3er Curs
D'ACTUALITZACIÓ EN
INSUFICIÈNCIA
CARDÍACA

Hotel Hilton Barcelona
 Avda Diagonal 589-591. 08014 BCN



Noves Guíes per a un nou temps, qué ens queda per saber?

Dr. Nicolás Manito

Cap Clínic de la Unitat d'Insuficiència Cardiaca Avançada i Trasplantament Cardiac Hospital Universitari de Bellvitge. L'Hospitalet del Llobregat – Barcelona







Conflictos de Interes

• Conferencias y AB:

Astra Zeneca, Boehringer Ingelheim, Bayer, Novartis, Vifor

qué ens queda per saber.....?

PHARMACOTHERAPY OF CHF

Order of adding DMT for HFrEF

Therapies for HFmrEF and HFpEF

HFrEF therapies in eGFR <30 mL/min/1.73m2

HF phenotypes: myocarditis, cardiotoxicity, inherited CMPs, PPCM, amyloidosis

Strategies for 'recovered LV' systolic function

Evidence on the effects of fluid restriction, dietary salt

Role of biomarkers RCT for HF stage B Diagnosis protocols for HFmrEF



DEVICES AND INTERVENTIONS

ICDs HFrEF/HFmrEF/HFpEF

CRT efficacy in AF

Outcomes of AF ablation

Percutaneous treatment of valve heart dis.

CCM & baroreceptor stimulation in HFrEF

DEFINITION AND EPIDEMIOLOGY

Research HFmrEF and HFpEF

Normal values/ranges of EF

'Recovered LV' systolic function

DISEASE MANAGEMENT

Remote monitoring in HF post COVID-19

Models for follow-up of stable HF patients

Specific options for palliative care

ADVANCED HF

Risk profiles according to INTERMACS

Outcomes of long-term MCS

Reduce the risk for LVAD of bleeding, Thromboembolic events, and infection

Medical treatment

AHF

Patient phenotypes

Imaging techniques and biomarkers

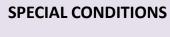
Impact on post-discharge outcomes

Strategies for congestión relief

Devices for short-term MCS

Therapeutic algorithms for cardiogenic shock

GAPS IN EVIDENCE



Treatment for PPCM

cancer therapies

injury with infection

Phenotyping of CMPs through genetic testing, biomarkers

NON-CV COMORBIDITIES

Outcomes of treatment of CSA

Cachexia and/or sarcopenia and/or frailty

Medical therapies or devices in severe CKD and HF

Medical treatment of electrolyte abnormalities

Prevention and treatment of cardiotoxicity of

Treatment of infections and prevention of cardiac

and imaging modalities, and tailoring of therapy

Treatment of different types of myocarditis, including immunosuppressive therapies

Treatments of different forms of cardiac amyloid

Definition and treatment of LA myopathy

CV COMORBIDITIES

Strategies for the treatment of ventricular arrhythmias

Coronary revascularization procedures in different patient subsets

Patients' outcomes and/or QOL of percutaneous treatment of mitral or tricuspid valve disease



Costs and healthcare utilisation of patients with heart failure in Spain

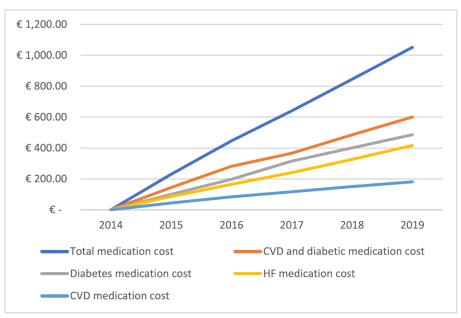


Carlos Escobar^{1*}, Luis Varela², Beatriz Palacios², Margarita Capel², Antoni Sicras³, Aram Sicras³, Antonio Hormigo⁴, Roberto Alcázar⁵, Nicolás Manito⁶ and Manuel Botana⁷

Patient cumulative hospital mean cost

€14,000 €12,000 €10,000 €8,000 €6,000 €4,000 €2,000 2014 2015 2016 2017 2018 2019 Cardiorenal cost ——HF cost CVD cost CKD cost Stroke cost MI cost PAD cost

Patient cumulative medication mean cost



The great burden for this cost was due to cardiorenal (HF and/or chronic kidney disease) hospitalizations (88.8% of the total cost), particularly HF (67.3% of the total cost)

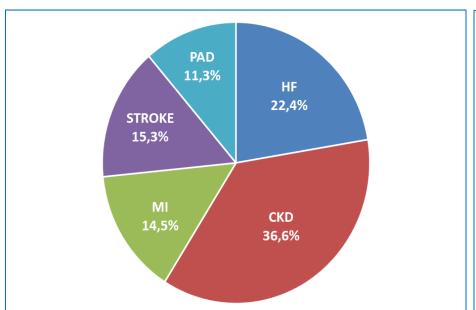
Escobar C, Manito N, et al. BMC Health Services Research 2020;20 (1):964.

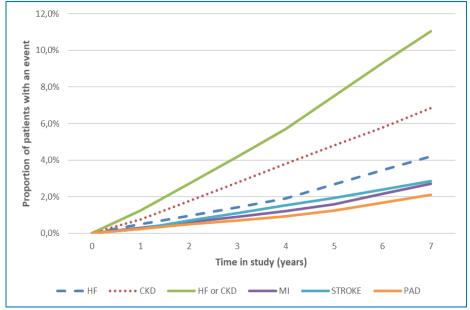
Epidemiology and resource use in Spanish type 2 diabetes patients without previous cardiorenal disease: CaReMe Spain study

First cardiovascular/renal manifestation during follow-up in type 2 diabetes patients (2013 –2019)

Percentage of events

Temporal evolution (first manifestation)







Universal definition and classification of heart failure: a report of the Heart Failure Society of America, Heart Failure Association of the European Society of Cardiology, Japanese Heart Failure Society and Writing Committee of the Universal Definition of Heart Failure

Endorsed by the Canadian Heart Failure Society, Heart Failure Association of India, Cardiac Society of Australia and New Zealand, and Chinese Heart Failure Association

New classification of HF according to LVEF

HF with reduced EF (HFrEF): HF with LVEF < 40% HF with mildly reduced EF (HFmrEF): HF with LVEF 41-49% **HF with preserved EF (HFpEF):** HF with LVEF > 50% HF with improved EF (HFimpEF): HF with a baseline LVEF ≤ 40%, a ≥ 10 point increase from baseline LVEF, and a second measurement of LVEF > 40%

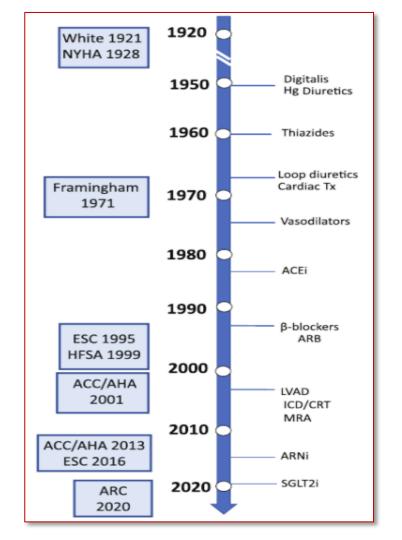
The path to universality

Eugene Braunwald* and Elliott M. Antman

TIMI Study Group, Division of Cardiovascular Medicine, Brigham and Women's Hospital, Department of Medicine, Harvard Medical School, Boston, MA, USA

Time-line for important treatments and definitions of heart failure between 1950 and 2020 (horizontal lines) and dates of heart failure definitions and/or practice guidelines (rectangles)

"An essential component of the trials that led to these recent advances was a clear definition of HF in the populations studied"



THE PRESENT AND FUTURE

HFpEF trials : LVEF inclusión criteria

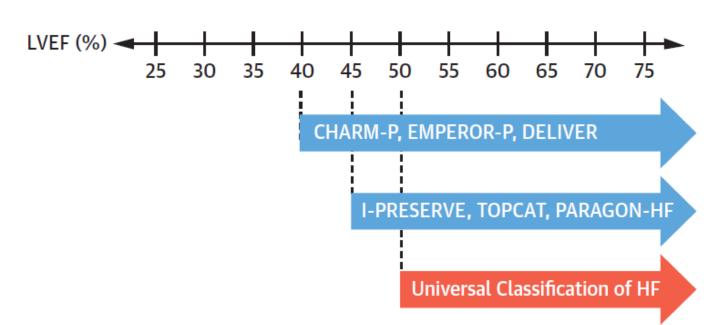
JACC REVIEW TOPIC OF THE WEEK

Classification of Heart Failure According to Ejection Fraction



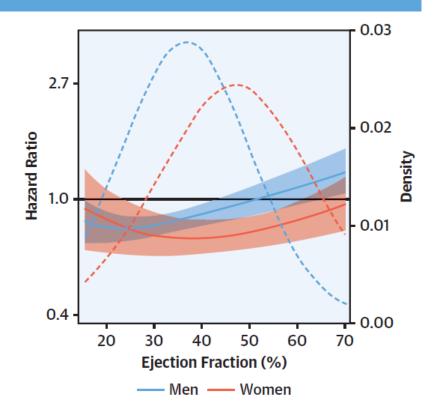
JACC Review Topic of the Week

Carolyn S.P. Lam, PhD, MBBS, a,b Scott D. Solomon, MDC

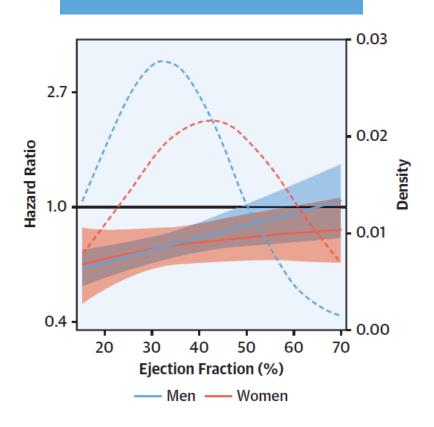


Interaction Between LVEF, Sex, and Neurohormonal Modulators in HF





Primary Outcome - MRA



THE PRESENT AND FUTURE

NEW PROPOSAL HEART FAILURE CLASSIFICATION

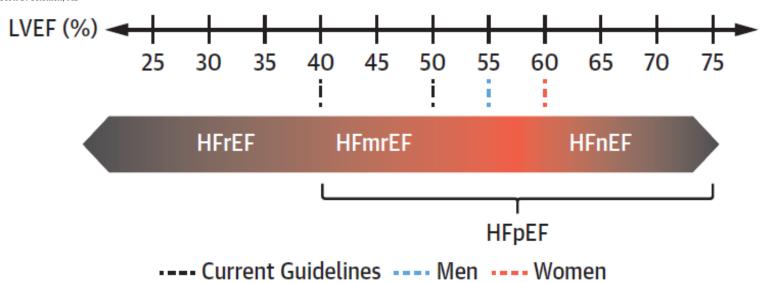
JACC REVIEW TOPIC OF THE WEEK

Classification of Heart Failure According to Ejection Fraction



JACC Review Topic of the Week

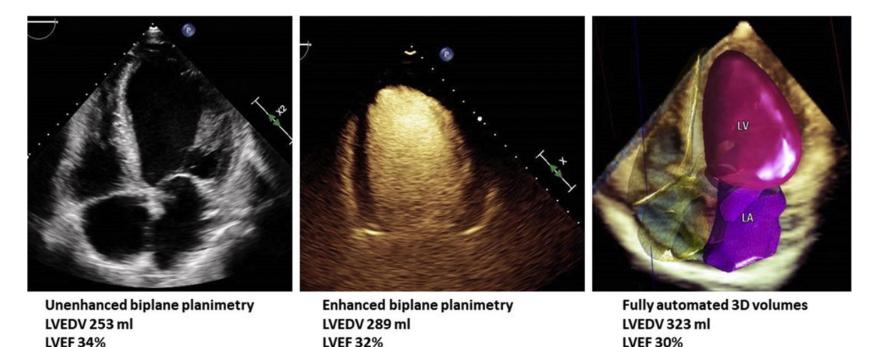
Carolyn S.P. Lam, PhD, MBBS, a,b Scott D. Solomon, MDC



