

Foundations of Cardiometabolic Health Certification Course

Certified Cardiometabolic Health Professional (CCHP)



Heart Failure

Alanna A. Morris MD, MSc

Associate Professor of Medicine

Director of Heart Failure Research

Emory Clinical Cardiovascular Research Institute

Emory University, Atlanta GA

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Definition, Epidemiology, and Pathophysiology of Heart Failure

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Heart Failure (HF):

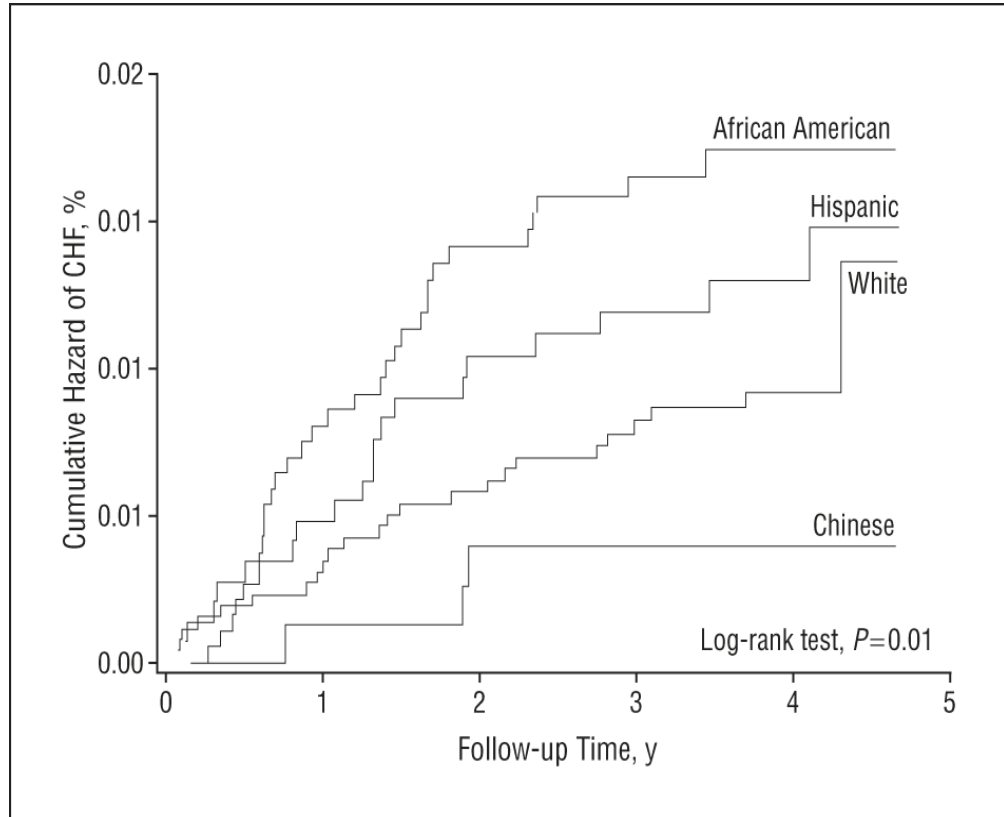
High Morbidity, Mortality, and Costs

Prevalence	Incidence	Mortality	Hospital Discharges	Cost
~6,500,000	~1,000,000	42-50% at 5 years	994,000	\$30.7 billion

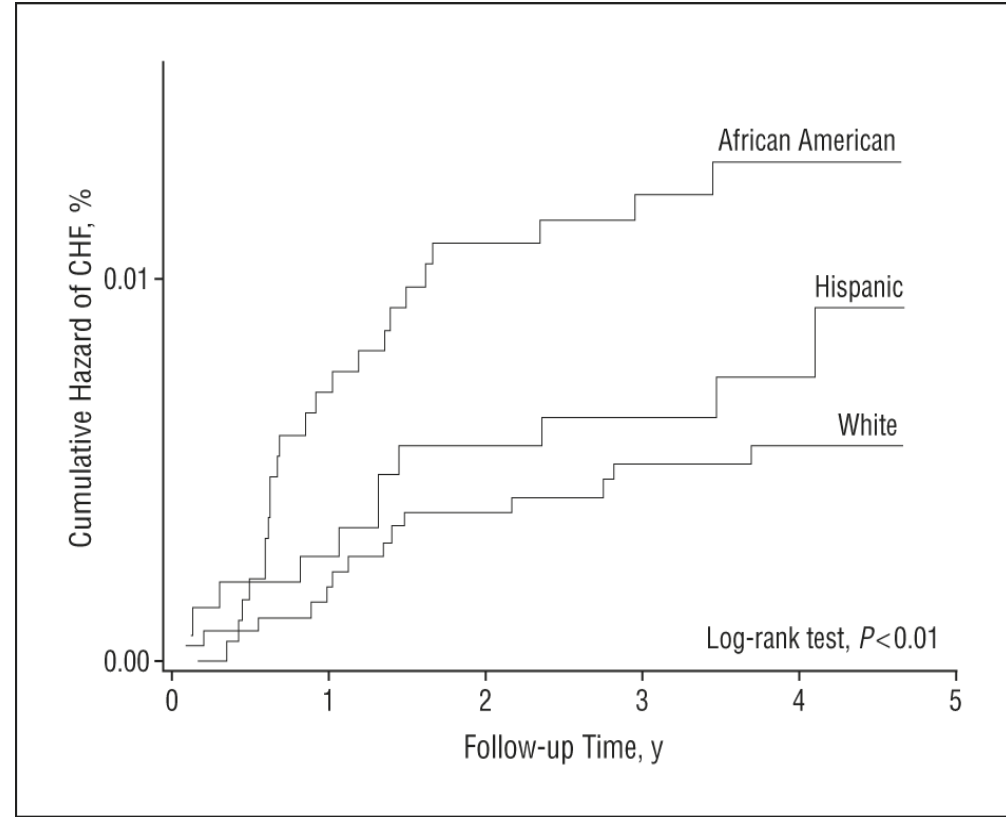
- Major cost-driver of HF is high incidence of hospitalizations

Incidence of HF Differs by Race and Ethnicity

Multi-Ethnic Study of Atherosclerosis (MESA) N=6814
Median follow-up 4.0 years



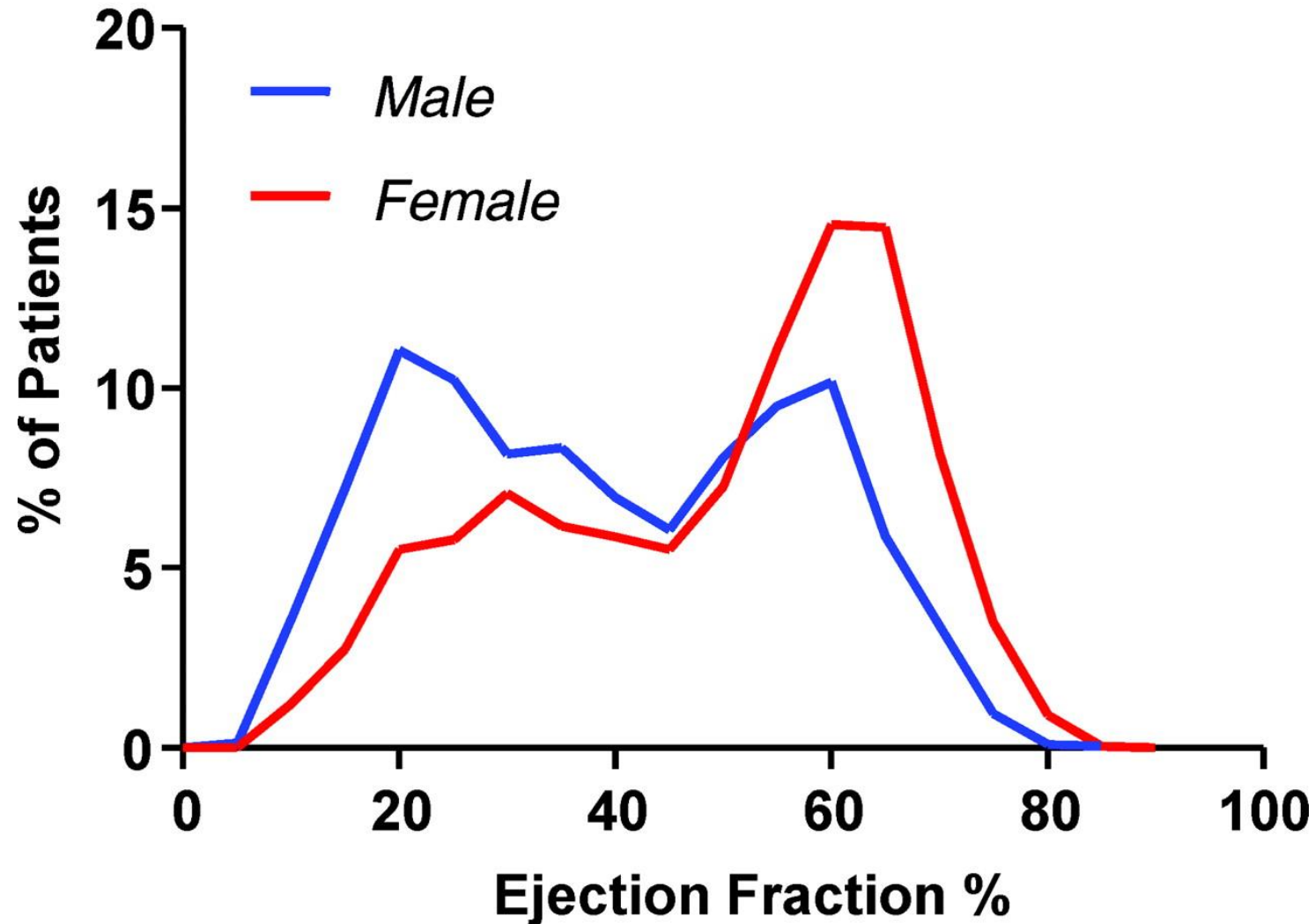
All Participants



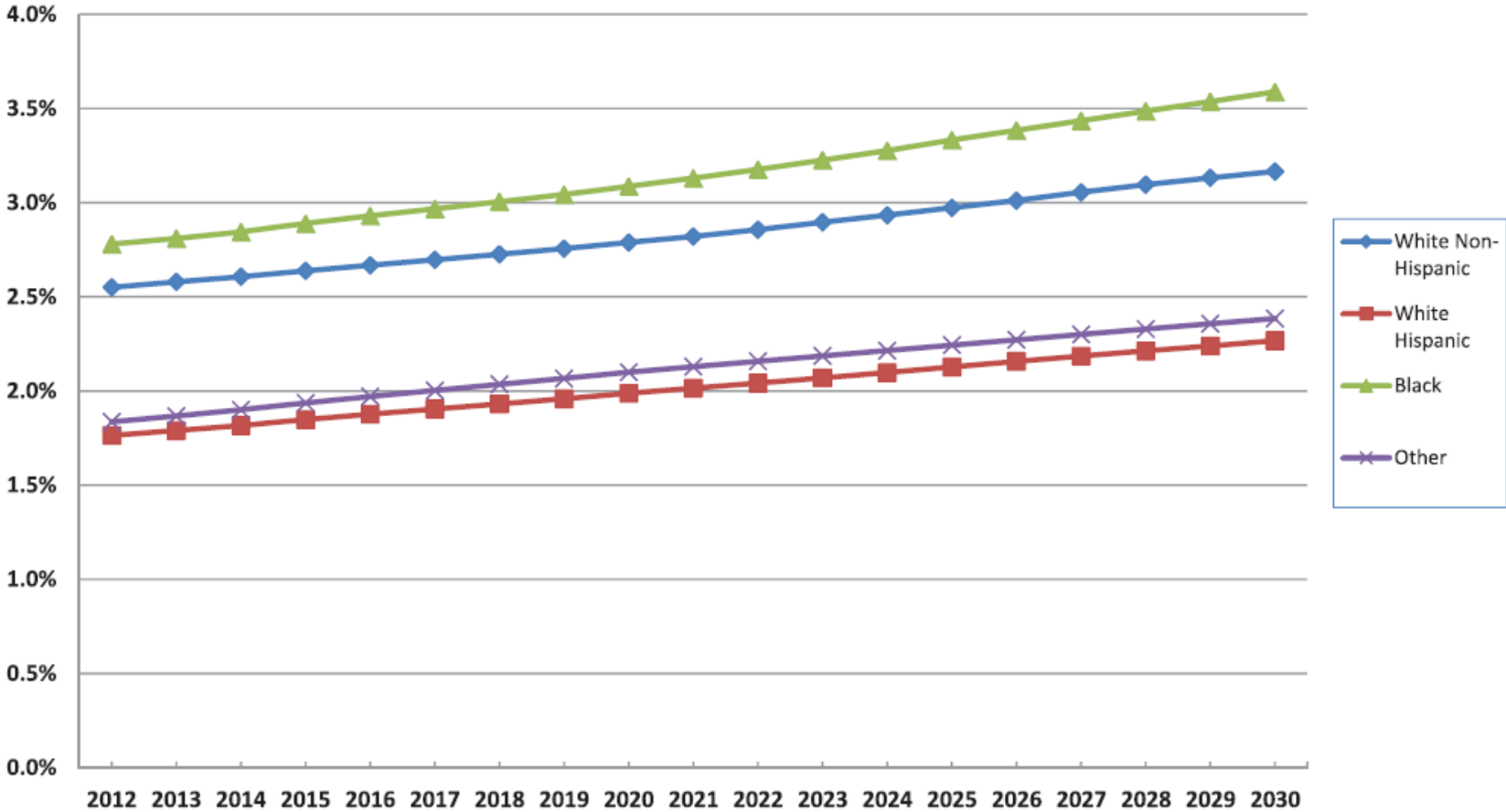
Participants without an Interim Myocardial Infarction

Incidence of HF Differs by Gender

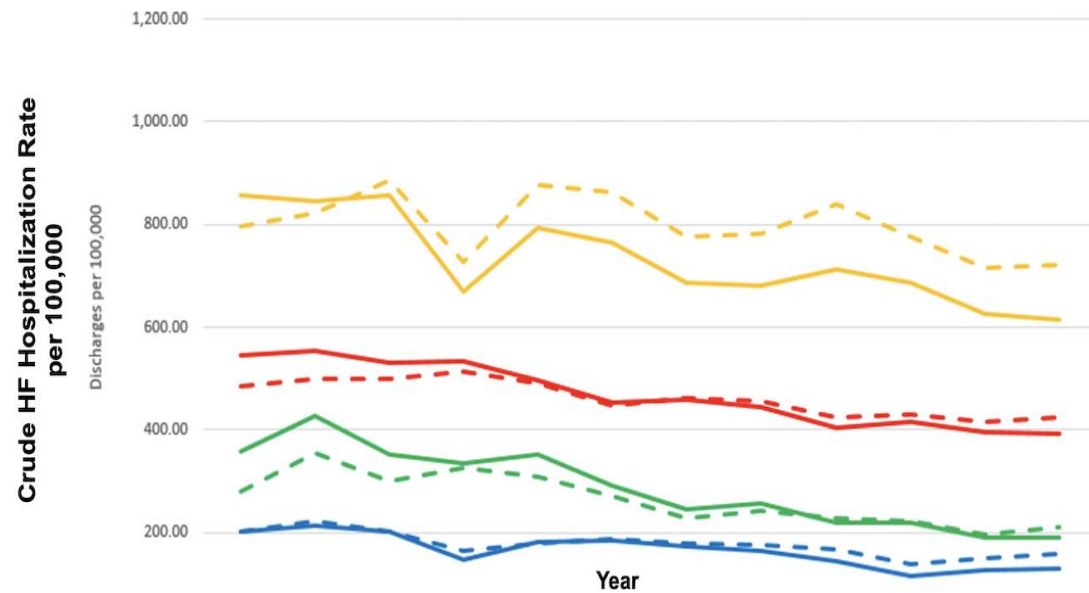
Hospital Based Sample (n = 4910)



