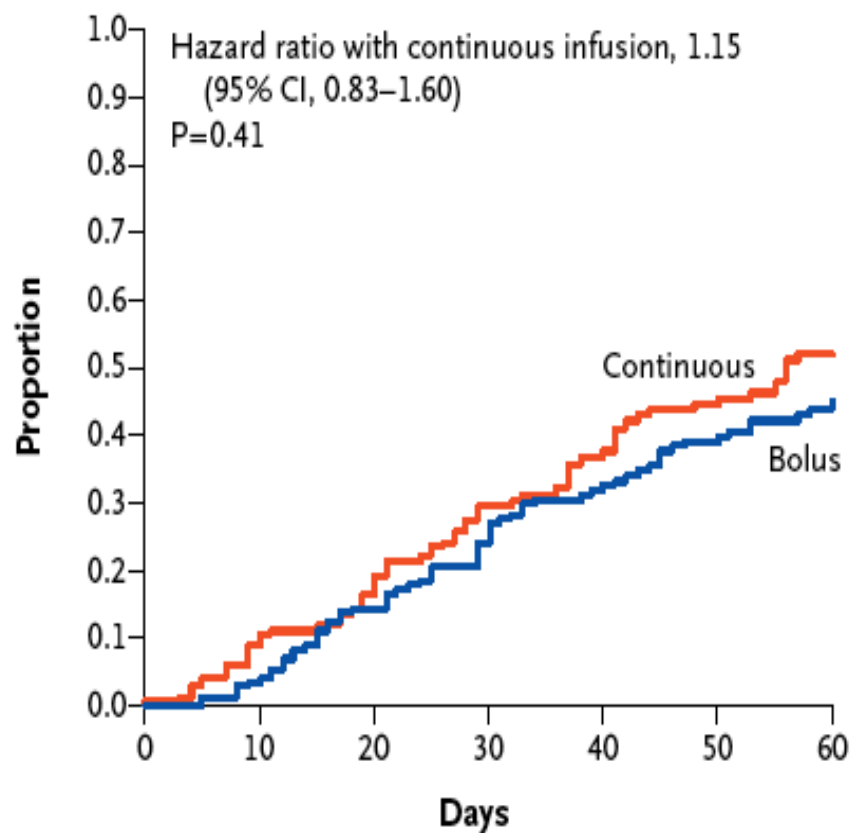


A las 48 horas el grupo dosis-bajas precisó duplicar la dosis con mayor frecuencia. ( 24 vs 9% -  $p= 0.01$  )

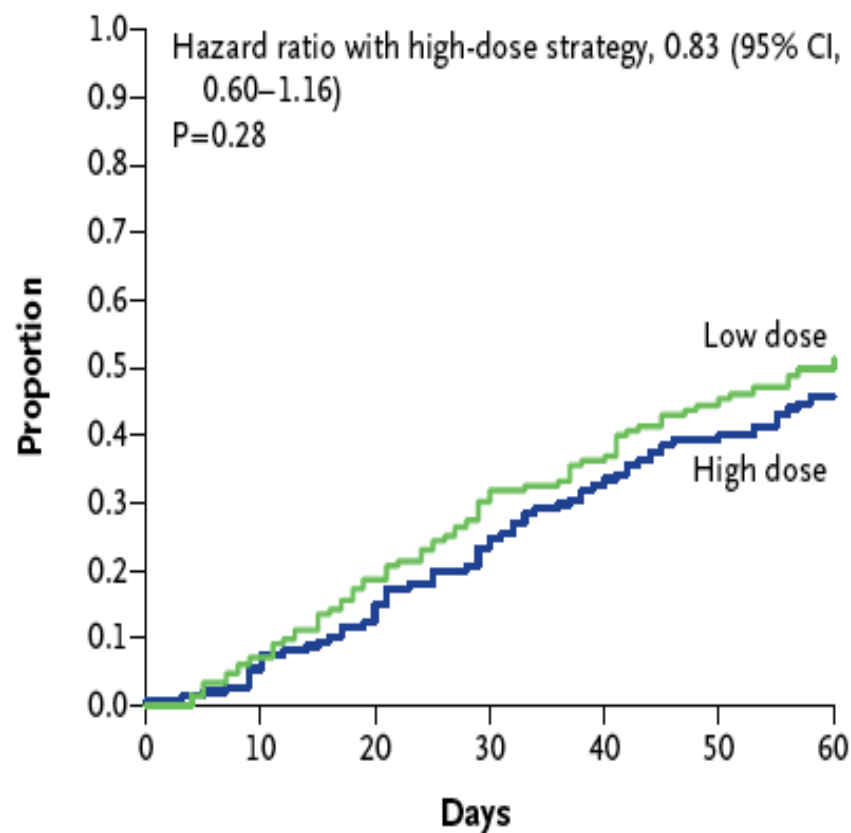
**Table 2. Secondary End Points for Each Treatment Comparison.\***

| End Point                                                                | Bolus Every 12 Hr<br>(N=156) | Continuous Infusion<br>(N=152) | P Value | Low Dose<br>(N=151) | High Dose<br>(N=157) | P Value |
|--------------------------------------------------------------------------|------------------------------|--------------------------------|---------|---------------------|----------------------|---------|
| AUC for dyspnea at 72 hr                                                 | 4456±1468                    | 4699±1573                      | 0.36    | 4478±1550           | 4668±1496            | 0.04    |
| Freedom from congestion at 72 hr —<br>no./total no. (%)                  | 22/153 (14)                  | 22/144 (15)                    | 0.78    | 16/143 (11)         | 28/154 (18)          | 0.09    |
| Change in weight at 72 hr — lb                                           | −6.8±7.8                     | −8.1±10.3                      | 0.20    | −6.1±9.5            | −8.7±8.5             | 0.01    |
| Net fluid loss at 72 hr — ml                                             | 4237±3208                    | 4249±3104                      | 0.89    | 3575±2635           | 4899±3479            | 0.001   |
| Change in NT-proBNP at 72 hr —<br>pg/ml                                  | −1316±4364                   | −1773±3828                     | 0.44    | −1194±4094          | −1882±4105           | 0.06    |
| Worsening or persistent heart failure<br>— no./total no. (%)             | 38/154 (25)                  | 34/145 (23)                    | 0.78    | 38/145 (26)         | 34/154 (22)          | 0.40    |
| Treatment failure — no./total no. (%)†                                   | 59/155 (38)                  | 57/147 (39)                    | 0.88    | 54/147 (37)         | 62/155 (40)          | 0.56    |
| Increase in creatinine of >0.3 mg/dl<br>within 72 hr — no./total no. (%) | 27/155 (17)                  | 28/146 (19)                    | 0.64    | 20/147 (14)         | 35/154 (23)          | 0.04    |
| Length of stay in hospital — days                                        | *                            |                                | 0.97    |                     |                      | 0.55    |
| Median                                                                   | 5                            | 5                              |         | 6                   | 5                    |         |
| Interquartile range                                                      | 3–9                          | 3–8                            |         | 4–9                 | 3–8                  |         |
| Alive and out of hospital — days                                         | *                            |                                | 0.36    |                     |                      | 0.42    |
| Median                                                                   | 51                           | 51                             |         | 50                  | 52                   |         |
| Interquartile range                                                      | 42–55                        | 38–55                          |         | 39–54               | 42–56                |         |

### A Bolus vs. Continuous Infusion



### B Low-Dose vs. High-Dose Strategy



**Figure 3.** Kaplan–Meier Curves for the Clinical Composite End Point of Death, Rehospitalization, or Emergency Department Visit.

# Diuretic Strategies in Patients with Acute Decompensated Heart Failure

¿Cómo ha afectado el poder cambiar el tratamiento a las 48 horas del inicio – sesgo?

Perfusión continua y dosis elevadas precisan menores ajustes de dosis a las 48 h

## Conclusiones:

- Cuando se comparan dosis altas vs bajas de furosemida:
  - \* **No hay diferencias significativas** en el alivio de los síntomas
  - \* **No hay diferencias significativas** en cuanto a cambios en la función renal
- Cuando se comparan bolus vs infusión continua:
  - \* **No hay diferencias significativas** en el alivio de los síntomas
  - \* **No hay diferencias significativas** en cuanto a cambios en la función renal
- Las **dosis elevadas** se asocian a una tendencia a la **mejoría más rápida de la disnea, pérdida de peso y volumen mayor , reducción de NT-proBNP mayor**
- No hay diferencias en cuanto a mortalidad y reingreso a los 60 días con ninguna de las estrategias.

# Betabloqueantes

# B-CONVINCED: Beta-blocker CONTinuation Vs. INTerruption in patients with Congestive heart failure hospitalizED for a decompensation episode

## β-Blockers in worsening heart failure: good or bad?



Eur Heart J 2008;29:2388–2442

'In patients admitted to hospital due to worsening HF, a reduction in the β-blocker dose may be necessary. In severe situations, temporary discontinuation can be considered. Low-dose therapy should be re-instituted and up-titrated as soon as the patient's clinical condition permits, preferably prior to discharge.'

-147 pacientes con FEVI<40% en tratamiento con BB durante al menos un mes previo a ingreso por agudización IC.

- Se randomiza a mantener BB con misma dosis vs retirada.

- ***Estudio de no inferioridad***

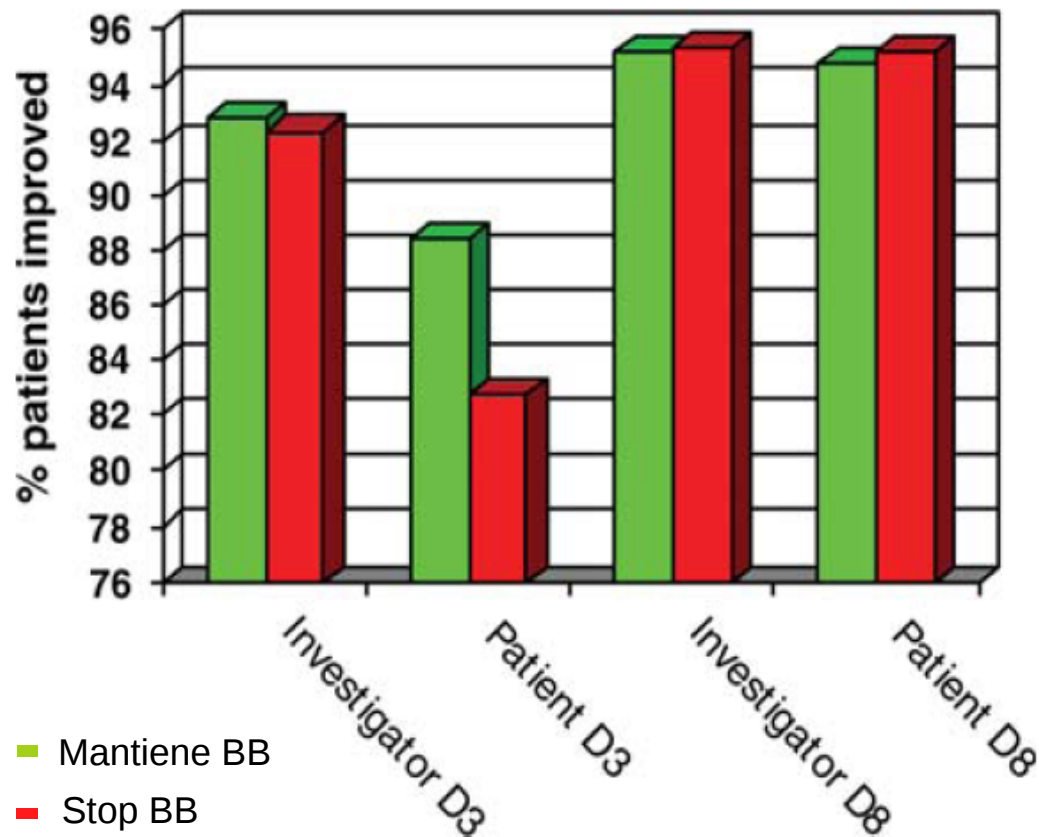
- ***Objetivo principal:*** valorar el efecto sobre el estado general y la mejoría de la disnea valorada por el clínico a los 3 días del ingreso ( Cuestionario )