The New Framework for Application of Cardiac Imaging to Patients With Heart Failure

Chamber volumes and remodeling Volumetric **Ejection fraction** Stroke volume Characterization Regurgitant volumes Doppler-derived velocity and flow Flow and Pressure measures Characterization Intracavitary flow dynamics Strain and strain rates **Tissue Tracking** Twist and untwist Myocardial stiffness Tissue Ischemia and hibernation Characterization Cardiac innervation Replacement fibrosis Extracellular expansion/deposition Edema Inflammation

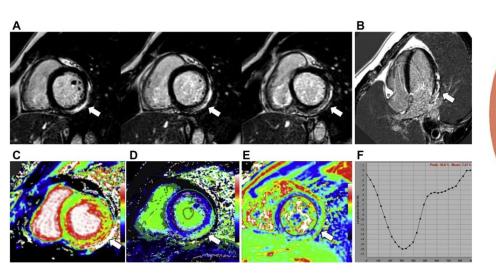
Application of Different Echocardiographic Methods for Volumetric Assessment

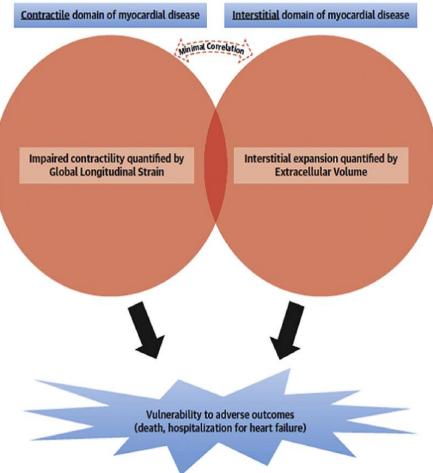


- In the same patients during the same study, unenhanced, contrast-enhanced, and fully automated 3-dimensional (3D) echocardiographic assessment yielded differing results. The differences are most prominent for left ventricular end-diastolic volume (LVEDV).
- Fully automated 3D echocardiographic assessment is more likely to provide volumes comparable with those provided by cardiac magnetic resonance. LA= left atrium; LV = left ventricle; LVEF = left ventricular ejection fraction.

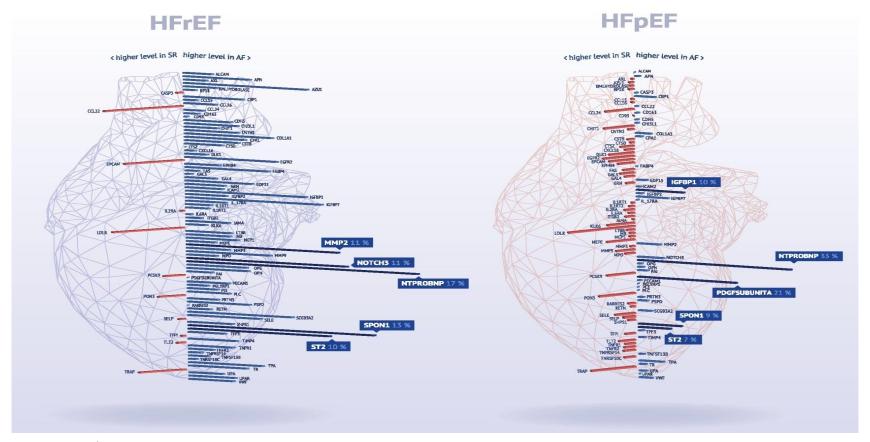
Longitudinal Systolic Function and Extracellular Volume Represent Distinct

Domains of Myocardial Vulnerability



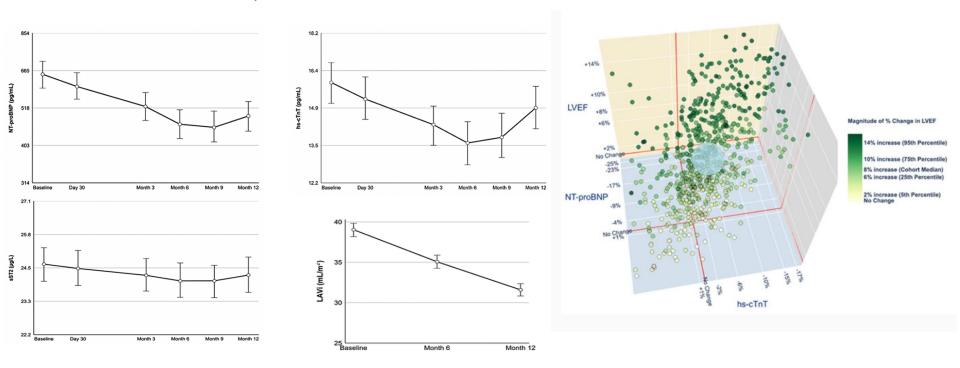


Comparing biomarker profiles of patients with heart failure: atrial fibrillation vs. sinus rhythm and reduced vs. preserved ejection fraction



Association Between Angiotensin Receptor-Neprilysin Inhibition, Cardiovascular Biomarkers, and Cardiac Remodeling in Heart Failure with Reduced Ejection Fraction

Reduction in NT-proBNP and hs-cTnT following initiation of sacubitril/valsartan may be a helpful indicator of reverse cardiac remodeling in the absence of repeat imaging shortly after initiating treatment with sacubitril/valsartan



Murphy SP, et al. Circulation Heart Fail. 2021. doi: 10.1161/CIRCHEARTFAILURE.120.008410. Online ahead of print

El fármaco ideal debería tener una evidencia científica sólida

VIII.

Capaz de dar marcha atrás y detener el paso del tiempo en el paciente con IC

- Muerte por IC
- Muerte súbita
- Muerte total
- Hospitalizaciones
- Rehospitalizaciones

- **Efecto independiente**

- **Efecto precoz**
 - Efecto sostenido
- **Ambulatorio**
- Hospitalizado

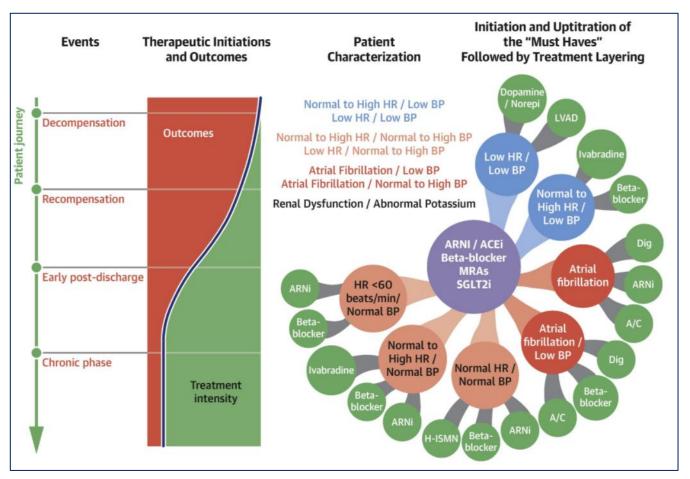
- Protección renal
- Protección metabólica
- Protección miocárdica
- Remodelado
- **Biomarcadores**

Coste-efectividad

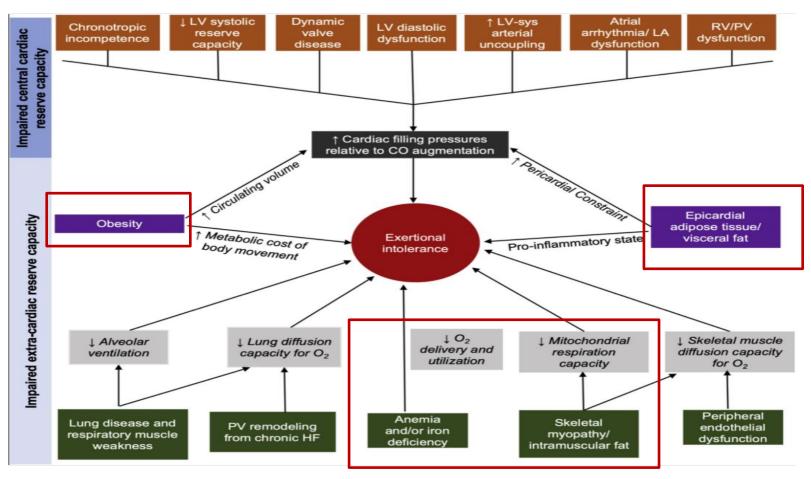
- **Tolerancia**
- Seguridad
- Calidad de vida
- **Síntomas**
- **Adherencia**



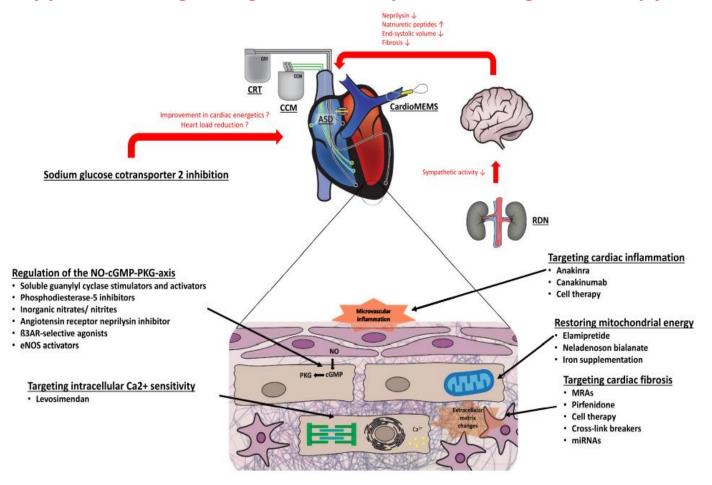
Fenotipos clínicos para el inicio del tratamiento en IC



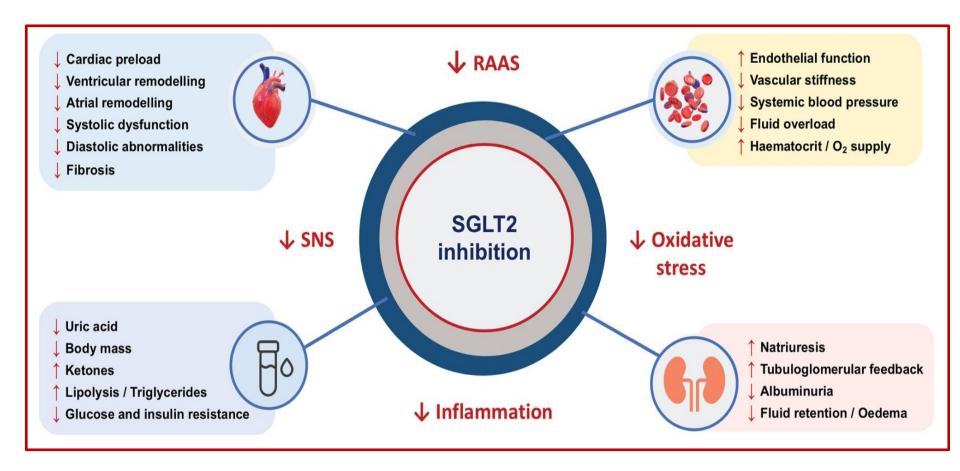
Pathophysiological Contributors to Exercise Intolerance in HFpEF