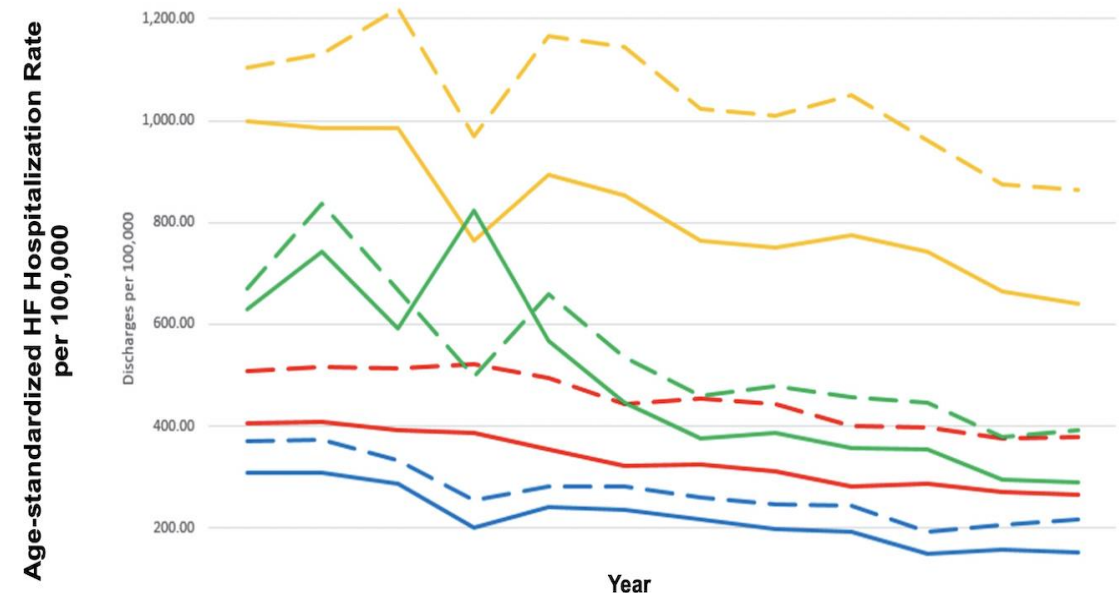
Prevalence of HF Differs by Race and Ethnicity



Black Americans Have Higher Rates of HF-related Hospitalization

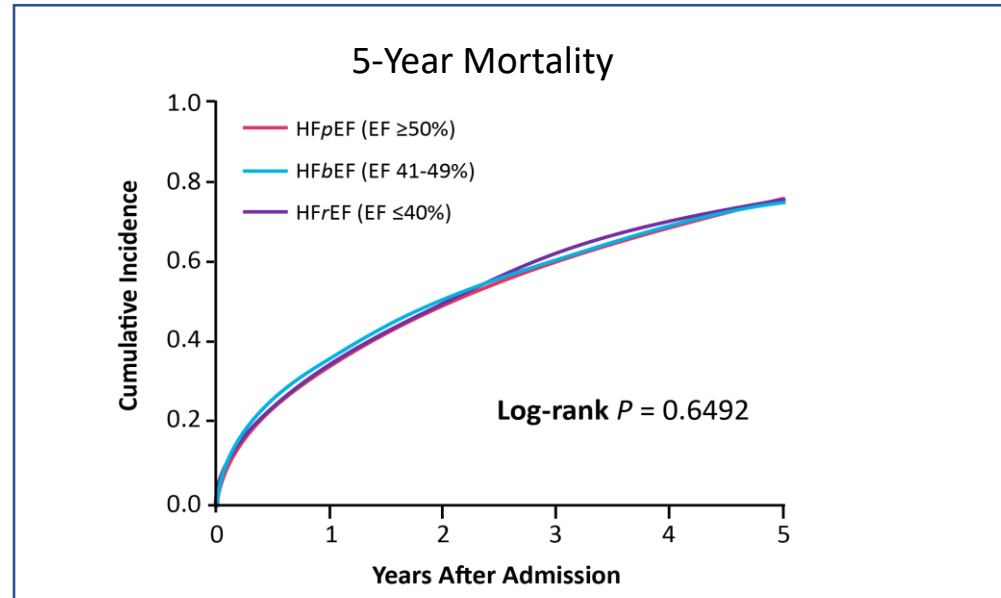
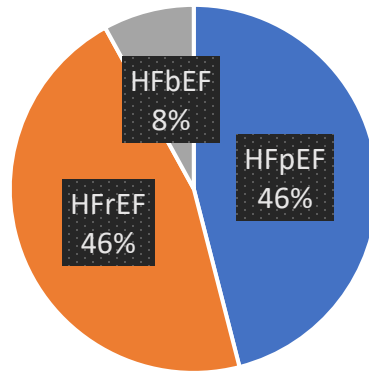


	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013
White Male	484.40	499.16	499.60	513.97	491.34	446.48	460.38	457.01	423.87	428.42	414.35	423.54
White Female	546.00	553.67	531.73	533.53	494.81	453.33	459.81	443.83	403.86	416.16	395.54	391.83
Black Male	797.24	821.17	887.15	727.78	876.40	864.05	775.70	782.31	840.97	776.95	716.85	720.75
Black Female	856.07	846.02	856.63	669.81	792.10	764.69	687.77	682.26	711.92	686.39	625.05	613.75
Hispanic Male	281.03	354.75	300.06	325.56	307.95	270.22	228.34	241.53	227.58	221.71	197.32	210.23
Hispanic Female	359.01	427.52	350.93	334.17	352.22	290.18	244.46	257.66	220.19	219.04	190.49	190.93
Asian & PI Male	202.75	222.46	201.23	163.21	178.46	188.20	179.59	175.73	166.13	137.06	150.51	158.99
Asian & PI Female	202.09	212.04	200.84	148.03	181.45	184.74	173.51	163.81	145.40	116.34	127.49	128.95



	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013
White Male	507.53	516.25	512.23	522.82	495.41	444.71	452.85	443.13	398.94	396.89	376.24	377.79
White Female	405.22	409.57	390.87	386.53	355.04	322.36	324.28	310.89	280.38	286.51	270.22	266.33
Black Male	1,104.07	1,131.19	1,221.23	968.52	1,164.84	1,144.04	1,021.79	1,010.83	1,050.09	960.94	875.29	863.45
Black Female	998.24	984.48	986.25	763.42	894.90	852.33	764.33	749.91	775.30	741.44	663.38	640.17
Hispanic Male	668.89	835.76	668.05	498.59	659.28	535.65	459.66	478.42	457.8	445.08	379.27	392.63
Hispanic Female	628.96	742.29	591.58	824.91	566.33	447.17	376.01	387.74	358.11	353.47	294.65	288.48
Asian & PI Male	371.14	373.78	333.58	253.94	282.08	282.87	259.17	246.22	242.99	192.63	206.82	215.53
Asian & PI Female	308.83	307.93	287.65	200.70	242.04	234.60	215.66	197.04	193.36	148.30	156.84	152.19

5-Year Outcomes of Elderly Patients Hospitalized with Heart Failure



Outcomes – 5-Year Event Rates (%)					
	Mortality	Readmission	CV Readmission	HF Readmission	Mortality/Readmission
HFrEF	75.3	82.2	63.9	48.5	96.4
HFbEF	75.7	85.7	63.3	45.2	97.2
HFpEF	75.7	84.0	58.9	40.5	97.3

HFrEF: heart failure with reduced ejection fraction; HFbEF: heart failure with borderline ejection fraction; HFpEF: heart failure with preserved ejection fraction

Universal Definition of Heart Failure

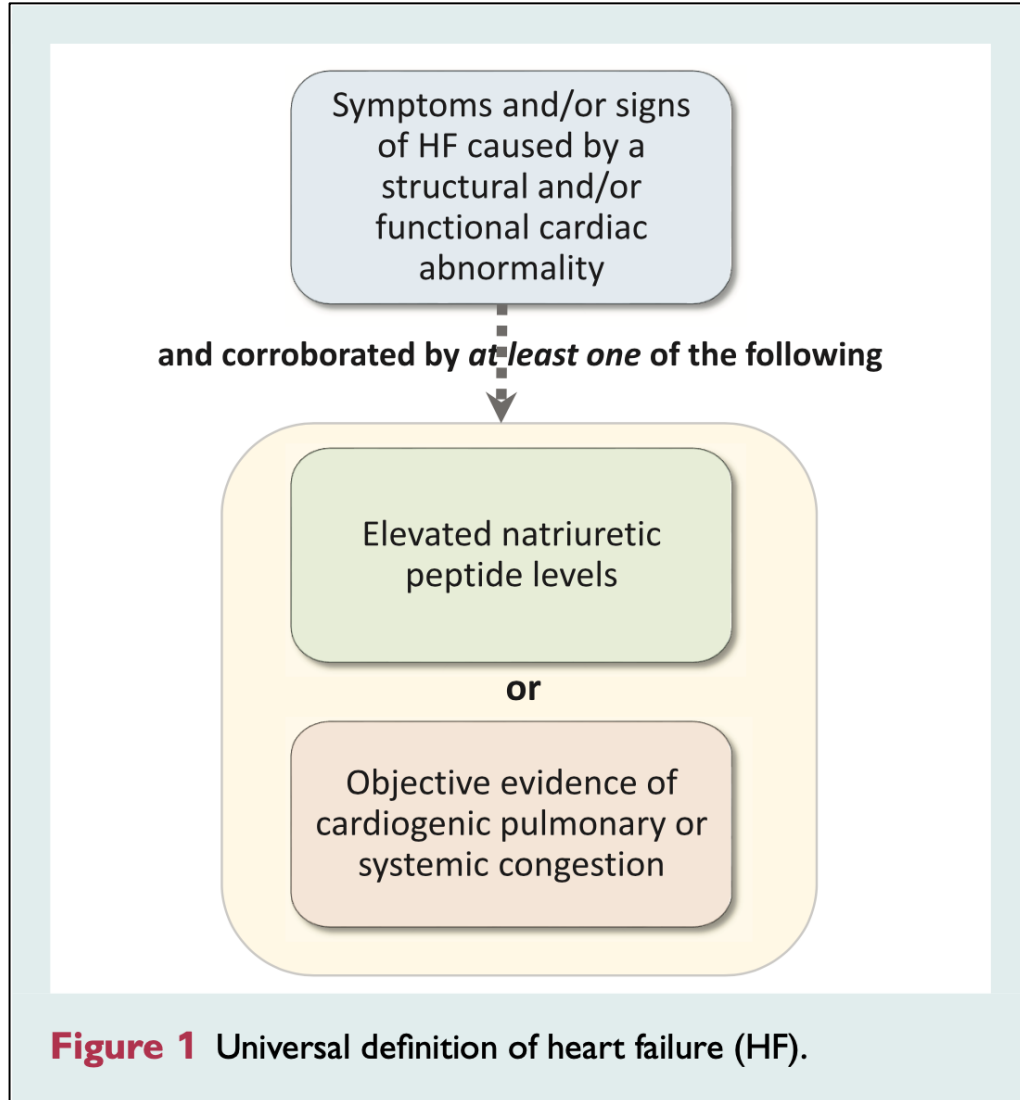
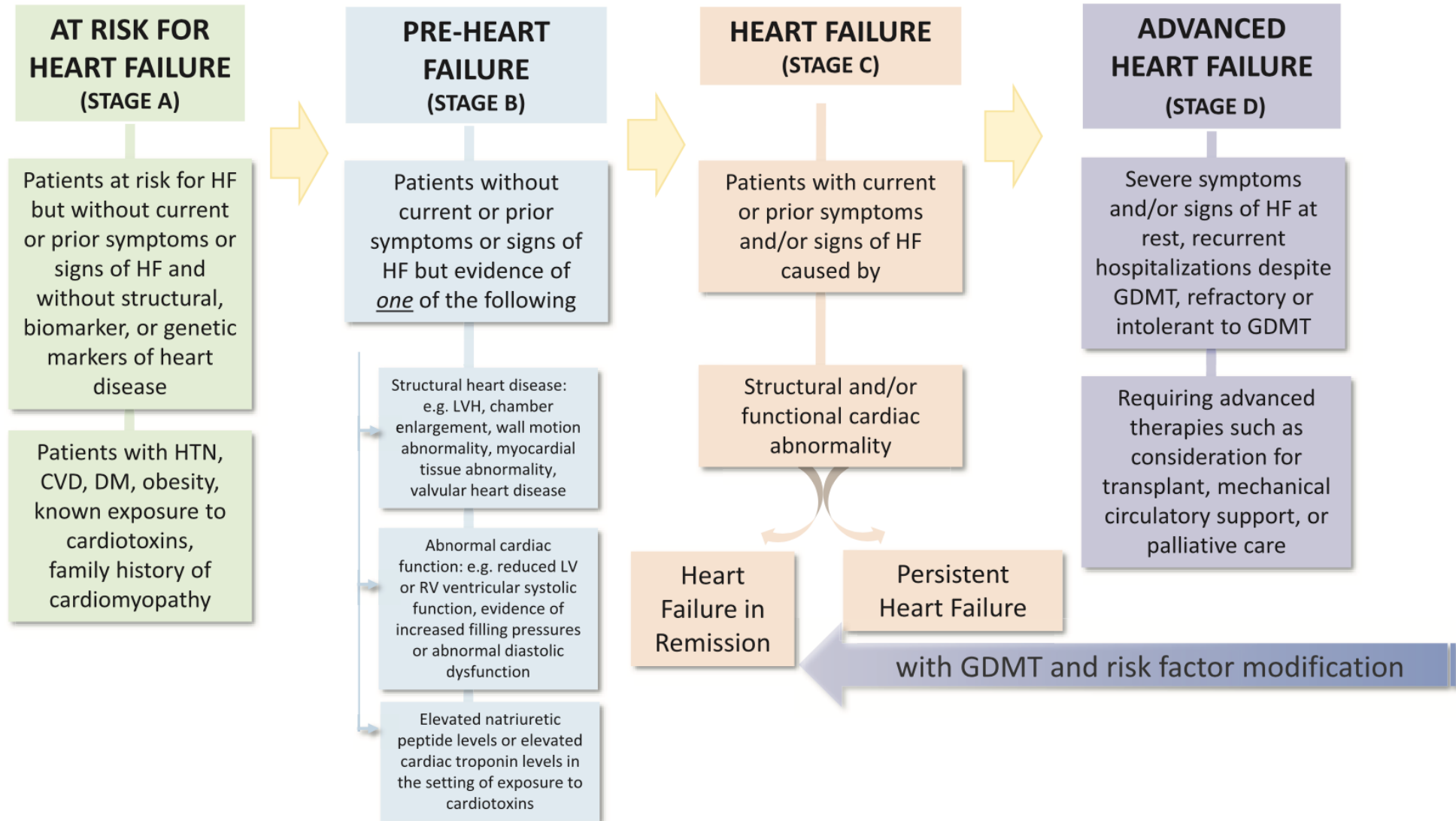


