

# MEDICAL MANAGEMENT OF HEART FAILURE: WHAT EVERY CLINICIAN SHOULD KNOW

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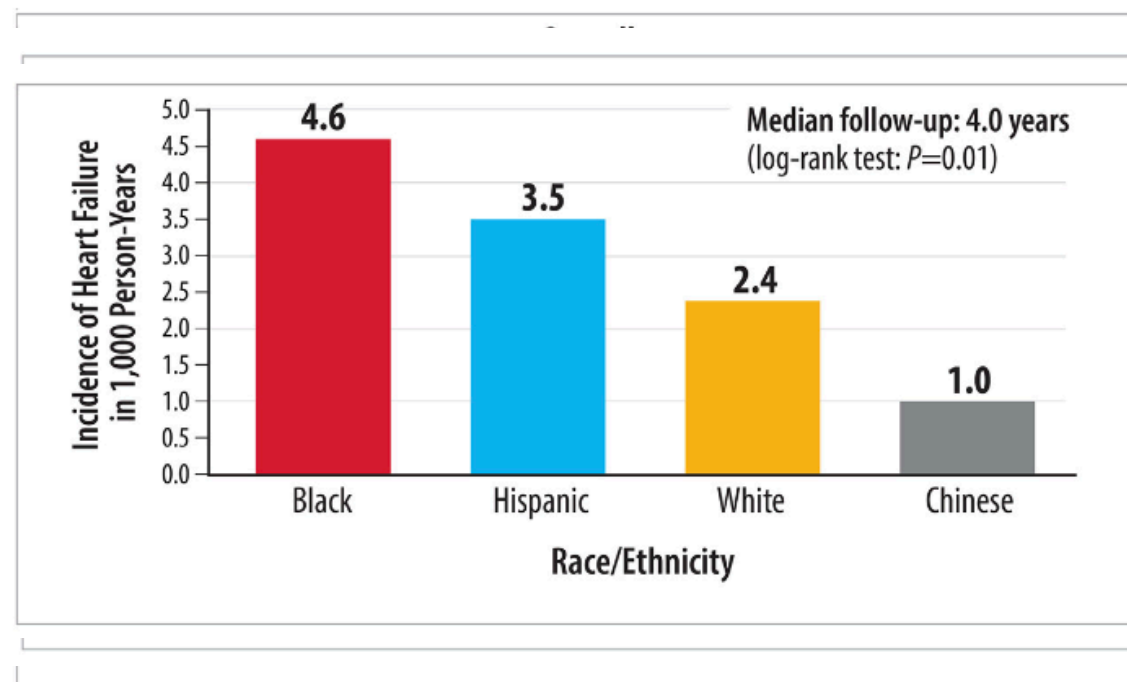
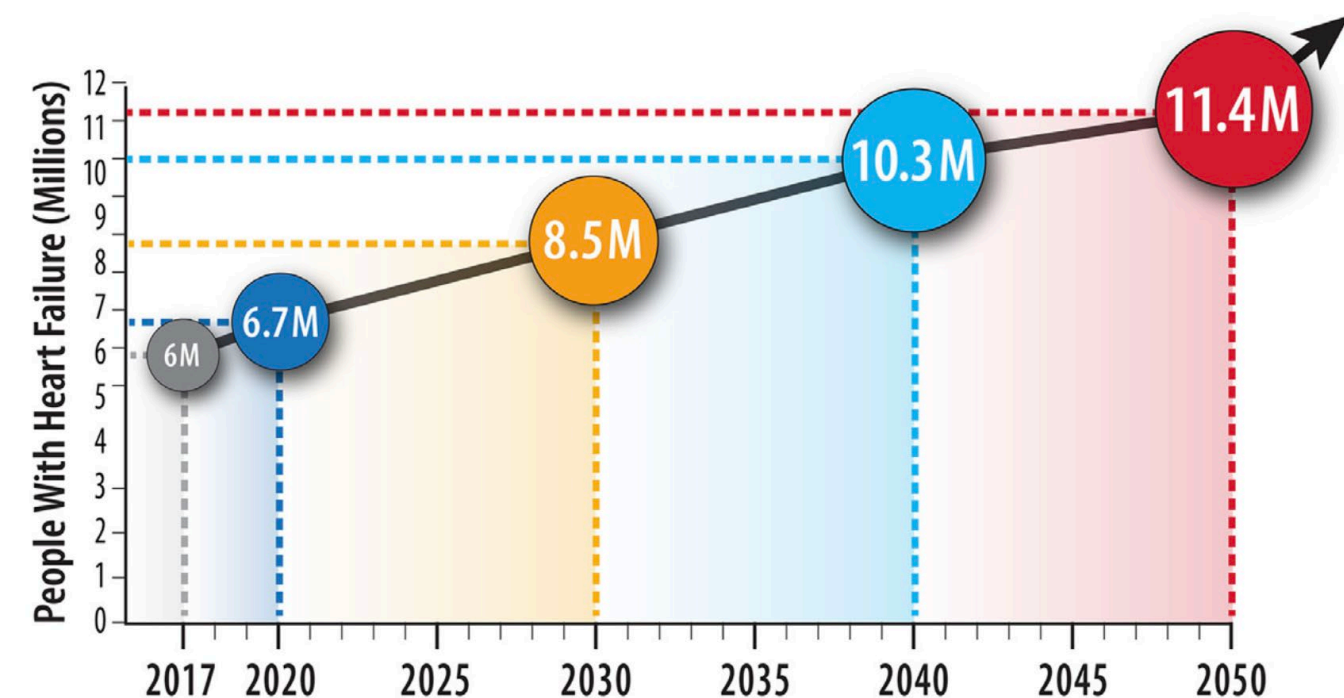
ADVANCED HEART FAILURE & TRANSPLANT CARDIOLOGY