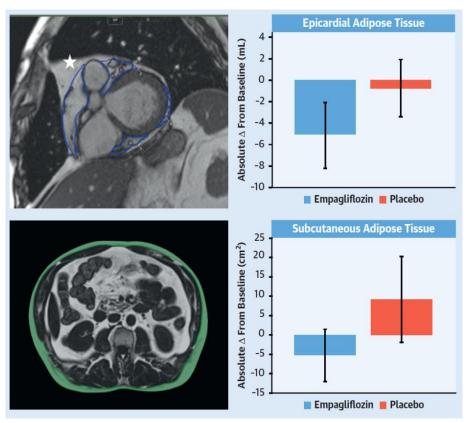
Main approaches regarding device and pharmacological therapy in HFpEF



Possible mechanisms detailing why SGLT2 inhibitors may work in HFpEF



Empagliflozin Reduces Epicardial and Subcutaneous Adipose Tissue in Nondiabetic Patients With Heart Failure With Reduced Ejection Fraction



JACC: HEART FAILURE

9 02021 BY THE AMERICAN COLLEGE OF CARDIOLOGY FOUNDATION
PUBLISHED BY ELSEVIER

EDITORIAL COMMENT

Empagliflozin-Induced Changes in Epicardial Fat



VOL. 9, NO. 8, 2021

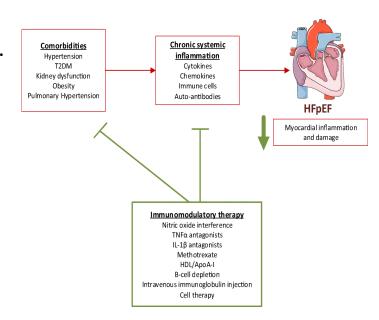
The Centerpiece for Myocardial Protection?*

Wilfried Mullens, MD, PHD, Pieter Martens, MD, PHD

	Major Function	Nominal P Value	BH P Value	Log2 Fold Change
TNFRSF10C	Inflammation; apoptosis	0.000049	0.004536	-0.084331
SELE	Inflammation	0.000289	0.013422	-0.046008
CHI3L1	Angiogenesis; inflammatory response	0.024798	0.193865	-0.144620
PAI	Inflammation; fibrinolysis	0.033353	0.193865	-0.090616
CCL16	Inflammation	0.030766	0.193865	0.093097
MPO	Inflammation; plaque formation and rupture	0.026782	0.193865	-0.089846
IL-1RT2	Inflammation	0.025677	0.193865	-0.039961
TR-AP	Binding inflammation receptor	0.025842	0.193865	-0.057800
GRN	Cell development, wound healing, tumorigenesis	0.001924	0.059630	-0.045100
PDGF subunit A	Cell proliferation, migration, chemotaxis	0.004022	0.093501	-0.213026
ICAM-2	Antigen-specific immune response, lymphocyte recirculation	0.007904	0.147009	-0.046557
PECAM-1	Cell adhesion and leukocyte migration	0.011300	0.175144	-0.060474
DLK-1	Inhibition of adipogenesis	0.023593	0.193865	0.068419
SHPS-1	Synaptic function, cell adhesion, phagocytosis, mast cell, and dendritic cell activation	0.028962	0.193865	-0.067255
MMP-9	Angiogenesis and wound repair	0.029667	0.193865	-0.096804
BLM hydrolase	Protection against bleomycin toxicity	0.031878	0.193865	-0.111773
CTSD	Angiogenesis	0.037319	0.204155	-0.101077

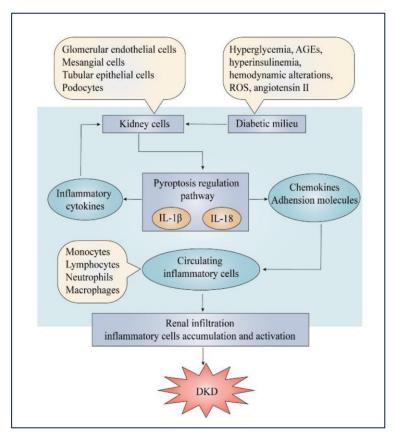
Immunomodulation in Heart Failure with Preserved Ejection Fraction: Current State and Future Perspectives

- 1. Pathophysiological mechanisms of heart failure with preserved ejection fraction (HFpEF) as targets for therapy.
- 2. **Comorbidities**, such as hypertension, type 2 diabetes mellitus (T2DM), and obesity, lead to chronic systemic inflammation and subsequently HFpEF, associated with myocardial inflammation and damage (red)
- 3. **Immunomodulation targeting** either comorbidities or underlying disease mechanisms (green) can decrease myocardial inflammation and damage and are currently under evaluation



PYROPTOSIS: New concepts

- Pyroptosis, a kind of programmed cell death, is induced through an executor protein of cysteine-aspartic proteases1 (caspase-1) by some immunoactivity cells stimulated by pathogens and danger signals
- The typical manifestations of pyroptosis are increased by expression of intracellular NLRP3 inflammasome and activated caspase-1.
- Podocyte loss in glomerulus is one of the early triggers of DKD. Pyroptosis has been reported as being correlated with the mechanism of podocyte loss
- Tubular epithelial cell pyroptosis is a risk factor to tubular injury in DKD. As mentioned earlier there is a ROS/TXNIP/NLRP3 inflammasome signalling pathway in tubular epithelial cells which leads to cell pyroptosis.
- SGLT2 inhibition target pyroptosis activity through NLRP3 inflammasome reduced expression in diabetic and nondiabetic milieu

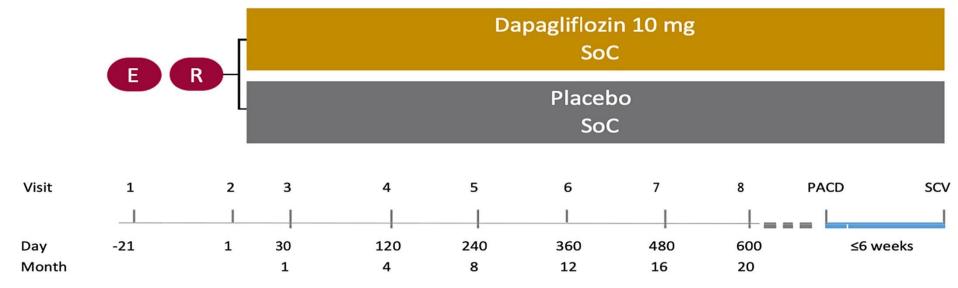


SGLT2 inhibitors are anti-inflammatory drugs

Dapagliflozin in heart failure with preserved and mildly reduced ejection fraction: rationale and design of the DELIVER trial

Symptomatic HF (NYHA class II–IV). LVEF >40% and evidence of structural heart disease

NT-proBNP ≥300 pg/mL at Visit 1 for patients without ongoing atrial fibrillation/flutter. If ongoing atrial fibrillation/flutter at Visit 1, NT-proBNP must be ≥600 pg/mL.



In person visits after 30 days; 4 months; thereafter every 4 months after randomization.

ONGOING TRIALS FOR HFPEF PATIENTS TREATED WITH IRON IV

1. Effect of IV Iron in Patients With Heart Failure With Preserved Ejection Fraction (FAIR-HFpEF)

Primary Outcome Measures: Exercise capacity [Time Frame: 52 weeks]The difference of 6-minute walking distance in meters from baseline to end of study in symptomatic patients with HFpEF with documented ID compared to the control group.

2. Effects of Iron Therapy in Heart Failure With Preserved Ejection Fraction and Iron Deficiency (PREFER-HF)

Primary Outcome Measures: Six minute walking test distance [Time Frame: 24 weeks] Change in meters traveled in six minute walking test from baseline to week 24. An increase in distance is related to an improvement in functional capacity.

- 1. NCT03074591
- 2. NCT03833336

HEART FAILURE COMPENDIUM

From Systemic Inflammation to Myocardial Fibrosis

The Heart Failure With Preserved Ejection Fraction Paradigm Revisited

Walter J. Paulus[®], Michael R. Zile[®]

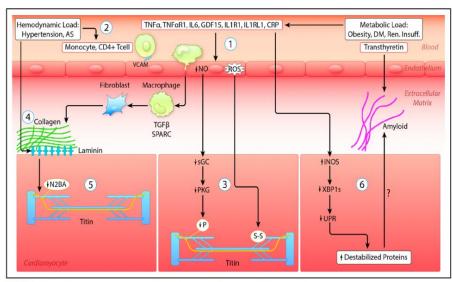
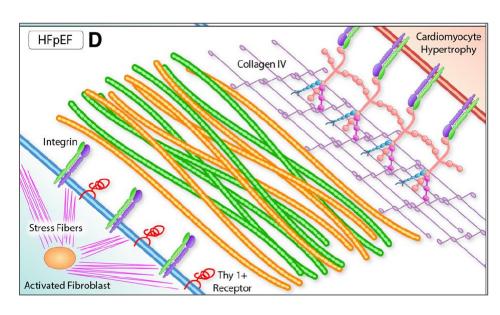


Figure 1. Pathophysiological mechanisms linking systemic inflammation to myocardial stiffness.

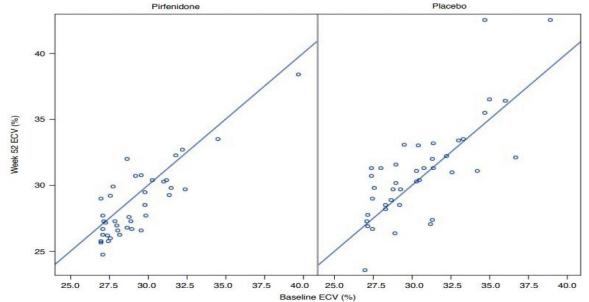


PIROUETTE trial. Pirfenidone in HFpEF: a randomized phase 2 trial

Myocardial fibrosis, defined as a myocardial extracellular volume of 27% or greater

Table 2 Primary outcome								
	Pirfenidone			Placebo				
	Baseline (n = 47)	52 weeks (n=39)	Δ from baseline to 52 weeks	Baseline (n = 47)	52 weeks (n = 41)	Δ from baseline to 52 weeks	Between-group difference (95% CI) ^a	P value
Myocardial ECV (%)	29.5 ± 2.5	28.6 ± 2.7	-0.7 ± 1.4	30.7 ± 2.9	31.1±3.8	0.5 ± 2.4	-1.21 (-2.12 to -0.31)	0.009

Data are mean ± s.d. *Analysis of covariance (ANCOVA), two-sided, adjusted for baseline myocardial ECV, sex and treatment group; F = 7.11, P = 0.009.