Table 7 Natriuretic peptide levels supporting definition of heart failure

	Ambulatory	Hospitalized/ decompensated
BNP pg/ml	≥35	≥ 100
NT-proBNP pg/ml	≥ 125	≥ 300

Universal Definition of Heart Failure: Stages



Classifications of HF According to Ejection Fraction (EF)

Normal:

- LVEF >55%

HF with reduced EF (HFrEF):

- HF with LVEF \leq 40%

HF with mildly reduced EF (HFmrEF):

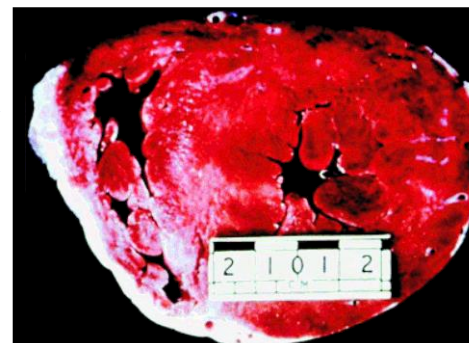
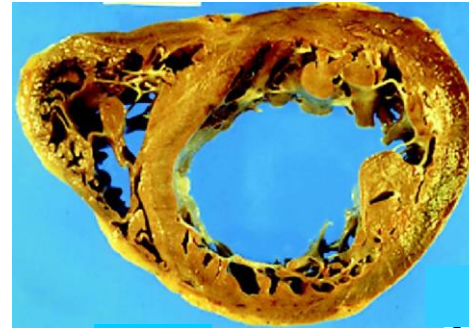
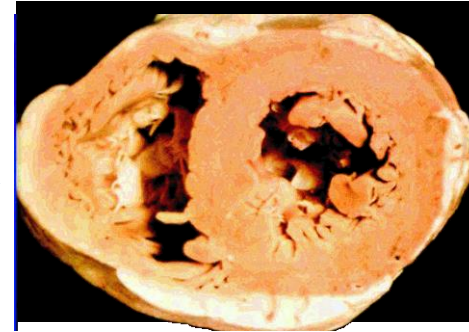
- HF with LVEF 41–49%

HF with preserved EF (HFpEF):

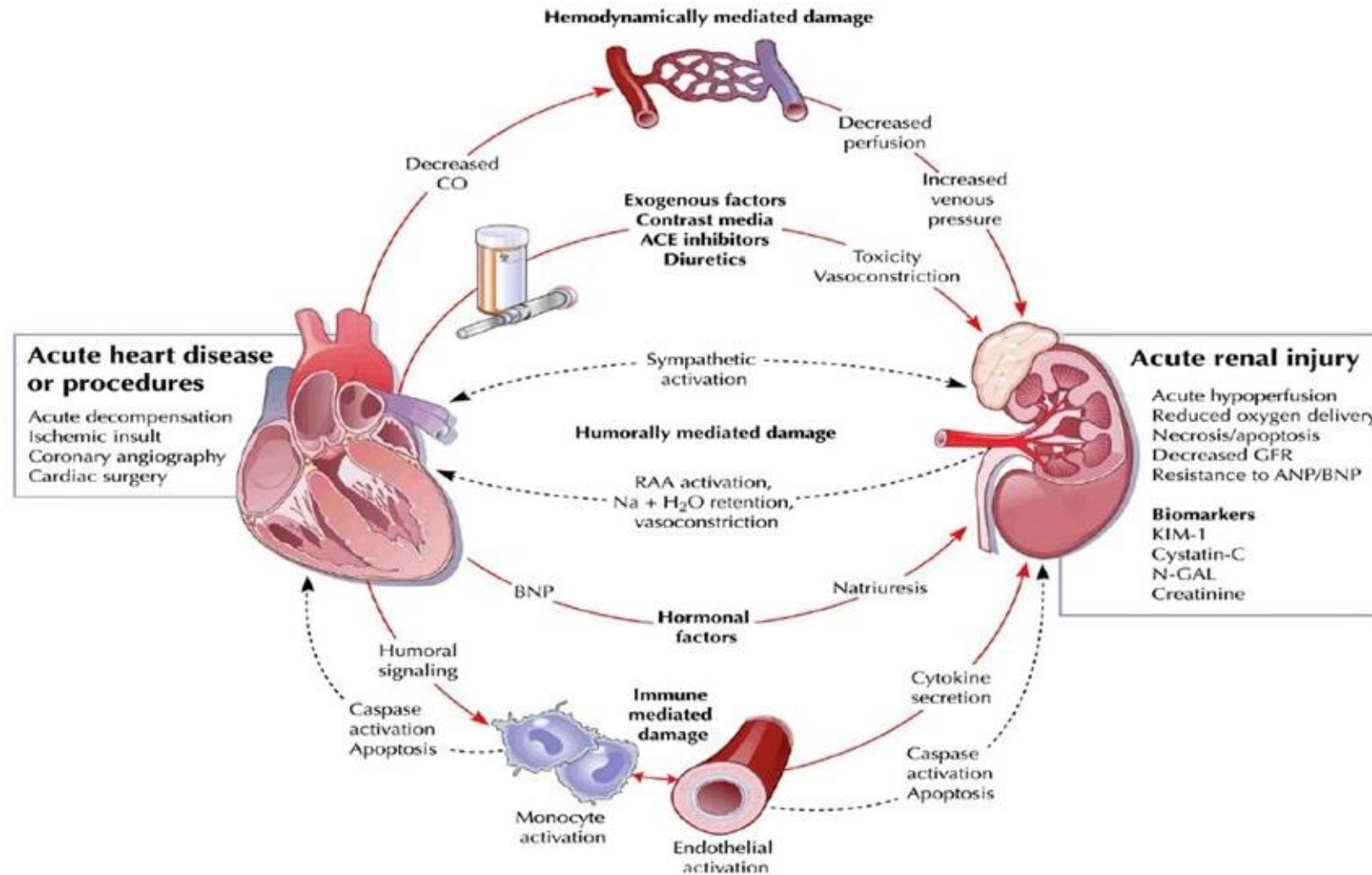
- HF with LVEF \geq 50%

HF with improved EF (HFimpEF):

- HF with a baseline LVEF \leq 40%, a \geq 10 point increase from baseline LVEF, and a second measurement of LVEF >40%

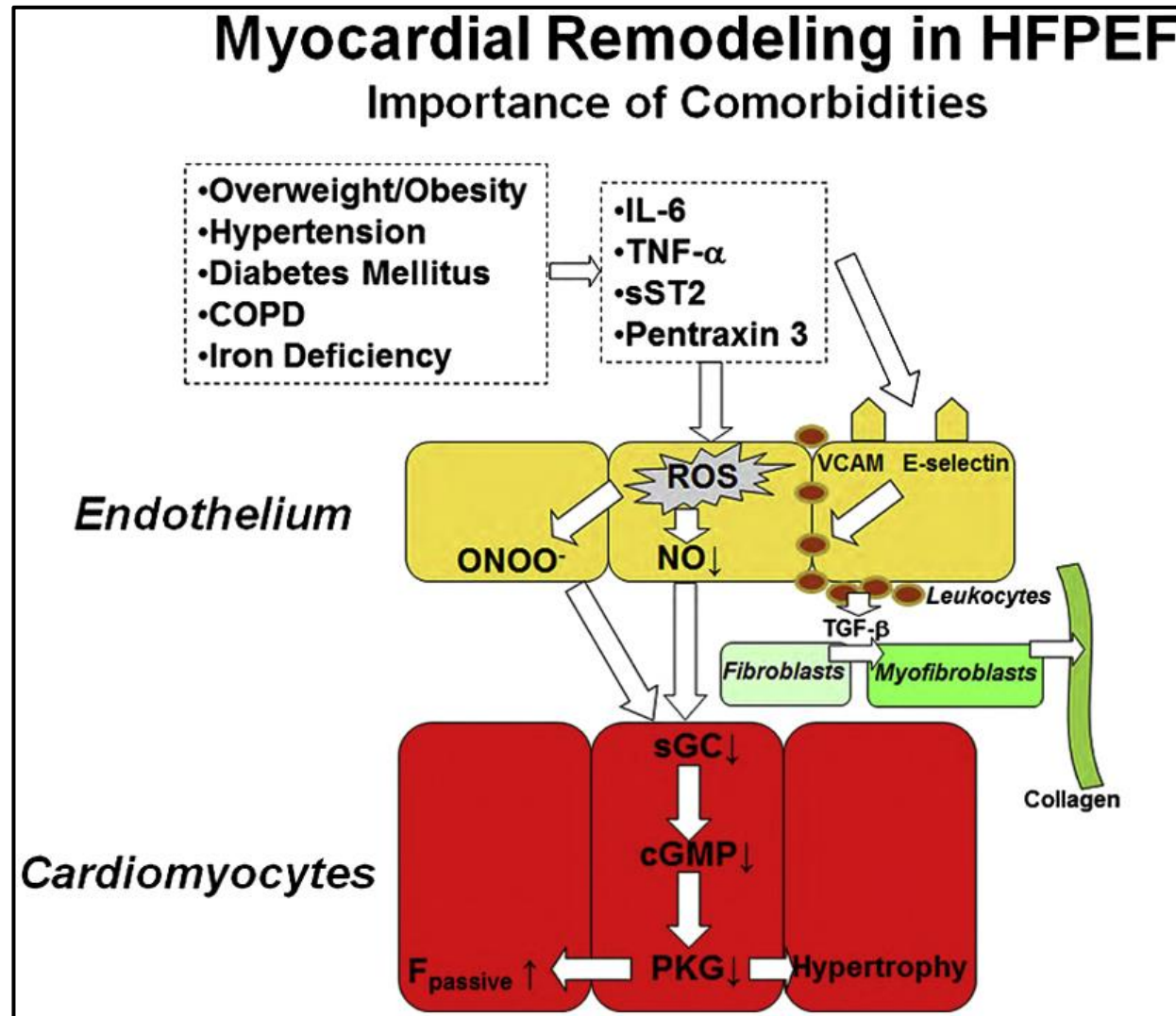


Pathophysiology of HFrEF



Pathophysiology of HFpEF:

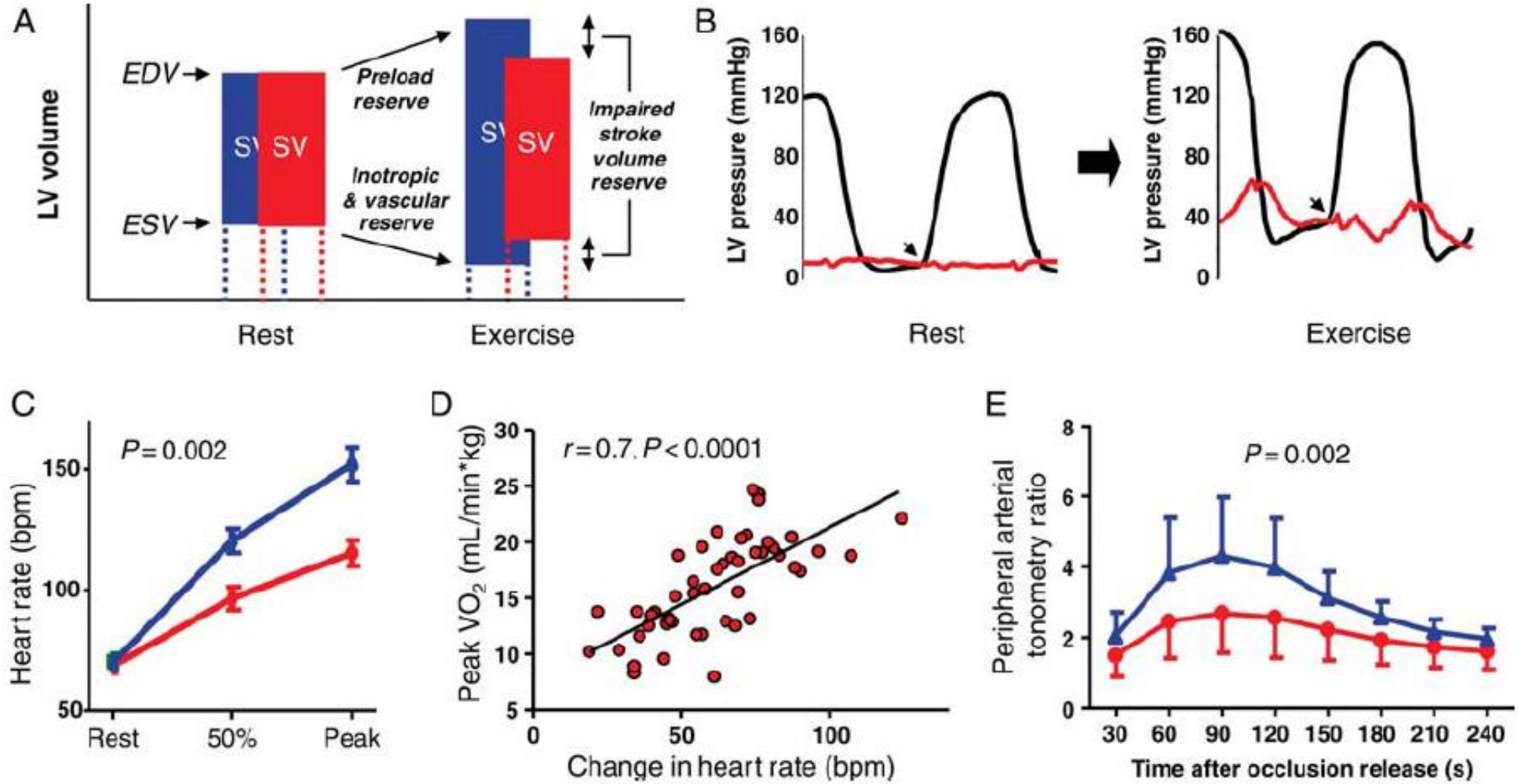
Comorbidities Drive Myocardial Dysfunction through Inflammation