GILL HEART & VASCULAR INSTITUTE

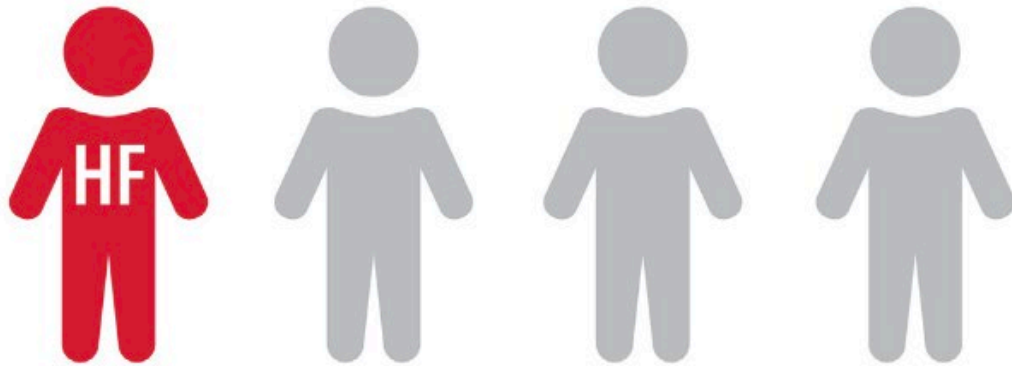
UNIVERSITY OF KENTUCKY



# Prevalence and Incidence of Heart Failure

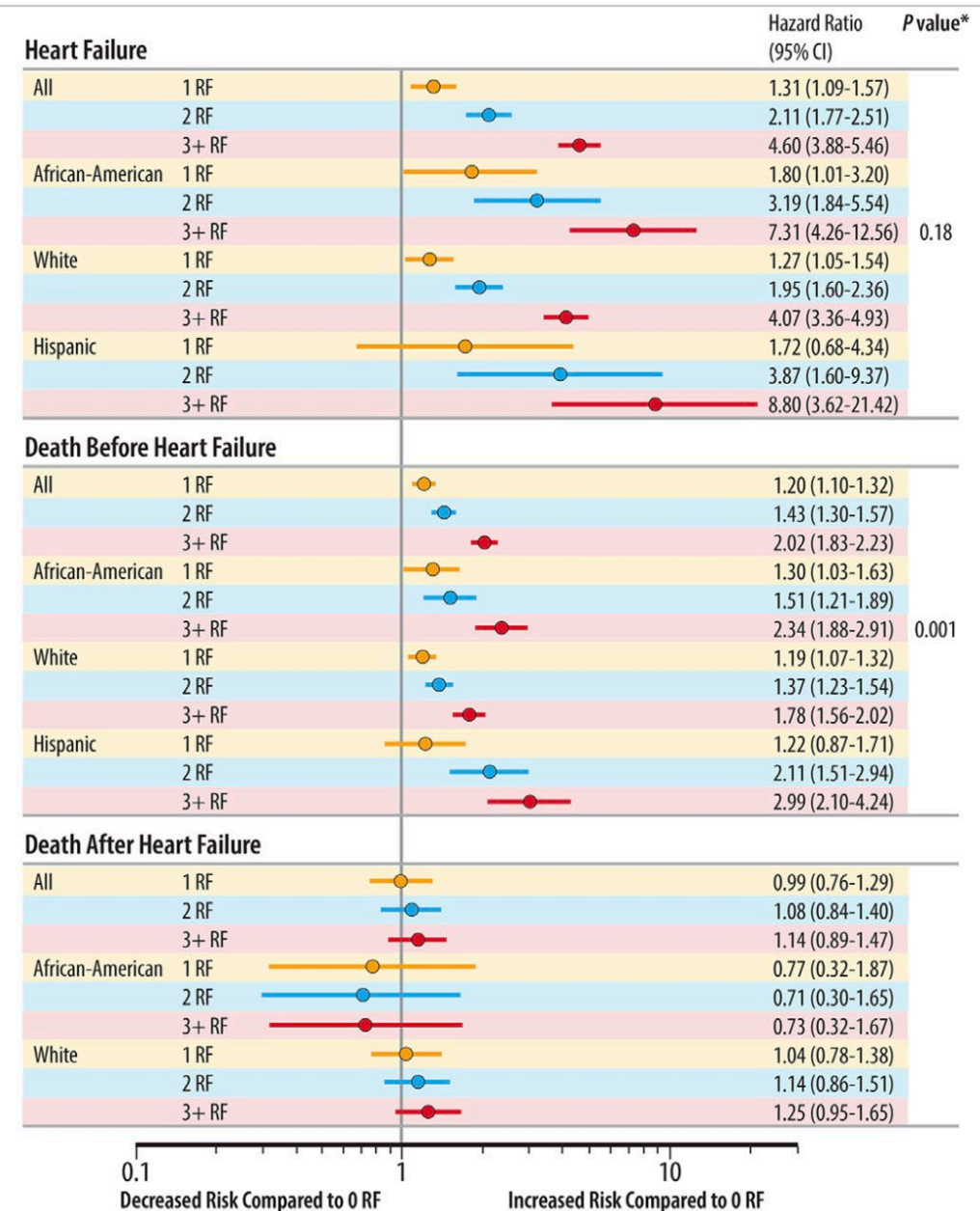


# Lifetime Risk of Heart Failure



**The lifetime risk of heart failure (HF)  
is 1 in 4 people.**

WRITING COMMITTEE  
MEMBERS. HF STATS 2025. *J  
Card Fail.* Published online  
August 29, 2025.



\*P value for interaction race/ethnicity by risk factor (RF) number.

The model is fully adjusted for menopausal hormone therapy status, age, and socioeconomic status.  
The reference is zero risk factors. CI indicates confidence interval.

