



Jeudi d'Unisanté

17.11.2022

Prise en charge ambulatoire de l'HFpEF

Dr. Julien Regamey, MD
FMH cardiologie et médecine interne
Spécialiste ESC en insuffisance cardiaque
Chef de clinique
Service de cardiologie HRC et CHUV

HFpEF: définition et critères diagnostiques

- Signes et symptômes d'insuffisance cardiaque
- **FEVG \geq 50%**
- Preuve d'anomalies **structurelles** ou **fonctionnelles** en faveur d'une élévation des pressions de remplissage gauche (mesures échocardiographiques, invasives, biologiques : BNP)

Parameter ^a	Threshold	Comments
LV mass index Relative wall thickness	≥ 95 g/m ² (Female), ≥ 115 g/m ² (Male) >0.42	Although the presence of concentric LV remodelling or hypertrophy is supportive, the absence of LV hypertrophy does not exclude the diagnosis of HFpEF
LA volume index ^a	>34 mL/m ² (SR)	In the absence of AF or valve disease, LA enlargement reflects chronically elevated LV filling pressure (in the presence of AF, the threshold is >40 mL/m ²)
E/e' ratio at rest ^a	>9	Sensitivity 78%, specificity 59% for the presence of HFpEF by invasive exercise testing, although reported accuracy has varied. A higher cut-off of 13 had lower sensitivity (46%) but higher specificity (86%). ^{71,259,274}
NT-proBNP BNP	>125 (SR) or >365 (AF) pg/mL >35 (SR) or >105 (AF) pg/mL Excellence valeur prédictive <u>négative</u> !	Up to 20% of patients with invasively proven HFpEF have NPs below diagnostic thresholds, particularly in the presence of obesity
PA systolic pressure TR velocity at rest ^a	>35 mmHg >2.8 m/s	Sensitivity 54%, specificity 85% for the presence of HFpEF by invasive exercise testing ^{259,261}

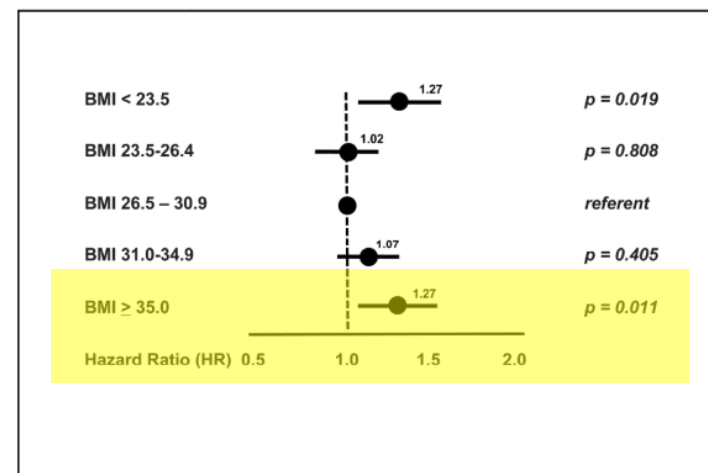
HFpEF: facteurs de risque

Table 2 | Epidemiological cohort studies reporting the incidence of HFpEF

Cohort study	Mean age at entry (years)	12-year cumulative incidence of HF (%)	Proportion with HFpEF (%)
Cardiovascular Health Study ¹¹³	73	13.7	53.3
Framingham Heart Study ^{114,115}	58	6.7	46.5
PREVEND ¹¹⁶	49	4.2	36.9

Risk factor for incident HFpEF	Subdistribution HR (95% CI)	P value
HFpEF		
Age (per 10 years)	1.90 (1.74–2.07)	<0.0001
Male sex	0.93 (0.78–1.11)	0.43
Systolic blood pressure (per 20 mmHg)	1.14 (1.05–1.24)	0.003
Body mass index (per 4 kg/m ²)	1.28 (1.21–1.37)	<0.0001
Antihypertensive treatment	1.42 (1.18–1.71)	0.0002
Previous myocardial infarction	1.48 (1.12–1.96)	0.006

Dunlay et al. Nat Rev Cardiol. 2017



Haass et al. Circ Heart Fail. 2011



HFpEF: facteurs de risque – probabilité

Table 1 Risk factors and findings consistent with heart failure with preserved ejection fraction in a symptomatic patient

Advanced age (age ≥ 70 in men or ≥ 75 in women)
 Overweight/obesity
 Metabolic syndrome/diabetes mellitus
 Physical inactivity/deconditioning
 Arterial hypertension
 Atrial fibrillation
 ECG abnormalities (beyond atrial fibrillation)
 Elevated natriuretic peptide levels (if available, BNP ≥ 35 pg/mL or NT-proBNP ≥ 125 pg/mL)

BNP, brain natriuretic peptide; NT-proBNP, N-terminal pro-brain natriuretic peptide.

ESC guidelines heart failure 2021

	Clinical Variable	Values	Points
H₂	H heavy	Body mass index > 30 kg/m ²	2
	H ypertensive	2 or more antihypertensive medicines	1
F	Atrial F ibrillation	Paroxysmal or Persistent	3
P	P ulmonary Hypertension	Doppler Echocardiographic estimated Pulmonary Artery Systolic Pressure > 35 mmHg	1
E	E lder	Age > 60 years	1
F	F illing Pressure	Doppler Echocardiographic E/e' > 9	1
H₂FPEF score			Sum (0-9)
<div> <div>Total Points</div> <div> <div>0</div> <div>1</div> <div>2</div> <div>3</div> <div>4</div> <div>5</div> <div>6</div> <div>7</div> <div>8</div> <div>9</div> </div> </div> <div> <div>Probability of HFpEF</div> <div> <div>0.2</div> <div>0.3</div> <div>0.4</div> <div>0.5</div> <div>0.6</div> <div>0.7</div> <div>0.8</div> <div>0.9</div> <div>0.95</div> </div> </div>			

Reddy et al. Circulation. 2018 Aug 28;138(9):861-870.