Eur Heart Journal (2009) 30, 2186–2192

## B-CONVINCED: Beta-blocker CONtinuation Vs. INterruption in patients with Congestive heart failure hospitalizED for a decompensation episode

|                                        | Keep BB, n = 69 | Stop BB, n = 78 | P-value |
|----------------------------------------|-----------------|-----------------|---------|
| Treatment before decompensation        |                 |                 |         |
| BB, n (%)                              |                 |                 |         |
| Max dose                               | 14 (20)         | 21 (27)         | 0.6     |
| 50% ≤ Dose < 100%                      | 18 (26)         | 17 (22)         |         |
| Dose < 50%                             | 37 (54)         | 40 (51)         |         |
| ACE-I, n (%)                           | 45 (65)         | 50 (64)         | 0.89    |
| Diuretics, n (%)                       | 56 (81)         | 66 (84)         | 0.57    |
| Treatment during decompensation, n (%) |                 |                 |         |
| Diuretics                              | 69 (100)        | 77 (99)         | 1.0     |
| Nitrates                               | 35 (51)         | 28 (36)         | 0.11    |

## **B-CONVINCED: Beta-blocker CONTinuation Vs. INTerruption in patients with Congestive heart failure hospitalizED for a decompensation episode**



-No existieron diferencias en la mejoría de la disnea ni del estado general percibida por los investigadores al tercer día de ingreso.

-Tampoco diferencias en los mismos items percibidos por los pacientes al tercer día.

-Tampoco al octavo día.

## B-CONVINCED: Beta-blocker CONTinuation Vs. INTerruption in patients with Congestive heart failure hospitalizED for a decompensation episode

**Table 3** Clinical events

|                            | Keep BB,<br><i>n</i> = 69 | Stop BB,<br><i>n</i> = 78 | <i>P</i> -value |
|----------------------------|---------------------------|---------------------------|-----------------|
| During hospitalization     |                           |                           |                 |
| Durations (days)           | 11.5 ± 8.3                | 10.4 ± 9.7                | 0.2             |
| Median, range              | 9 (1–50)                  | 8 (1–62)                  |                 |
| Deaths ( <i>n</i> )        | 1 (HF)                    | 2 (HF)                    |                 |
| Dobutamine ( <i>n</i> )    | 3                         | 1                         |                 |
| After 3 months             |                           |                           |                 |
| Deaths, <i>n</i> (%)       | 6 (9)                     | 6 (8)                     | 0.83            |
| Rehospit, <i>n</i> (%)     | 27 (40)                   | 36 (47)                   | 0.43            |
| For HF                     | 15 (22)                   | 24 (32)                   | 0.19            |
| For arrhythmia             | 2 (3)                     | 3 (4)                     | 1               |
| Receiving BB, <i>n</i> (%) | 61 (90)                   | 58 (76)                   | 0.04            |

Rehospit, rehospitalization; HF, heart failure; BB, beta-blocker.

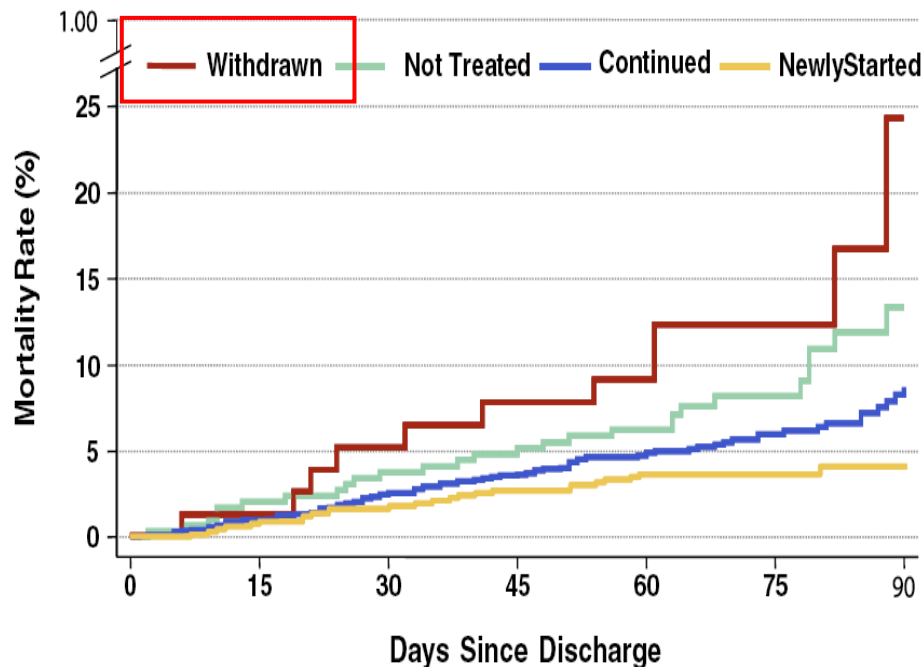
El número de pacientes que recibía BB 3 meses después del ingreso era menor en el grupo al que se le había suspendido el tratamiento.

## B-CONVINCED: Beta-blocker CONTinuation Vs. INTerruption in patients with Congestive heart failure hospitalizED for a decompensation episode

### Conclusiones:

\* Queda establecida la seguridad de continuar el tto BB durante las fases de agudización de la IC ( *en pacientes sin necesidad de dobutamina* )

\* El mantener el tto BB supone un mejor cumplimiento del tto BB a los 3 meses de la agudización



### Influence of Beta-Blocker Continuation or Withdrawal on Outcomes in Patients Hospitalized With Heart Failure

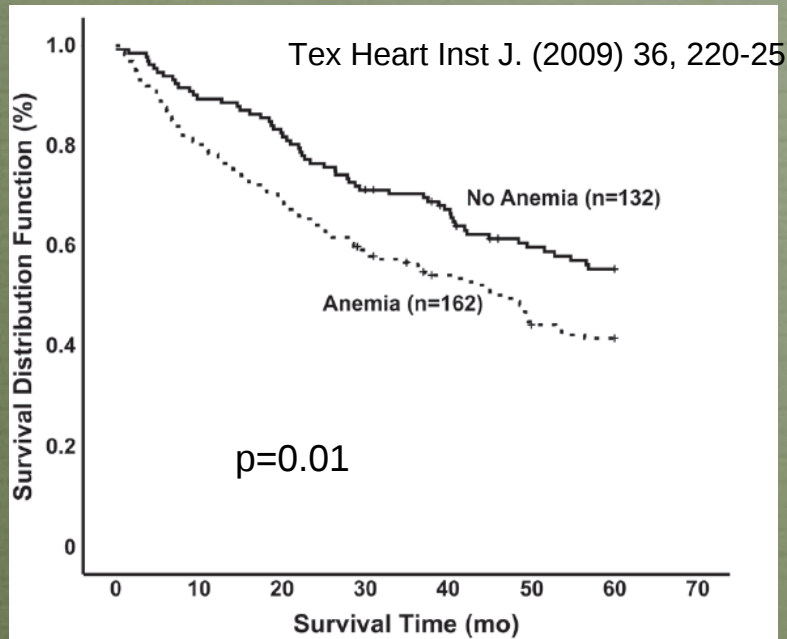
Findings From the OPTIMIZE-HF Program

J Am Coll Cardiol 2008;52:190–9

No diseñado para valorar mortalidad

# Hierro intravenoso

# Ferric Carboxymaltose in Patients with Heart Failure and Iron Deficiency



459 pacientes ICC en clase II-III NYHA + FEVI < 45% + deficit de hierro

Deficit de hierro: ferritina < 100 µg/L o < 300 µg/L e ISAT < 20%

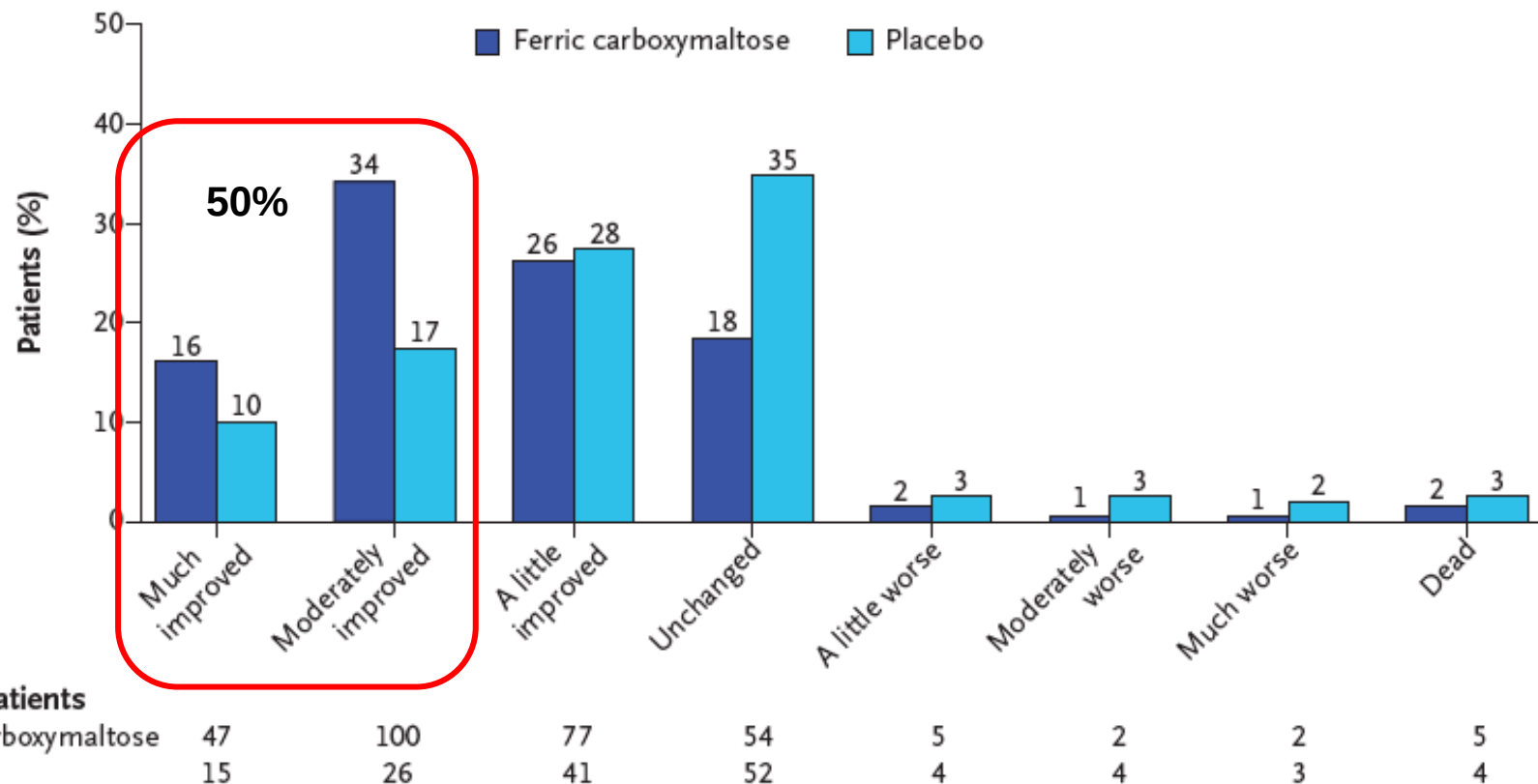
**Objetivo principal:** Patient Global Assessment + NYHA semana 24 .