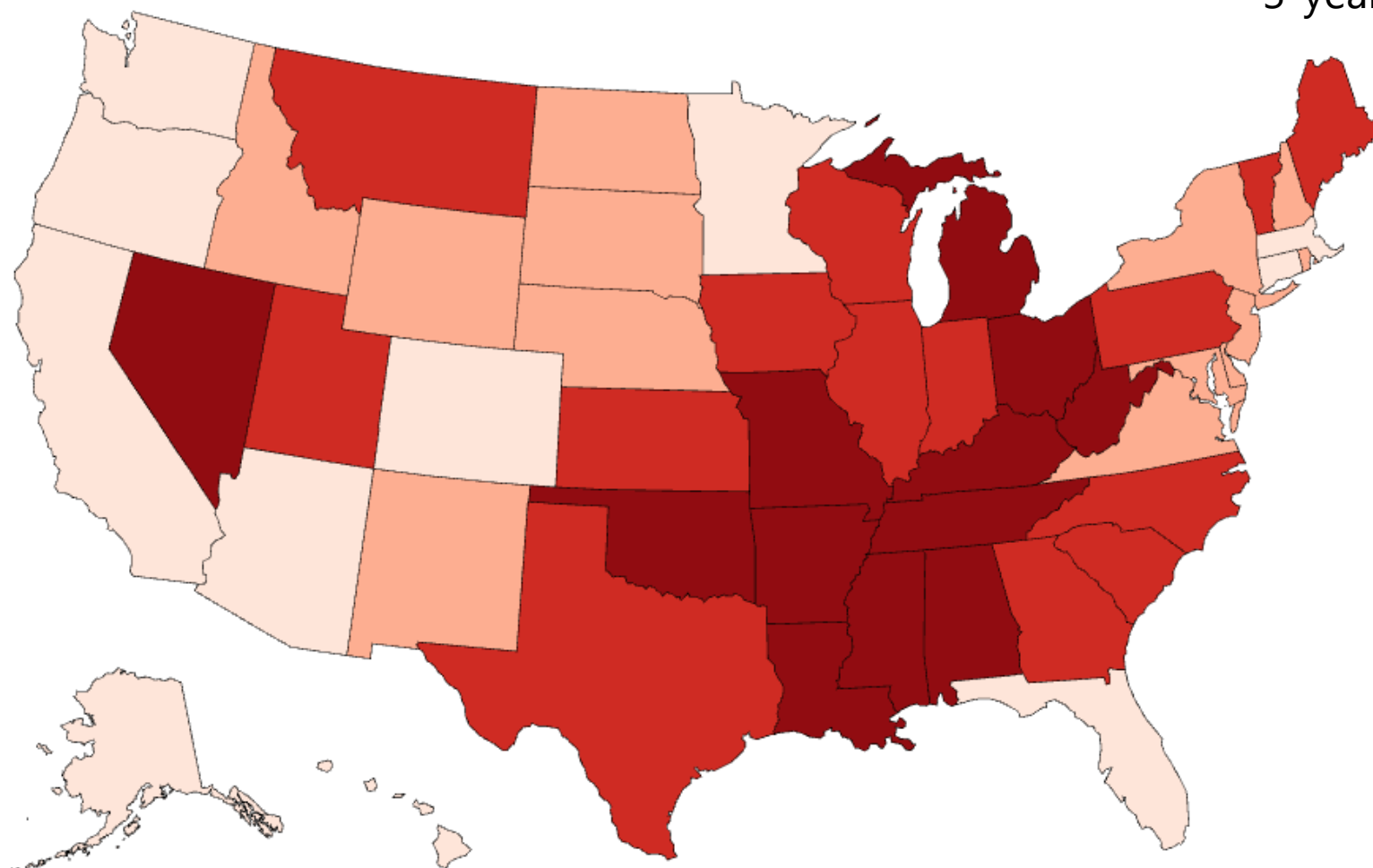


How important is the family physician in the care of heart failure patients?



# Heart Disease Death Rates in the United States

3-year