Lewis GA, et al. Nature Med. 2021;27:1477–1482

NCT02932566

Technologies for Treating Left Atrial Decompression in Heart Failure

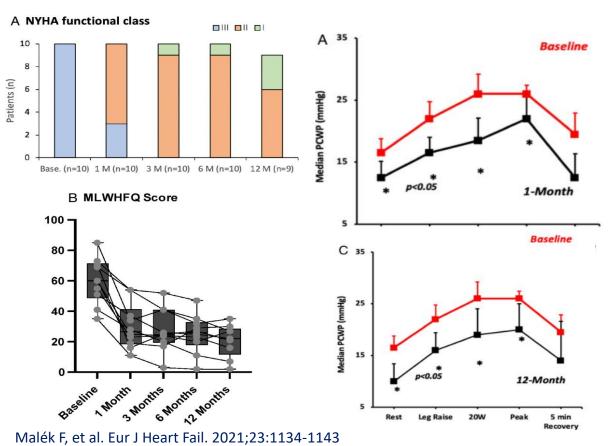
Total number patients treated	V-Wave device ⁵	Interatrial Shunt Device ^{6, 7} 86	Atrial Flow Regulator ⁸ 36	Levoatrial-Coronary Sinus Shunt ⁹
HFpEF / HFrEF	8 / 30	86/0	20/16	6/2
PAP (mmHg)	\Longrightarrow			\Longrightarrow
PCWP at rest (mmHg)		\Longrightarrow	*/>	*
PCWP at exercise (mmHg)	_	1*	_	_
6MWT-distance (m)	_	*		
NYHA functional class	_	*	*	*
NT-pro BNP (pg/ml)	-	$\stackrel{\blacktriangledown}{\Longrightarrow}$	_	▼ ⇒

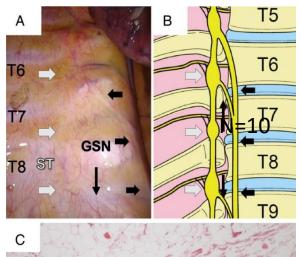
Felix Mahfoud et al. J Am Coll Cardiol Intv 2020; 13:1248-1250.

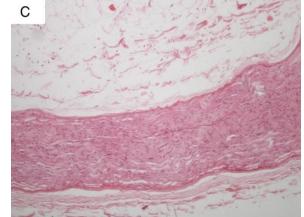
PRELOAD MODULATION

Surgical ablation of the right greater splanchnic nerve for the treatment of

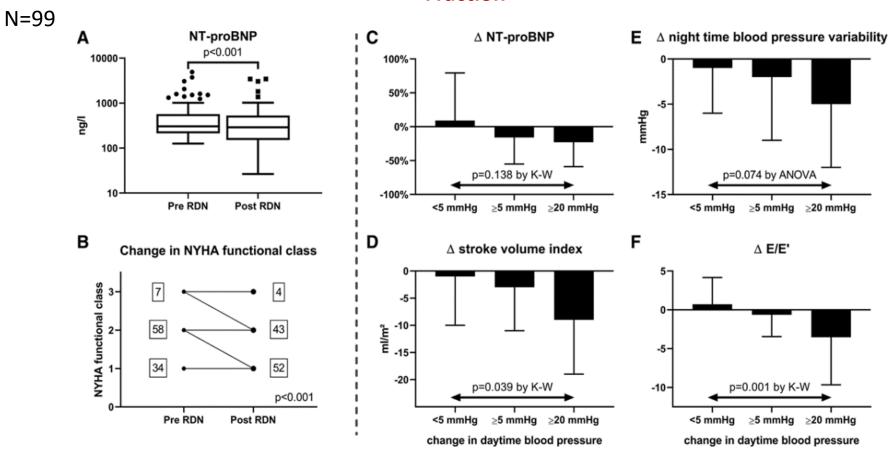
HFpEF: first-in-human clinical trial







Renal Sympathetic Denervation in Patients With Heart Failure With Preserved Ejection Fraction



Kresoja KP, et al. Circulation: Heart Failure. 2021. DOI:10.1161/CIRCHEARTFAILURE.120.007421

Proposed High-Priority Extracardiac Areas for Future Research in HFpEF

Translation

Discovery



Implementation



Identify geroscience targets for treatment:

- · Senescent cellular pathways
- Epigenetic pathways
- Adipose biology
- Skeletal muscle microvascular and mitochondrial dysfunction

Comprehensive phenotypic characterization:

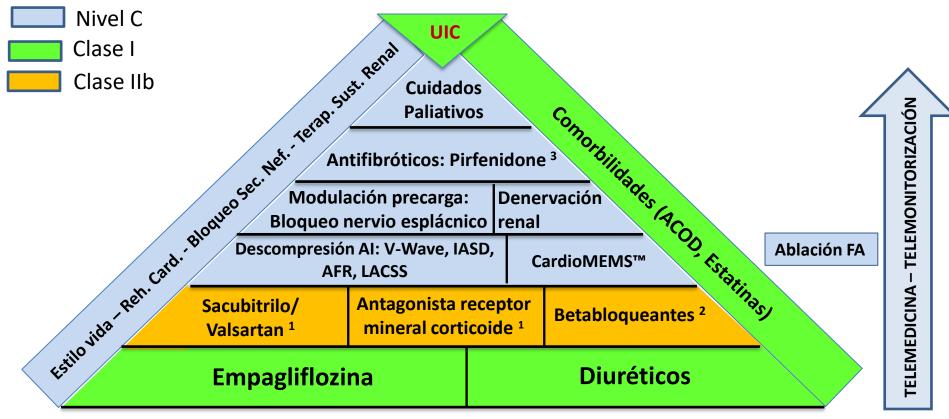
- Human biospecimens
- Biomarkers

Proof of concept intervention trials to modify the geroscience targets



- Evaluate multimorbidity management approaches
- Test promising geroscience interventions
- Improve adherence to effective lifestyle interventions
- Disseminate successful strategies into community practice

Alternativas terapéuticas para el paciente con ICFEp



1. FDA recomendación. 2. Si HTA y/o EAC y/o frecuencia cardiaca (RSR) elevada y en ausencia de incompetencia cronotrópica. 3. Estudios Fase II.

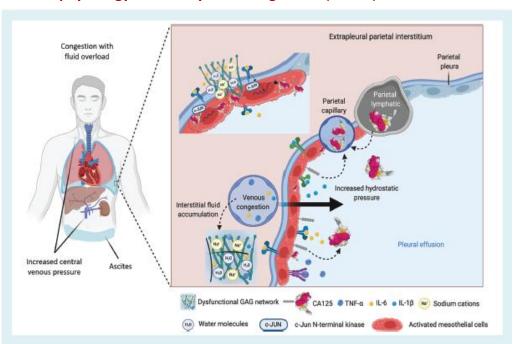
Al: aurícula izquierda; EAC: Enfermedad arterial coronaria; FA: fibrilación auricular; ACOD; anticoagulante acción directa IASD: Interatrial shunt device Corvia®; AFR: Atrial Flow regulator; LACSS: Levo-atrial coronary sinus shunt

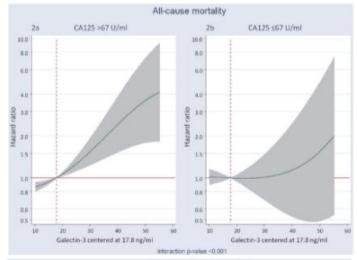


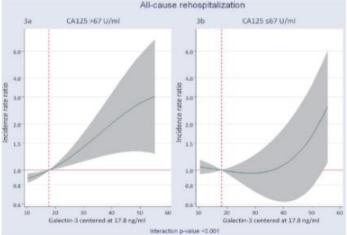
Antigen carbohydrate 125 as a biomarker in heart failure: a narrative review

Julio Núñez^{1,2,3}, Rafael de la Espriella^{1,2}, Gema Miñana^{1,2,3}, Enrique Santas^{1,2}, Pau Llácer⁴, Eduardo Núñez¹, Patricia Palau², Vicent Bodí^{1,2,3}, Francisco J. Chorro^{1,2,3}, Juan Sanchis^{1,2,3}, Josep Lupón^{2,5,6,7}, and Antoni Bayés-Genís^{3,5,6,7}*

Pathophysiology of carbohydrate antigen 125 (CA125) in heart failure







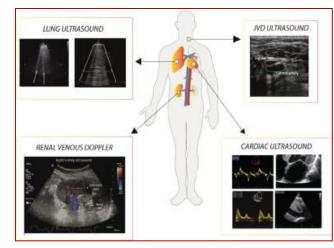
Nuñez J, et al. Eur J Heart Fail. 2021



Ultrasound imaging of congestion in heart failure: examinations beyond the heart

Pierpaolo Pellicori¹*†, Elke Platz^{2†}, Jeroen Dauw^{3,4†}, Jozine M. ter Maaten^{3,5}, Pieter Martens^{3,4}, Emanuele Pivetta⁶, John G.F. Cleland¹, John J.V. McMurray⁷, Wilfried Mullens^{3,8}, Scott D. Solomon², Faiez Zannad^{9,10}, Luna Gargani^{11‡}, and Nicolas Girerd^{9,10‡}

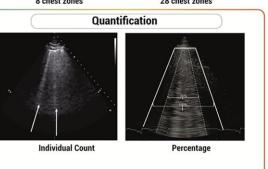


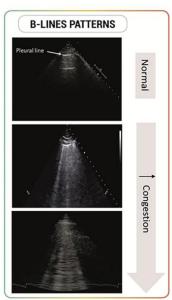


LUNG ULTRASOUND

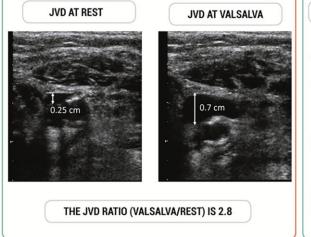


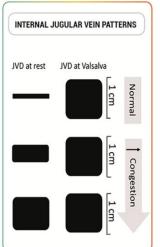
8 chest zones 28 chest zones





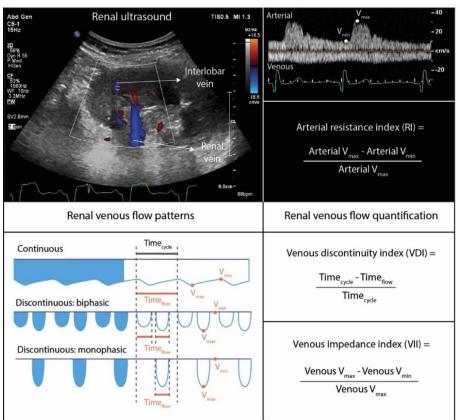
JVD ULTRASOUND

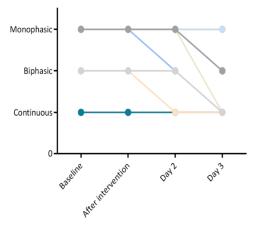




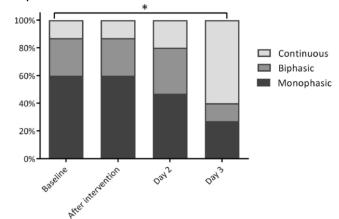
The Effect of Decongestion on Intrarenal Venous Flow Patterns in Patients With Acute Heart Failure

Assessment and calculation of renal ultrasound parameters





Changes in percentages of continuous venous flow following hemodynamic alterations. *P < .05.