**Objetivos secundarios:** Walking test + Calidad de vida

**Iron deficiency may impair aerobic performance.** This study aimed to determine whether treatment with intravenous iron (ferric carboxymaltose) would improve symptoms in patients who had heart failure, reduced left ventricular ejection fraction, and iron deficiency, either with or without anemia.

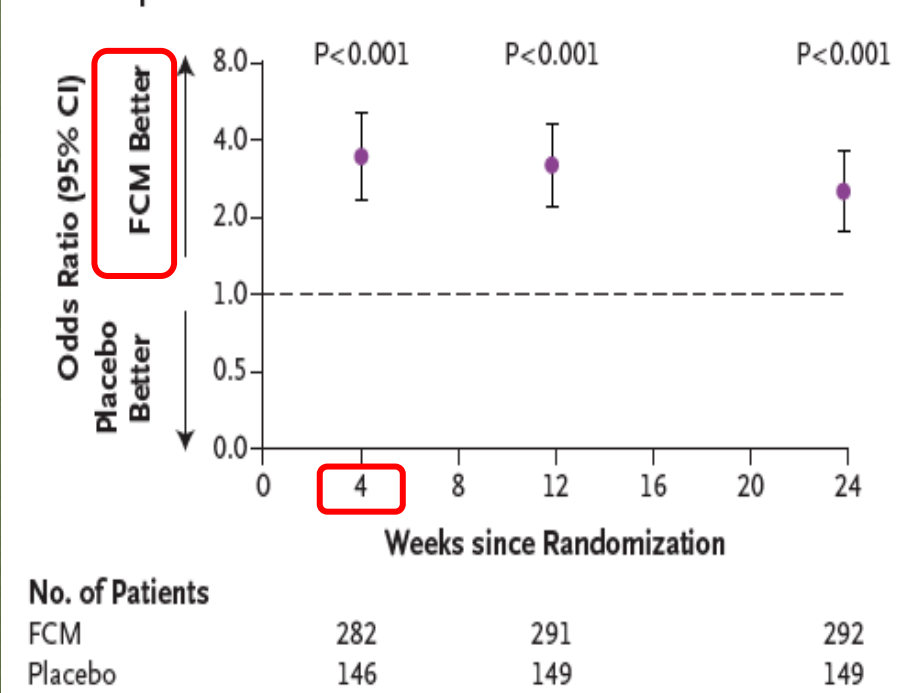
# Ferric Carboxymaltose in Patients with Heart Failure and Iron Deficiency

## A Self-Reported Patient Global Assessment at Wk 24

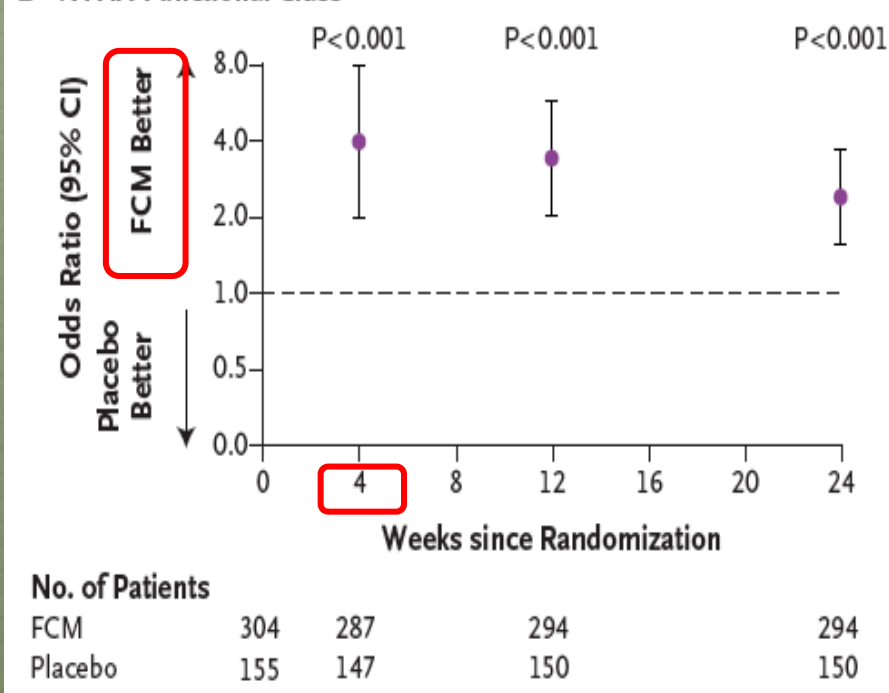


# Ferric Carboxymaltose in Patients with Heart Failure and Iron Deficiency

**A Self-Reported Patient Global Assessment**

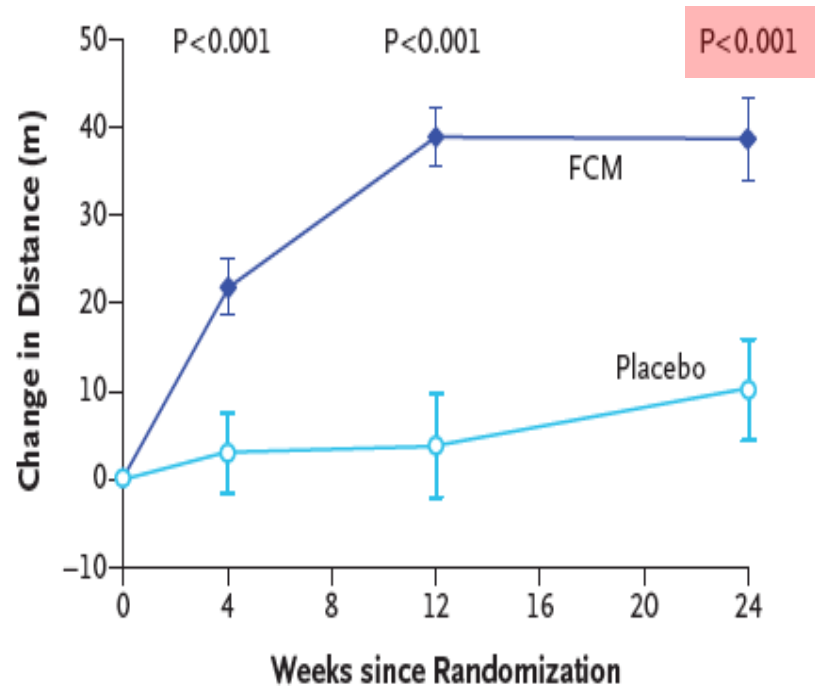


**B NYHA Functional Class**

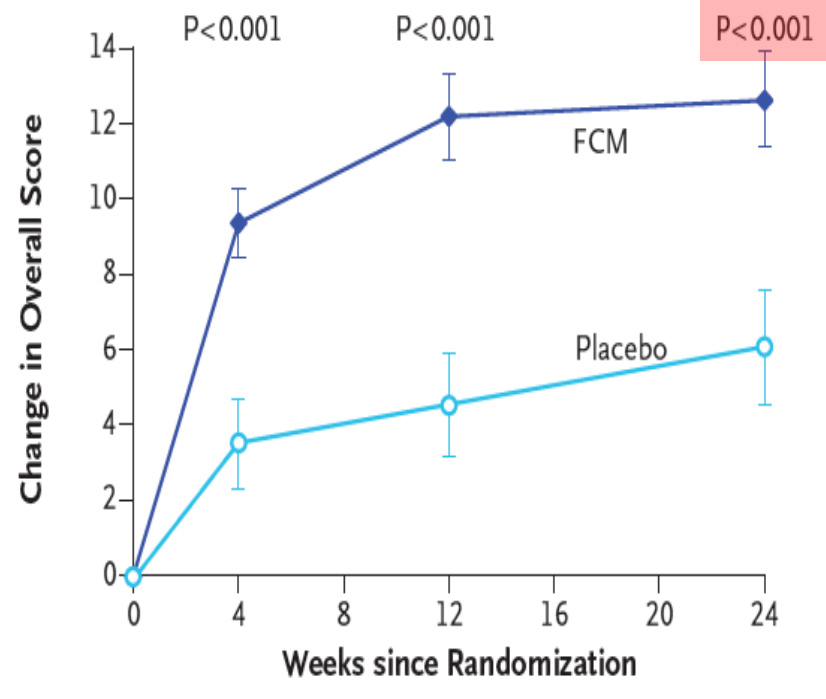


| Subgroup       | Self-Reported Patient Global Assessment   |         |                                                                                   |                            | NYHA Functional Class                     |         |                                                                                     |                            |
|----------------|-------------------------------------------|---------|-----------------------------------------------------------------------------------|----------------------------|-------------------------------------------|---------|-------------------------------------------------------------------------------------|----------------------------|
|                | Ferric Carboxy-maltose<br>no. of patients | Placebo | Odds Ratio<br>(95% CI)                                                            | P Value for<br>Interaction | Ferric Carboxy-maltose<br>no. of patients | Placebo | Odds Ratio<br>(95% CI)                                                              | P Value for<br>Interaction |
| Hemoglobin     |                                           |         |                                                                                   | 0.98                       |                                           |         |                                                                                     | 0.51                       |
| ≤120 (g/liter) | 146                                       | 74      |  |                            | 148                                       | 74      |  |                            |
| >120 (g/liter) | 146                                       | 75      |  |                            | 146                                       | 76      |  |                            |