averages, 2021-2023, ages 35+



United States by State

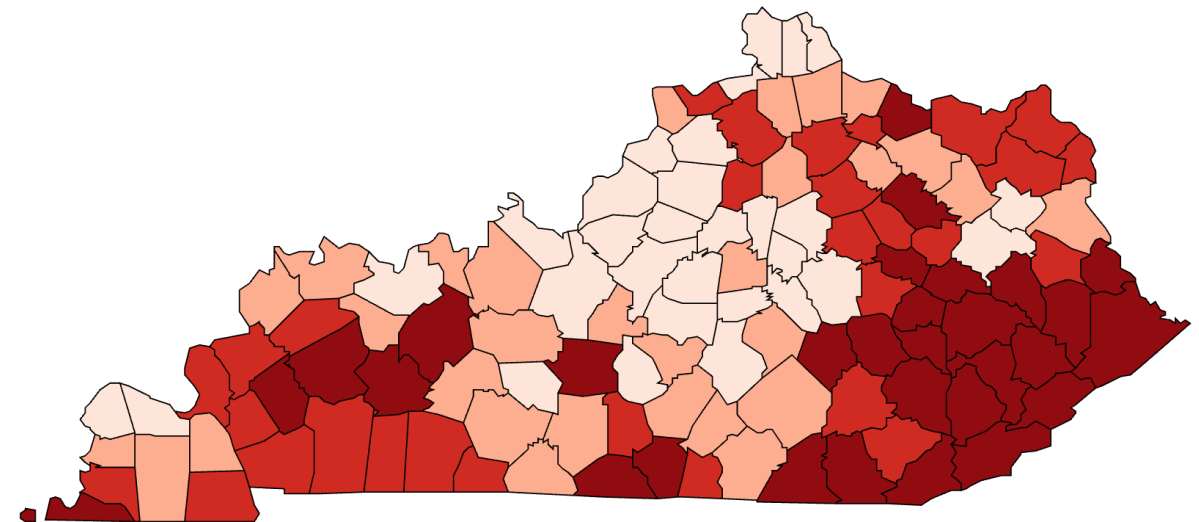
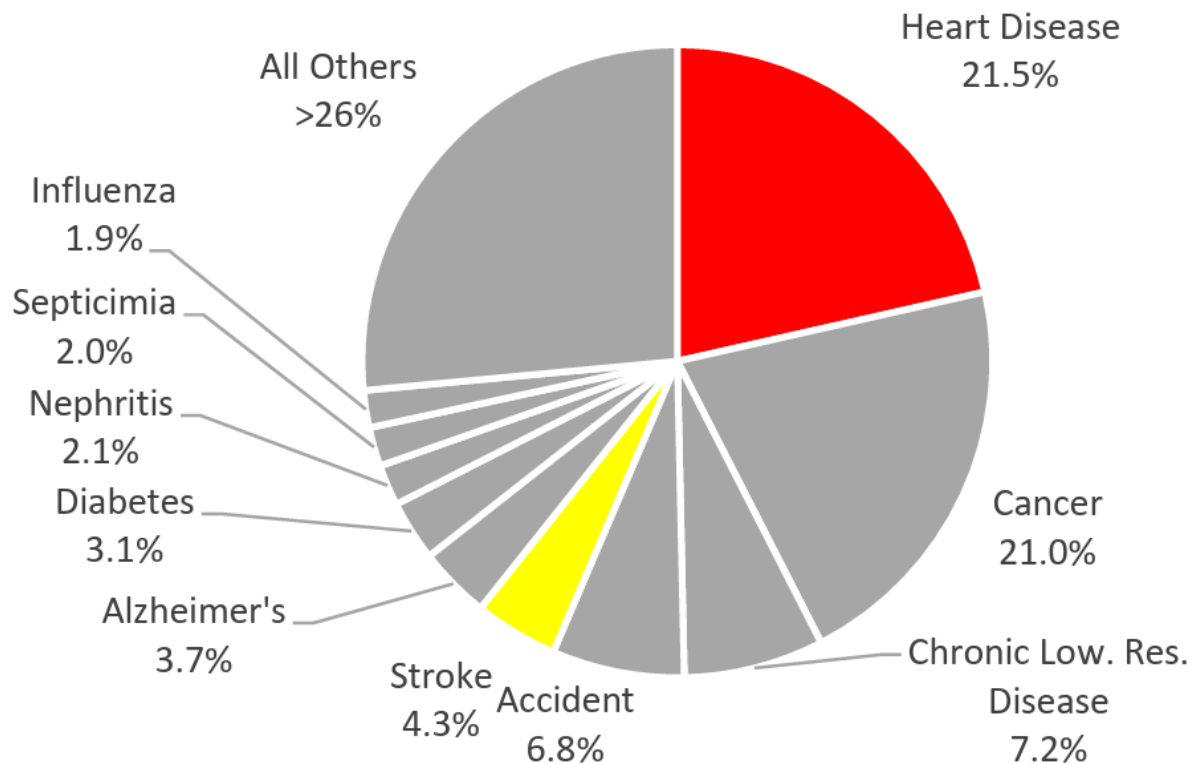
Heart Disease  
Death Rate per 100,000\*