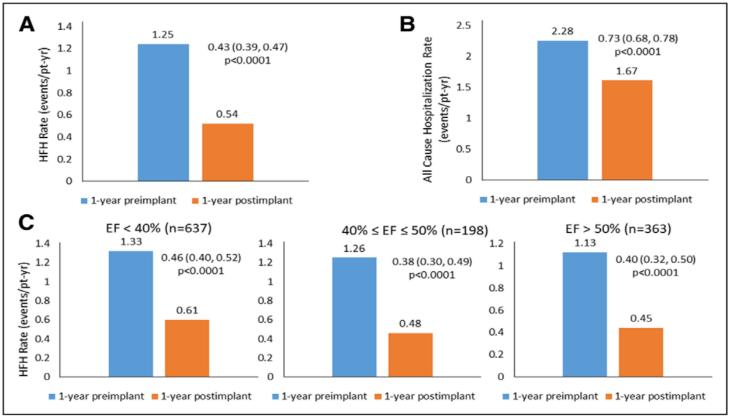


CardioMEMS

Heart Failure. Lower Rates of Heart Failure and All-Cause Hospitalizations During Pulmonary Artery Pressure-Guided Therapy for Ambulatory Heart Failure

CardioMEMS Post-Approval Study Investigators

N = 1200 patients



Haemodynamic-guided management of heart failure (GUIDE-HF): a randomised controlled trial



JoAnn Lindenfeld, Michael R Zile, Akshay S Desai, Kunjan Bhatt, Anique Ducharme, Douglas Horstmanshof, Selim R Krim, Alan Maisel, Mandeep R Mehra, Sara Paul, Samuel F Sears, Andrew J Sauer, Frank Smart, Marcel Zughaib, Paige Castaneda, Jean Kelly, Nessa Johnson, Poornima Sood, Greg Ginn, John Henderson, Philip B Adamson, Maria Rosa Costanzo

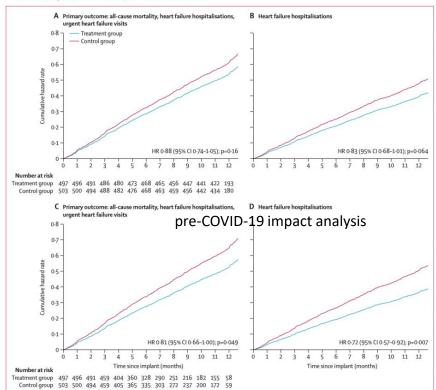


Figure 2: Cumulative hazard rate curves and 95% CIs for the primary composite endpoint and heart failure hospitalisations

Lindenfeld J, et al. Lancet. 2021;398:991-1001

GUIDE-HF trial







1000 patients with implantable pulmonary artery pressure

nonitor	n	Treatment events (rate)	Control events (rate)		Hazard ratio (95% CI)	Subgroup p value	P _{interacti}
Overall	1000	253 (0.563)	289 (0-640)	-	0-88 (0-74-1-05)	0-16	
NYHA class							
II*	296	53 (0-401)	75 (0-554)	-	0.72 (0.50-1.05)	0.086	0.095
III	650	171 (0.589)	198 (0-677)	-	0-87 (0-70-1-08)	0.21	
IV*	54	29 (1.527)	16 (0-910)	-	1.68 (0.88-3.20)	0.12	
II and III*	946	224 (0-525)	273 (0-633)	-	0.83 (0.69-1.00)	0.050	0.046
Qualification							
Heart failure hospitalisation in past year*	557	181 (0-757)	212 (0-820)		0-92 (0-74-1-14)	0-47	0.71
Elevated BNP/NT-proBNP only*	442	72 (0-350)	77 (0-409)		0.86 (0.62-1.19)	0.36	
Ejection fraction							
HFpEF (ejection fraction >40%)	469	90 (0-442)	114 (0-518)		0.85 (0.64-1.14)	0.28	0.90
HFrEF (ejection fraction ≤40%)	531	163 (0-677)	175 (0-773)		0.88 (0.70-1.10)	0-26	
Ejection fraction (additional)							
HFpEF (ejection fraction ≥50%)*	492	154 (0-693)	168 (0-810)	-	0.86 (0.68-1.08)	0-20	0.75
HFmrEF (≤40% ejection fraction <50%)*	110	26 (0.540)	25 (0-513)		1.05 (0.60-1.86)	0.86	
HFrEF (ejection fraction <40%)*	398	73 (0-420)	96 (0-505)		0.83 (0.61-1.14)	0-25	
Age (years)							
Below median (<71)	492	156 (0.730)	173 (0-758)	_	0.96 (0.76-1.22)	0.75	0.30
Median and above (≥71)	508	97 (0-420)	116 (0-529)		0-79 (0-60-1-05)	0-11	
Sex							
Male	625	178 (0-656)	171 (0-625)		1-05 (0-84-1-31)	0-67	0-010
Female	375	75 (0-434)	118 (0-681)		0.64 (0.47-0.87)	0-004	
Race				0.00			
White	807	189 (0.516)	194 (0-531)	_	0-97 (0-79-1-20)	0-78	0-095
Black	179	60 (0.811)	94 (1-184)		0-68 (0-48-0-97)	0-035	
Ethnicity				N. 50			
Hispanic	33	18 (1-257)	21 (1-383)		0.91 (0.45-1.83)	0-79	0.95
Non-Hispanic	960	232 (0.544)	264 (0-615)	-	0.88 (0.73-1.07)	0-20	
Ischaemic cardiomyopathy							
Ischaemic	397	99 (0.545)	112 (0-670)		0.81 (0.61-1.08)	0.16	0.40
Non-ischaemic	560	142 (0.578)	160 (0-607)		0.95 (0.75-1.21)	0-69	
Device implant							
With CRT-D/CRT-P/ICD	562	162 (0-633)	188 (0-759)		0.83 (0.67-1.04)	0.11	0.48
Without CRT-D/CRT-P/ICD	438	91 (0-482)	101 (0-508)		0-95 (0-71-1-28)	0.73	
				0.5 1	2 4		

THE PRESENT AND FUTURE

JACC REVIEW TOPIC OF THE WEEK: POINT

In-Hospital Initiation of Sodium-Glucose Cotransporter-2 Inhibitors for Heart Failure With Reduced Ejection Fraction



EMPULSE (NCT04157751) Randomized, double-blind study; 530 patients hospitalized for HF on day 2-5 randomized 1:1 to empagliflozin 10 mg or placebo. Patients must be stabilized with SBP >100 mm Hg, no IV vasodilators or increase in IV loop diuretics in 6 hours and no IV inotropes in 24 hours. Includes patients with HFpEF and HFrEF. eGFR ≥20 mL/min per 1.73 m2.

Primary endpoint is composite of death, number of heart failure events, time to first heart failure event, and ≥5 point KCCQ-TSS score change at 90 days using a "win-ratio" approach.

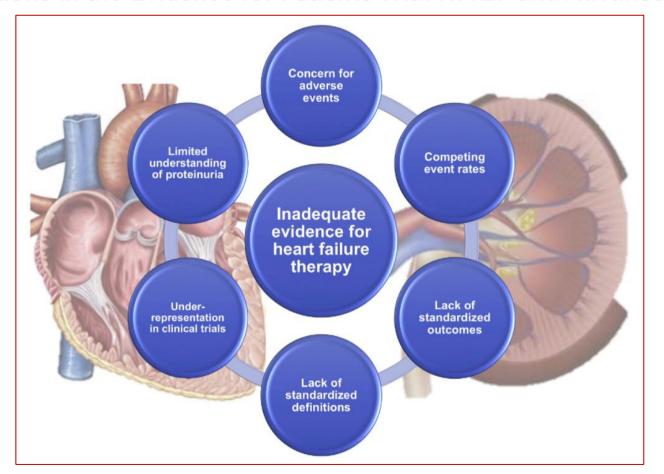
Eiection Fraction Patient-Centered Benefits Safety and Tolerability Early clinical benefit within days to weeks Favorable blood pressure and kidney of initiation Deferring in-hospital initiation exposes patients Minimal to no effect on blood pressure, and no to excess risk of early post-discharge clinical excess risk of symptomatic hypotension. No worsening, readmission, and death. adverse renal effects (instead preserves kidney function and prevents dialysis). Deferred in-hospital initiation of GDMT Favorable glycemic safety profile is associated with never initiating No excess risk of hypoglycemia in clinical trials. Among patients eligible for therapy, discharging without medication associated with >75% No excess risk of DKA in HFrEF trials chance will not be started within 1 year. (†absolute risk of DKA < 0.2% across all SGLT2i trials). Potential improved tolerance to other Well-tolerated and safe, including evidence-based therapies among high-risk subgroups MRA Hospitalized population vulnerable to in-hospital Numerically fewer serious adverse events than and post-discharge discontinuation of GDMT may placebo. Rarely symptomatic side effects and particularly benefit from + risk of hyperkalemia well tolerated among older patients. and worsening renal function.

CENTRAL ILLUSTRATION: Rationale for Routine In-Hospital Initiation of

Sodium-Glucose Cotransporter-2 Inhibitors for Heart Failure With Reduced

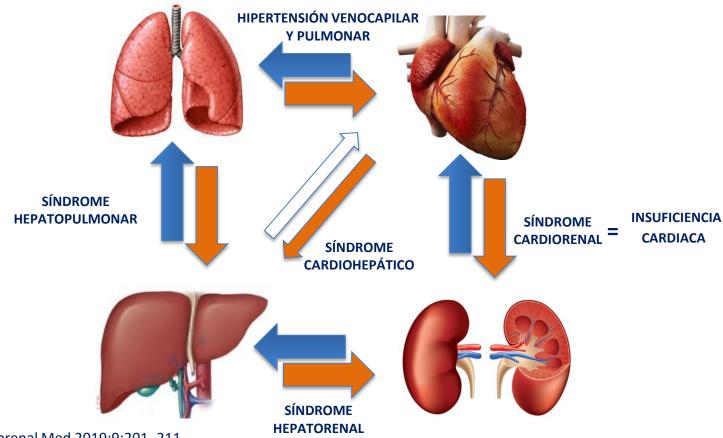
Rao, V.N. et al. J Am Coll Cardiol. 2021;78(20):2004-2012.

Limitations in the Evidence for Patients With HFrEF and Advanced CKD



INTERRELACIÓN MULTIORGÁNICA EN LA INSUFICIENCIA CARDIACA

The "organ crosstalk" is the complex biological communication between different body systems, mediated via cellular, subcellular, molecular, neural, endocrine and paracrine factors through numerous feedback ¹



1. Virzi GM, et al. Cardiorenal Med 2019;9:201–211

Multi-Omics Approach: New Potential Key Mechanisms Implicated in Cardiorenal Syndromes

Multiomic Data: genome, epigenome, metabolome, transcriptome and proteome

New mechanisms implicated in CRS crosstalk

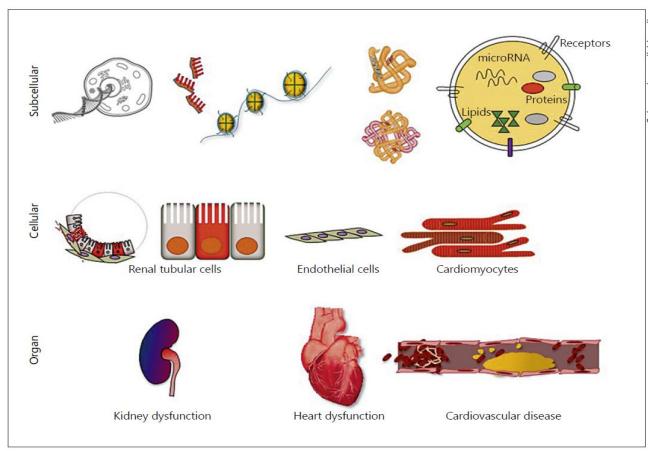
Gene expression

Epigenetic Mechanisms

Prenatal programming

Small non-coding RNA

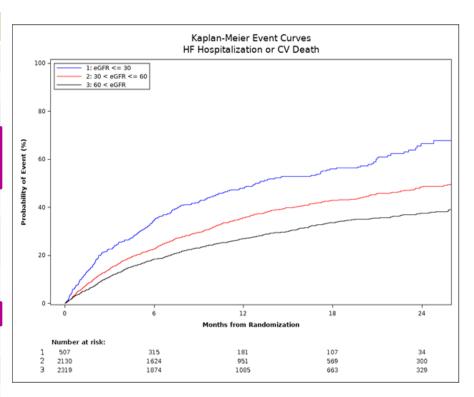
Extracellular vesicles



Vericiguat Global Study in Subjects with HFrEF

Baseline Clinical Characteristics in VICTORIA¹

Key baseline features of VICTORIA (N=5050)					
EF at screening, mean (SD), %	28.9 (8.3)				
NYHA class at baseline, No. (%)					
n	5046				
Ш	2975/ 5046 (59.0%)				
III	2003 (39.7%)				
IV	66 (1.3%)				
eGFR category at randomisation (ml/min/1.73m²)					
n	4959				
≤30	506 (10.2%)				
>30 to ≤60	2118 (42,7%)				
>60	2335 (47.1%)				
Mean (SD)	61.5 (27.2)				