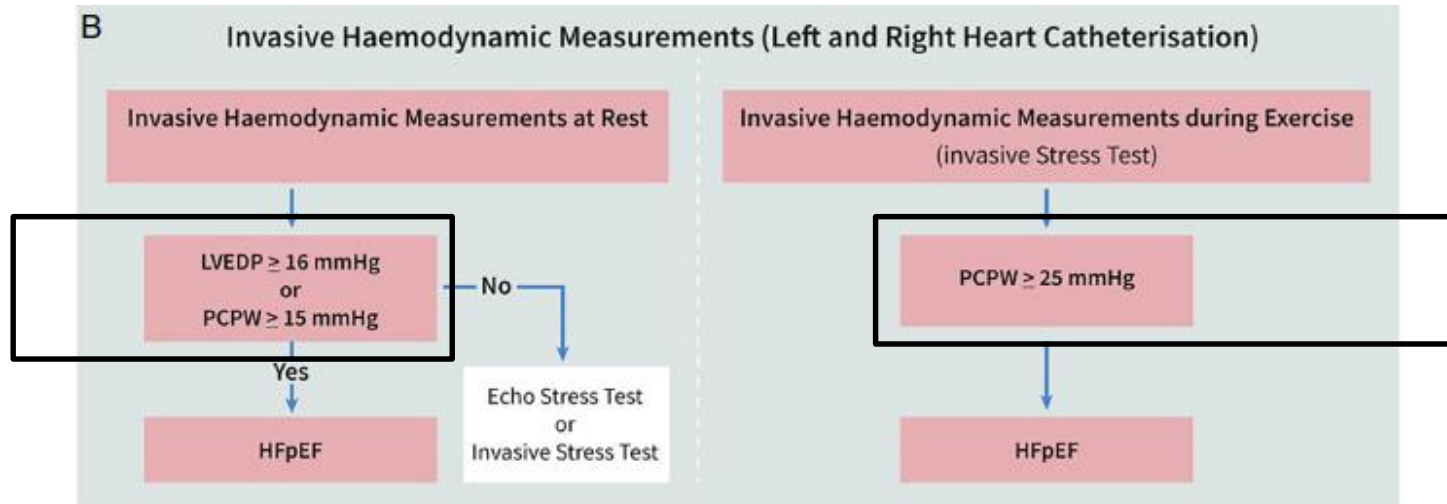
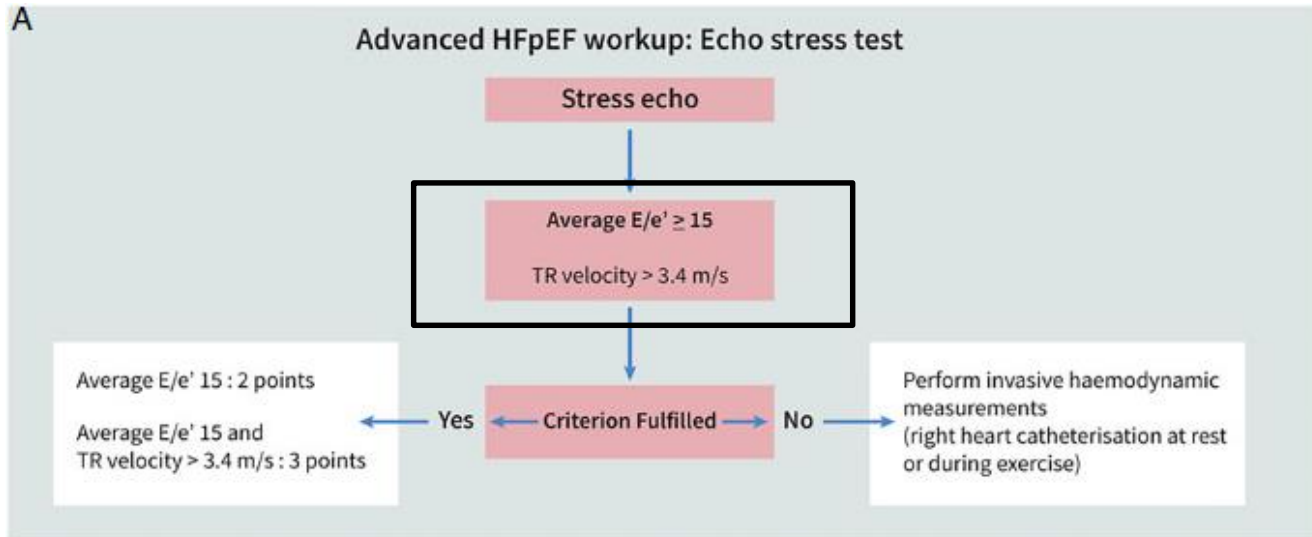
Diagnostic de l'HFpEF : un challenge – aussi pour le spécialiste

HFA-PEFF score

	Functional	Morphological	Biomarker (SR)	Biomarker (AF)
Major	septal $e' < 7$ cm/s or lateral $e' < 10$ cm/s or Average $E/e' \geq 15$ or TR velocity > 2.8 m/s (PASP > 35 mmHg)	LAVI > 34 ml/m ² or LVMI $\geq 149/122$ g/m ² (m/w) and RWT $> 0,42$ #	NT-proBNP > 220 pg/ml or BNP > 80 pg/ml	NT-proBNP > 660 pg/ml or BNP > 240 pg/ml
Minor	Average $E/e' 9 -14$ or GLS < 16 %	LAVI 29-34 ml/m ² or LVMI $> 115/95$ g/m ² (m/w) or RWT $> 0,42$ or LV wall thickness ≥ 12 mm	NT-proBNP 125-220 pg/ml or BNP 35-80 pg/ml	NT-proBNP 365-660 pg/ml or BNP 105-240 pg/ml
Major Criteria: 2 points	≥ 5 points: HFpEF			
Minor Criteria: 1 point	2-4 points: Diastolic Stress Test or Invasive Haemodynamic Measurements			

Pieske et al. Eur Heart J. 2019 Oct 21;40(40):3297-3317.

Diagnostic de l'HFpEF : examens avancés ?

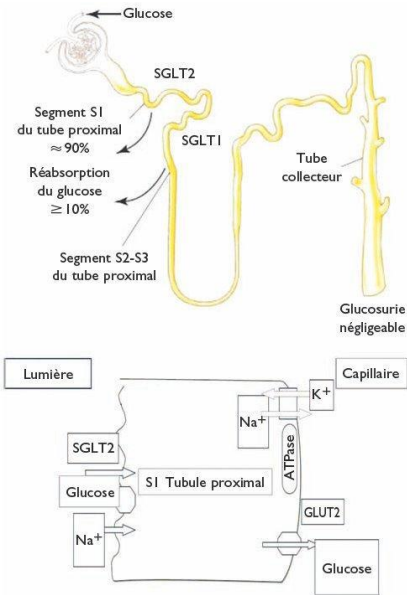


Suspicion d'HFpEF : bilan général

- **ECG 12 pistes** (FA ? HVG ? Ondes Q ...)
- **Bilan biologique :**
 - FSS (Hb !)
 - Bilan martial (fer, ferritine, transferrine, coef. sat. transferrine)
 - Fonction rénale, électrolytes, tests hépatiques
 - (NT-pro)-BNP
 - TSH
- **Rx Thorax**
- **HTA – MAPA ?**
- **+/- Fonctions pulmonaires** (particulièrement si tabagisme chronique)

*Anémie ? Carence martiale ? Pneumopathie ?
HTA ? Bien contrôlée ?*

Traitement de l'HFpEF: Empagliflozine (EMPEROR-PRESERVED)



The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

OCTOBER 14, 2021

VOL. 385 NO. 16

Empagliflozin in Heart Failure with a Preserved Ejection Fraction

S.D. Anker, J. Butler, G. Filippatos, J.P. Ferreira, E. Bocchi, M. Böhm, H.-P. Brunner-La Rocca, D.-J. Choi, V. Chopra, E. Chuquiere-Valenzuela, N. Giannetti, J.E. Gomez-Mesa, S. Janssens, J.L. Januzzi, J.R. Gonzalez-Juanatey, B. Merkely, S.J. Nicholls, S.V. Perrone, I.L. Piña, P. Ponikowski, M. Senni, D. Sim, J. Spinar, I. Squire, S. Taddei, H. Tsutsui, S. Verma, D. Vinereanu, J. Zhang, P. Carson, C.S.P. Lam, N. Marx, C. Zeller, N. Sattar, W. Jamal, S. Schnaidt, J.M. Schnee, M. Brueckmann, S.J. Pocock, F. Zannad, and M. Packer, for the EMPEROR-Preserved Trial Investigators*

Empagliflozine 10 mg/j vs placebo chez n=5988 patients :

- IC symptomatique et chronique (min. 3 mois)
- FEVG > 40% (env. 1/3 avec FEVG < 50%)
- NT-proBNP > 300 pg/ml ou > 900 pg/ml si FA
- Anomalies structurales ou s/p hospitalisation pour IC
- env. 50% de diabétiques

Exclus : eGFR < 20 ml/min, TAs < 100 mmHg

Issue primaire : mortalité CV ou hospitalisation pour IC
Suivi moyen = 26 mois

Traitement de l'HFpEF: Empagliflozine (EMPEROR-PRESERVED)