197 - 291 292 - 313 314 - 375  
376 - 554 Insufficient Data

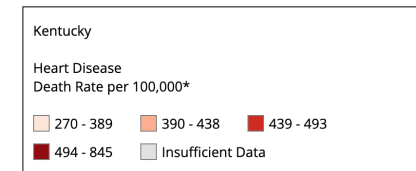
National Rate: 325.5

# How are we doing in Kentucky with regards to heart disease?

## Leading Causes of Death in Kentucky 2017



CDC.gov

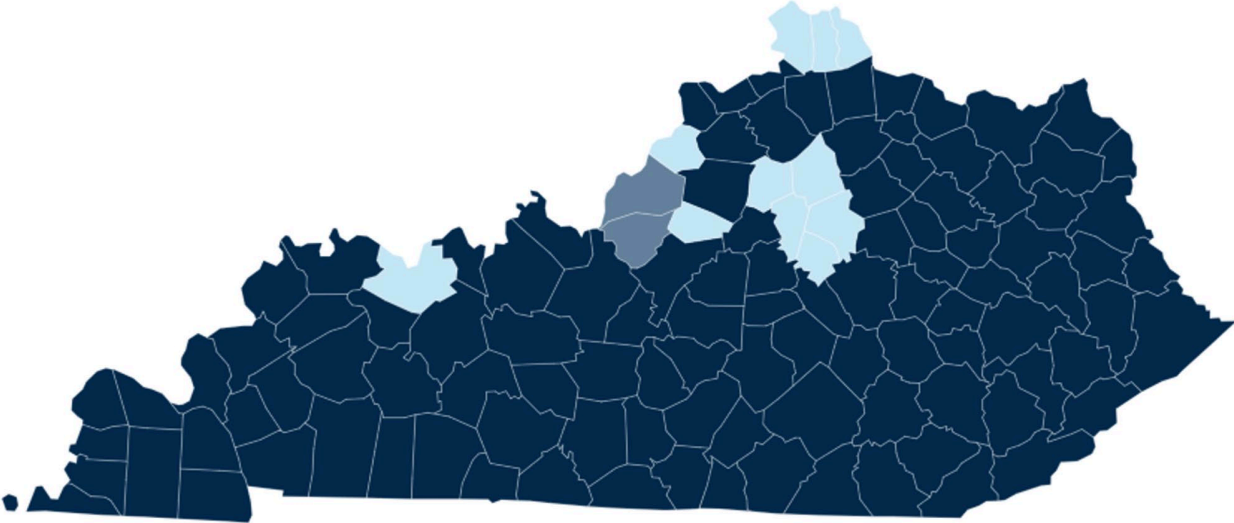