1. Armstrong PW et al. N Engl J Med. 2020; doi:10.1056/NEJMoa1915928.

Renal function and the effects of vericiguat in patients with worsening heart failure with reduced ejection fraction: insights from the VICTORIA (Vericiguat Global Study in Subjects with HFrEF) trial

	eGFR≤30			eGFR>30			
		Placebo		Vericiguat	Placebo		
	Vericiguat	Rate		Rate	Rate		Interaction
Clinical Outcome	Rate* (Events)	(Events)	HR (95% CI)	(Events)	(Events)	HR (95% CI)	P-value
HF hospitalization or	64.9% (144)	61.3% (127)	1.06 (0.84 - 1.35)	30.8% (737)	35.5% (826)	0.88 (0.79 - 0.97)	0.143
CV death							
HF hospitalization or	70.4% (156)	70.0% (145)	1.01 (0.80 - 1.26)	32.7% (783)	37.2% (867)	0.89 (0.80 - 0.98)	0.306
All-cause death							
CV death	21.3% (67)	23.7% (67)	0.90 (0.64 - 1.26)	11.9% (336)	12.9% (365)	0.92 (0.80 - 1.07)	0.882

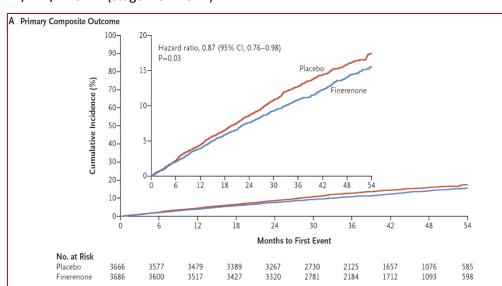
Finerenone in Reducing CV Mortality and Morbidity in Diabetic Kidney Disease - FIGARO-DKD

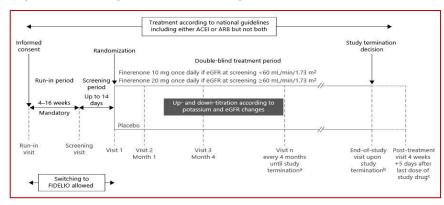
ORIGINAL ARTICLE

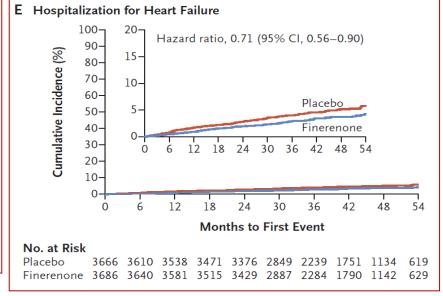
Cardiovascular Events with Finerenone in Kidney Disease and Type 2 Diabetes

B. Pitt, G. Filippatos, R. Agarwal, S.D. Anker, G.L. Bakris, P. Rossing, A. Joseph, P. Kolkhof, C. Nowack, P. Schloemer, and L.M. Ruilope, for the FIGARO-DKD Investigators*

7437 patients DM+CKD: urinary albumin-to creatinine ratio of 30 to less than 300 and an eGFR of 25 to 90 ml/min/1.73 m2 (stage 2 to 4 CKD) or a urinary albumin-to-creatinine ratio of 300 to 5000 and an eGFR of at least 60 ml/min/1.73 m2 (stage 1 or 2 CKD)







Pitt B, et al. New Egl J Med. 2021. DOI: 10.1056/NEJMoa2110956



CARDIO-NEFROLOGIA

Review

Developing the subspecialty of cardio-nephrology: The time has come. A position paper from the coordinating committee from the Working Group for Cardiorenal Medicine of the Spanish Society of Nephrology

Javier Díez^{a,b,*}, Juan F. Navarro-González^{c,d}, Alberto Ortiz^{d,e}, Rafael Santamaría ^{d,f,g}, Patricia de Sequera ^h

UNIDADES CARDIORRENALES

REC: CardioClinics

www.reccardioclinics.org

Artículo especial

Bases para la creación de las unidades clínicas cardiorrenales. Documento de consenso de los grupos de trabajo cardiorrenal de la SEC y la SEN

Rafael de la Espriella^a, Miguel González^b, José Luis Górriz^{b,c,d}, María José Soler^{c,e}, Javier Díez^{f,g}, Patricia de Sequera^h, Alberto Ortiz Arduan^{c,i}, Juan F. Navarro-González^{c,j}, Rafael Santamaría^{c,k}, Marta Cobo^{g,l} y Julio Núñez^{a,d,g,*}

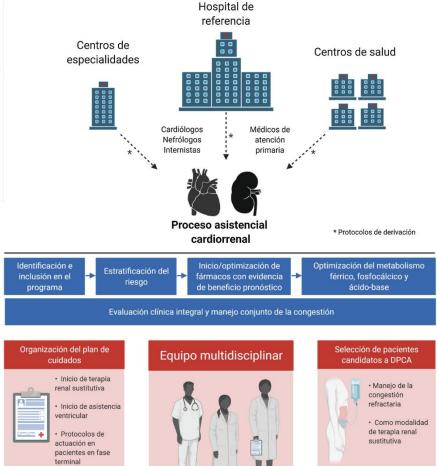
REC: CardioClinics

www.reccardioclinics.org

Artículo especial

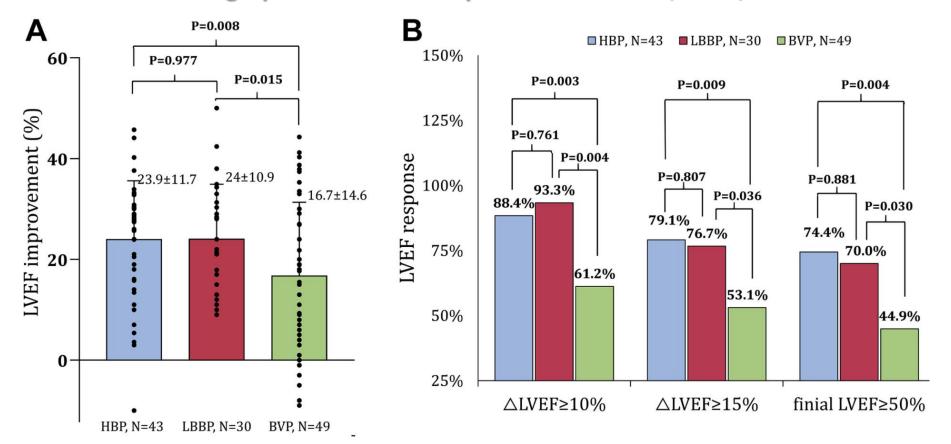
Bases para la creación de las unidades clínicas cardiorrenales. Documento de consenso de los grupos de trabajo cardiorrenal de la SEC y la SEN

Rafael de la Espriella , Miquel González , José Luis Górriz b,c,d, María José Soler c,e, Javier Díez f.g., Patricia de Sequerah, Alberto Ortiz Arduan c.i. Juan F. Navarro-González (1), Rafael Santamaría (1), Marta Cobo (9) y Julio Núñez (1), **



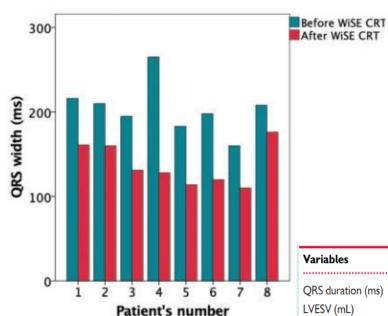


Echocardiographic outcomes in patients with HBP, LBBP, and BVP



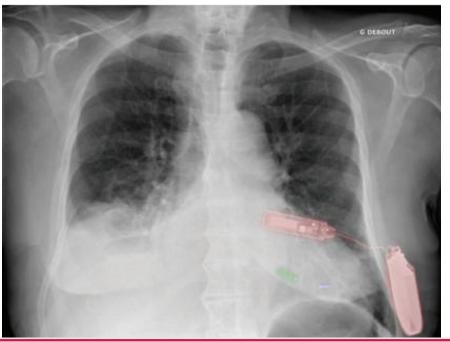
Micra and WiSECRT systems

European experience with a first totally leadless cardiac resynchronization therapy pacemaker system



Europace (2021) 23, 740-747

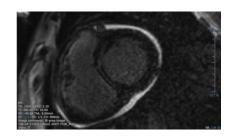
European Society doi:10.1093/europace/euaa342

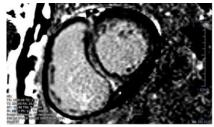


Variables	Before WiSE-CRT implantation	After WiSE-CRT implantation	Change	P-value
QRS duration (ms)	204.37 ± 30.26	137.50 ± 24.75	-66.88 ±31.58	0.012
LVESV (mL)	117.33 ± 35.61	91.86 ± 48.43	-23 ± 27.77	0.24
LVEDV (mL)	160 ± 22.69	129.4 ± 40.70	-30.60 ± 29.30	0.22
LVEF (%)	28.43 ± 8.01	39.71 ± 11.89	$+11.29 \pm 8.46$	0.018
NYHA	2.63 ± 0.51	2.29 ± 0.95		0.18

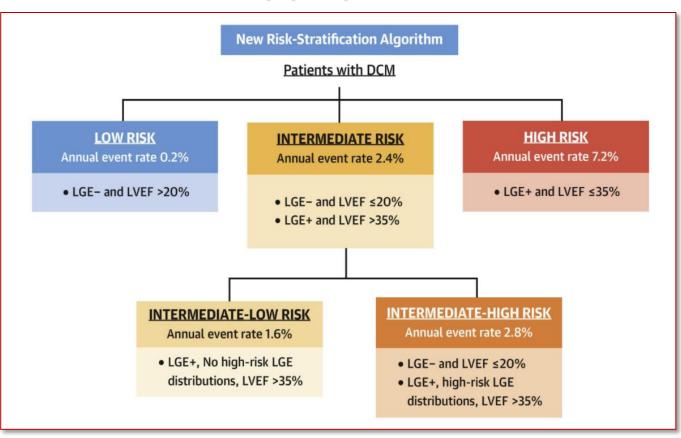
Improved Risk Stratification for Ventricular Arrhythmias and Sudden Death in Patients With Nonischemic Dilated Cardiomyopathy

Schematic Representation of the Proposed New Algorithm

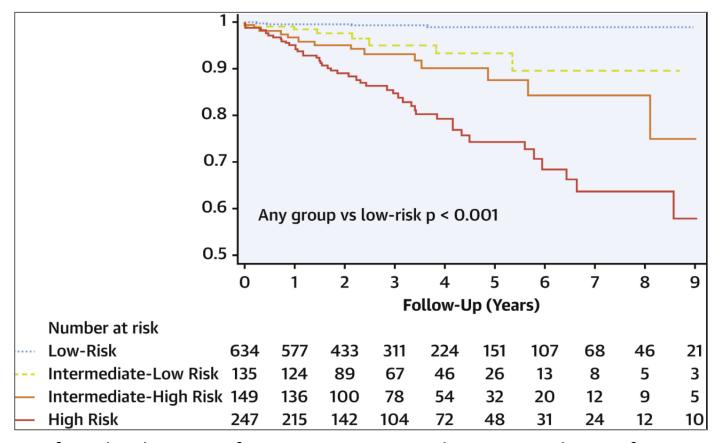




high-risk LGE: epicardial LGE, transmural LGE, or combined septal and free-wall LGE