Pathophysiology of HFpEF

Blue=controls

Red=HFpEF



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HF & Cardiometabolic Risk connections

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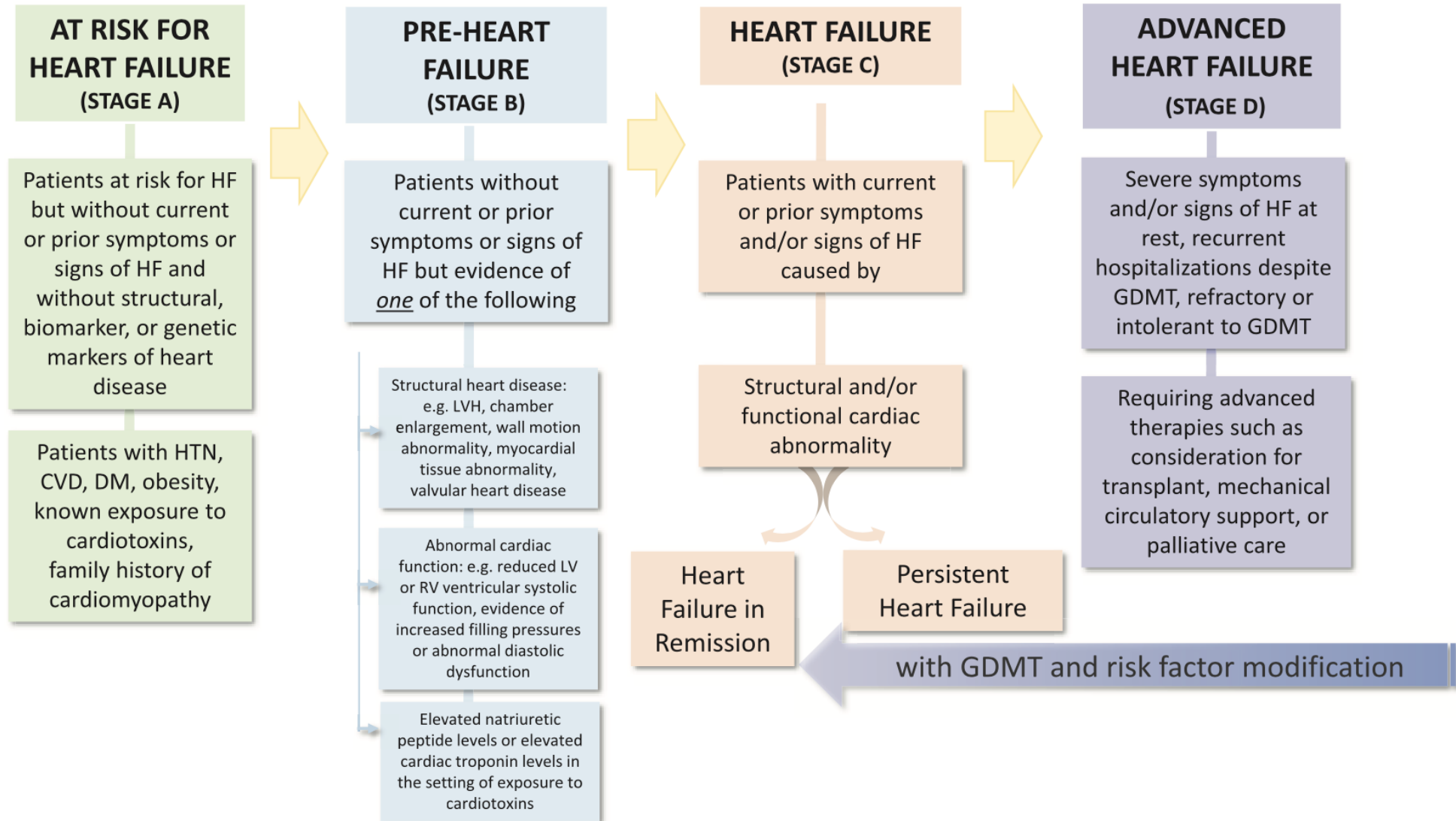
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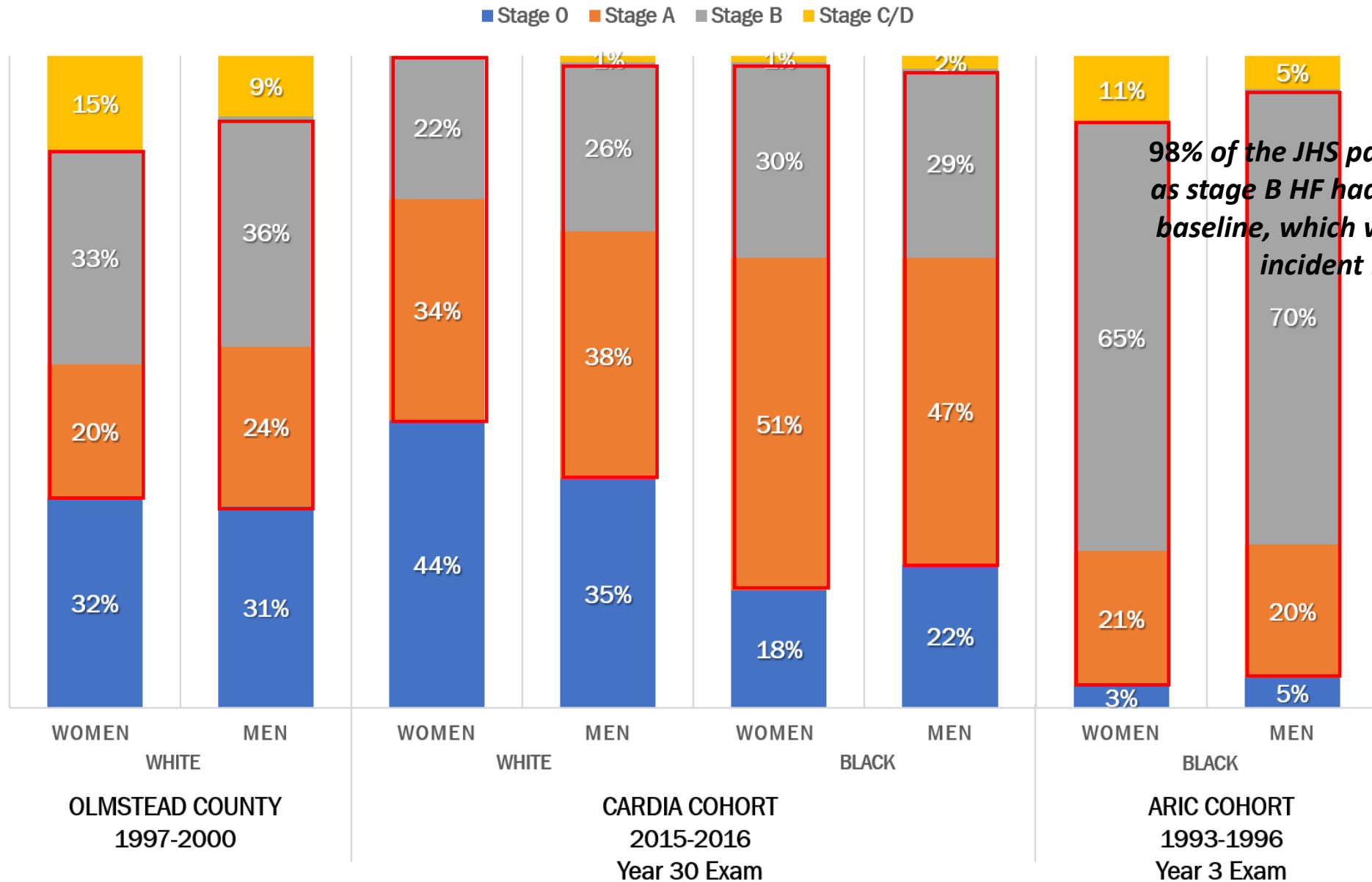
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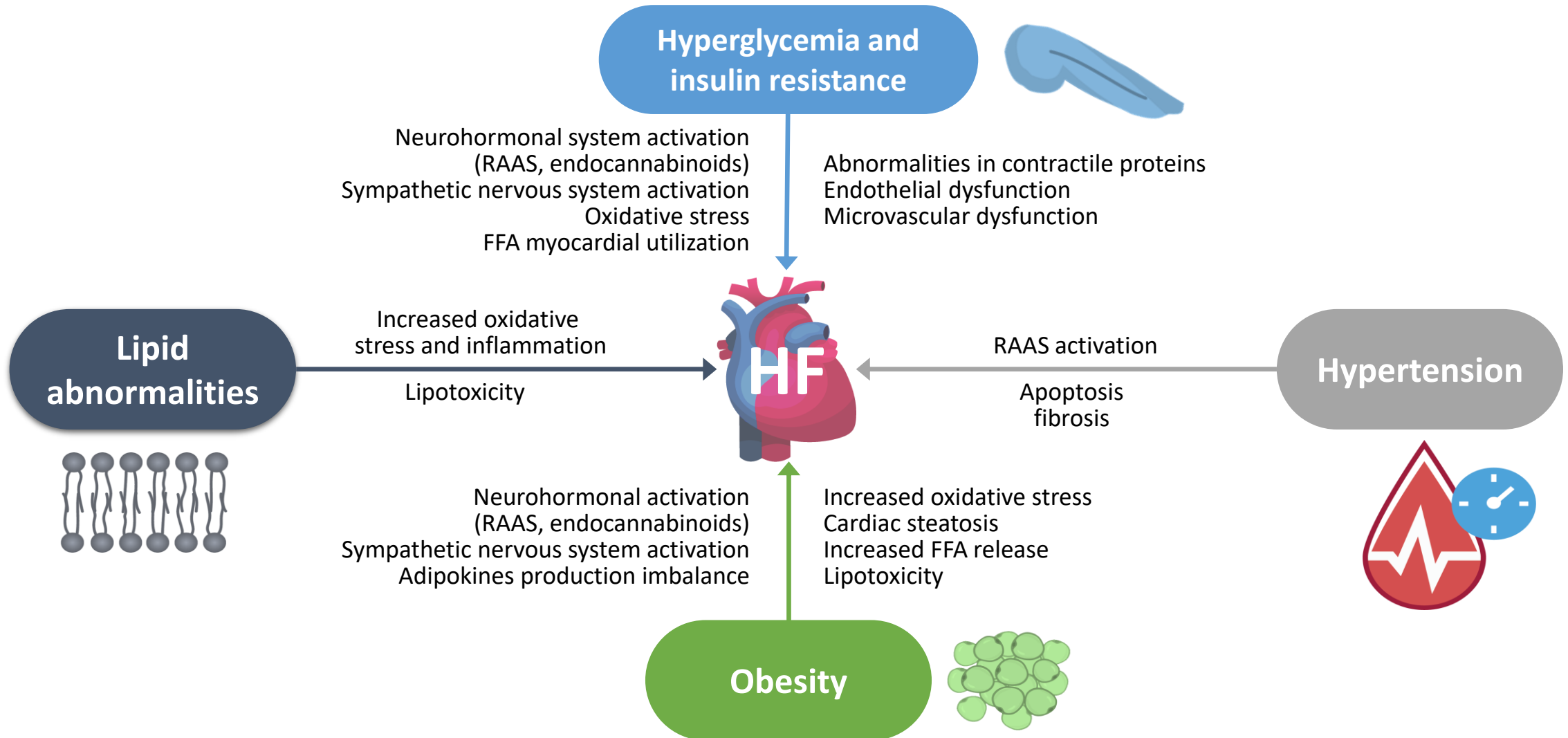
Universal Definition of Heart Failure: Stages



Comparison of HF Stages Across Community Cohorts



At Risk for HF (Stage A): Cardiometabolic Risk Factors



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Screening and Diagnostic Approaches

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Risk Equations for Incident ASCVD

Age (years)	<input type="text" value="40-79"/>
Gender	<input checked="" type="radio"/> Male <input type="radio"/> Female
Race	<input type="radio"/> African American <input checked="" type="radio"/> Other
Total cholesterol (mg/dL)	<input type="text" value="130-320"/>
HDL cholesterol (mg/dL)	<input type="text" value="20-100"/>
Systolic blood pressure (mmHg)	<input type="text" value="90-200"/>
Diastolic blood pressure (mmHg)	<input type="text" value="30-140"/>
Treated for high blood pressure	<input checked="" type="radio"/> No <input type="radio"/> Yes
Diabetes	<input checked="" type="radio"/> No <input type="radio"/> Yes
Smoker	<input checked="" type="radio"/> No <input type="radio"/> Yes
<input type="button" value="Calculate"/>	

Calculate your 10-year risk of heart disease or stroke using the ASCVD algorithm published in [2013 ACC/AHA Guideline on the Assessment of Cardiovascular Risk](#).

This calculator assumes that you have not had a prior heart attack or stroke. If you have, generally it is recommended that you discuss with your doctor about starting aspirin and a statin. Furthermore, if you have an LDL-cholesterol (bad cholesterol) greater than 190, it is also generally recommended that you discuss with your doctor about starting aspirin and a statin.

Unfortunately, there is insufficient data to reliably predict risk for those less than 40 years of age or greater than 79 years of age and for those with total cholesterol greater than 320.

UPDATE (11/21/17) -- The ACC/AHA has released their [2017 Guideline for the Prevention, Detection, Evaluation, and Management of High Blood Pressure in Adults](#). At a high level, the new guidelines redefine hypertension as blood pressure >130/80 and recommend starting anti-hypertensives based on ASCVD risk score of >10%. This will be a significant change from JNC-8. Please let us know if you would like us to incorporate the new guidelines into [cvriskcalculator.com](#) by completing [this 1-question survey](#).

UPDATE (6/30/16) -- The calculator has been vetted against the [final guidelines from the USPSTF](#) for initiating aspirin therapy.