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

OCTOBER 14, 2021

VOL. 385 NO. 16

Empagliflozin in Heart Failure with a Preserved Ejection Fraction

Table 2. Primary and Secondary Cardiovascular Outcomes.*

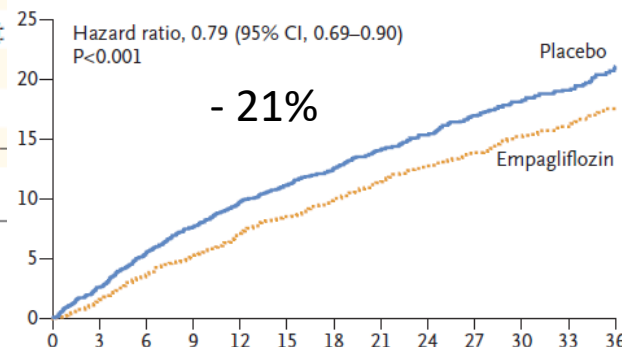
Variable	Empagliflozin (N=2997)		Placebo (N=2991)		Hazard Ratio or Difference (95% CI)	P Value
	events per 100 patient-yr		events per 100 patient-yr			
Primary composite outcome — no. (%)	415 (13.8)	6.9	511 (17.1)	8.7	0.79 (0.69–0.90)	<0.001
Hospitalization for heart failure	259 (8.6)	4.3	352 (11.8)	6.0	0.71 (0.60–0.83)	
Cardiovascular death	219 (7.3)	3.4	244 (8.2)	3.8	0.91 (0.76–1.09)	
Secondary outcomes specified in hierarchical testing procedure						
Total no. of hospitalizations for heart failure	407	—	541	—	0.73 (0.61–0.88)	<0.001
eGFR (CKD-EPI) mean slope change per year — ml/ min/1.73 m ² †	-1.25±0.11	—	-2.62±0.11	—	1.36 (1.06–1.66)	<0.001
Other prespecified analyses						
Change in KCCQ clinical summary score at 52 wk‡	—		—		1.32 (0.45–2.19)	
Total no. of hospitalizations for any cause	—		—		0.93 (0.85–1.01)	
Composite renal outcome — no. (%)	2.2		7.4		0.95 (0.73–1.24)	
Onset of new diabetes in patients with prediabetes	7.4		7.4		0.84 (0.65–1.07)	
Death from any cause — no. (%)	6.7		6.7		1.00 (0.87–1.15)	

Hazard ratio, 0.79 (95% CI, 0.69–0.90)
P<0.001

- 21%

Placebo

Empagliflozin



Traitement de l'HFpEF: Empagliflozine (EMPEROR-PRESERVED)

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

OCTOBER 14, 2021

VOL. 385 NO. 16

Empagliflozin in Heart Failure with a Preserved Ejection Fraction

S.D. Anker, J. Butler, G. Filippatos, J.P. Ferreira, E. Bocchi, M. Böhm, H.-P. Brunner-La Rocca, D.-J. Choi, V. Chopra, E. Chuquique-Valenzuela, N. Giannetti, J.E. Gomez-Mesa, S. Janssens, J.L. Januzzi, J.R. Gonzalez-Juanatey, B. Merkely, S.J. Nicholls, S.V. Perrone, I.L. Piña, P. Ponikowski, M. Senni, D. Sim, J. Spinar, I. Squire, S. Taddei, H. Tsutsui, S. Verma, D. Vinereanu, J. Zhang, P. Carson, C.S.P. Lam, N. Marx, C. Zeller, N. Sattar, W. Jamal, S. Schnaidt, J.M. Schnee, M. Brueckmann, S.J. Pocock, F. Zannad, and M. Packer, for the EMPEROR-Preserved Trial Investigators*

Subgroup	Empagliflozin no. of patients with events/total no.	Placebo no. of patients with events/total no.	Hazard Ratio (95% CI)
Overall	415/2997	511/2991	0.79 (0.69–0.90)
Diabetes at baseline			
Yes	239/1466	291/1472	0.79 (0.67–0.94)
No	176/1531	220/1519	0.78 (0.64–0.95)
LVEF at baseline			
<50%	145/995	193/988	0.71 (0.57–0.88)
≥50% to <60%	138/1028	173/1030	0.80 (0.64–0.99)
≥60%	132/974	145/973	0.87 (0.69–1.10)
Age			
<70 yr	134/1066	152/1084	0.88 (0.70–1.11)
≥70 yr	281/1931	359/1907	0.75 (0.64–0.87)
Sex			
Male	253/1659	297/1653	0.81 (0.69–0.96)
Female	162/1338	214/1338	0.75 (0.61–0.92)

Traitement de l'HFpEF: Empagliflozine (EMPEROR-PRESERVED)

TABLE S6. SELECTED ADVERSE EVENTS OF INTEREST

	Empagliflozin (n=2996) N (%)	Placebo (n=2989) N (%)
Patients with any adverse event	2574 (85.9)	2585 (86.5)
Patients with any serious adverse event	1436 (47.9)	1543 (51.6)
Selected adverse events of interest		
Hypotension	311 (10.4)	257 (8.6)
Symptomatic hypotension ^a	197 (6.6)	156 (5.2)
Acute renal failure	363 (12.1)	384 (12.8)
Ketoacidosis ^b	4 (0.1)	5 (0.2)
Hepatic injury	115 (3.8)	155 (5.2)
Hypoglycemic events ^c	73 (2.4)	78 (2.6)
In patients with diabetes mellitus	63 (4.3)	66 (4.5)
In patients without diabetes mellitus	10 (0.7)	12 (0.8)
Urinary tract infections	297 (9.9)	243 (8.1)
Complicated urinary tract infections	57 (1.9)	45 (1.5)
Genital infections	67 (2.2)	22 (0.7)
Complicated genital infections	8 (0.3)	8 (0.3)

Systolic blood pressure (mm Hg) – mean (SE)

– 1.8 ± 0.3

– 0.6 ± 0.3

– 1.2 (–2.1 to –0.3)

Traitement de l'HFpEF: Dapagliflozine (DELIVER-HF)

ORIGINAL ARTICLE

Dapagliflozin in Heart Failure with Mildly Reduced or Preserved Ejection Fraction

S.D. Solomon, J.J.V. McMurray, B. Claggett, R.A. de Boer, D. DeMets, A.F. Hernandez, S.E. Inzucchi, M.N. Kosiborod, C.S.P. Lam, F. Martinez, S.J. Shah, A.S. Desai, P.S. Jhund, J. Belohlavek, C.-E. Chiang, C.J.W. Borleffs, J. Comin-Colet, D. Dobreanu, J. Drozd, J.C. Fang, M.A. Alcocer-Gamba, W. Al Habeeb, Y. Han, J.W. Cabrera Honorio, S.P. Janssens, T. Katova, M. Kitakaze, B. Merkely, E. O'Meara, J.F.K. Saraiva, S.N. Tereshchenko, J. Thierer, M. Vaduganathan, O. Vardeny, S. Verma, V.N. Pham, U. Wilderäng, N. Zaozerska, E. Bachus, D. Lindholm, M. Petersson, and A.M. Langkilde, for the DELIVER Trial Committees and Investigators*

Dapagliflozine 10 mg/j vs placebo chez n=6263 patients :

- IC chronique symptomatique (min. 3 mois)
- FEVG > 40% (env. 1/3 avec FEVG < 50%)
 - 18% avec FE préalablement < 40% («HFimpEF»)
- NT-proBNP > 300 pg/ml ou > 600 pg/ml si FA
- Anomalies structurelles (OG, VG)
- Env. 50% de diabétiques

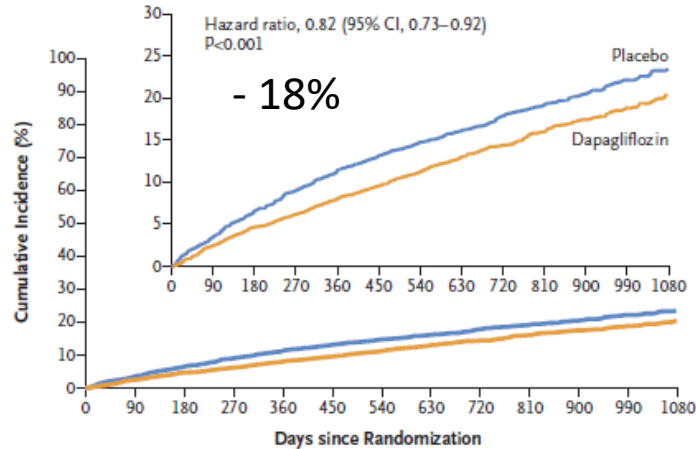
Exclus : eGFR <25 ml/min, TAs <95 mmHg

Issue primaire : mortalité CV ou hospit. pour IC ou [visite urgente](#)

Suivi moyen = 2.3 ans

Traitement de l'HFpEF: Dapagliflozine (DELIVER-HF)