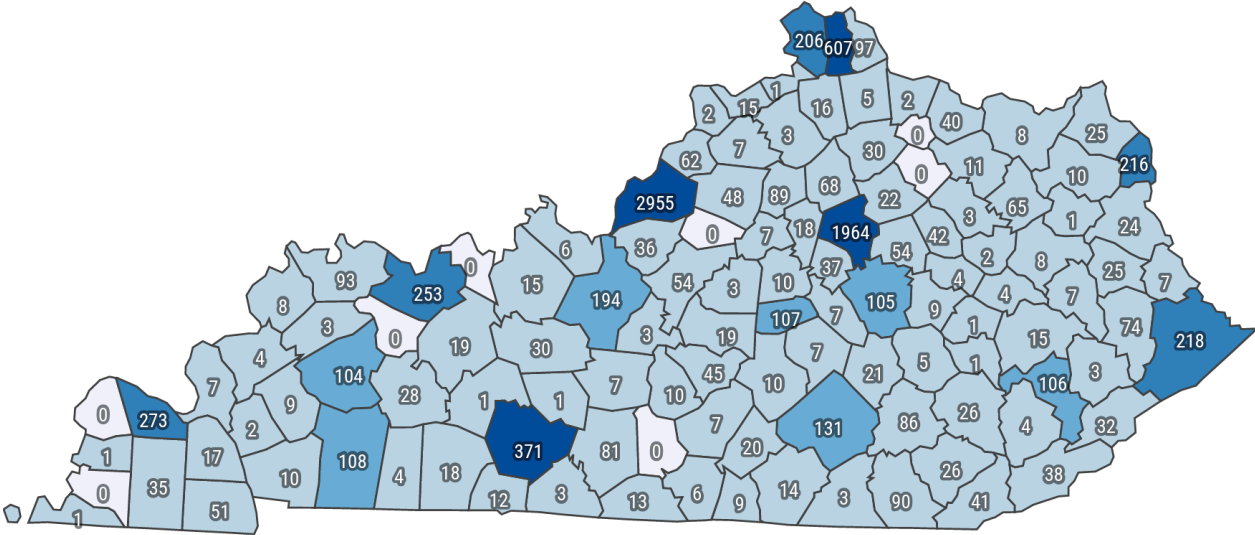
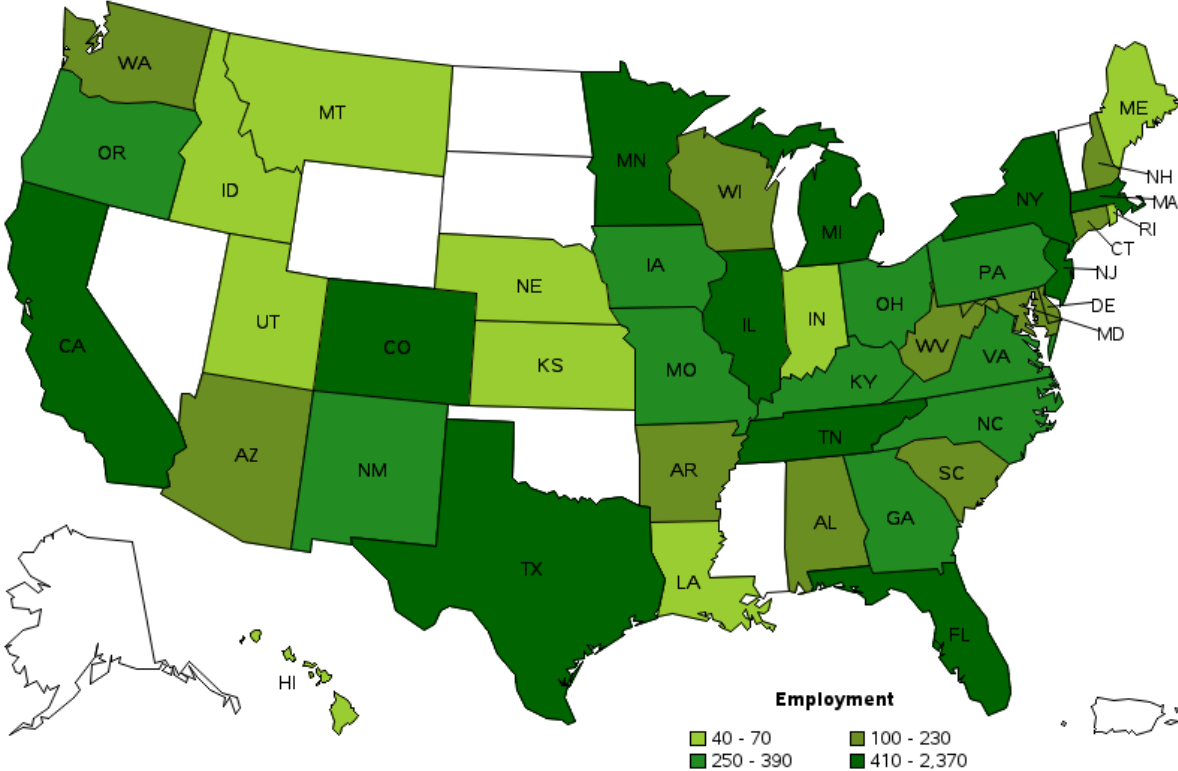


Kentucky has the 8th highest death rate from cardiovascular disease in the country.

Distribution of Physicians By County

What is the physician population of Kentucky?