Risk Equations for Incident HF

CENTRAL ILLUSTRATION: Examples of Predicted 10-Year HF Risk Using the Online PCP-HF Tool

10-Year Predicted HF Risk = 9.5%				10-Year Predicted HF Risk = 7.8%			
10-Year Heart Failure Risk Calculator Pooled Cohort Equations to Prevent HF (PCP-HF)				10-Year Heart Failure Risk Calculator Pooled Cohort Equations to Prevent HF (PCP-HF)			
Age: (30-79 years)	50	Hypertension treatment?	<input checked="" type="checkbox"/> YES <input type="checkbox"/> NO	Age: (30-79 years)	65	Hypertension treatment?	<input checked="" type="checkbox"/> YES <input type="checkbox"/> NO
Gender:	<input checked="" type="checkbox"/> M <input type="checkbox"/> F	Fasting Glucose:	120	Gender:	<input type="checkbox"/> M <input checked="" type="checkbox"/> F	Fasting Glucose:	120
Race:	<input checked="" type="checkbox"/> WHITE <input type="checkbox"/> BLACK	Diabetes treatment?	<input type="checkbox"/> YES <input checked="" type="checkbox"/> NO	Race:	<input checked="" type="checkbox"/> WHITE <input type="checkbox"/> BLACK	Diabetes treatment?	<input type="checkbox"/> YES <input checked="" type="checkbox"/> NO
Currently smoke?	<input checked="" type="checkbox"/> Y <input type="checkbox"/> N	Total Cholesterol: (80-300 mg/dL)	200	Currently smoke?	<input type="checkbox"/> Y <input checked="" type="checkbox"/> N	Total Cholesterol: (80-300 mg/dL)	200
BMI:	35	HDL Cholesterol (15-100 mg/dL)	45	BMI:	35	HDL Cholesterol (15-100 mg/dL)	45
Systolic Blood Pressure: (80-200 mm Hg)	140	QRS Duration: (ms)	90	Systolic Blood Pressure: (80-200 mm Hg)	140	QRS Duration: (ms)	100
<input type="button" value="Calculate Risk"/>				<input type="button" value="Calculate Risk"/>			



Similar 10-year HF risk estimates in a 50-year-old black man and a 65-year-old white woman



Predictors of HF in T2D: A Practical Approach to Prevention

For patients with T2D without clinical ASCVD or HF

Clinical factors associated with increased incidence of HF

Diabetes-specific risk factors

- Insulin, sulfonylurea, DPP4i, TZD use
- Duration of diabetes > 10 years
- A1C \geq 7%
- Microvascular disease

Cardiovascular risk factors

- Hypertension
- Obesity
- Atrial fibrillation

Non-cardiovascular risk factors

- Increasing age
- Obstructive sleep apnea

Consider measuring Natriuretic Peptides

NT-proBNP < 50 pg/mL

Continue to monitor

NT-proBNP 50–124 pg/mL

NT-proBNP \geq 125 pg/mL
BNP \geq 35 pg/mL

Consider SGLT2 inhibitors

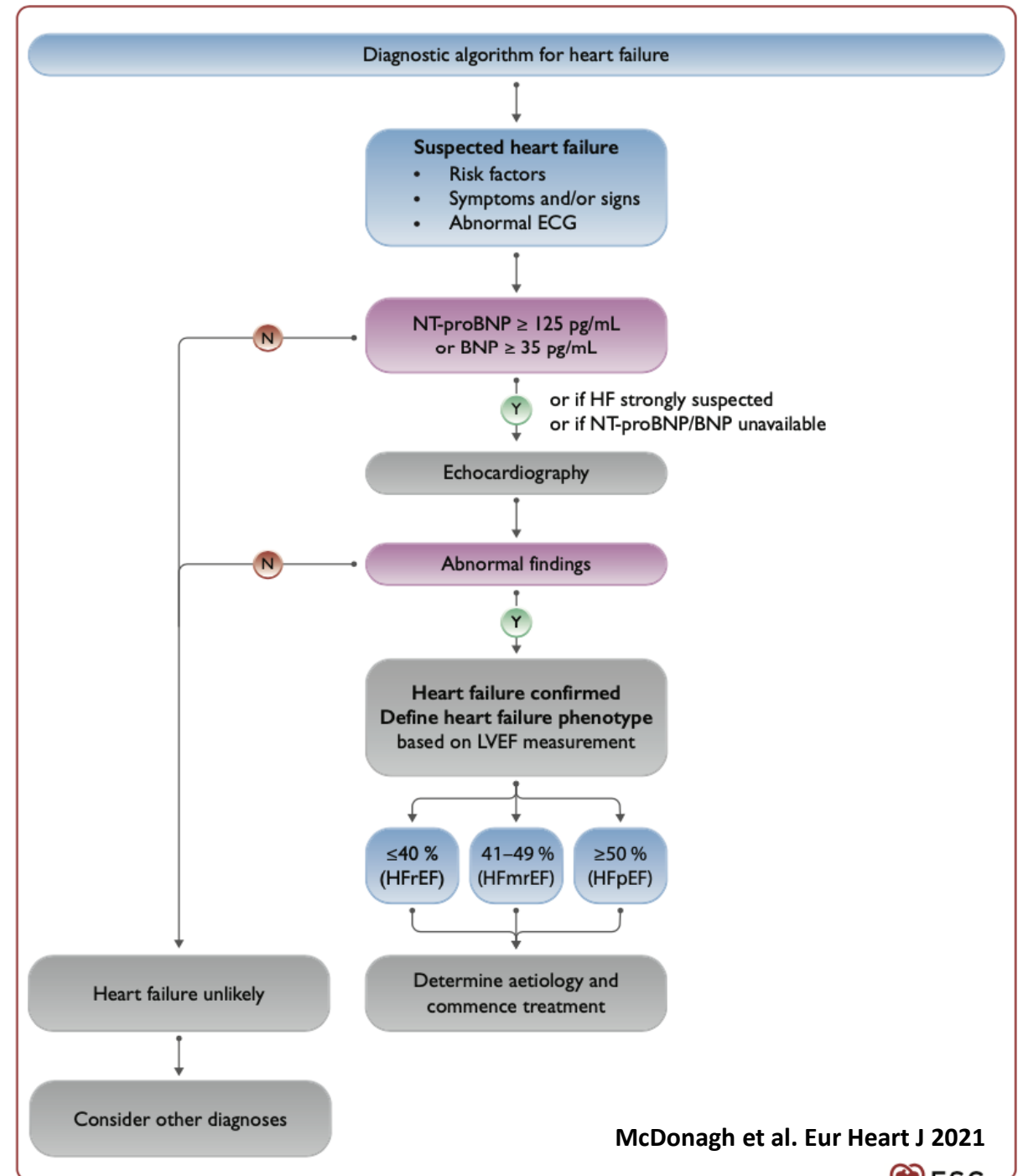
Using NT-proBNP in combination with clinical risk factors may help identify patients with T2D who may benefit from SGLT2 inhibitor

ASCVD, atherosclerotic cardiovascular disease; DPP4i, dipeptidyl peptidase 4 inhibitor; A1C, glycated haemoglobin A_{1c}; NT-proBNP, N-terminal pro-B-type natriuretic peptide; SGLT2, sodium-glucose cotransporter 2; TZD, thiazolidinedione.

Diagnosis of HF

- **History**

- Traditional risk factors, dietary habits, new medications (including OTC and holistic/alternative therapies), substance use
- 3-generation family history
- Recent hospitalizations or ED visits



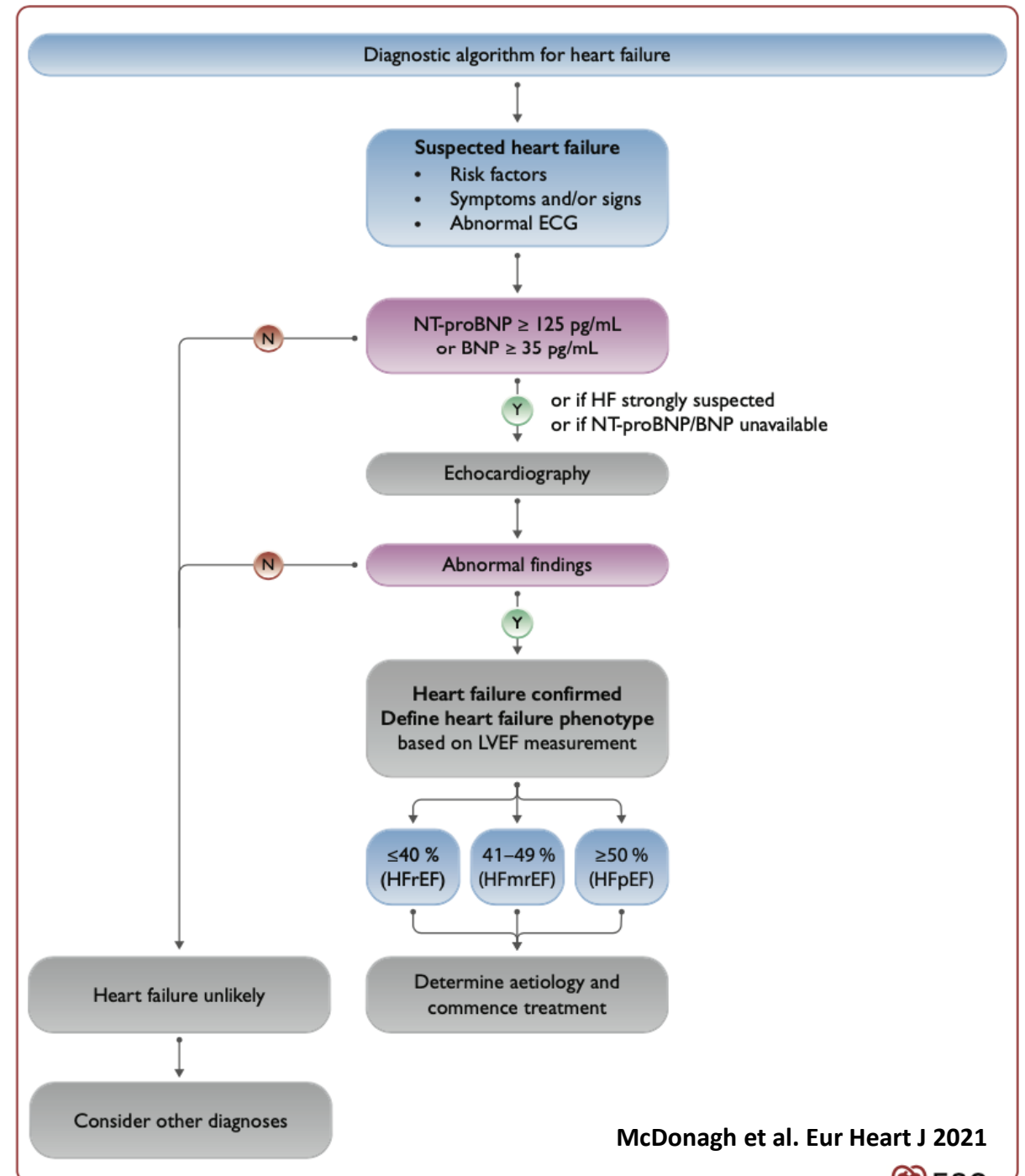
Diagnosis of HF

• History

- Traditional risk factors, dietary habits, new medications (including OTC and holistic/alternative therapies), substance use
- 3-generation family history
- Recent hospitalizations or ED visits

• Physical – Symptoms and Signs

- Dyspnea, orthopnea, paroxysmal nocturnal dyspnea
- Fatigue/exercise intolerance
- Weight gain, peripheral edema, abdominal distention
- Early satiety



Diagnosis of HF

• History

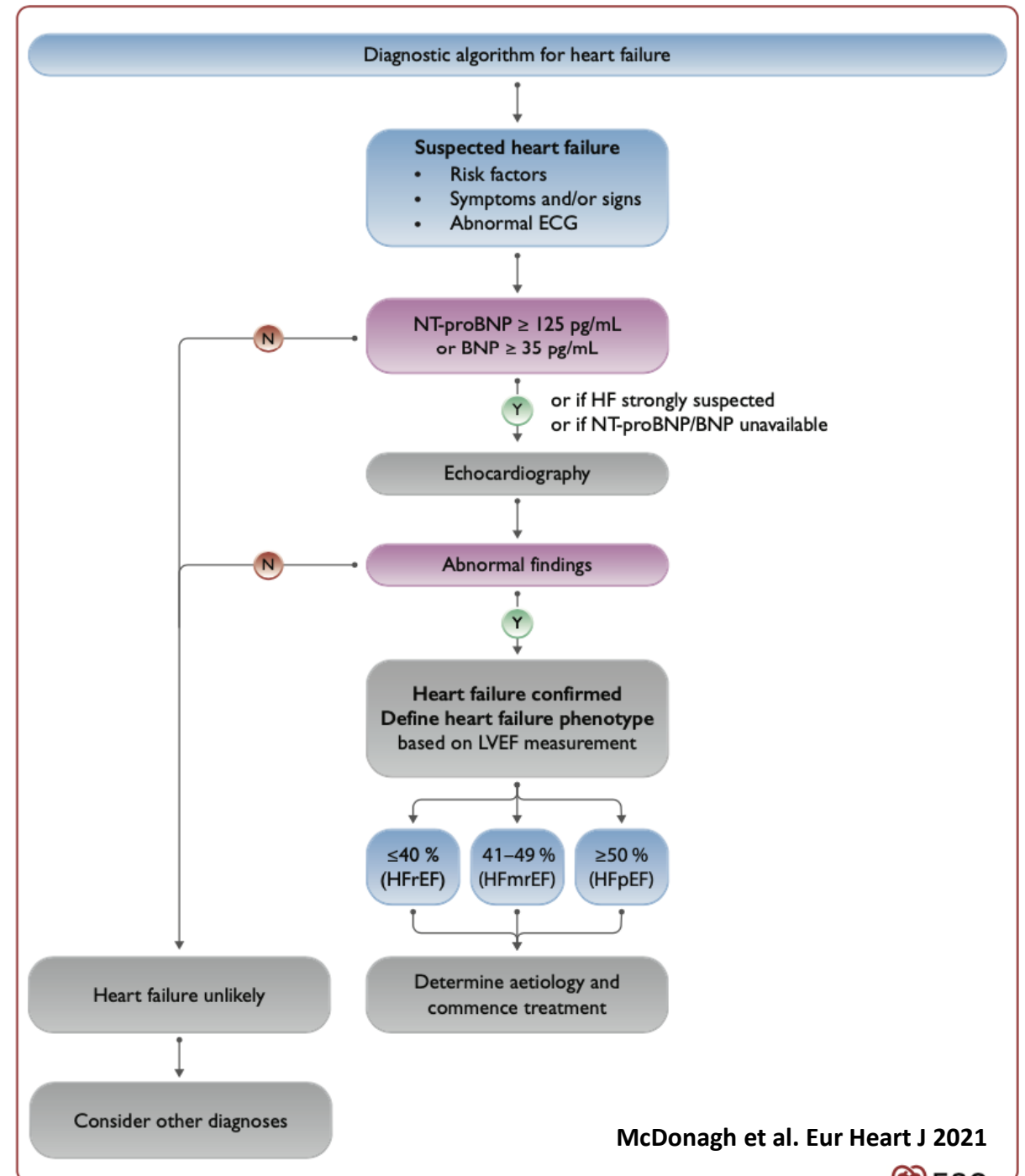
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• Physical – Symptoms and Signs

- Dyspnea, orthopnea, paroxysmal nocturnal dyspnea
- Fatigue/exercise intolerance
- Weight gain, peripheral edema, abdominal distention
- Early satiety

• Diagnostic Tests

- 12-lead ECG
- Labs: CBC, urinalysis, basic metabolic panel, fasting lipid profile, liver function tests, and thyroid-stimulating hormone (Class IC)
- Biomarkers
 - Natriuretic peptides (BNP and NT-proBNP)
 - Other biomarkers (troponins) as applicable



Diagnosis of HF

History

- Traditional risk factors, dietary habits, new medications (including OTC and holistic/alternative therapies), substance use
- 3-generation family history
- Recent hospitalizations or ED visits