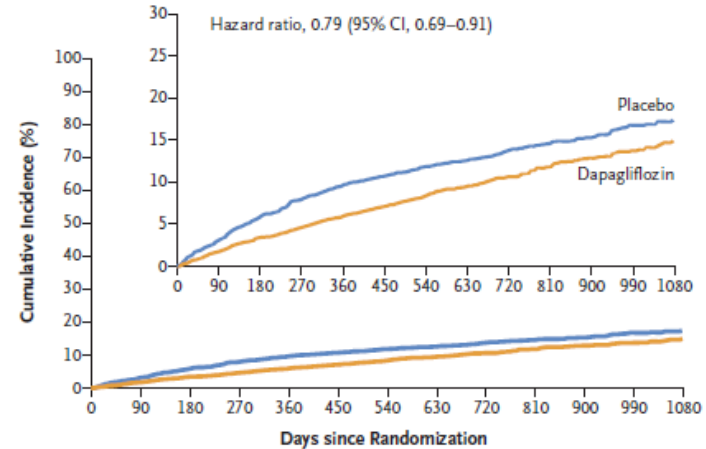
A Primary Outcome



No. at Risk

Placebo	3132	3007	2896	2799	2710	2608	2318	2080	1923	1554	1140	772	383
Dapagliflozin	3131	3040	2949	2885	2807	2716	2401	2147	1982	1603	1181	801	389

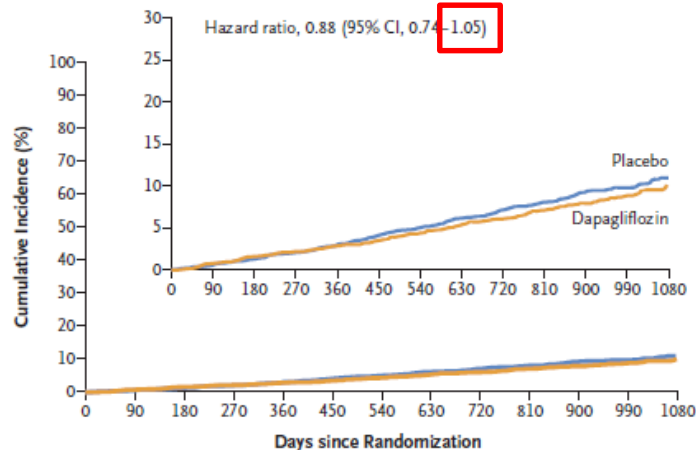
B Worsening Heart Failure Event



No. at Risk

Placebo	3132	3007	2896	2799	2710	2608	2318	2080	1923	1554	1140	772	383
Dapagliflozin	3131	3040	2949	2885	2807	2716	2401	2147	1982	1603	1181	801	389

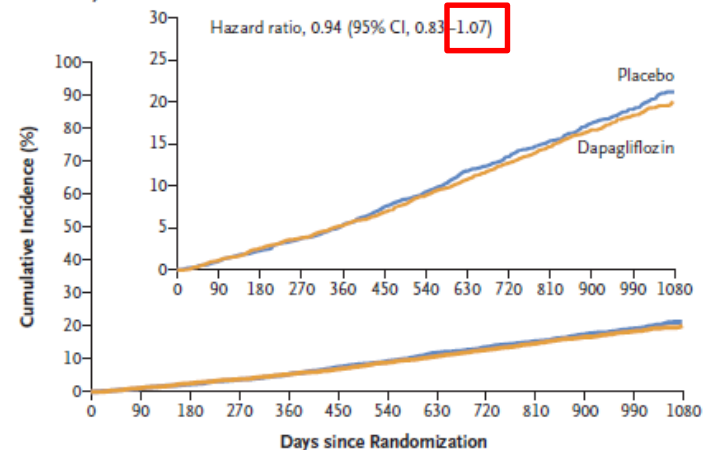
C Death from Cardiovascular Causes



No. at Risk

Placebo	3132	3096	3054	3008	2957	2872	2570	2314	2157	1759	1306	910	451
Dapagliflozin	3131	3091	3046	3006	2960	2892	2584	2339	2171	1775	1312	903	441

D Death from Any Cause



No. at Risk

Placebo	3132	3097	3058	3012	2962	2877	2575	2319	2161	1762	1309	910	451
Dapagliflozin	3131	3093	3048	3009	2962	2895	2587	2342	2174	1778	1314	905	443

Traitement de l'HFpEF: Dapagliflozine (DELIVER-HF)

ORIGINAL ARTICLE

Dapagliflozin in Heart Failure with Mildly Reduced or Preserved Ejection Fraction

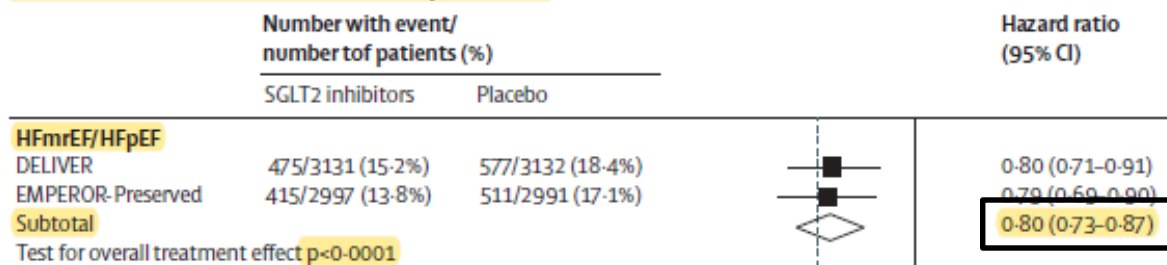
Subgroup	Dapagliflozin no. of patients with events/total no.	Placebo no. of patients with events/total no.	Hazard Ratio (95% CI)
All patients	512/3131	610/3132	0.82 (0.73–0.92)
Age			
≤72 yr	247/1545	306/1604	0.82 (0.69–0.97)
>72 yr	265/1586	304/1528	0.81 (0.69–0.96)
Sex			
Female	195/1364	243/1383	0.81 (0.67–0.97)
Male	317/1767	367/1749	0.82 (0.71–0.96)
Race			
Asian	97/630	106/644	0.91 (0.69–1.21)
Black	21/81	19/78	1.08 (0.58–2.01)
White	372/2214	461/2225	0.79 (0.69–0.90)
Other	22/206	24/185	0.83 (0.46–1.48)
Geographic region			
Europe or Saudi Arabia	261/1494	309/1511	0.83 (0.70–0.98)
Asia	92/607	103/619	0.89 (0.67–1.18)
Latin America	70/602	87/579	0.78 (0.57–1.07)
North America	89/428	111/423	0.75 (0.57–1.00)
NYHA class at enrollment			
II	331/2314	411/2399	0.81 (0.70–0.94)
III or IV	181/817	198/732	0.80 (0.65–0.98)
LVEF at enrollment			
≤49%	207/1067	229/1049	0.87 (0.72–1.04)
50–59%	174/1133	211/1123	0.79 (0.65–0.97)
≥60%	131/931	170/960	0.78 (0.62–0.98)

Traitement de l'HFpEF: meta-analyse EMPEROR-P + DELIVER

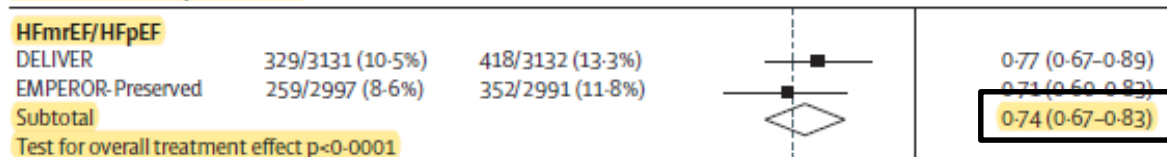
SGLT-2 inhibitors in patients with heart failure: a comprehensive meta-analysis of five randomised controlled trials

Muthiah Vaduganathan*, Kieran F Docherty*, Brian L Claggett, Pardeep S Jhund, Rudolf A de Boer, Adrian F Hernandez, Silvio E Inzucchi, Mikhail N Kosiborod, Carolyn S P Lam, Felipe Martinez, Sanjiv J Shah, Akshay S Desai, John J V McMurray†, Scott D Solomon†

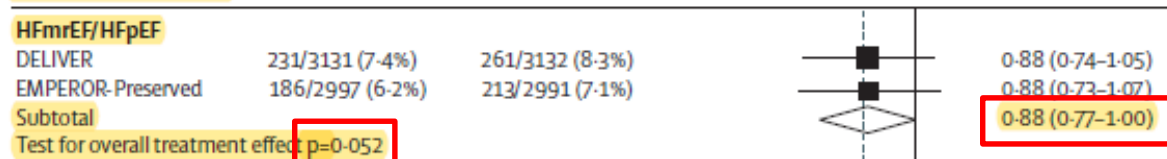
Cardiovascular death or heart failure hospitalisation



Heart failure hospitalisation



Cardiovascular death



Traitement de l'HFpEF: sacubitril-valsartan (PARAGON-HF) ?