### C 6-Minute-Walk Test



### E Kansas City Cardiomyopathy Questionnaire



# Ferric Carboxymaltose in Patients with Heart Failure and Iron Deficiency

## Conclusiones:

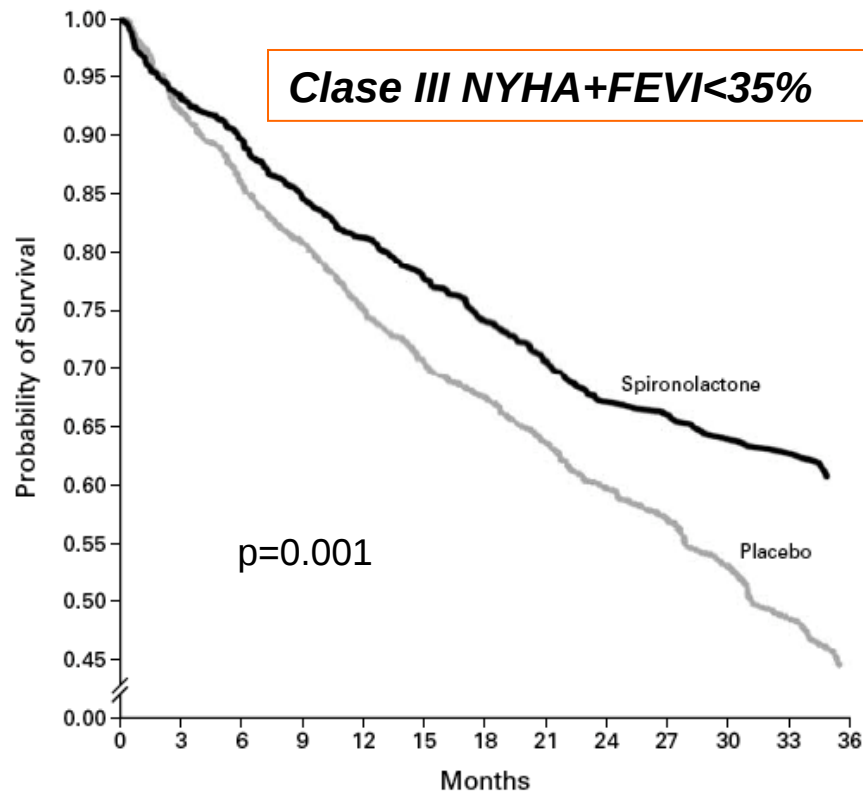
- El tratamiento con hierro intravenoso consigue mejoría de forma global y en la clase funcional.
- Mejora la capacidad funcional medida mediante walking test.
- Mejora la autopercepción de la calidad de vida. .
- *Sus efectos beneficiosos se observan en aquellos pacientes con ferropenia pero sin anemia.*
- No efectos sobre mortalidad o reingresos

**Tratamiento con hierro intravenoso puede ser un nuevo objetivo terapeutico en pacientes con ferropenia y FEVI deprimida**

# **Antialdosterónicos**

# THE EFFECT OF SPIRONOLACTONE ON MORBIDITY AND MORTALITY IN PATIENTS WITH SEVERE HEART FAILURE

## RANDOMIZED ALDACTONE EVALUATION STUDY



| CHARACTERISTIC        | PLACEBO GROUP<br>(N=841) | SPIRONOLACTONE<br>GROUP (N=822) |
|-----------------------|--------------------------|---------------------------------|
| Medications — %       |                          |                                 |
| Loop diuretics        | 100                      | 100                             |
| ACE inhibitors        | 94                       | 95                              |
| Digitalis             | 72                       | 75                              |
| Aspirin               | 37                       | 36                              |
| Potassium supplements | 27                       | 29                              |
| Beta-blockers         | 10                       | 11                              |

| PLACEBO GROUP<br>(N=841) | SPIRONOLACTONE<br>GROUP (N=822) |
|--------------------------|---------------------------------|
|--------------------------|---------------------------------|

|     |     |
|-----|-----|
| 100 | 100 |
| 94  | 95  |
| 72  | 75  |
| 37  | 36  |
| 27  | 29  |
| 10  | 11  |

N Engl J Med 1999;341:709-17.

# Eplerenone in Patients with Systolic Heart Failure and Mild Symptoms

EMPHASIS-HF Study

- 2737 pacientes con FEVI < 35% en clase II NYHA estables durante 6 meses y con tto óptimo.

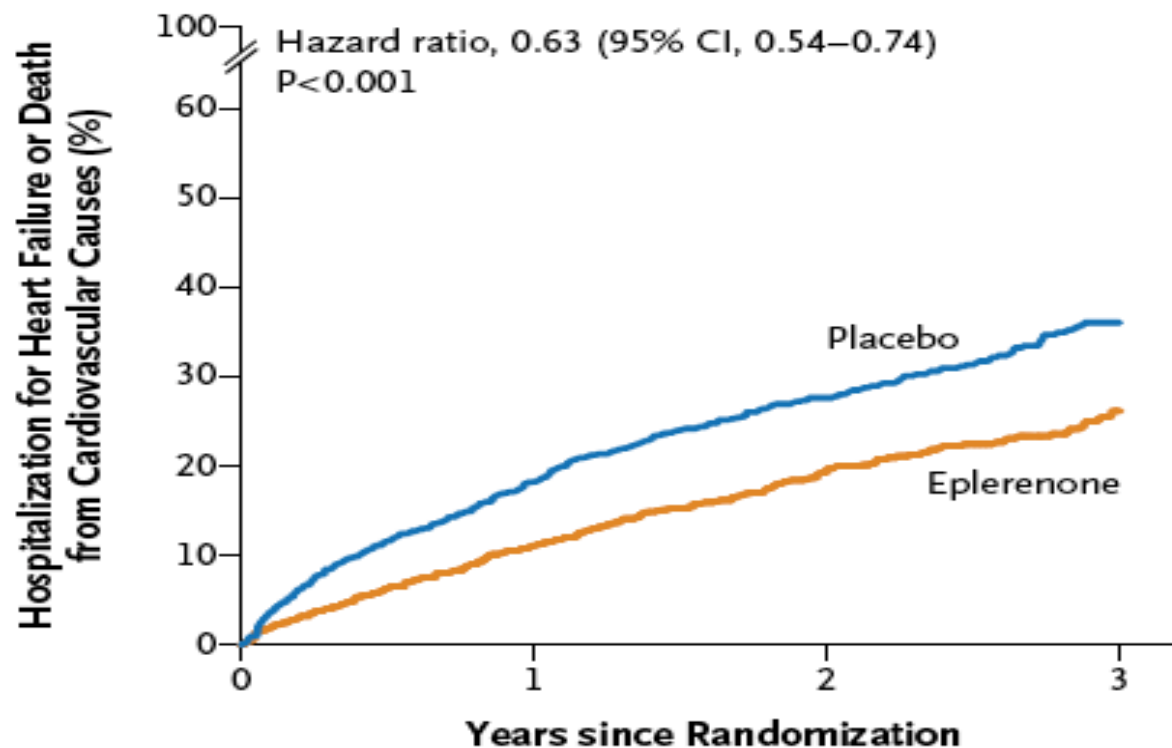
-Se excluyen hiperpotasemia y FG < 30 ml/min.

-**Objetivos:** Mortalidad cardiovascular o reingreso por insuficiencia cardíaca

| Characteristic                                           | Eplerenone<br>(N= 1364) | Placebo<br>(N= 1373) |
|----------------------------------------------------------|-------------------------|----------------------|
| Medication at randomization visit — no. (%)              |                         |                      |
| Diuretic                                                 | 1150 (84.3)             | 1176 (85.7)          |
| ACE inhibitor                                            | 1068 (78.3)             | 1055 (76.8)          |
| ARB                                                      | 261 (19.1)              | 266 (19.4)           |
| ACE inhibitor, ARB, or both                              | 1282 (94.0)             | 1275 (92.9)          |
| Beta-blocker                                             | 1181 (86.6)             | 1193 (86.9)          |
| Digitalis glycosides                                     | 363 (26.6)              | 377 (27.5)           |
| Antiarrhythmic drug                                      | 196 (14.4)              | 192 (14.0)           |
| Antithrombotic drug (antiplatelet or oral anticoagulant) | 1205 (88.3)             | 1214 (88.4)          |
| Lipid-lowering agent                                     | 857 (62.8)              | 856 (62.3)           |

# Eplerenone in Patients with Systolic Heart Failure and Mild Symptoms

**A**



**No. at Risk**

|            |      |     |     |     |
|------------|------|-----|-----|-----|
| Placebo    | 1373 | 848 | 512 | 199 |
| Eplerenone | 1364 | 925 | 562 | 232 |

# Eplerenone in Patients with Systolic Heart Failure and Mild Symptoms

| Outcome                                                       | Eplerenone<br>(N = 1364) | Placebo<br>(N = 1373) | Adjusted<br>Hazard Ratio<br>(95% CI) | Adjusted<br>P Value | Unadjusted<br>Hazard Ratio<br>(95% CI) | Unadjusted<br>P Value |
|---------------------------------------------------------------|--------------------------|-----------------------|--------------------------------------|---------------------|----------------------------------------|-----------------------|
| Prespecified adjudicated secondary outcomes                   |                          |                       |                                      |                     |                                        |                       |
| Death from any cause or hospitalization for heart failure     | 270 (19.8)               | 376 (27.4)            | 0.65 (0.55–0.76)                     | <0.001              | 0.68 (0.58–0.79)                       | <0.001                |
| Death from any cause                                          | 171 (12.5)               | 213 (15.5)            | 0.76 (0.62–0.93)                     | 0.008               | 0.78 (0.64–0.95)                       | 0.01                  |
| Death from cardiovascular causes                              | 147 (10.8)               | 185 (13.5)            | 0.76 (0.61–0.94)                     | 0.01                | 0.77 (0.62–0.96)                       | 0.02                  |
| Hospitalization for any reason                                | 408 (29.9)               | 491 (35.8)            | 0.77 (0.67–0.88)                     | <0.001              | 0.78 (0.69–0.89)                       | <0.001                |
| Hospitalization for heart failure                             | 164 (12.0)               | 253 (18.4)            | 0.58 (0.47–0.70)                     | <0.001              | 0.61 (0.50–0.75)                       | <0.001                |
| Hospitalization for cardiovascular causes                     | 304 (22.3)               | 399 (29.1)            | 0.69 (0.60–0.81)                     | <0.001              | 0.72 (0.62–0.83)                       | <0.001                |
| Fatal or nonfatal myocardial infarction                       | 45 (3.3)                 | 33 (2.4)              | 1.32 (0.84–2.06)                     | 0.23                | 1.34 (0.86–2.10)                       | 0.20                  |
| Death from any cause or hospitalization for any reason        | 462 (33.9)               | 569 (41.4)            | 0.75 (0.66–0.85)                     | <0.001              | 0.76 (0.68–0.86)                       | <0.001                |
| Death from heart failure or hospitalization for heart failure | 170 (12.5)               | 262 (19.1)            | 0.58 (0.48–0.70)                     | <0.001              | 0.61 (0.51–0.74)                       | <0.001                |

# Eplerenone in Patients with Systolic Heart Failure and Mild Symptoms

## Conclusiones:

- El tratamiento con eplerenona disminuye la mortalidad cardiovascular y los reingresos por insuficiencia cardiaca
- Se asocia a mayor incidencia de hiperpotasemia.