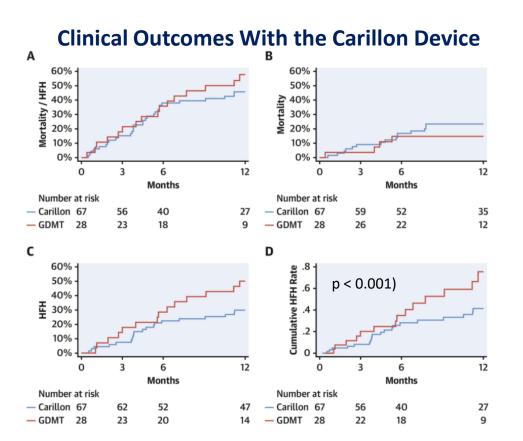


Survival Free From the Combined Arrhythmic Endpoint



LGE was found to be a significant, consistent, and strong predictor of sustained VA and sudden death

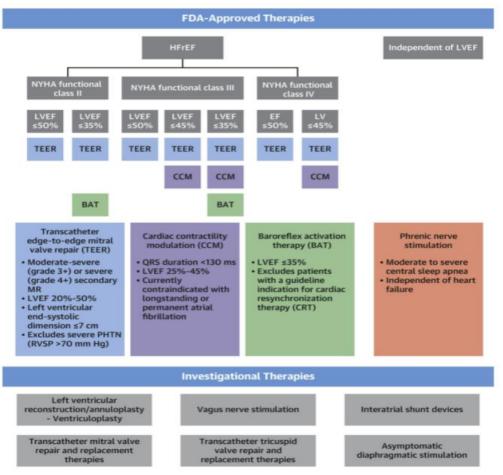
Percutaneous Mitral Valve Annuloplasty in Patients With Secondary Mitral Regurgitation and Severe Left Ventricular Enlargement (LVEDD >6.5 cm)



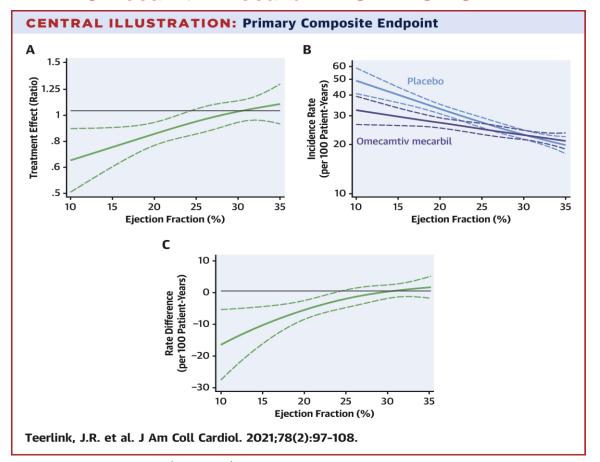


Regurgitant volume decreased by 12 ml (p < 0.001), MR grade decreased by 0.6 U (p < 0.001), LV end-diastolic volume decreased by 25 cm3 (p = 0.005), and LV end-systolic volume decreased by 21 cm3 (p = 0.01)

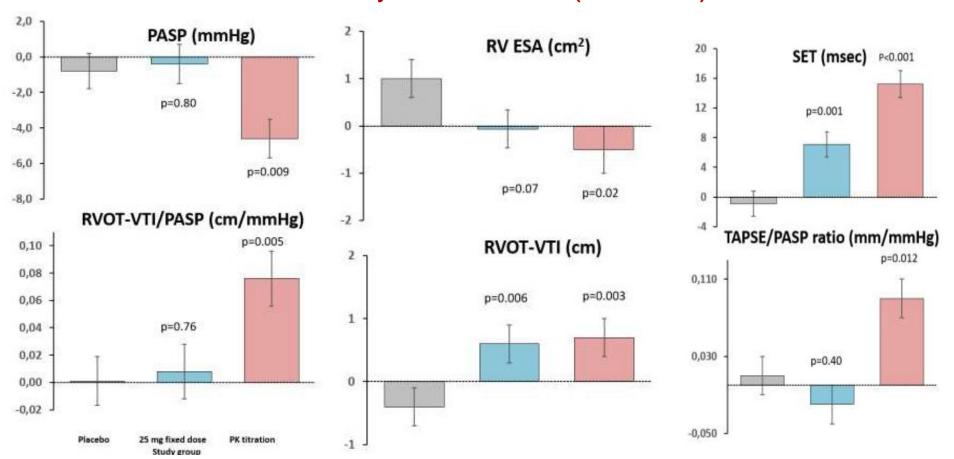
U.S. Food and Drug Administration-Approved and Breakthrough Designated Device Therapies



Effect of Ejection Fraction on Clinical Outcomes in Patients treated with Omecamtiv Mecarbil in GALACTIC-HF



Effect of the Cardiac Myosin Activator, Omecamtiv Mecarbil, on Right Ventricular Structure and Function in Chronic Systolic Heart Failure (COSMIC -HF)



Biering-Sørensen T, et al. Eur J Heart Fail. 2021. epub ahead of print

STS-INTERMACS 2020 ANNUAL REPORT



↑ Implant volumes (2019 highest annual)

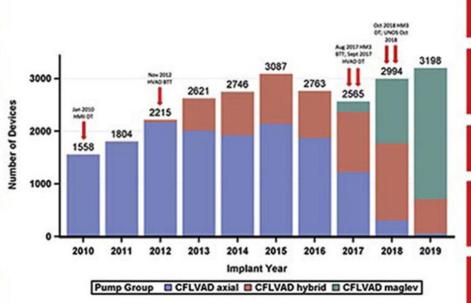
↑ African-American (27% in 2015-2019 era)

↑ temporary MCS (36.8% in 2015-2019 era)

↑ INTERMACS Profiles 1-2 (50% in 2019)

↑ Destination Therapy (73% in 2019)

↑ MagLev Technology (77% in 2019) 25,551 patients undergoing primary isolated CF-LVAD implantation between 2010-2019



Contemporary Outcomes

Improved 1- and 2-year survival: 82.3% and 73.1%

Major bleeding and infection are the leading adverse events

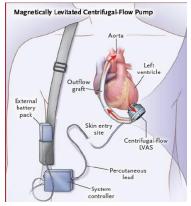
Incident stroke ↓ to 12.7% at 1-year

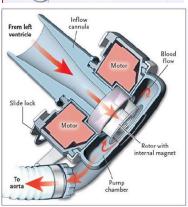
Readmission rates remain high: 38.6% at 90 days and 72.2% at 12 months

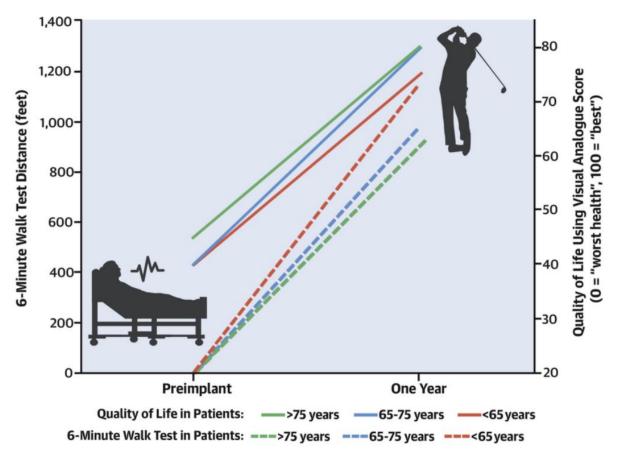
> Withdrawal of care represents a rising cause of death

Quality of Life and Functional Capacity Before and After Left Ventricular

Assist Device







Trasplante cardíaco de donantes en asistolia tipo III

CITY EAST

Central Sydney

World-first dead heart transplant at Sydney's St Vincent's Hospital a game changer

ROBBIE PATTERSON WENTWORTH COURIER OCTOBER 24, 2014 10:01AM

Adult heart transplantation with distant procurement and ex-vivo preservation of donor hearts after circulatory death: a case series



Kumud K Dhital, Arjun Iyer, Mark Connellan, Hong C Chew, Ling Gao, Aoife Doyle, Mark Hicks, Gayathri Kumarasinghe, Claude Soto, Andrew Dinale, Bruce Cartwright, Priya Nair, Emily Granger, Paul Jansz, Andrew Jabbour, Eugene Kotlyar, Anne Keogh, Christopher Hayward, Robert Graham, Phillip Spratt, Peter Macdonald

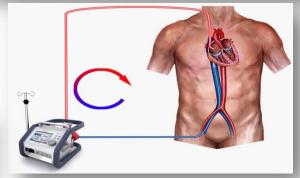
www.thelancet.com Vol 385 June 27, 2015

Success of new innovative heart transplant technique celebrated at Papworth Hospital

By CambridgeNews | Posted: December 30, 2015

By Freya Leng





Trasplante cardíaco de donantes en asistolia tipo III

LAVANGUARDIA

9 de junio 2021

Sociedad

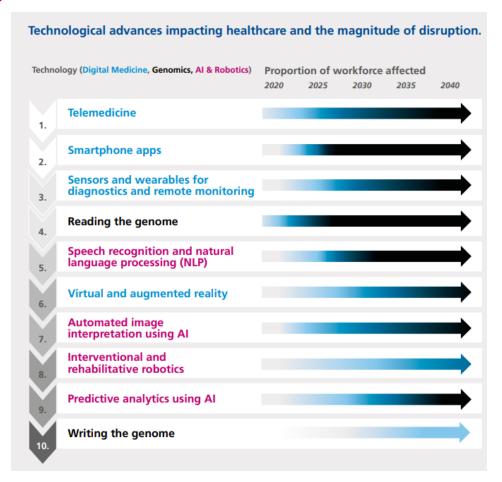
El Hospital de Bellvitge hace el primer trasplante cardíaco con corazón parado en Catalunya

• Es la octava intervención que se realiza en España con este procedimiento



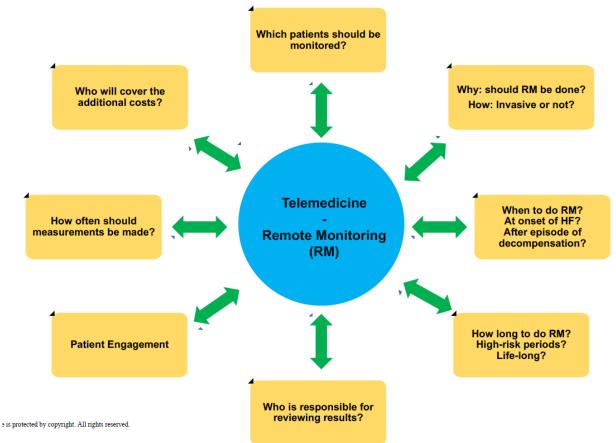


Top 10 digital healthcare technologies and their projected impact on the NHS workforce from 2020 to 2040





A Current and Future Outlook on Upcoming Technologies in the Remote Monitoring of Patients with Heart Failure



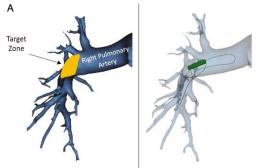


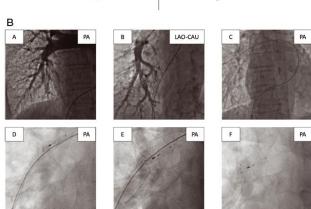
European Journal of Heart Failure (2020) doi:10.1002/eihf.1870

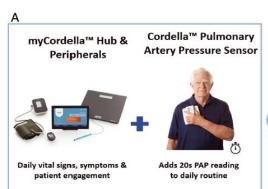


Digital health care solution for proactive heart failure management with the Cordella Heart Failure System: results of the SIRONA first-in-human study

Wilfried Mullens ^1,2 ϕ ^†, Faisal Sharif 2 †, Matthias Dupont^2, Alexander M.K. Rothman^4, and William Wijns^5

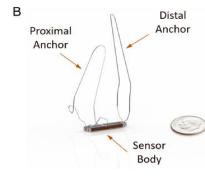


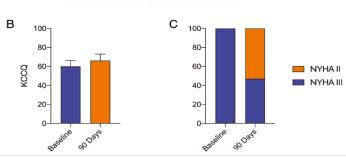












EPIC-HF: HYPOTHESIS

Patients who have been "activated" prior to a clinic appointment will be more likely to engage their provider around their HFrEF medication plan, which in turn will prompt greater optimization of HFrEF medications.