The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

OCTOBER 24, 2019

VOL. 381 NO. 17

Angiotensin–Neprilysin Inhibition in Heart Failure with Preserved Ejection Fraction

S.D. Solomon, J.J.V. McMurray, I.S. Anand, J. Ge, C.S.P. Lam, A.P. Maggioni, F. Martinez, M. Packer, M.A. Pfeffer, B. Pieske, M.M. Redfield, J.L. Rouleau, D.J. van Veldhuisen, F. Zannad, M.R. Zile, A.S. Desai, B. Claggett, P.S. Jhund, S.A. Boytsov, J. Comin-Colet, J. Cleland, H.-D. Düngen, E. Goncalvesova, T. Katova, J.F. Kerr Saraiva, M. Lelonek, B. Merkely, M. Senni, S.J. Shah, J. Zhou, A.R. Rizkala, J. Gong, V.C. Shi, and M.P. Lefkowitz, for the PARAGON-HF Investigators and Committees*

Sac-val 200 mg 2x/j vs valsartan 160 mg 2x/j
n=4822

- LVEF $\geq 45\%$ + dilatation OG ou HVG
- NT-proBNP > 300 ng/l (>900 si FA)
et > 200 ng/l (>600 si FA) si hosp derniers 9 mois

Excl: eGFR < 30 ml/min; K⁺ ≥ 5.2 mM

Issue primaire : mortalité CV + hosp. pour IC

Suivi moyen : 3.

Table 2. Primary and Secondary Outcomes.*

Outcome	Sacubitril–Valsartan (N=2407)	Valsartan (N=2389)	Ratio or Difference (95% CI)
Primary composite outcome and components			
Total hospitalizations for heart failure and death from cardiovascular causes†			RR, 0.87 (0.75–1.01)
Total no. of events	894	1009	
Rate per 100 patient-yr	12.8	14.6	
Total no. of hospitalizations for heart failure	690	797	RR, 0.85 (0.72–1.00)
Death from cardiovascular causes — no. (%)	204 (8.5)	212 (8.9)	HR, 0.95 (0.79–1.16)
Secondary outcomes			
Change in NYHA class from baseline to 8 mo — no./total no. (%)			OR, 1.45 (1.13–1.86)
Sex			
Improved	Male	980/2317	1.03 (0.85–1.25)
Unchanged	Female	923/2479	0.73 (0.59–0.90)
Worsened	Left ventricular ejection fraction		
Change in KCCQ clinical summary score at 8 mo‡	≤Median (57%)	1048/2495	0.78 (0.64–0.95)
Renal composite outcome — no. (%)§	>Median (57%)	855/2301	1.00 (0.81–1.23)
Death from any cause — no. (%)	342 (14.2)	349 (14.6)	HR, 0.97 (0.84–1.13)

p=0.056

Traitement de l'HFpEF: spironolactone (TOPCAT) ?

Spironolactone 12.5-50 mg vs placebo chez n=3445

- LVEF $\geq 45\%$
- NT-proBNP > 360 ng ou hosp. 60 jours avant randomisation (env. 70%!)

Excl: eGFR < 30 ml/min; K⁺ ≥ 5 mM

Issue primaire : mortalité CV + hospit. Pour IC + mort subite avortée

Suivi moyen : 3.5 ans

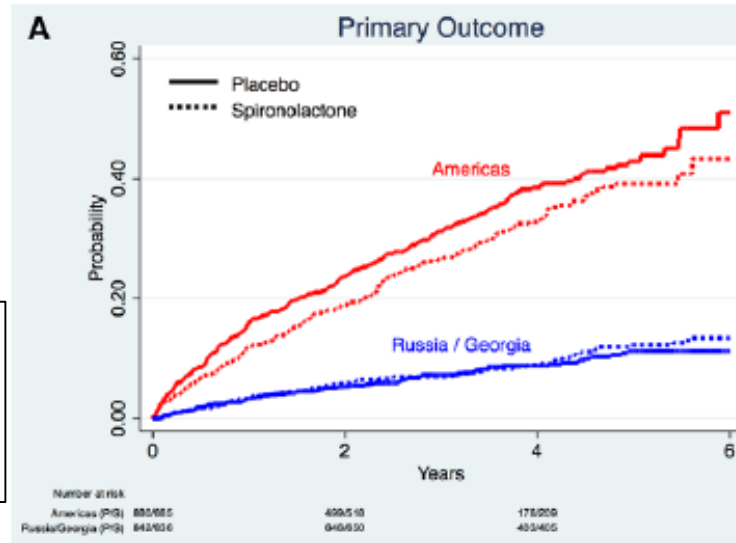
Table 2. Incidence Rates of the Primary Composite Outcome, Its Components, and Additional Secondary Outcomes.*

Outcome	Spironolactone (N=1722)		Placebo (N=1723)		Hazard Ratio with Spironolactone (95% CI)†	P Value
	Participants with Event	Incidence Rate	Participants with Event	Incidence Rate		
	no. (%)	no./100 person-yr	no. (%)	no./100 person-yr		
Primary outcome	320 (18.6)	5.9	351 (20.4)	6.6	0.89 (0.77–1.04)	0.14
Components of the primary outcome						
Death from cardiovascular causes	160 (9.3)	2.8	176 (10.2)	3.1	0.90 (0.73–1.12)	0.35
Aborted cardiac arrest	3 (0.2)	0.05	5 (0.3)	0.09	0.60 (0.14–2.50)	0.48
Hospitalization for heart failure	206 (12.0)	3.8	245 (14.2)	4.6	0.83 (0.69–0.99)	0.04
Additional secondary outcomes						
Death from any cause	252 (14.6)	4.2	274 (15.9)	4.6	0.91 (0.77–1.08)	0.29
Hospitalization for any reason	766 (44.5)	18.8	792 (46.0)	20.0	0.94 (0.85–1.04)	0.25
Myocardial infarction	65 (3.8)	1.2	64 (3.7)	1.1	1.00 (0.71–1.42)	0.98
Stroke	57 (3.3)	1.0	60 (3.5)	1.1	0.94 (0.65–1.35)	0.73