Physical – Symptoms and Signs

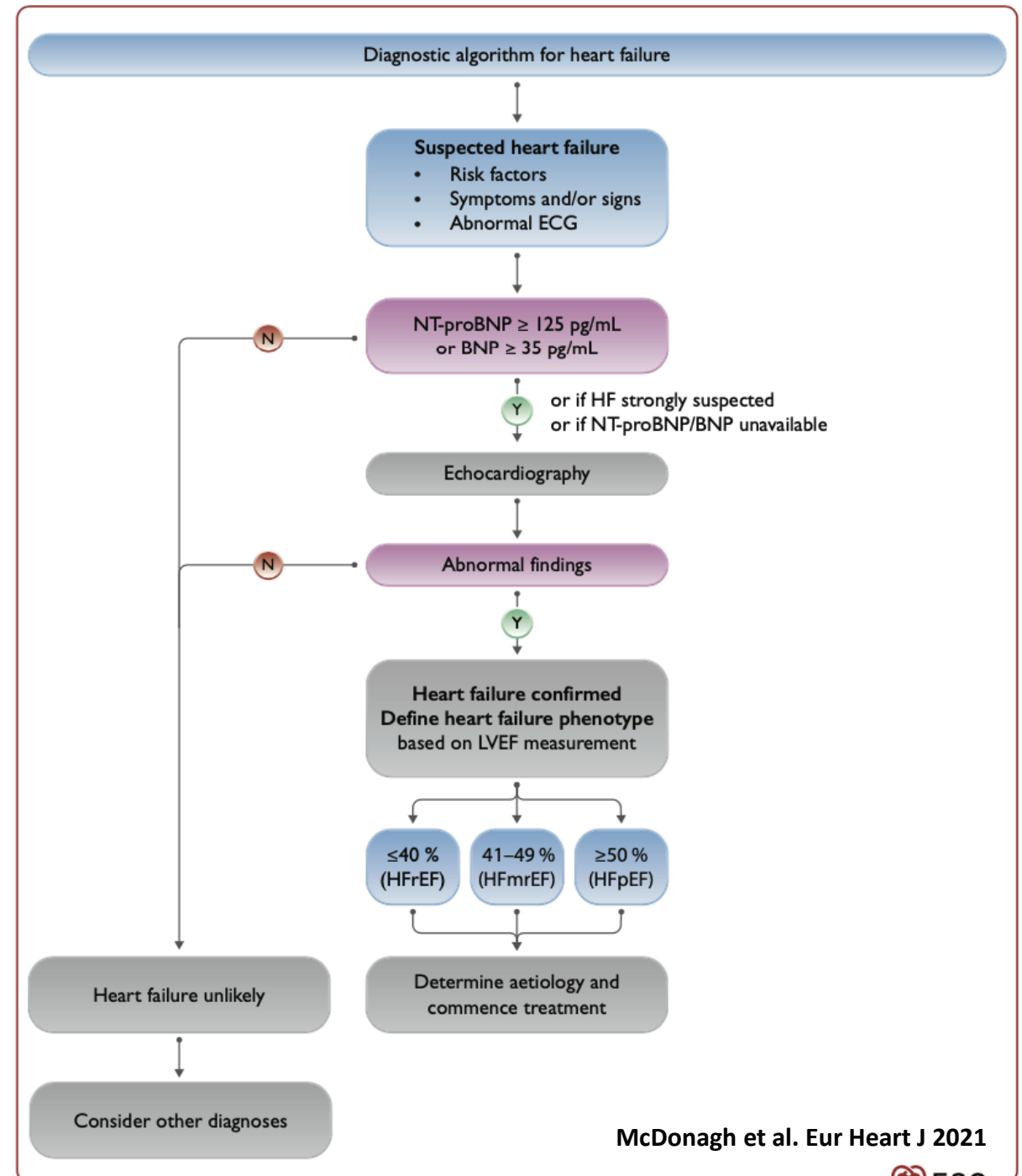
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Diagnostic Tests

- 12-lead ECG
- Labs: CBC, urinalysis, basic metabolic panel, fasting lipid profile, liver function tests, and thyroid-stimulating hormone (Class IC)
- Biomarkers
 - Natriuretic peptides (BNP and NT-proBNP)
 - Other biomarkers (troponins) as applicable

Other tests (etiology)

- Cardiac catheterization
- Genetic testing
- Advanced Imaging: cardiac MRI, Technetium pyrophosphate ($^{99m}\text{Tc-PYP}$) scintigraphy, etc)



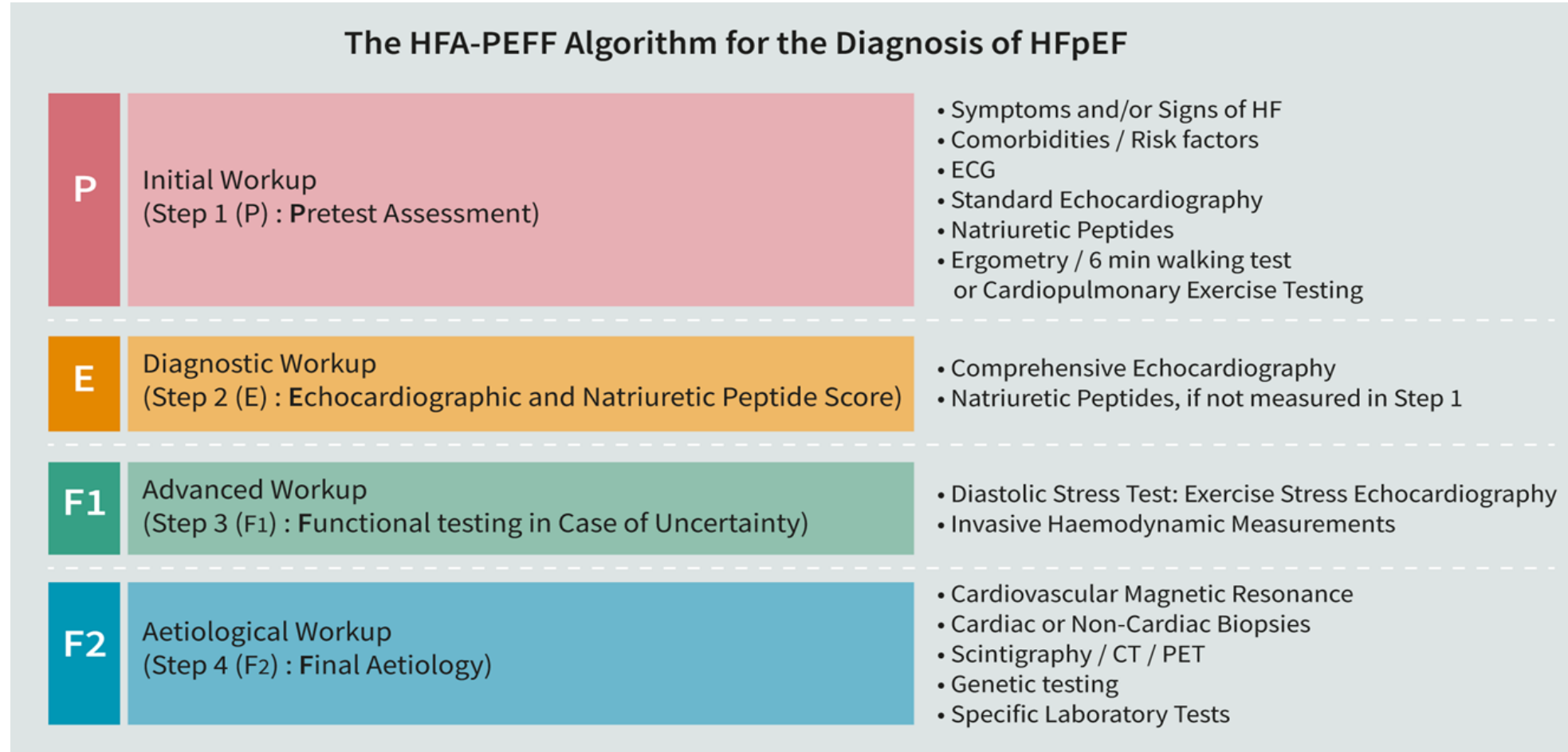
Evaluation and Diagnosis of HFpEF

- HFpEF diagnosis may be more challenging to diagnose than HFrEF since:
 - Laboratory parameters (e.g., BNP, E/e' ratio, ECG data) normal in some cases
- Two scoring systems (H2FPEF, HFA-PEFF)
 - Available to evaluate HFpEF based on:
 - Clinical characteristics + Diagnostic data
(↑ Sensitivity to detect HFpEF)

H₂FPEF Scoring System

	Clinical Variable	Values	Points
H ₂	<u>H</u> heavy	BMI > 30 kg/m ²	2
	<u>H</u> ypertensive	Two or more antihypertensive medications	1
F	Atrial <u>F</u> ibrillation	Paroxysmal or Persistent	3
P	<u>P</u> ulmonary Hypertension	Doppler echocardiographic estimated pulmonary artery systolic pressure > 35 mmHg	1
E	<u>E</u> lder	Age > 60 years	1
F	<u>F</u> illing Pressure	Doppler Echocardiographic E/e' > 9	1
H₂FPEF score (≥ 6 or greater = ≥ 90% of HFpEF probability)			Total (0-9)

Heart Failure Association Algorithm (HFA-PEFF)

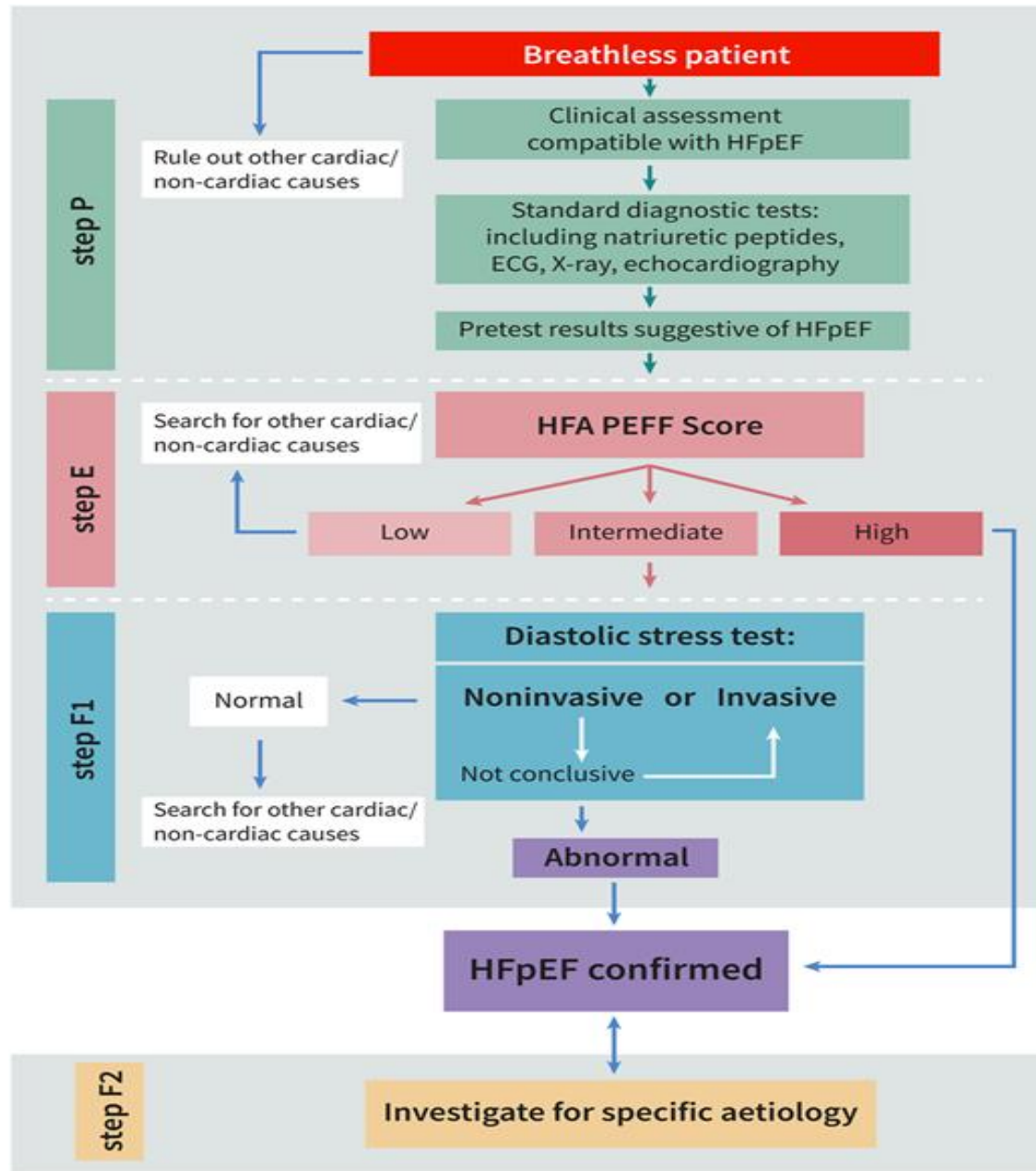


CT, computed tomography; PET, positron emission tomography

Heart Failure Association Algorithm (HFA-PEFF)

	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				

AF: atrial fibrillation, **BNP:** B-type natriuretic peptide, **GLS:** global longitudinal strain, **LAVI:** left atrial volume index, **LV:** left ventricle, **LVMI:** left ventricular mass index, **NT-proBNP:** N-terminal pro b-type natriuretic peptide, **PASP:** pulmonary artery systolic pressure, **RWT:** relative wall thickness, **SR:** sinus rhythm, **TR:** tricuspid regurgitation



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HF Case Presentation

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Case #1: Pt AM

- AM is a 48F with a history of hypertension, pre-DM, class II obesity
- Her EKG shows some LVH
- Current medications: lisinopril 20 mg daily
- On exam: BP 142/85, HR 70 bpm, BMI 38 kg/m²



Case #1: Pt AM

- AM is a 48F with a history of hypertension, pre-DM, class II obesity
- Her EKG shows some LVH
- Current medications: lisinopril 20 mg daily
- On exam: BP 142/85, HR 70 bpm, BMI 38 kg/m²
- You order an echo – grade I diastolic dysfunction, LV posterior wall diameter 1.2 cm (+mild LVH)
- Labs – NT pro-BNP 100 pg/mL, eGFR 65 ml/min/1.73 m²
- She reports compliance with all of her medications, and attends all of her scheduled office visits