## Retrospectivo de 144 ingresos con criterios

Table II. Use of heart failure medications by different age groups before and after admission

| Age (y) | Furosemide (loop diuretics)   |                                   | ACE Inhibitors                |                                   | BB                            |                                   | AA                            |                                   |
|---------|-------------------------------|-----------------------------------|-------------------------------|-----------------------------------|-------------------------------|-----------------------------------|-------------------------------|-----------------------------------|
|         | home care<br>[no. of pts (%)] | hospital care<br>[no. of pts (%)] | home care<br>[no. of pts (%)] | hospital care<br>[no. of pts (%)] | home care<br>[no. of pts (%)] | hospital care<br>[no. of pts (%)] | home care<br>[no. of pts (%)] | hospital care<br>[no. of pts (%)] |
| 21-30   | 5 (100)                       | 5 (100)                           | 5 (100)                       | 5 (100)                           | 4 (80)                        | 3 (60)                            | 4 (80)                        | 5 (10)                            |
| 31-40   | 13 (81)                       | 16 (100)                          | 13 (81)                       | 16 (100)                          | 14 (88)                       | 12 (75)                           | 9 (56)                        | 14 (88)                           |
| 41-50   | 26 (96)                       | 26 (96)                           | 23 (85)                       | 26 (96)                           | 17 (63)                       | 19 (70)                           | 14 (52)                       | 16 (59)                           |
| 51-60   | 38 (83)                       | 45 (98)                           | 34 (74)                       | 41 (89)                           | 30 (65)                       | 31 (67)                           | 17 (37)                       | 27 (59)                           |
| 61-70   | 15 (75)                       | 18 (90)                           | 12 (60)                       | 19 (95)                           | 12 (60)                       | 14 (70)                           | 4 (20)                        | 12 (60)                           |
| 71-80   | 14 (78)                       | 18 (100)                          | 11 (61)                       | 15 (83)                           | 14 (78)                       | 13 (72)                           | 6 (33)                        | 8 (44)                            |
| 81-90   | 7 (78)                        | 9 (100)                           | 7 (78)                        | 8 (89)                            | 4 (44)                        | 4 (44)                            | 2 (22)                        | 2 (22)                            |
| 91-100  | 2 (67)                        | 3 (100)                           | 2 (67)                        | 3 (100)                           | 1 (33)                        | 1 (33)                            | 0 (0)                         | 0 (0)                             |
| Total   | 120 (83)                      | 140 (97)                          | 107 (74)                      | 133 (92)                          | 96 (67)                       | 97 (67)                           | 39%<br>84 (58)                | 58%<br>84 (58)                    |

AA = aldosterone antagonists; BB =  $\beta$ -adrenoceptor antagonists ( $\beta$ -blockers); pts = patients.

¿ Existen diferencias entre espironolactona y eplerenona?

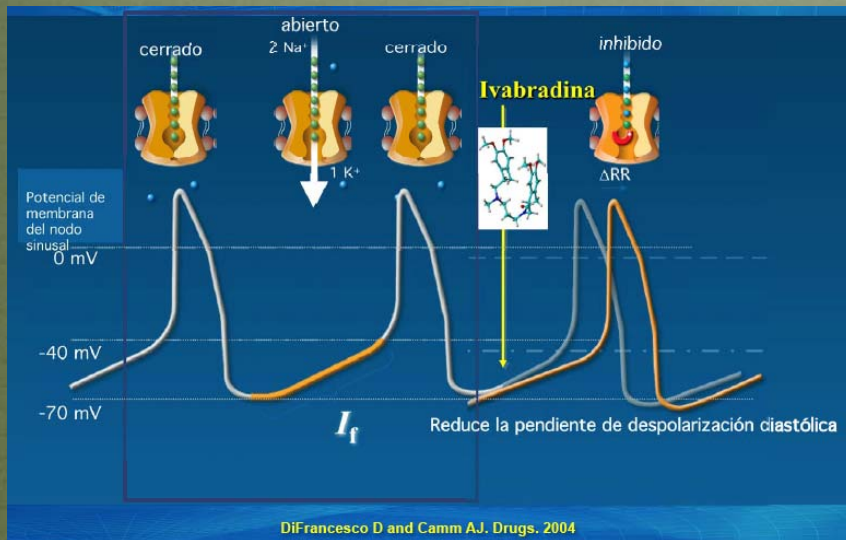
Guglin M et al Am J Cardiovasc Drugs 2007; 7 (1): 75-79

N Engl J Med 2011;364:11-21.

# **Frecuencia Cardiaca -** ***Ivabradina***

# Ivabradine and outcomes in chronic heart failure (SHIFT): a randomised placebo-controlled study

**Background** Chronic heart failure is associated with high mortality and morbidity. Raised resting heart rate is a risk factor for adverse outcomes. We aimed to assess the effect of heart-rate reduction by the selective sinus-node inhibitor ivabradine on outcomes in heart failure.



6558 pacientes ICC estable + FEVI < 35%  
+ tto óptimo

Ritmo sinusal > 70 x'

Ivabradina vs placebo

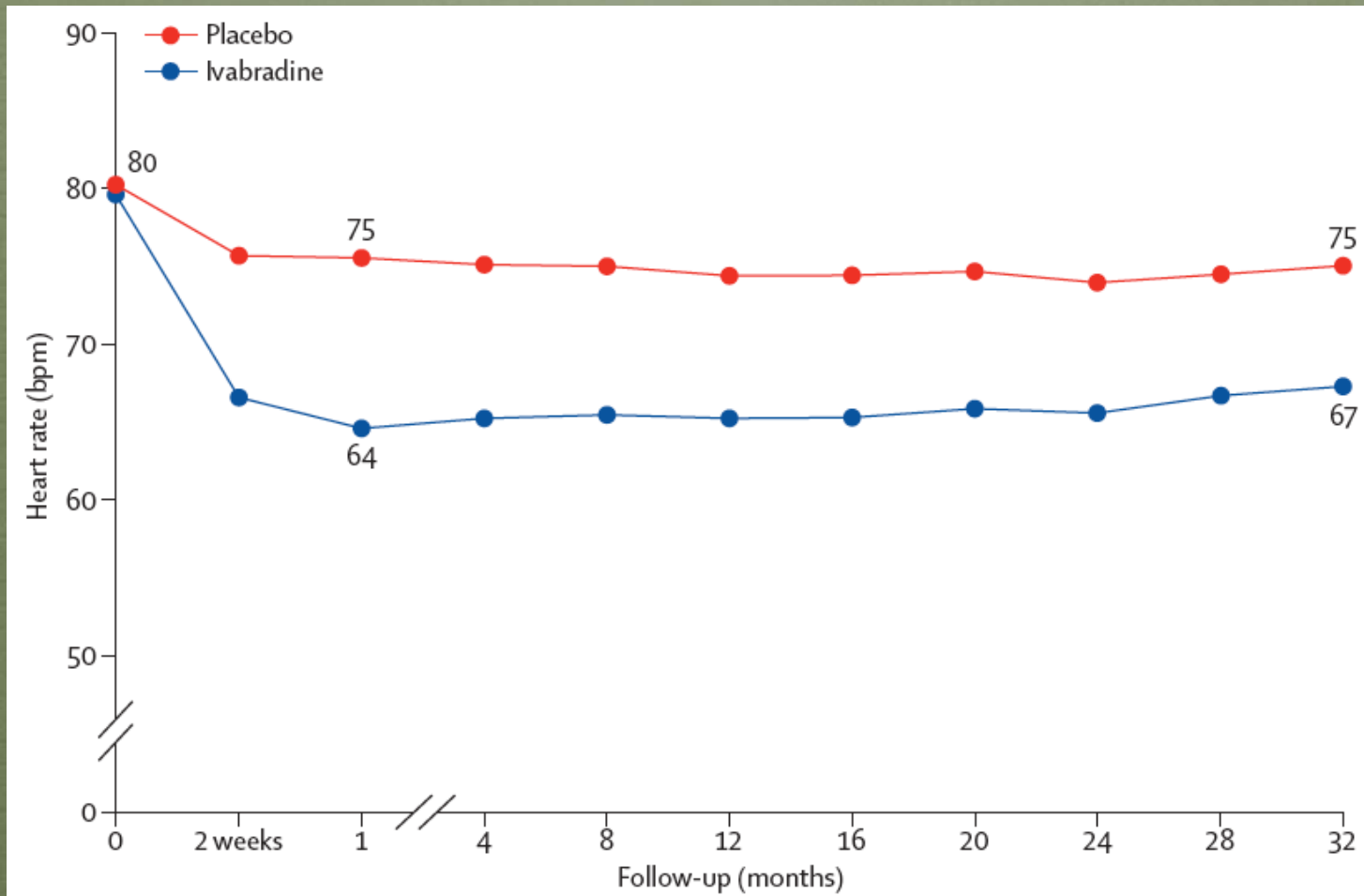
**Objetivo principal:** Combinación de muerte cardiovascular u hospitalización por ICC.

**Objetivos secundarios:** combinado en tratados con >50% dosis objetivo de betabloqueante.

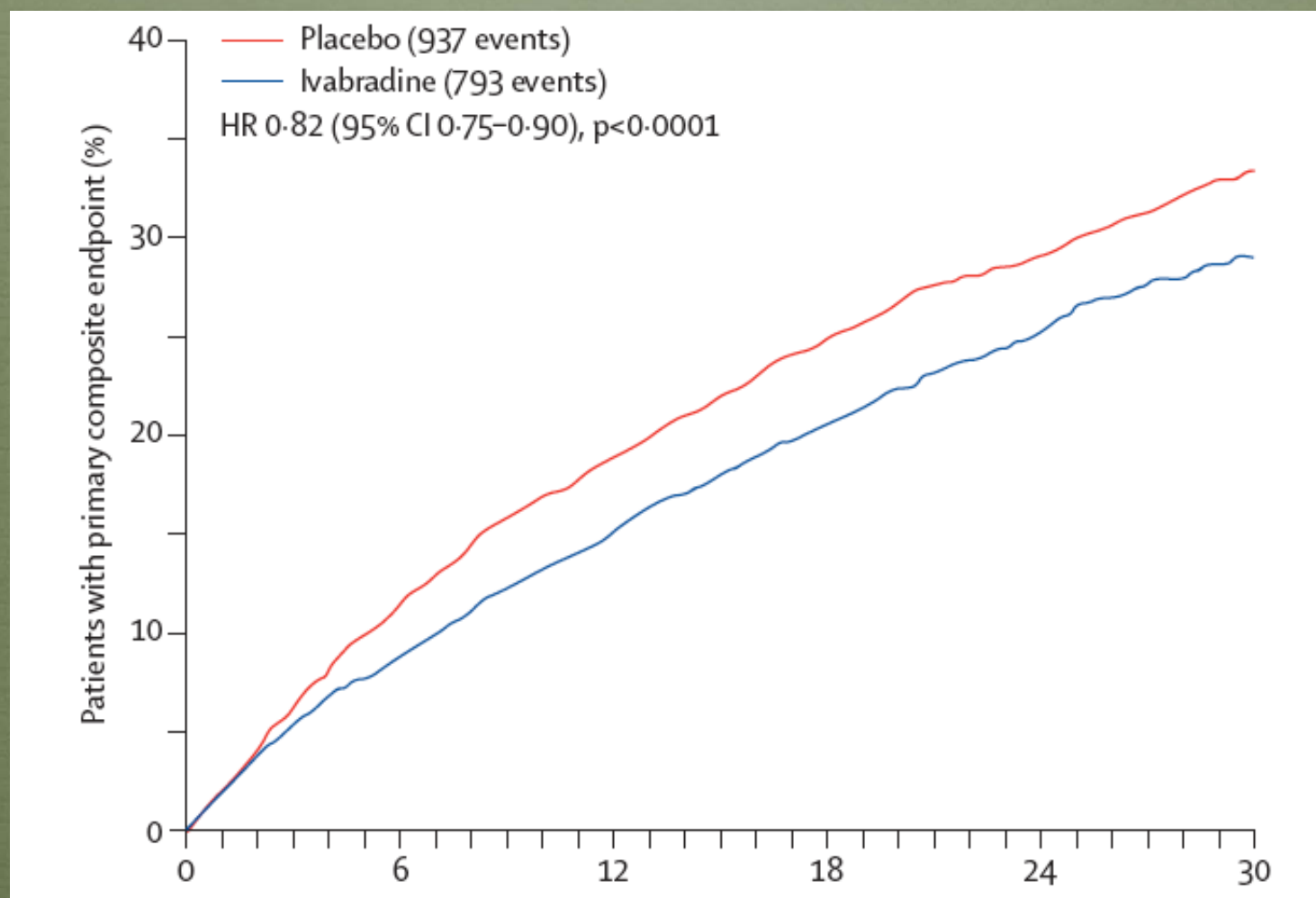
# Ivabradine and outcomes in chronic heart failure (SHIFT): a randomised placebo-controlled study