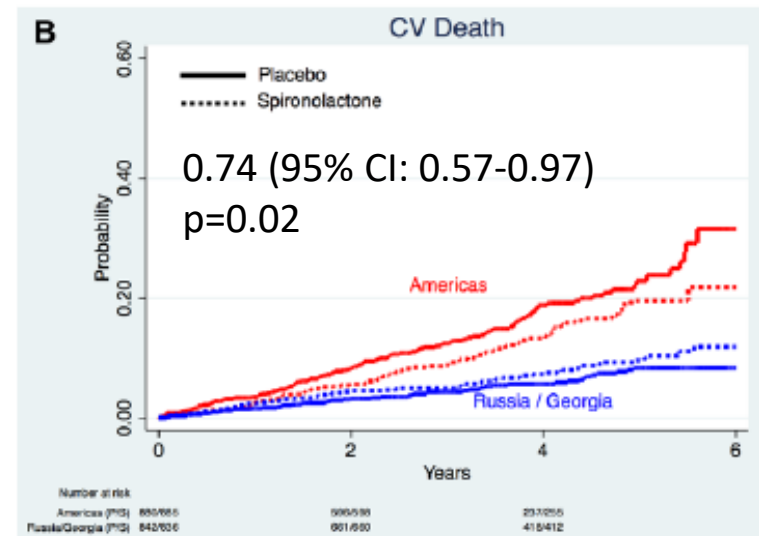
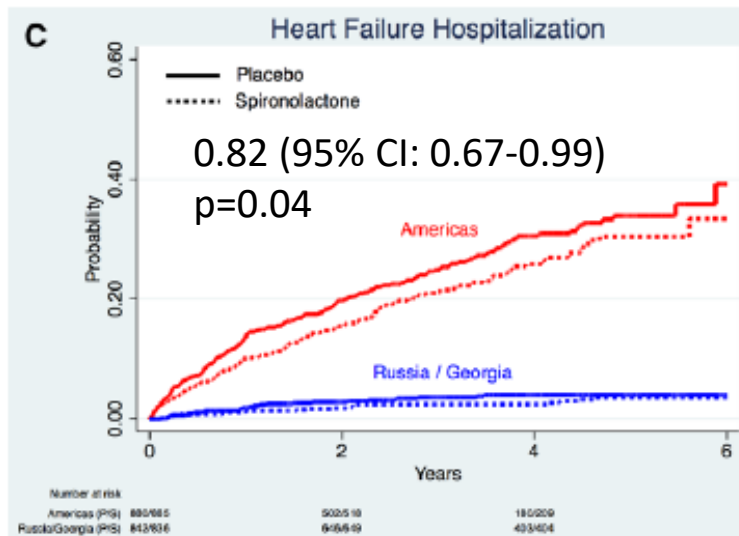
Traitement de l'HFpEF: spironolactone (TOPCAT)

Analyse post-hoc USA vs Russie/Georgie

N.B.: Efficacité plus importante pour les FEVG sur le versant bas.
Solomon et al. Eur Heart J. 2016 Feb 1;37(5):455-62.



0.82 (95% CI: 0.69-0.98)
p=0.02



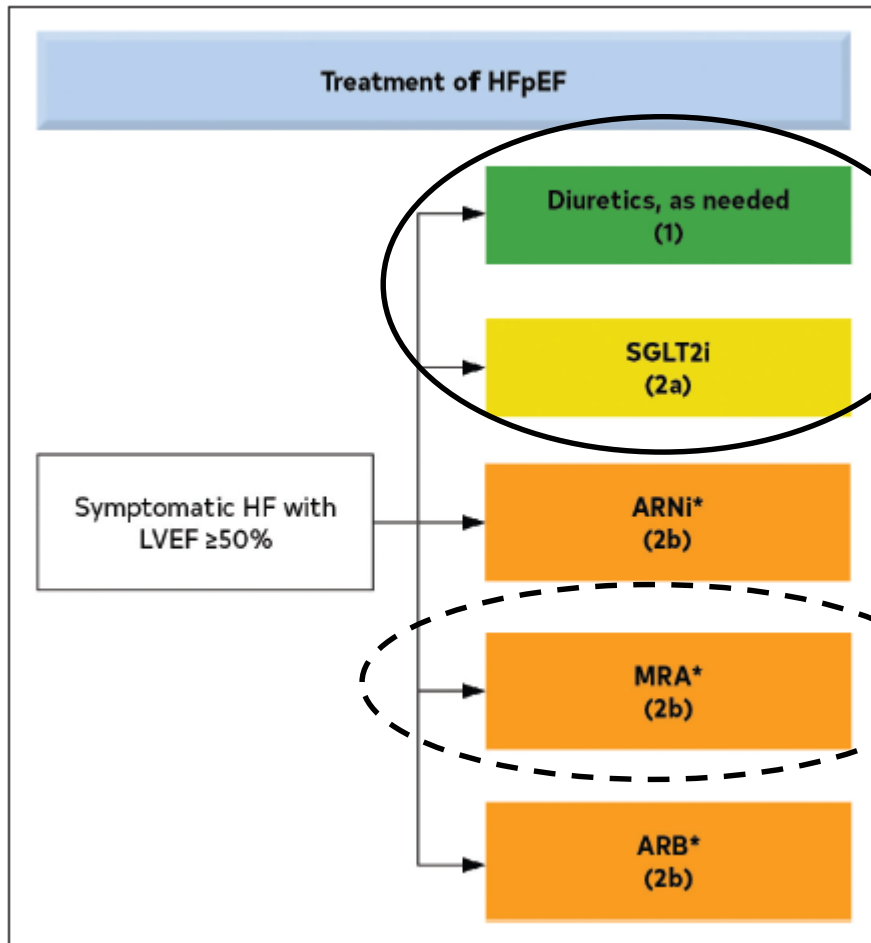
Traitement pharmacologique de l'HFpEF: CH: 2022: résumé

HFpEF (FEVG ≥ 50%)	Hospitalisations + mortalité CV	Hospitalisations seules	Mortalité CV seule	Remboursement CH
Torsemide (Torem®)	-	✓	-	✓
Empagliflozine (Jardiance®)	-21%	-29%	ns*	✓
Dapagliflozine (Forxiga®)	-18%	-21%	ns*	! Uniquement si diabétique !
Spironolactone (Aldactone®)	ns/(-18% USA)	-17%/(-18% USA)	ns/(-26% USA)	✓
Sacubitril- valsartan (Entresto®)	ns	ns°	ns	X

*Tendance à une réduction de mortalité CV dans méta-analyse: 0.88 (95% CI: 0.77-**1.00**; p= 0.052)

°Tendance à une réduction des hospitalisations (p= 0.056) : remboursé par la FDA aux USA !

Traitement pharmacologique de l'HFpEF: USA 2022



! Non-remboursé en Suisse pour l'HFpEF !

Si accord préalable de l'assurance, discutable pour réduire les hospitalisations pour les *FEVG versant bas*, *femmes* ou si *HTA sévère*...

A considérer pour réduire les hospitalisations. En particulier en cas d'HTA «résistante».

CAVE: IRC et hyperkaliémie !

Faible réduction des hospitalisations sous candesartan dans CHARM-Preserved (FEVG > 40%): HR 0.84; 95% CI, 0.70–**1.00**; p=0.047

SGLT2-i: fiche pratique

Jardiance® 10 mg ou combiné **Jardiance met®** (5/500, 5/750, 5/1000 mg)

Forxiga® 10 mg/j ou combiné **Xigduo XR®** (5/500, 5/1000 mg)

Posologie unitaire dans l'indication insuffisance cardiaque : 10 mg/j.

Contre-indications principales :

- **Diabète de type I ou diabète «insulinopénique»** : risque d'acido-cétose diabétique euglycémique.
- Jeûn prolongé (p.ex. péri-opératoire, alcoolisme, malnutrition)
- eGFR < 20 ml/min (absence de données...)

Précautions à l'introduction :

- Risque de mycose génitale >> infection urinaire : informer le patient et l'encourager à une bonne hygiène intime.
- Effet diurétique synergique :
 - Suivi rapproché +/- adaptation des diurétiques (CAVE: IRC sévère)
- Avis diabétologique si doute quant à l'origine du diabète (insulinopénie ?)
- Faible effet hypotenseur...majoré en cas d'hypovolémie

Spironolactone : fiche pratique

Aldactone® 25 ou 50 mg

Posologie dans l'IC : 12.5-50 mg/j

Contre-indications principales :

- K⁺ de base ≥ 5.0 mM
- **Prudence si eGFR < 30 ml/min** : nécessite un suivi biologique régulier et év. mise en suspens en situation favorisant l'hypovolémie (canicule, fièvre, diarrhées, vomissements, patient âgé sans réflexe de soif, etc...)
- Antécédent de gynécomastie

Précautions :