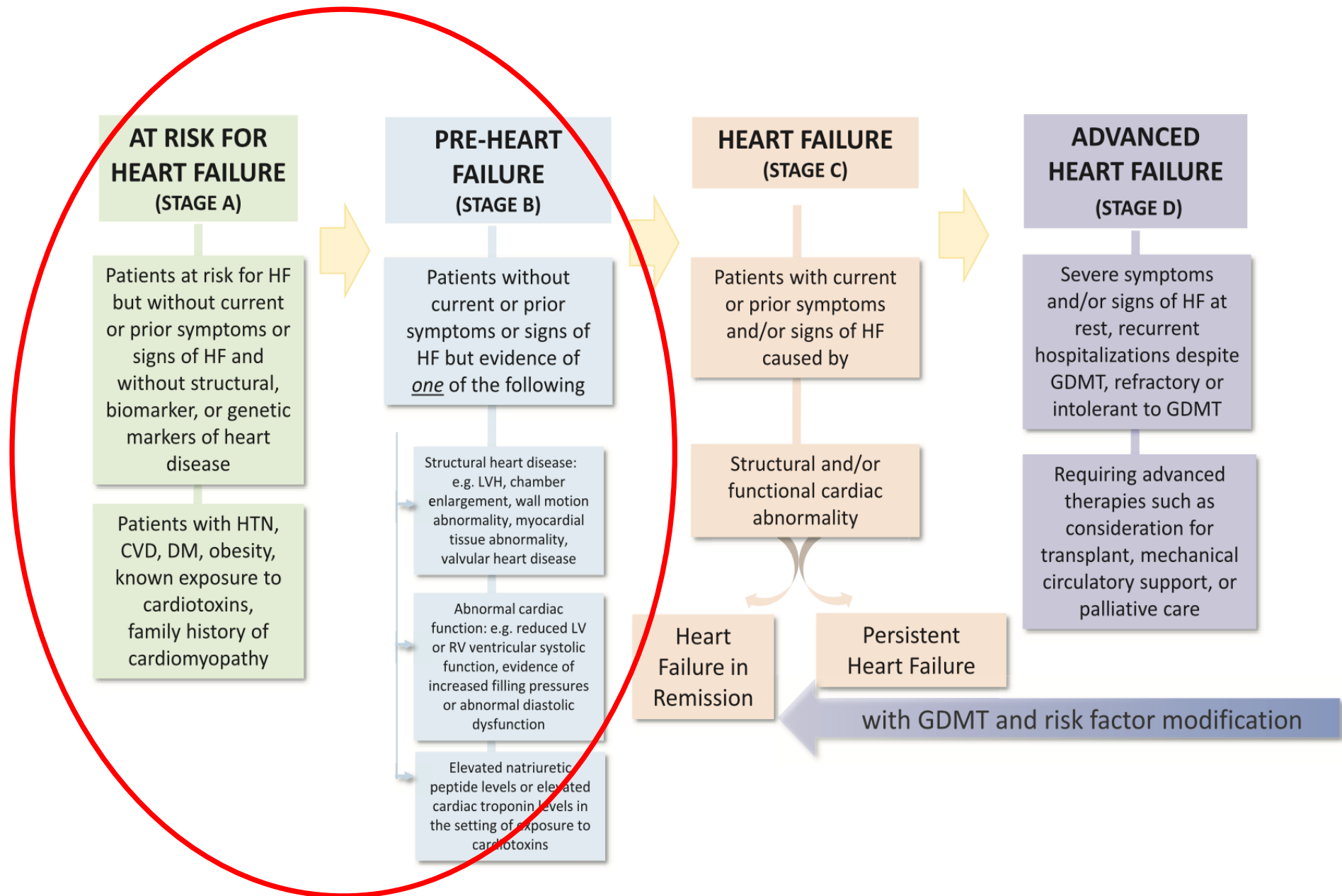
Case #1: Pt AM

- How would you optimize her medical management?
 - A. Increase lisinopril to 40 mg qd
 - B. Increase lisinopril to 40 mg qd and add semaglutide 2.4 mg qd
 - C. Add empagliflozin 10 mg qd
 - D. Add isosorbide mononitrate 30 mg qd

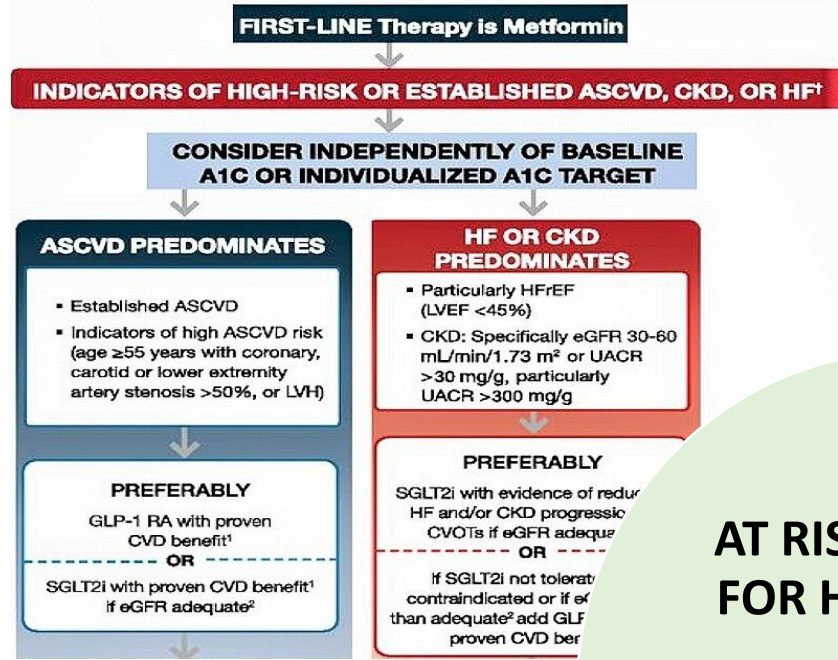


Case #1: Pt AM

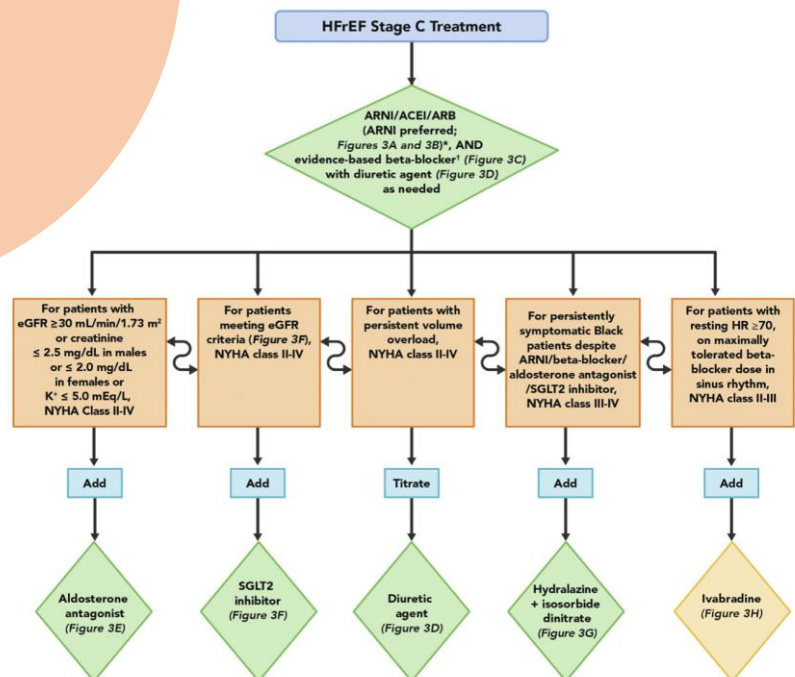
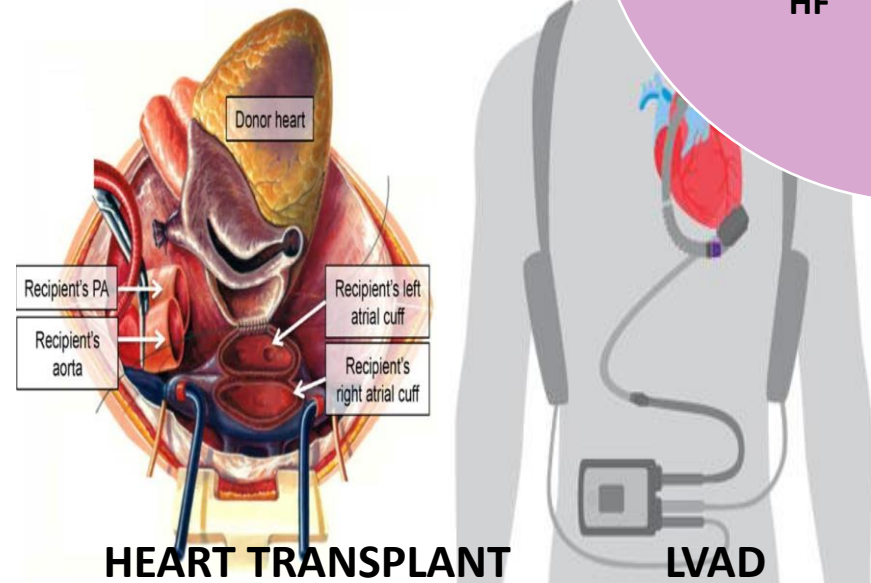
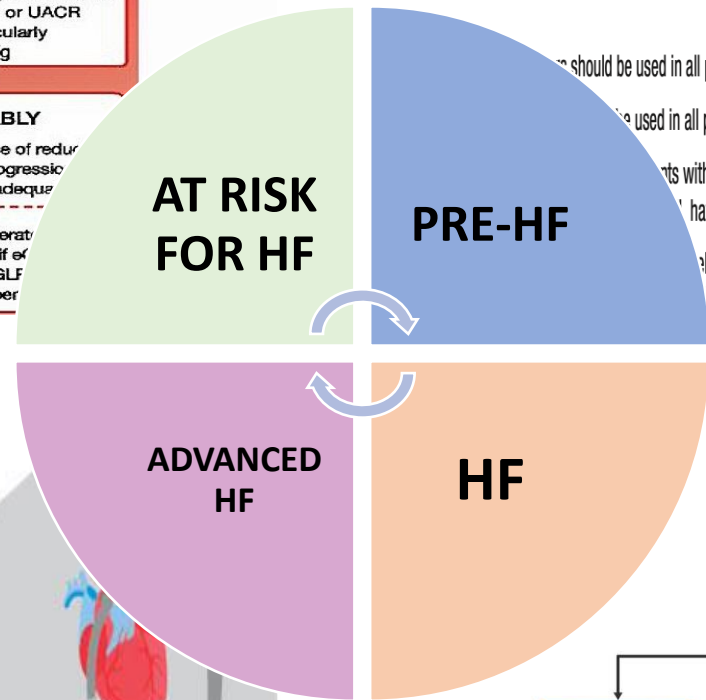
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Treatment Landscape for HF

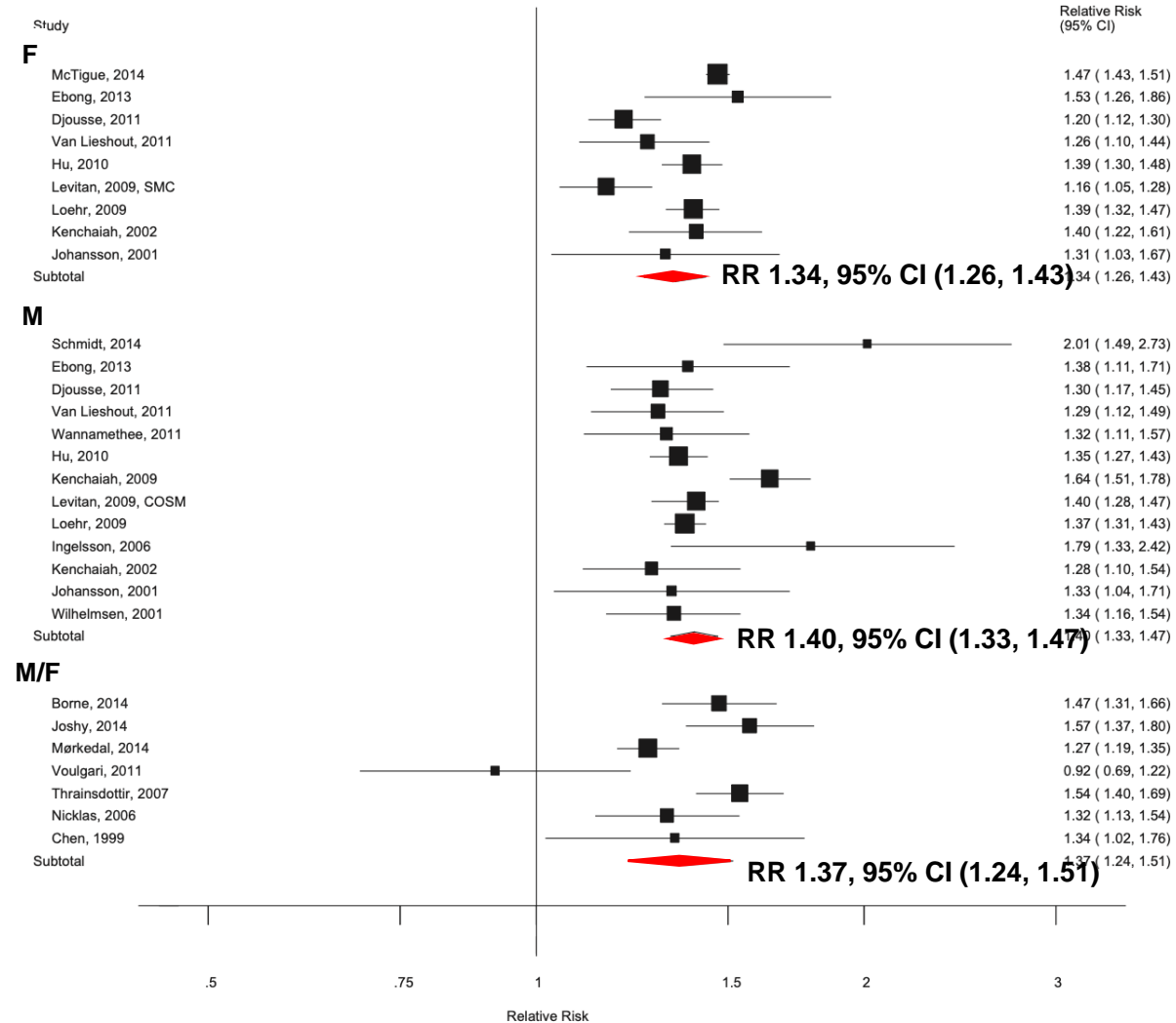


Recommendations	COR	LOE
In patients with a history of MI and reduced EF, ACE inhibitors or ARBs should be used to prevent HF	I	A
In patients with MI and reduced EF, evidence-based beta blockers should be used to prevent HF	I	B
In patients with MI, statins should be used to prevent HF	I	A
Blood pressure should be controlled to prevent symptomatic HF	I	A
ARNI should be used in all patients with a reduced EF to prevent HF	I	A
ARNI should be used in all patients with a reduced EF to prevent HF	I	C
ARNI should be used in patients with asymptomatic ischemic cardiomyopathy who have an LVEF $\leq 30\%$, and on GDMT	IIa	B
ARNI should be used in patients with asymptomatic ischemic cardiomyopathy who have an LVEF $\leq 30\%$, and on GDMT	III: Harm	C