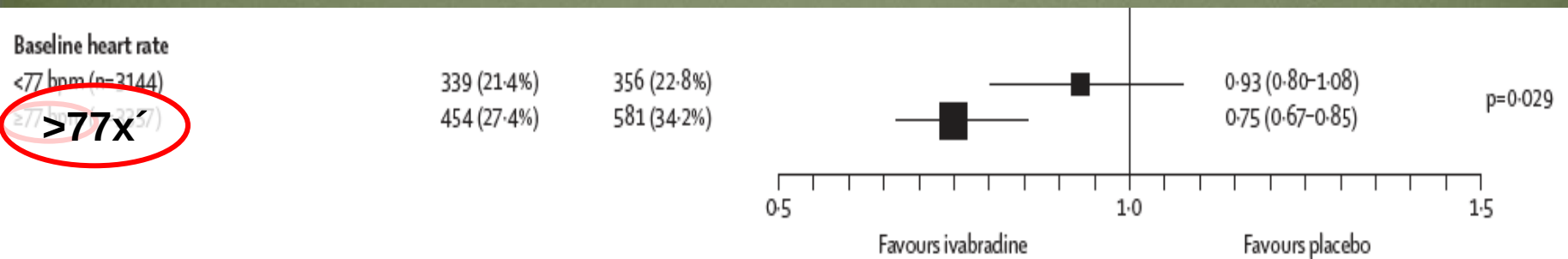
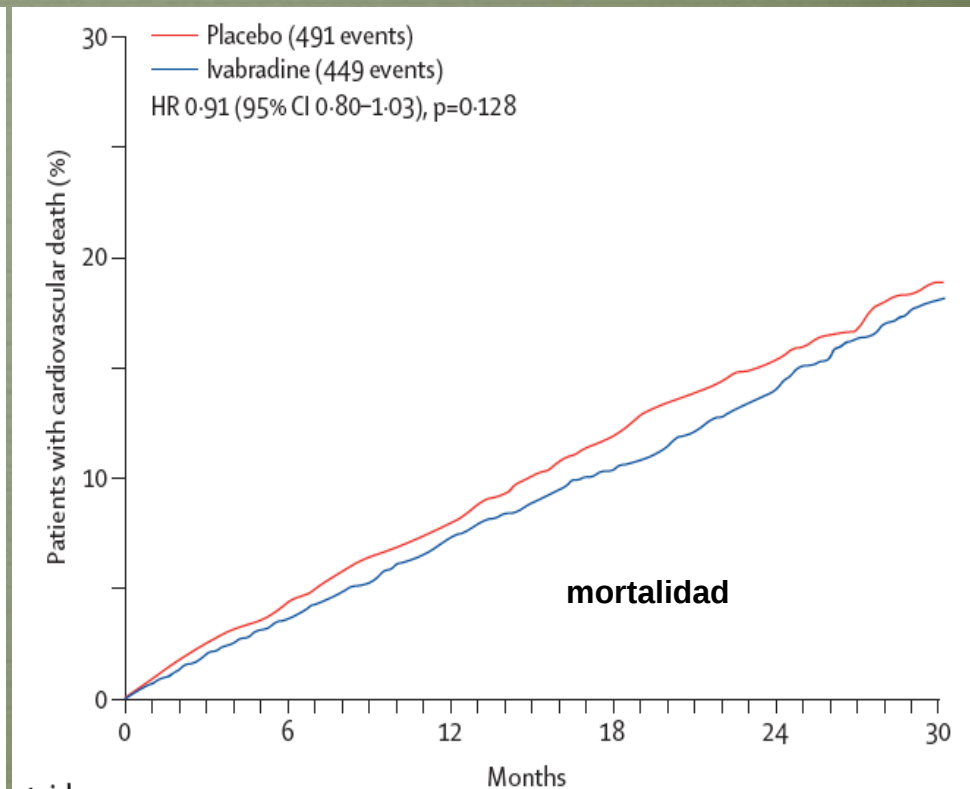
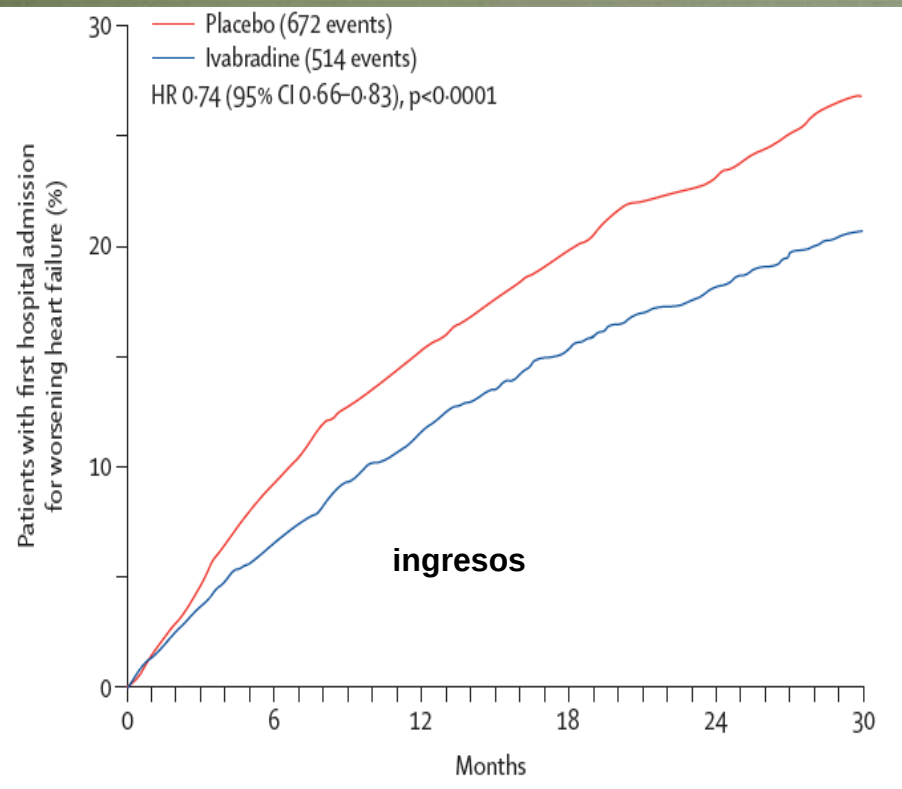
|                                                           | Ivabradine group<br>(n=3241) | Placebo group<br>(n=3264) |
|-----------------------------------------------------------|------------------------------|---------------------------|
| Patients receiving $\beta$ blocker                        | 2897 (89%)                   | 2923 (90%)                |
| Patients at target dose of $\beta$ blocker* ✖             | 743 (26%)                    | 745 (26%)                 |
| Patients at $\geq 50\%$ target dose of $\beta$ blocker* ✖ | 1581 (56%)                   | 1600 (56%)                |
| Reasons for failure to reach target dose*†                |                              |                           |
| Hypotension                                               | 933 (44%)                    | 952 (45%)                 |
| Fatigue                                                   | 676 (32%)                    | 670 (32%)                 |
| Dyspnoea                                                  | 284 (14%)                    | 302 (14%)                 |
| Dizziness                                                 | 267 (13%)                    | 245 (12%)                 |
| Bradycardia                                               | 134 (6%)                     | 125 (6%)                  |
| Other                                                     | 199 (9%)                     | 219 (10%)                 |

# Ivabradine and outcomes in chronic heart failure (SHIFT): a randomised placebo-controlled study

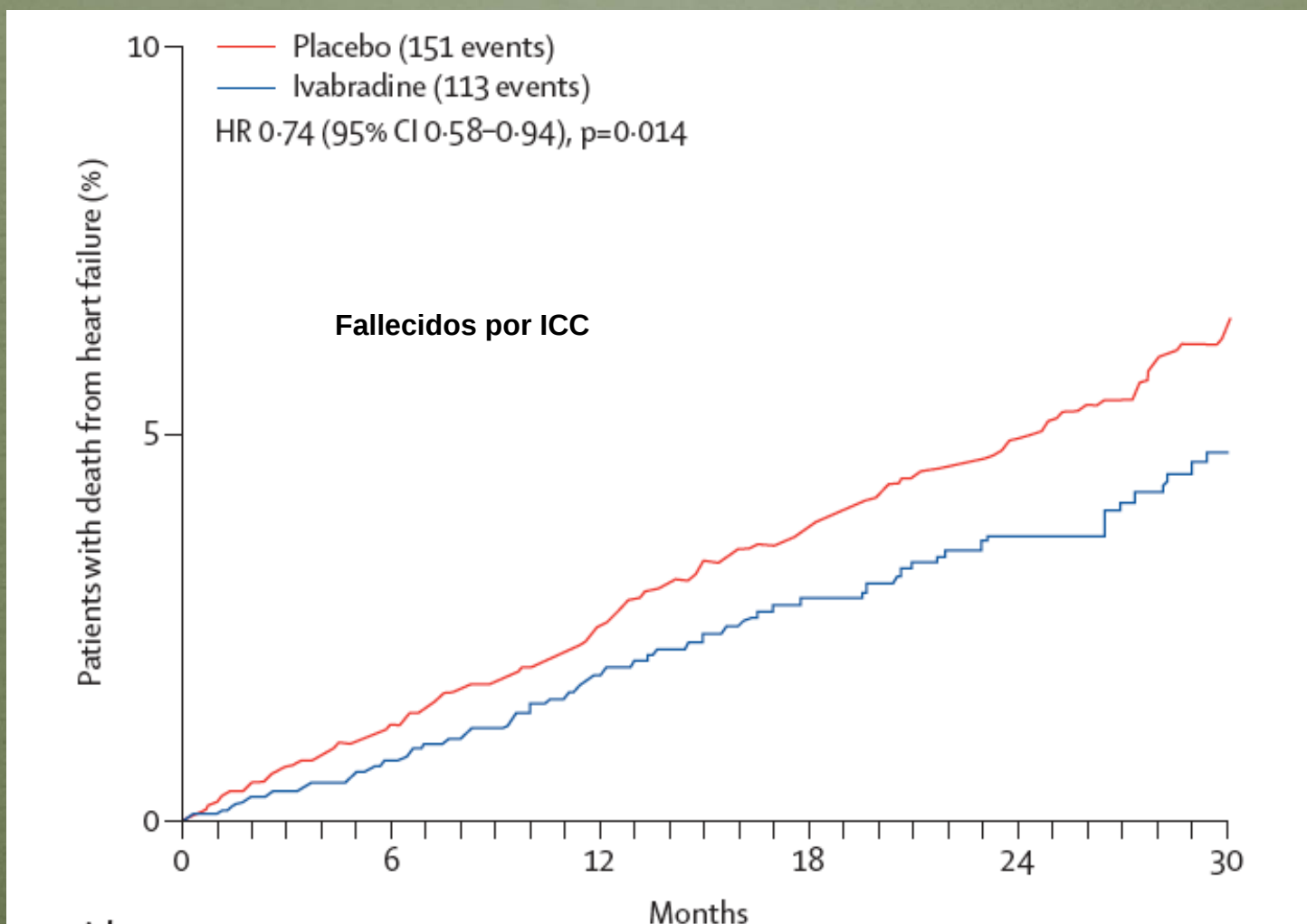


# Ivabradine and outcomes in chronic heart failure (SHIFT): a randomised placebo-controlled study





# Ivabradine and outcomes in chronic heart failure (SHIFT): a randomised placebo-controlled study



# Ivabradine and outcomes in chronic heart failure (SHIFT): a randomised placebo-controlled study

## Conclusiones:

- El tratamiento con Ivabradina en pacientes con ICC y FEVI deprimida reduce los reingresos por IC.
- Reduce la mortalidad por IC en el reingreso.
- Beneficio demostrado en frecuencia cardiaca  $> 77$  x' en reposo.