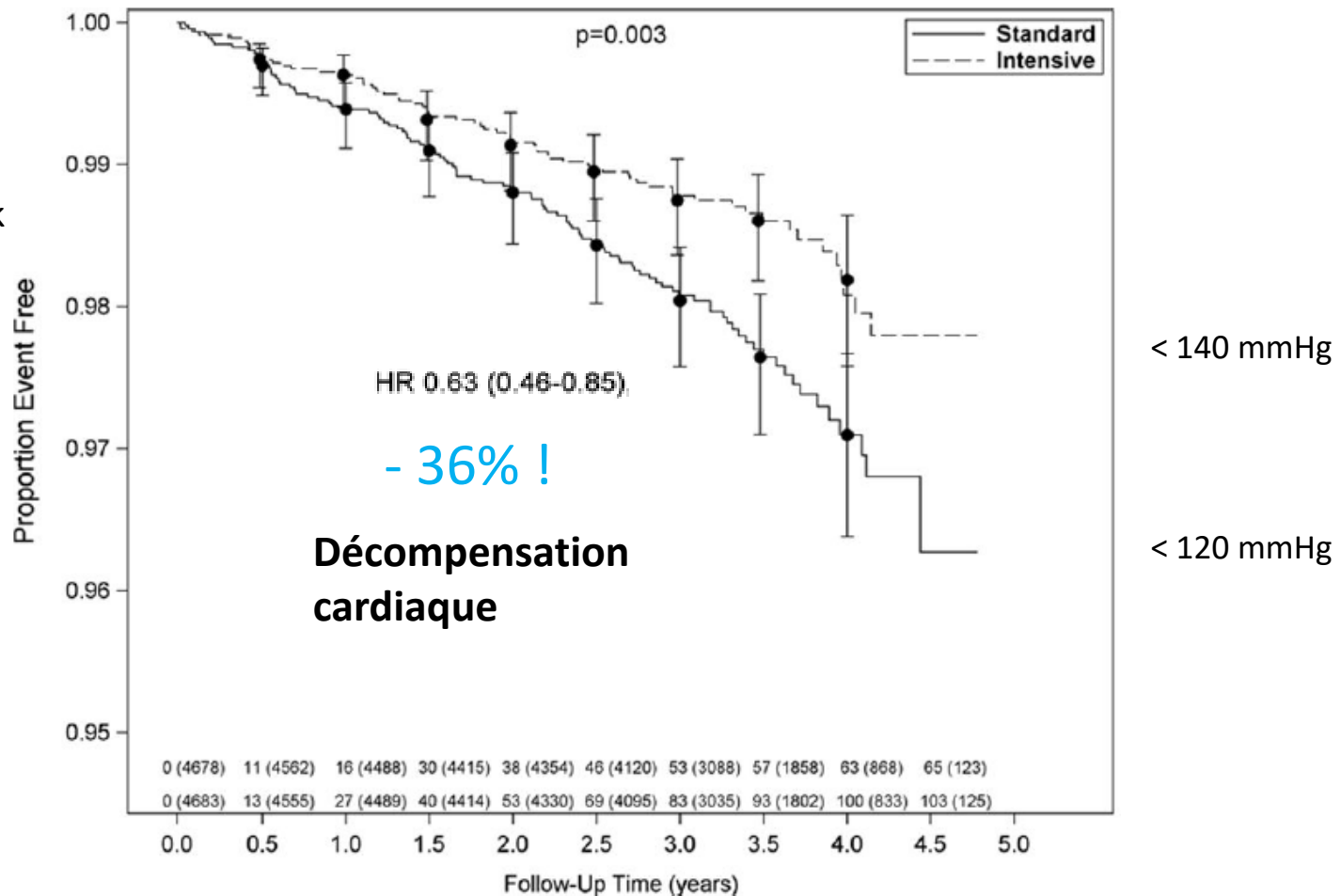
- Contrôle du K⁺ à 10-15 jours de l'introduction :
 - Si K⁺ 5.0-5.0 : pas d'adaptation, suivi régulier.
 - Si K⁺ > 5.5 mM : réduction de posologie (12.5 mg/j ou 1 jour sur 2)
 - Si K⁺ > 6.0 mM : stop et traitement spécifique

HFpEF : importance du contrôle de l'hypertension

SPRINT

N=9361

CVD or
High CV risk



Excluded: symptomatic HF or LVEF <35%, history of stroke or diabetes.

HFpEF : cible de pression artérielle

Recommendation for Prevention

COR	LOE	Recommendations	Comment/Rationale
I	B-R	In patients at increased risk, stage A HF, the optimal blood pressure in those with hypertension should be less than 130/80 mm Hg.^{189–193}	NEW: Recommendation reflects new RCT data.
See Online Data Supplements E and F.			
A large RCT demonstrated that in those with increased cardiovascular risk (defined as age >75 years, established vascular disease, chronic renal disease, or a Framingham Risk Score >15%), control of blood pressure to a goal systolic pressure of <120 mm Hg, as determined by blood pressure assessment as per research protocol, was associated with a significant reduction in the incidence of HF ¹⁹¹ and an overall decrease in cardiovascular death. Blood pressure measurements as generally taken in the office setting are typically 5 to 10 mm Hg higher than research measurements; thus, the goal of <130/80 mm Hg is an approximation of the target blood pressure in conventional practice. <i>Targeting a significant reduction in systolic blood pressure in those at increased risk for cardiovascular disease is a novel strategy to prevent HF.</i>			

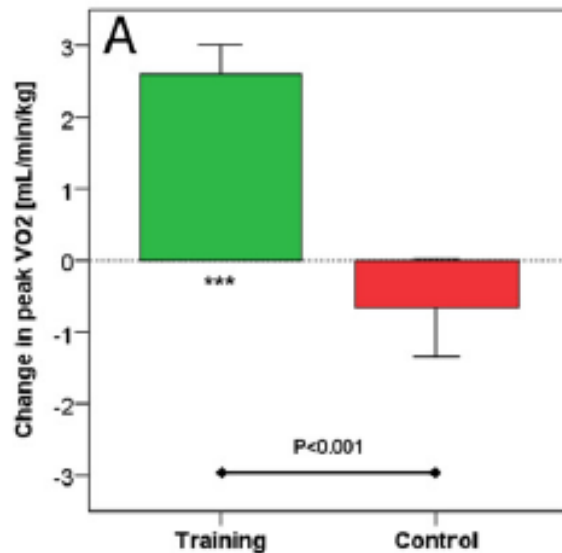
Recommendation for Hypertension in Stage C HFpEF

COR	LOE	Recommendation	Comment/Rationale
I	C-LD	Patients with HFpEF and persistent hypertension after management of volume overload should be prescribed GDMT titrated to attain systolic blood pressure less than 130 mm Hg.^{9,167,169,170,195–199}	NEW: New target goal blood pressure based on updated interpretation of recent clinical trial data.
See Online Data Supplements E and F.			
The use of nitrates in the setting of HFpEF is associated with a signal of harm and, in most situations, should be avoided. For many common antihypertensive agents, including alpha blockers, beta blockers, and calcium channel blockers, there are limited data to guide the choice of antihypertensive therapy in the setting of HFpEF. ¹⁷² Nevertheless, RAAS inhibition with ACE inhibitor, ARB (especially mineralocorticoid receptor antagonists), and possibly ARNI would represent the preferred choice. A shared decision-making discussion with the patient influenced by physician judgment should drive the ultimate choice of antihypertensive agents.			