Obesity Increases the Risk for HF

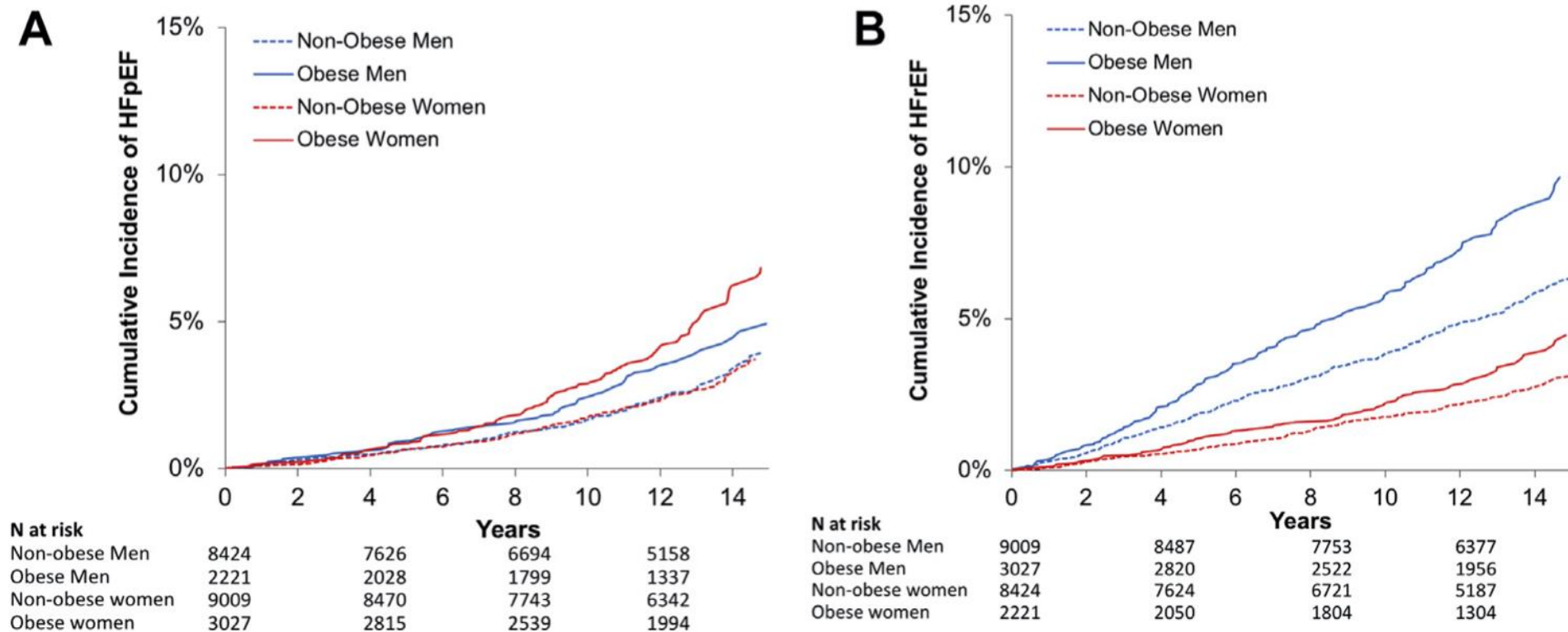
BMI and heart failure incidence by sex, dose-response analysis, per 5 units



Relation of HF Subtypes among Obese and Non-Obese

N=22,681 participants from 4 community-based cohorts in the US
(23% from CHS, 15% from FHS, 29% from MESA, and 32% from PREVEND)

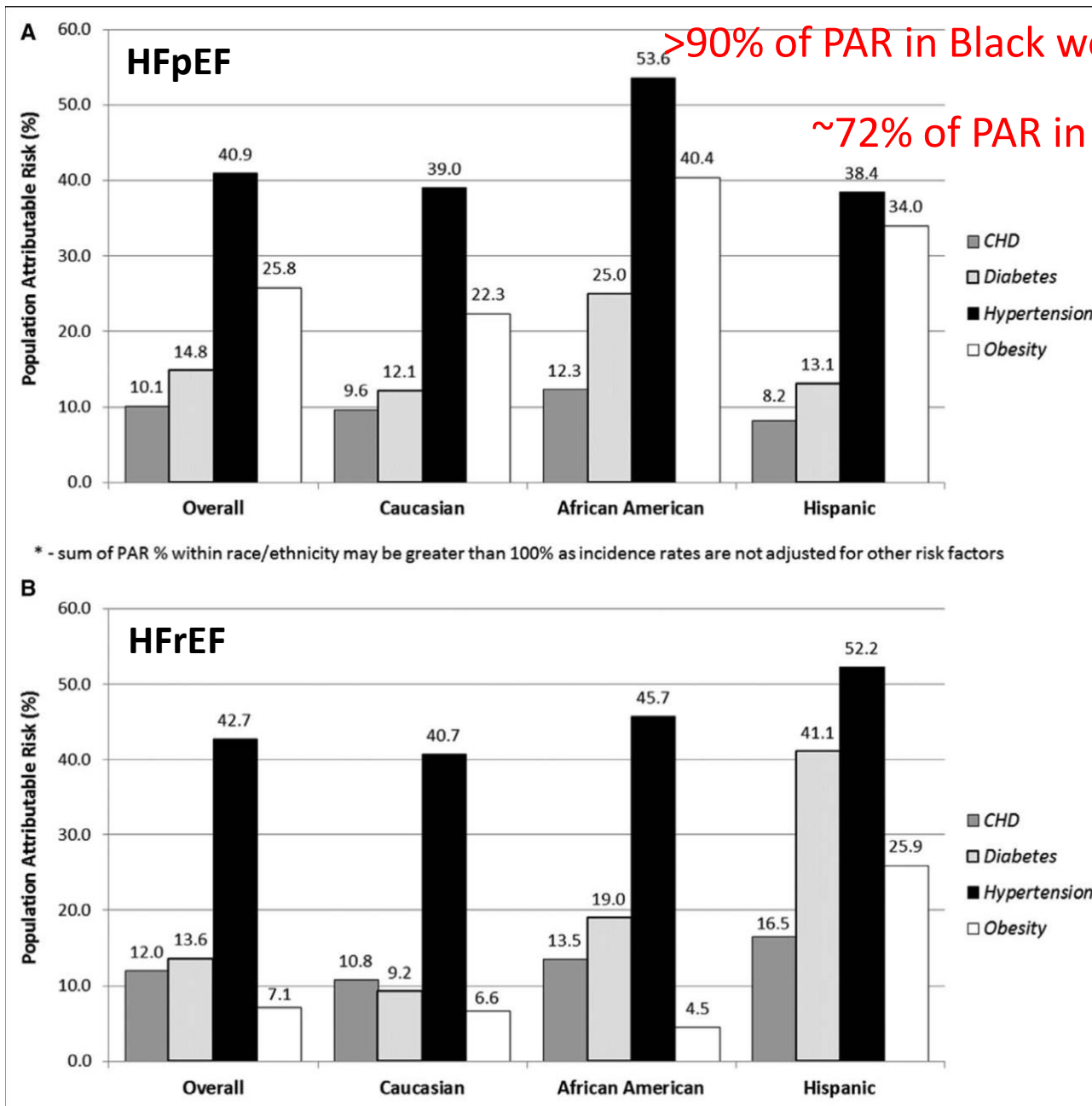
FIGURE 1 Cumulative Incidence of HF Subtypes Among Obese and Nonobese Men and Women



Obesity Harbors a Greater risk for HFpEF in Women From Race-Ethnic Minority Groups

N= 42170 postmenopausal women mean follow-up 13.2 yrs incident hospitalized HF

Risk factor	HFpEF HR (95% CI)* Total	HFrEF HR (95% CI) Total	HFpEF HR (95% CI) white	HFrEF HR (95% CI) White	HFpEF HR (95% CI) African American	HFrEF HR (95% CI) African American	HFpEF HR (95% CI) Hispanic	HFrEF HR (95% CI) Hispanic
Age (ref=50–59 y)	†	‡	†	‡	§			
60–69	2.46 (1.95–3.10)	1.48 (1.11–1.97)	2.82 (2.05–3.87)	1.97 (1.30–2.97)	2.03 (1.40–2.94)	1.16 (0.75–1.81)	2.74 (1.25–6.04)	0.93 (0.34–2.57)
70–69	5.22 (4.05–6.73)	2.76 (2.01–3.79)	6.24 (4.49–8.67)	3.80 (2.48–5.84)	4.03 (2.61–6.21)	1.74 (1.01–3.01)	2.46 (0.75–8.02)	2.08 (0.62–6.98)
BMI (ref=BMI <25 kg/m ²)	†		†		†			
25–<30	1.11 (0.88–1.40)	0.91 (0.68–1.21)	1.00 (0.78–1.29)	0.87 (0.62–1.21)	3.57 (1.40–9.08)	1.10 (0.60–2.03)	1.39 (0.43–4.43)	1.10 (0.21–5.65)
30–<35	1.35 (1.06–1.72)	1.00 (0.74–1.36)	1.08 (0.81–1.43)	1.12 (0.79–1.60)	6.27 (2.49–15.77)	0.81 (0.41–1.59)	0.90 (0.23–3.44)	1.75 (0.32–9.66)
≥35	2.36 (1.84–3.03)	0.87 (0.61–1.24)	2.10 (1.57–2.80)	0.69 (0.43–1.11)	7.50 (2.96–18.98)	1.09 (0.56–2.13)	4.29 (1.24–14.90)	3.09 (0.48–19.80)
Physical activity (ref=<1.25 MET h/wk)								
1.25–<6.25	0.91 (0.75–1.11)	0.92 (0.70–1.20)	0.94 (0.74–1.20)	0.95 (0.68–1.34)	0.77 (0.52–1.13)	0.81 (0.49–1.33)	1.62 (0.53–4.12)	1.08 (0.30–3.88)
6.25–<15.3	0.81 (0.66–1.00)	0.72 (0.54–0.96)	0.83 (0.65–1.07)	0.72 (0.50–1.03)	0.73 (0.48–1.11)	0.75 (0.44–1.27)	0.87 (0.29–2.60)	0.45 (0.08–2.47)
≥15.3	0.74 (0.59–0.93)	0.74 (0.54–1.00)	0.75 (0.57–0.98)	0.77 (0.53–1.12)	0.65 (0.41–1.03)	0.59 (0.32–1.09)	1.32 (0.46–3.82)	1.34 (0.30–6.08)

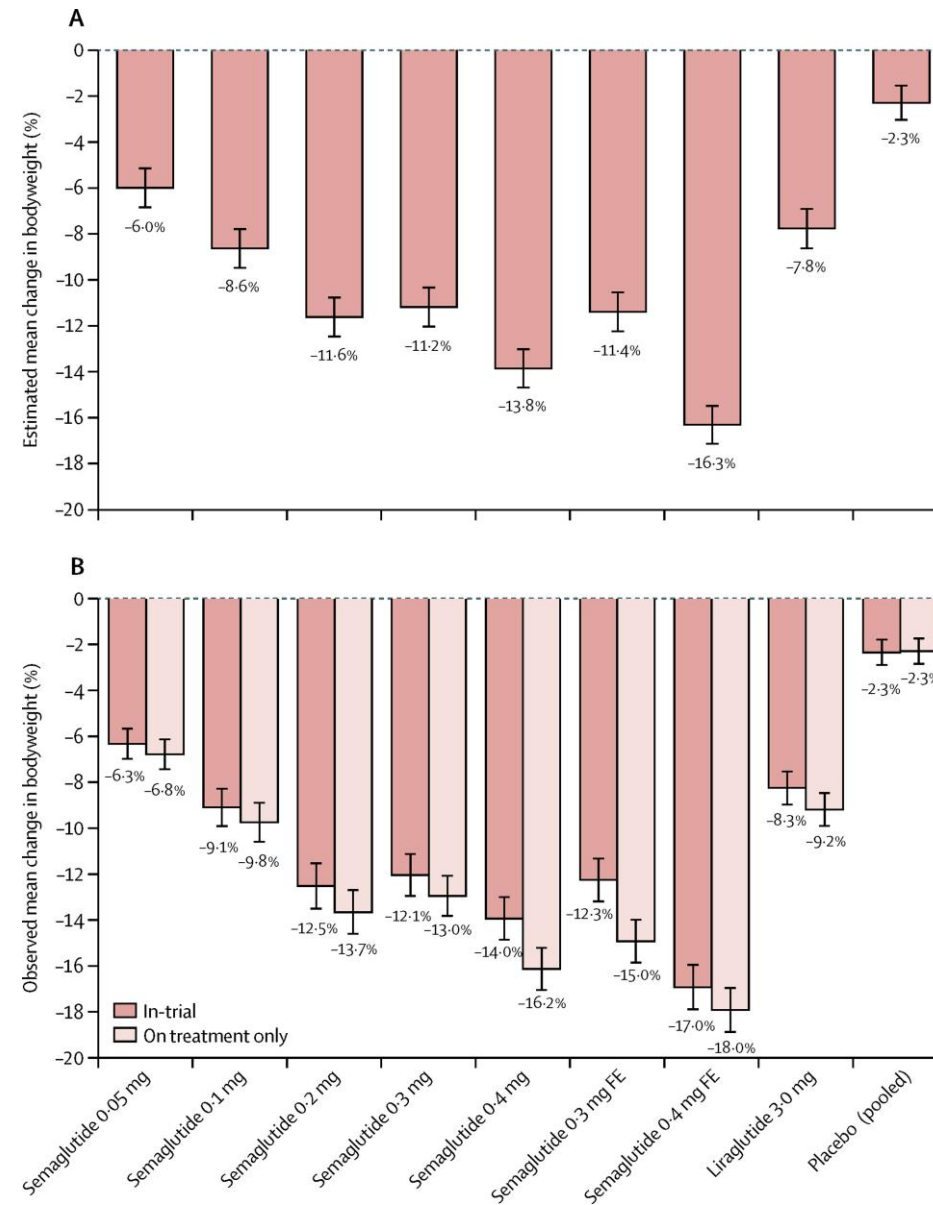


>90% of PAR in Black women due to HTN and obesity

~72% of PAR in Hispanic women due to HTN and obesity

- N= 42170 postmenopausal women
- mean follow-up 13.2 yrs
- incident hospitalized HFpEF and HFrEF

Opportunities for Prevention – Obesity



Conclusions

- Patients ***at risk for HF (Stage A)*** or with ***pre-HF (stage B)*** constitute the largest group
 - Patients *at risk* should be treated with medical therapy that can prevent HF
 - *Pre-HF* patients, such as asymptomatic patients with elevated natriuretic peptide levels, may require referral to a cardiologist for further diagnostic and treatment strategies to prevent progression of HF
- Patients with ***HF (Stage C)*** can receive a timely diagnosis using a combination of elevated natriuretic peptide levels, or recognition of evidence of systemic or pulmonary congestion/elevated filling pressures