Employment of cardiologists, by state, May 2022

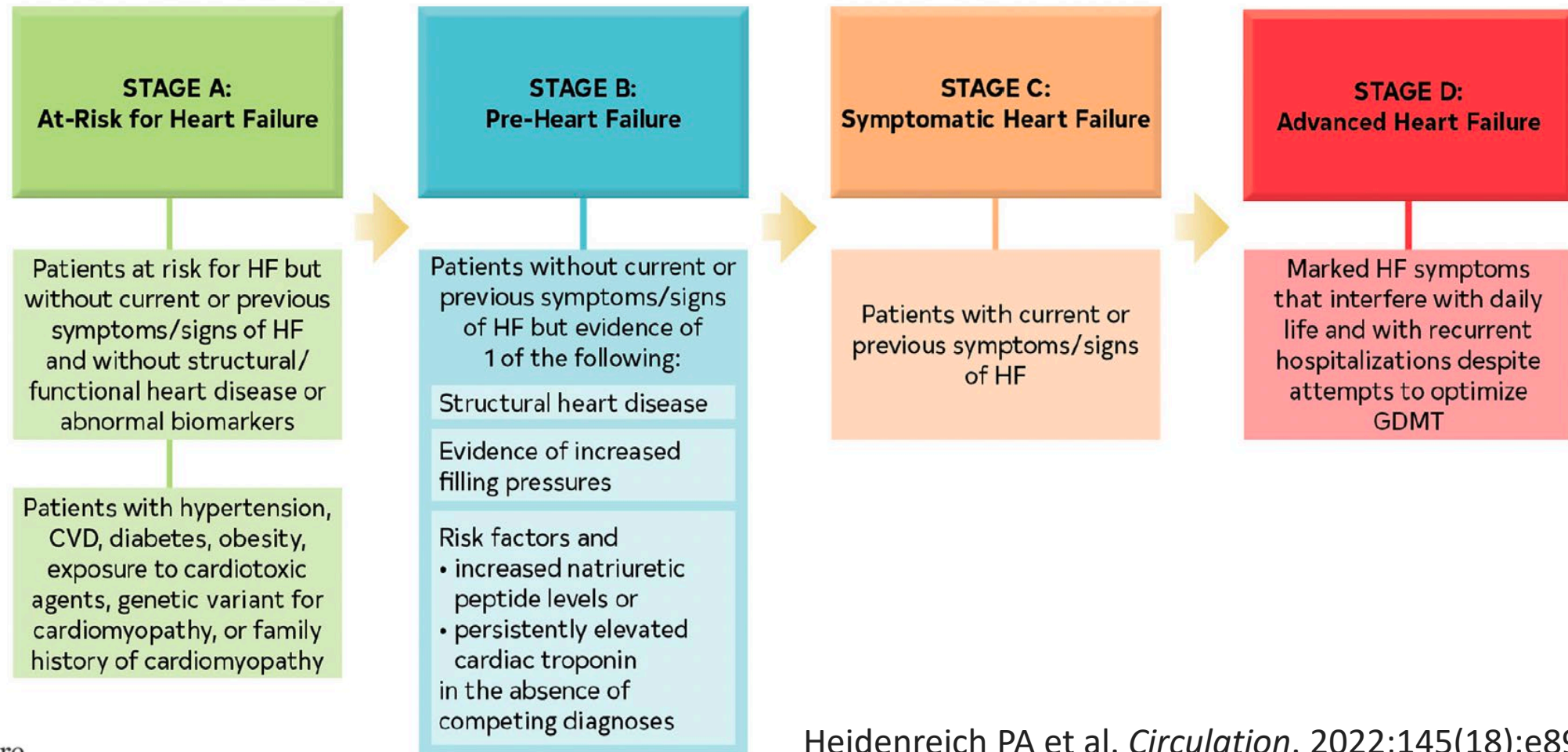


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## Primary Prevention of Heart Failure



# Stages of Heart Failure



Heidenreich PA et al. *Circulation*. 2022;145(18):e895-e1032.