¿ Se podría haber optimizado el tratamiento betabloquante ?  
¿ Mismo efecto con otros fármacos controladores de frecuencia cardiaca ?

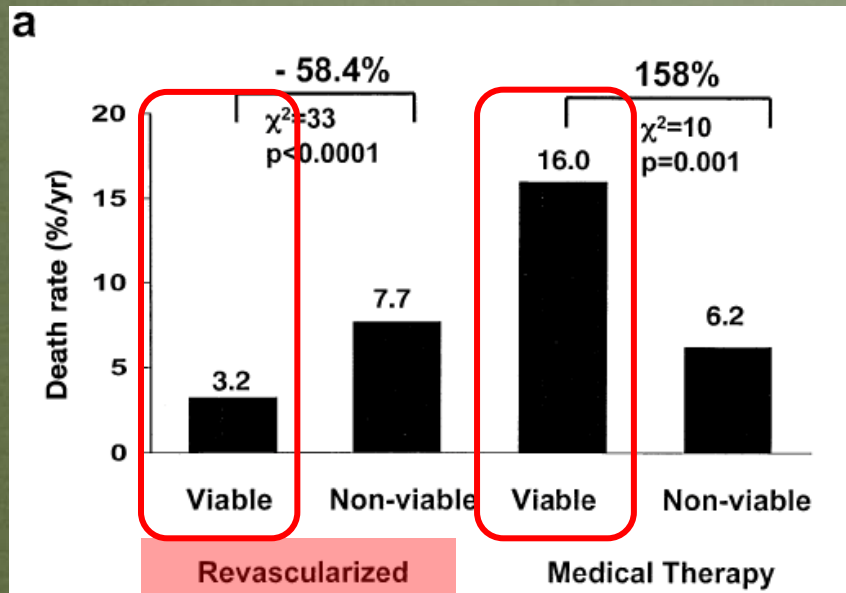
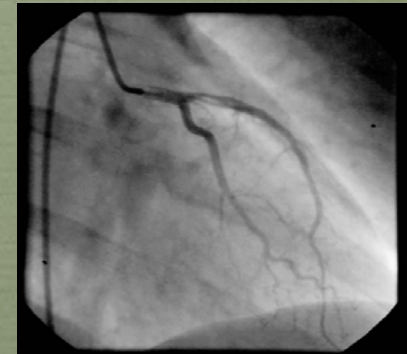
# **Revascularización coronaria**

## A RANDOMIZED TRIAL OF CORONARY ARTERY BYPASS SURGERY

### Survival of Patients with a Low Ejection Fraction

N Engl J Med 1985;312:1665–1671.

Myocardial Viability Testing and Impact of Revascularization on Prognosis in Patients With Coronary Artery Disease and Left Ventricular Dysfunction: A Meta-Analysis



ESC GUIDELINES

#### Key evidence

There are no data from multicentre trials assessing the value of revascularization procedures for the relief of HF symptoms.

However, single-centre, observational studies on HF of ischaemic origin suggest that revascularization may lead to symptomatic improvement and potentially improve cardiac function. Clinical trials are ongoing that address the effect of intervention on clinical outcomes.<sup>134</sup>

J Am Coll Cardiol 2002;39:1151– 8

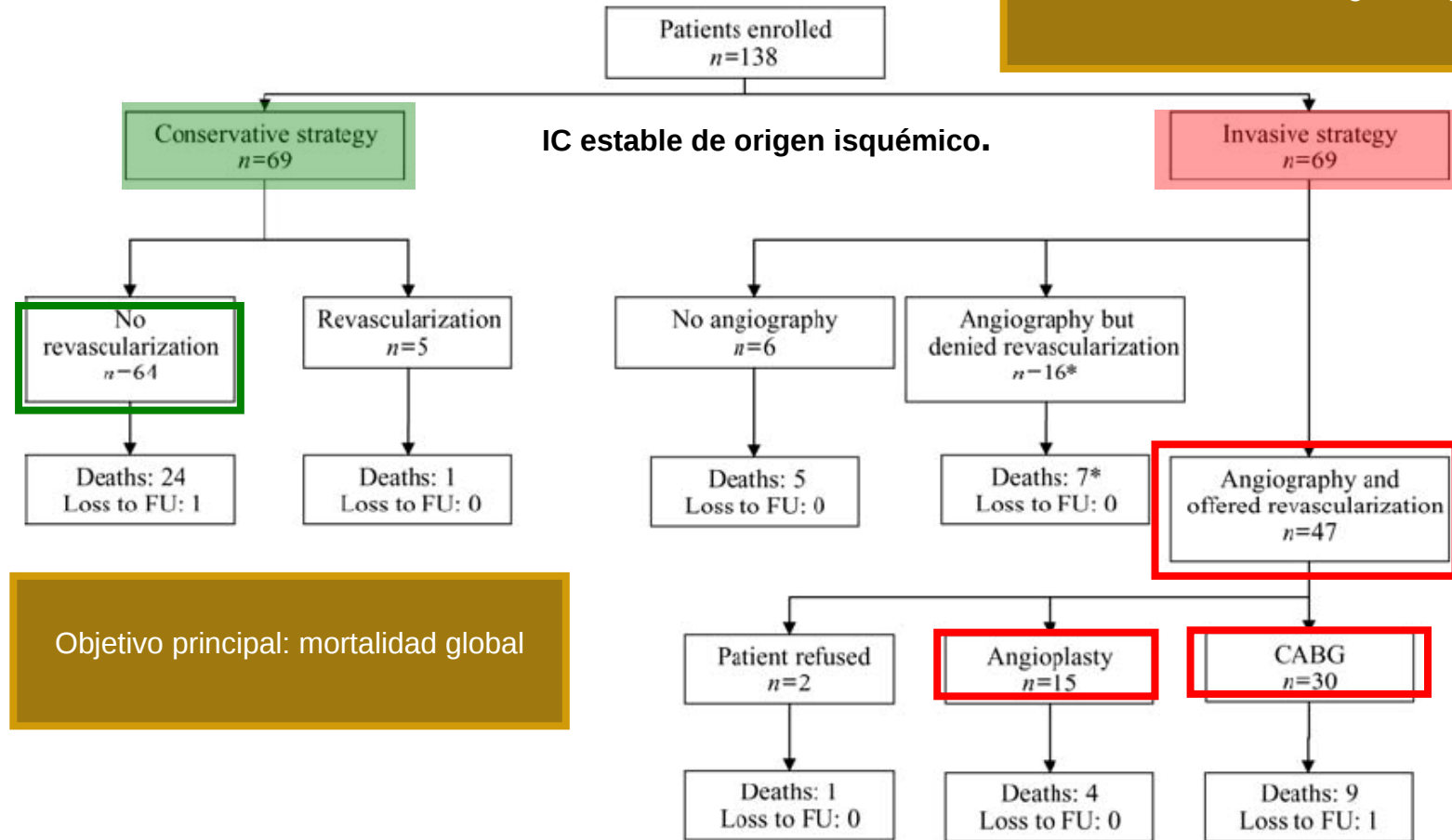
Eur Heart Journal (2011) 32, 820–828

# **The Heart Failure Revascularisation Trial (HEART)**

¿ Y si a pesar de miocardio viable optamos por el tratamiento farmacológico ?

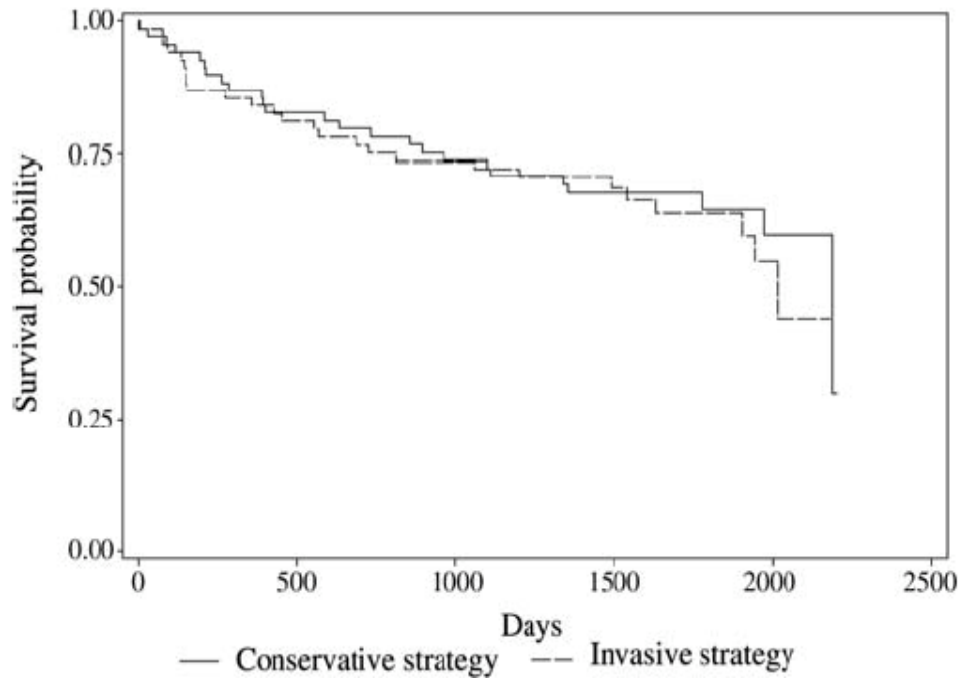
# The Heart Failure Revascularisation Trial (HEART)

Viabilidad miocárdica demostrada mediante eco-stress o gammagrafía



Objetivo principal: mortalidad global

# The Heart Failure Revascularisation Trial (HEART)



-Mortalidad global sin diferencias.

Estrategia Invasiva- 38%

Estrategia conservadora- 36%

- Calidad de vida 6 meses (Minnessota) sin diferencias.