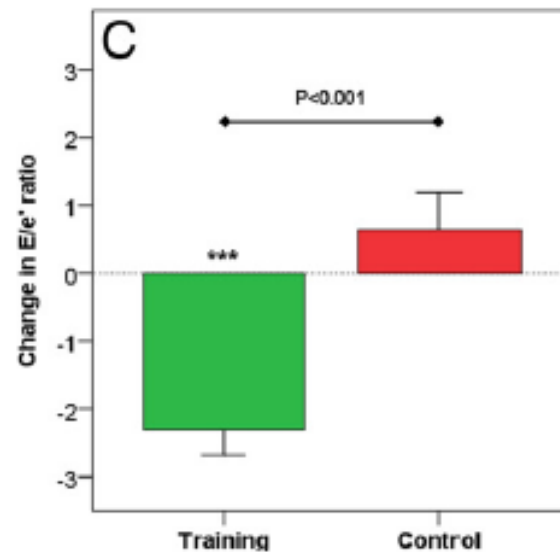
HFpEF : bénéfice de l'exercice physique

Ex-DHF
Prospectif; N=64

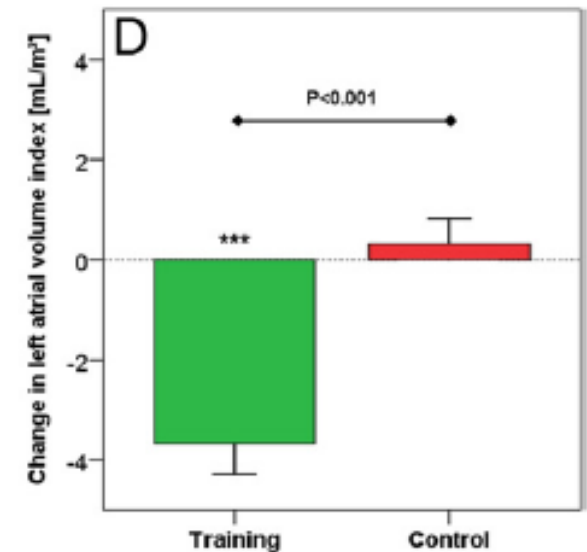
Endurance + resistance training (32 séances; 4 semaines)



Amélioration de la pVO2



Amélioration E/E'



Réduction volume OG

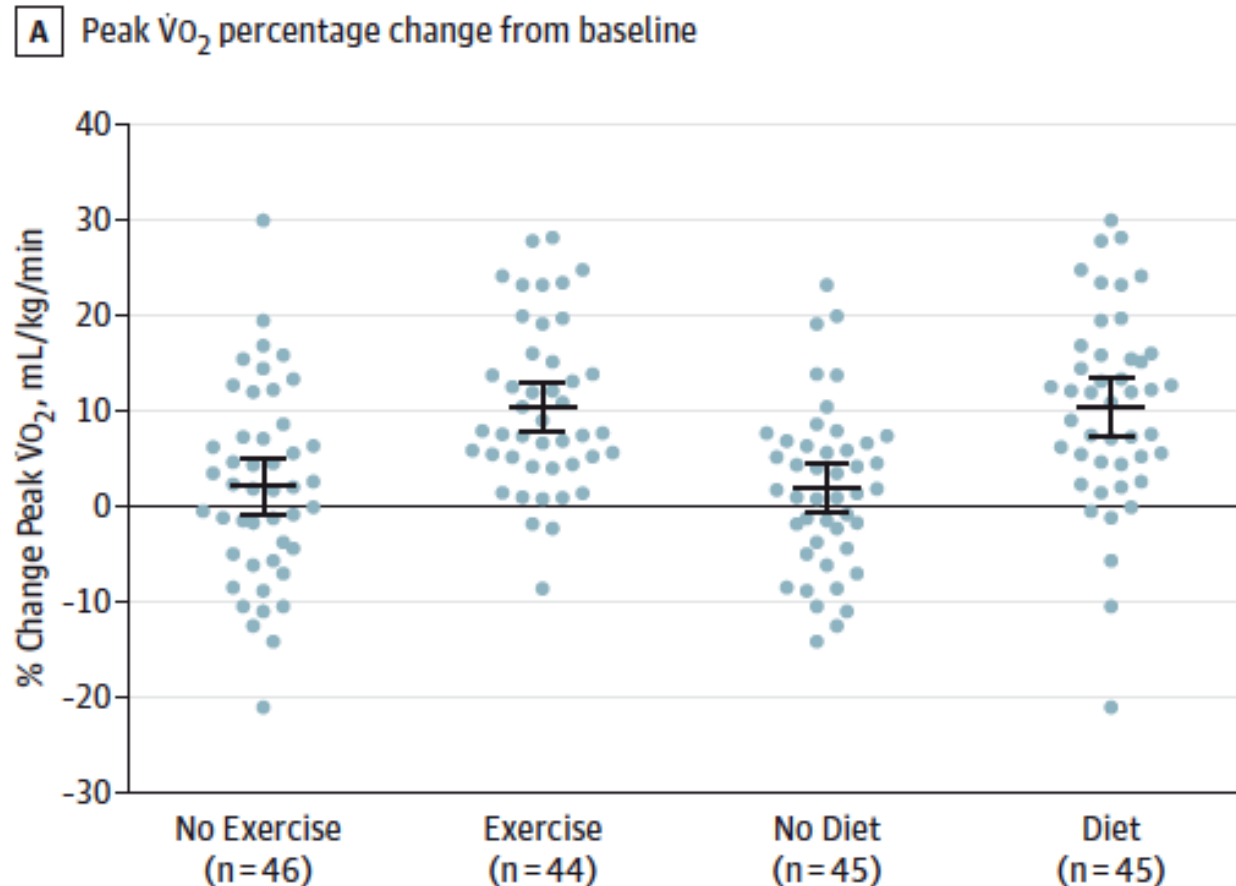
HFpEF : bénéfice de l'exercice physique + régime hypocalorique

Prospectif

N=100

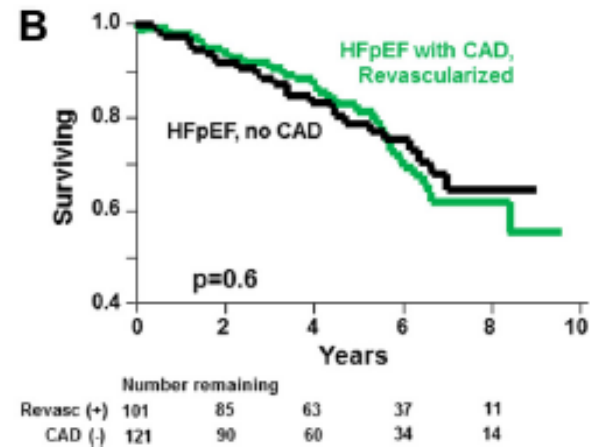
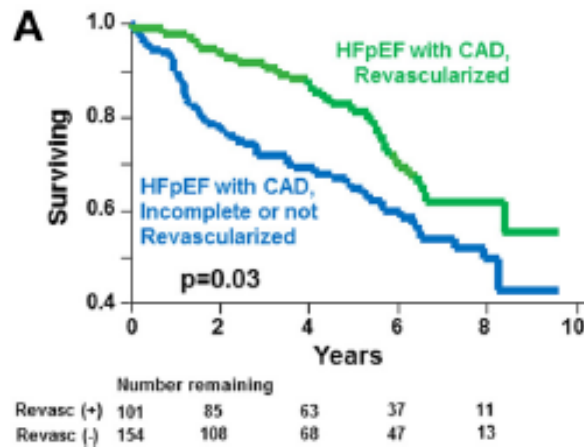
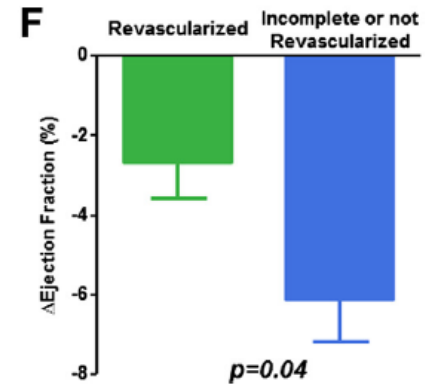
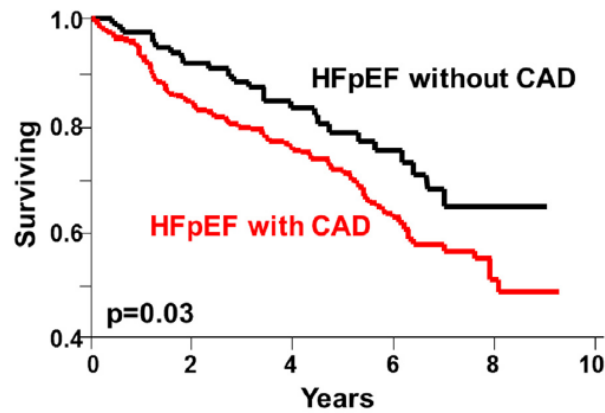
BMI médian : 39.3

20 semaines de
régime ou exercice
ou les 2



HFpEF : maladie coronarienne

n=376
HFpEF
(68% avec CAD)



Merci de votre attention !



julien.regamey@chuv.ch
julien.regamey@hopitalrivierachablais.ch