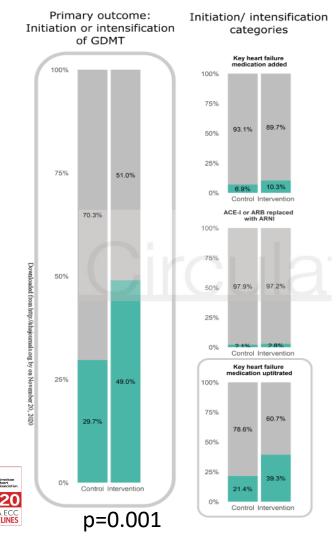






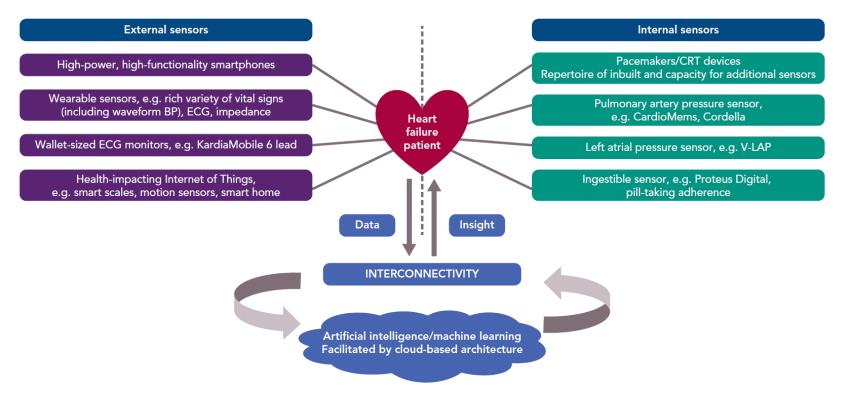


- 290 patients with heart failure with HFrEF randomized to receive UC vs additional patient engagement and education tools: A three-minute video and a one-page medication checklist delivered electronically one week, three days and 24 hours prior to a visit at a cardiology clinic.
- The median age was 65 years, 29% were female, and the median LVEF was 32%.

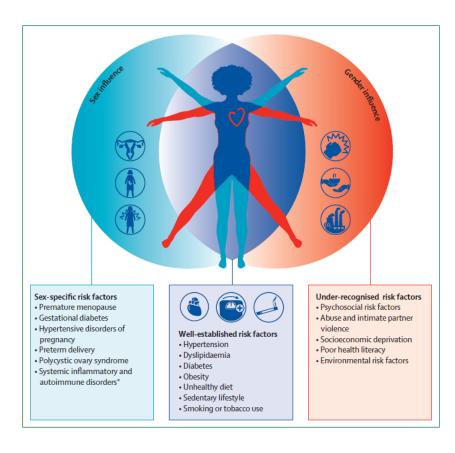


Artificial Intelligence, Data Sensors and Interconnectivity: Future Opportunities for Heart Failure

Overview of External and Internal Sensors Relevant to Heart Failure Patients



Cardiac Failure Review 2020;6:e11. **DOI**: https://doi.org/10.15420/cfr.2019.14



The Lancet women and cardiovascular disease Commission: reducing the global burden by 2030

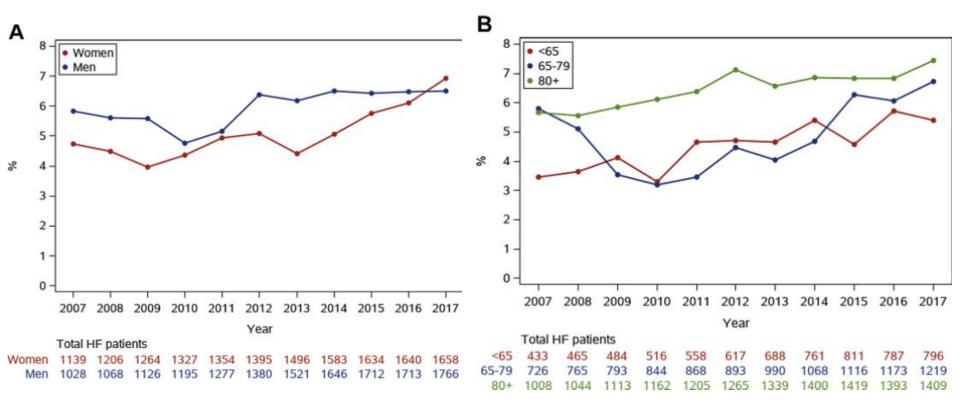


Heart failure

- The overwhelming increase in the **incidence of heart failure with preserved ejection fraction in women** with few therapeutic options underlines the importance of further research in this area
- Evidence points towards sex-specific **target doses in heart failure therapies** and should be validated in prospective, sex-specific, dose-finding studies
- Cardiac resynchronisation therapy should be offered to women with a clinical indication
- Women are more susceptible than men to **cardiogenic shock after myocardial infarction**; further research is urgently needed to investigate the underlying mechanisms
- Further research is needed to better understand the observed sex differences in the transplantation field

Advanced Heart Failure Epidemiology and Outcomes: A Population-Based Study

Prevalence of Advanced HF in Patients With HF



Dunlay SM et al. J Am Coll Cardiol HF 2021;9:722-732

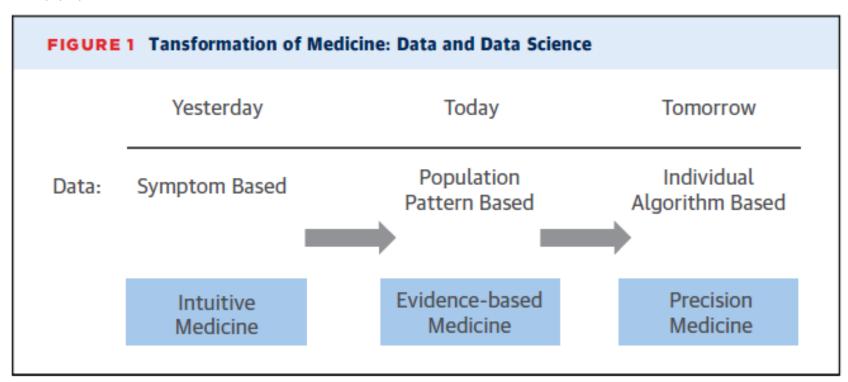
LEADERSHIP PAGE



Reading the Tea Leaves Where Will Cardiology Be in 2050?



Richard A. Chazal, MD, FACC, ACC President

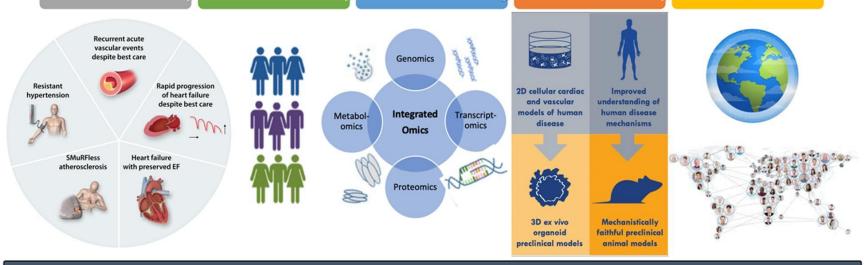


Call to action: new global approaches to cardiovascular disease drug solutions

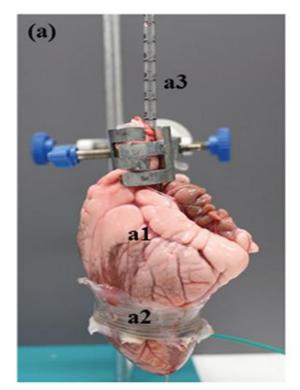


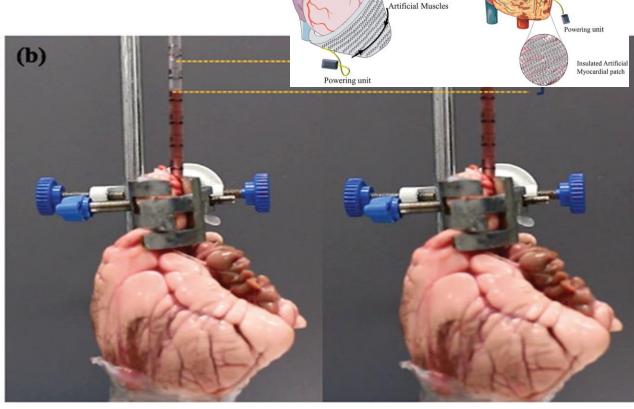
A Call to Action for New Global Approaches to Cardiovascular Disease Drug Solutions

Improved definition and focus on unmet clinical need Patient stratification in large wellphenotyped cohorts "omics" to unravel new mechanisms & drug targets in large cohorts Preclinical models reflecting human disease subtypes Clinical trial networks with innovative design for fasttrack development



High Performance Artificial Muscles to Engineer a Ventricular Cardiac Assist Device and Future Perspectives of a Cardiac Sleeve





Heart Sleeve with

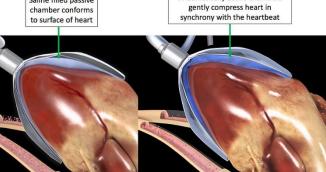
insulation material

Kongahage D, et al. Advanced Materials Technologies. 2021; DOI: (10.1002/admt.202000894)



CORINNOVA

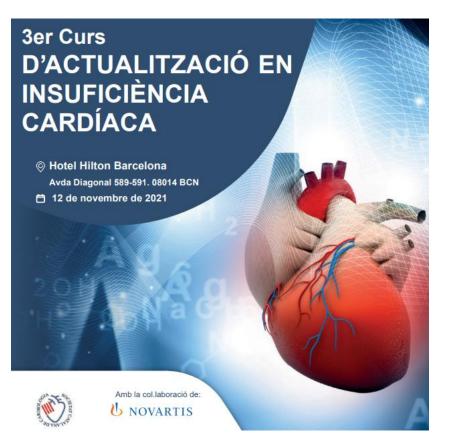
Presented November 5th at TCT 2021 session "Emerging Technological Trends III: Interventional Heart Failure and Neuromodulation Therapies"







Moltes Gracies!!





I ens veiem en el 4 Curs D'ACTUALITZACIÓ EN INSUFICIÈNCIA CARDÍACA