## Coronary-Artery Bypass Surgery in Patients with Left Ventricular Dysfunction

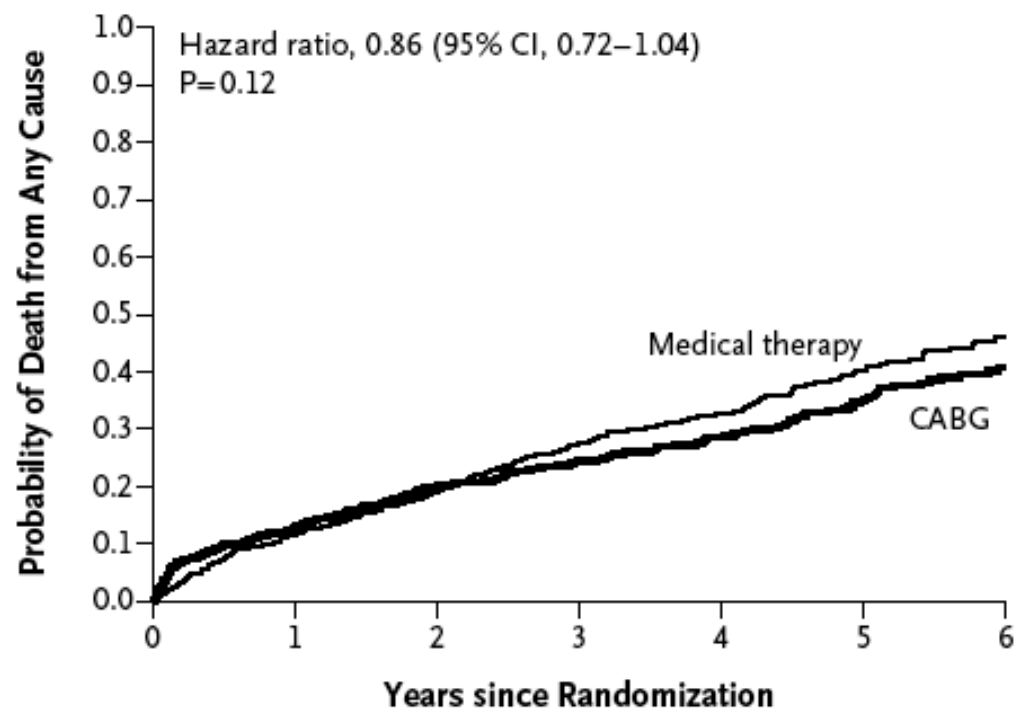
**CABG + medical vs Medical**

### BACKGROUND

The role of coronary-artery bypass grafting (CABG) in the treatment of patients with coronary artery disease and heart failure has not been clearly established.

### METHODS

Between July 2002 and May 2007, a total of 1212 patients with an ejection fraction of 35% or less and coronary artery disease amenable to CABG were randomly assigned to medical therapy alone (602 patients) or medical therapy plus CABG (610 patients). The primary outcome was the rate of death from any cause. Major secondary outcomes included the rates of death from cardiovascular causes and of death from any cause or hospitalization for cardiovascular causes.



**No. at Risk**

|                 |     |     |     |     |     |     |    |
|-----------------|-----|-----|-----|-----|-----|-----|----|
| Medical therapy | 602 | 532 | 487 | 435 | 312 | 154 | 80 |
| CABG            | 610 | 532 | 486 | 459 | 340 | 174 | 91 |

**Figure 1.** Kaplan–Meier Curves for the Probability of Death from Any Cause. CABG denotes coronary-artery bypass grafting.

## Coronary-Artery Bypass Surgery in Patients with Left Ventricular Dysfunction

| Outcome                                                   | Medical<br>Therapy<br>(N=602)<br><br><i>no. (%)</i> | CABG<br>(N=610)<br><br><i>no. (%)</i> | Hazard Ratio<br>with CABG<br>(95% CI) | P Value† |
|-----------------------------------------------------------|-----------------------------------------------------|---------------------------------------|---------------------------------------|----------|
| Primary outcome: rate of death from any cause             | 244 (41)                                            | 218 (36)                              | 0.86 (0.72–1.04)                      | 0.12     |
| Secondary outcomes                                        |                                                     |                                       |                                       |          |
| Death from any cause within 30 days after randomization   |                                                     |                                       |                                       |          |
| Logistic-regression model                                 | 7 (1)                                               | 22 (4)                                | 3.19 (1.35–7.52)‡                     | 0.008    |
| Cox proportional-hazards model                            | 7 (1)                                               | 22 (4)                                | 3.12 (1.33–7.31)                      | 0.006    |
| Death from cardiovascular causes                          | 201 (33)                                            | 168 (28)                              | 0.81 (0.66–1.00)                      | 0.05     |
| Death from any cause or hospitalization for heart failure | 324 (54)                                            | 290 (48)                              | 0.84 (0.71–0.98)                      | 0.03     |

## Myocardial Viability and Survival in Ischemic Left Ventricular Dysfunction

**CABG + medical      vs      Medical**

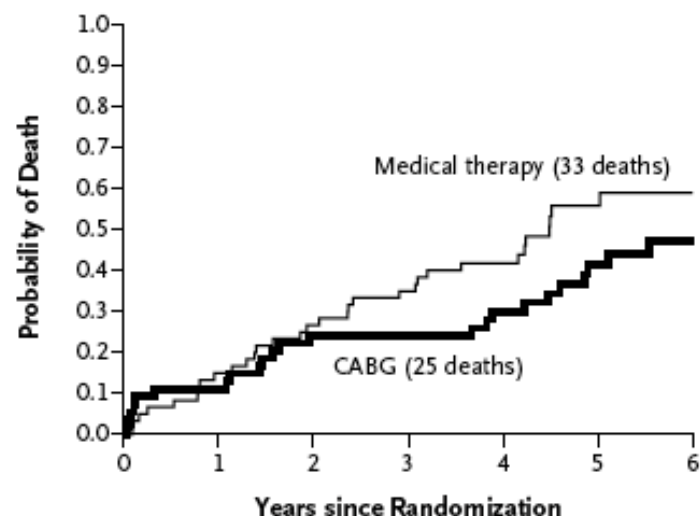
### BACKGROUND

The assessment of myocardial viability has been used to identify patients with coronary artery disease and left ventricular dysfunction in whom coronary-artery bypass grafting (CABG) will provide a survival benefit. However, the efficacy of this approach is uncertain.

### METHODS

In a substudy of patients with coronary artery disease and left ventricular dysfunction who were enrolled in a randomized trial of medical therapy with or without CABG, we used single-photon-emission computed tomography (SPECT), dobutamine echocardiography, or both to assess myocardial viability on the basis of pre-specified thresholds.

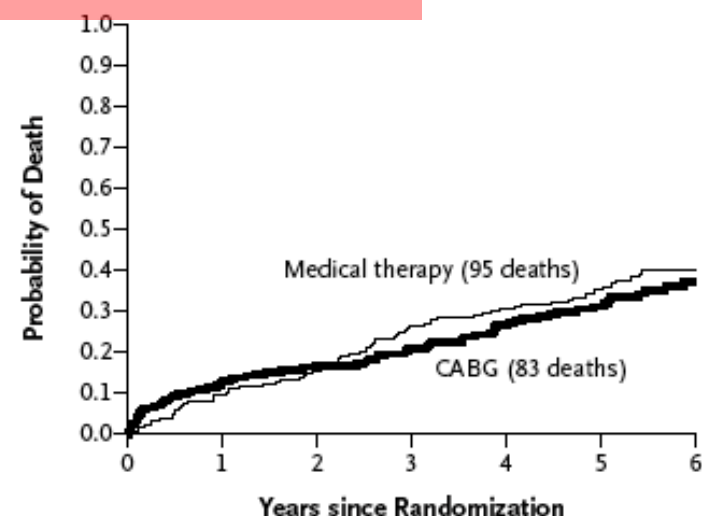
### A Without Myocardial Viability



#### No. at Risk

|                 |    |    |    |    |    |    |    |
|-----------------|----|----|----|----|----|----|----|
| Medical therapy | 60 | 51 | 44 | 39 | 29 | 14 | 4  |
| CABG            | 54 | 48 | 41 | 41 | 34 | 22 | 12 |

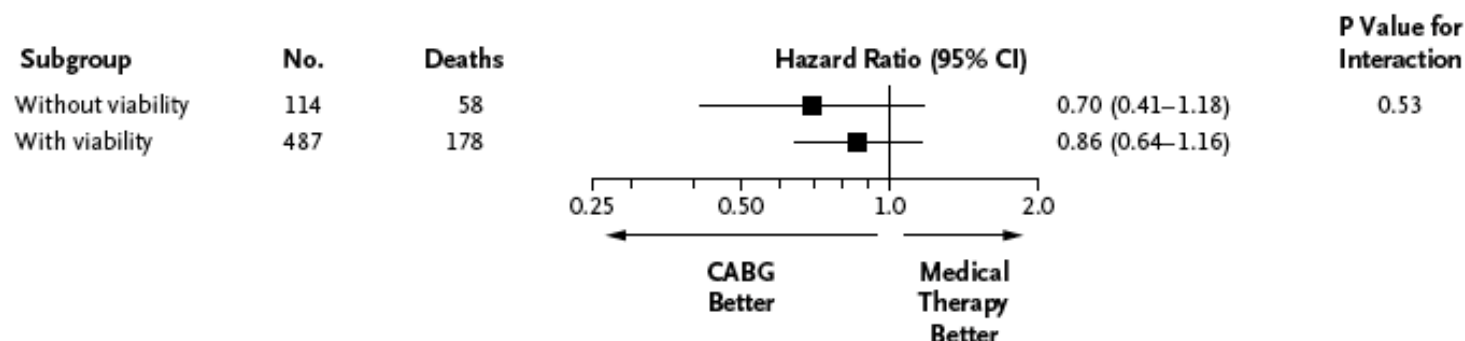
### B With Myocardial Viability



#### No. at Risk

|                 |     |     |     |     |     |    |    |
|-----------------|-----|-----|-----|-----|-----|----|----|
| Medical therapy | 243 | 219 | 206 | 179 | 146 | 94 | 51 |
| CABG            | 244 | 213 | 203 | 192 | 148 | 94 | 51 |

### C



**Figure 2. Kaplan–Meier Analysis of the Probability of Death According to Myocardial-Viability Status and Treatment.**

At 5 years in the intention-to-treat analysis, the rates of death for patients without myocardial viability were 41.5% in the group assigned to undergo coronary-artery bypass grafting (CABG) and 55.8% in the group assigned to receive medical therapy (Panel A). Among patients with myocardial viability, the respective rates were 31.2% and 35.4% (Panel B). There was no significant interaction between viability status and treatment assignment with respect to mortality ( $P=0.53$ ) (Panel C).

## The Heart Failure Revascularisation Trial (HEART)

ORIGINAL ARTICLE

### Myocardial Viability and Survival in Ischemic Left Ventricular Dysfunction

Sharma S, et al. N Engl J Med. 2014;371:130-40.

ORIGINAL ARTICLE

### Coronary-Artery Bypass Surgery in Patients with Left Ventricular Dysfunction

Sharma S, et al. N Engl J Med. 2014;371:130-40.

#### Conclusiones:

- Estrategia de revascularización coronaria no es superior al tratamiento farmacológico óptimo en pacientes con insuficiencia cardiaca estable de etiología isquémica.
- No es superior en pacientes con viabilidad coronaria.
- Mortalidad precoz asociada al procedimiento es mayor.
- No mejoría en la calidad de vida

**Muestra pequeña de pacientes con viabilidad estudiada ( <50% STICH1)**

**Empleo de distintos métodos para evaluar viabilidad miocárdica**

**No se especifican complicaciones derivadas del procedimiento**

# CONCLUSIONES

Diuréticos

Dosis elevadas – seguridad y eficacia  
No superioridad de infusión continua

Betabloqueantes

Mantener en la fase aguda

Hierro intravenoso

Mejora capacidad funcional / calidad de vida

Antialdosterónicos

Menor mortalidad cardiovascular y reingresos por Insuficiencia Cardíaca

Ivabradina

Menor mortalidad cardiovascular y reingresos por Insuficiencia Cardíaca

Revascularización

Sin claros beneficios en ausencia de clínica anginosa a pesar de miocardio viable



*Gracias por vuestra atención*