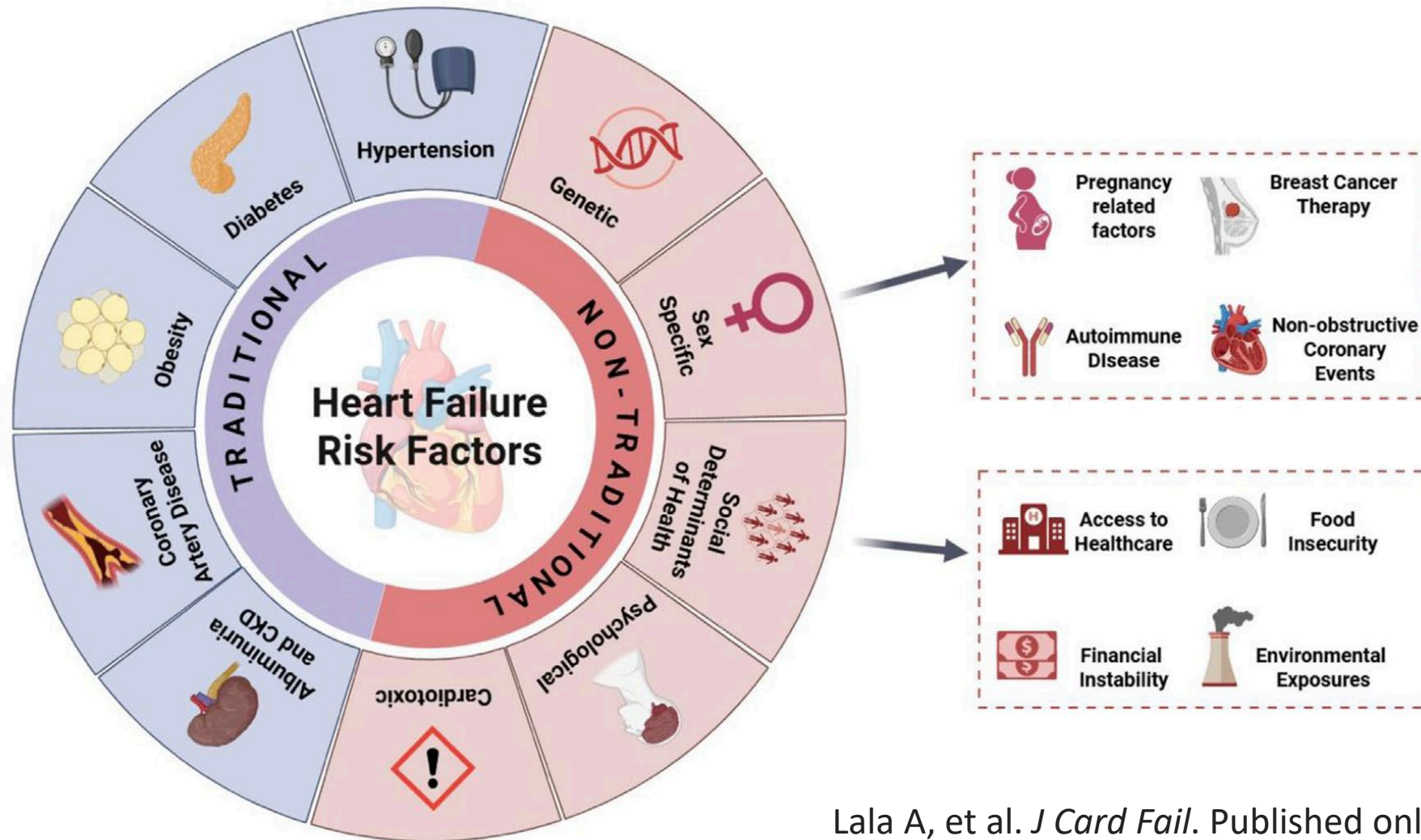
# HEART FAILURE PREVENTION

| PRIMARY PREVENTION   |  | SECONDARY PREVENTION                  |                                    | TERTIARY PREVENTION   |
|--|--|---------------------------------------|------------------------------------|---|
| STAGE A<br>(At Risk)   | STAGE B<br>(Pre-HF)                                      | STAGE C <sup>1</sup><br>(Symptomatic) | Stage D <sup>1</sup><br>(Advanced) | POST-ADVANCED<br>THERAPY<br>(OHT, VAD)                      |
|  | ICD IF LVEF≤30%<br>(>40 DAYS POST MI<br>IN ICM)          | ICD/CRT (when appropriate)            |                                    | ASA, LIPID LOWERING<br>THERAPIES (CAV<br>PREVENTION IN OHT) |
|  |  | ARNi (LVEF<normal); ACE/ARB           |                                    |   |
|  |  | MRA (LVEF<normal, eGFR>30)            |                                    |   |
|  | BETA BLOCKER,<br>ACEi/ARB if<br>LVEF≤40%; SGLT2i<br>(DM) | BETA BLOCKERS (LVEF≤40%)              |                                    | ANTICOAGULATION<br>(LVAD)                                   |
|  |  | SGLT2i                                |                                    | SGLT2i (DM, CKD)  |
| nsMRA (CKD+DM)   | nsMRA (CKD+DM, LVEF≥ 40%)                                |                                       |                                    |   |
| GLP-1RAs (CKD+ DM) (? benefit if LVEF ≤ 40%)   |  |                                       |                                    |   |
| HF Prevention<br>RISK SCORES   | CARDIAC REHAB  |                                       |                                    |   |
| GENETIC SCREENING/COUNSELING   |  |                                       |                                    |   |
| BIOMARKERS: NATRIURETIC PEPTIDE ASSESSMENT   |  |                                       |                                    |   |
| PSYCHOLOGICAL WELL-BEING   |  |                                       |                                    |   |
| LIFE'S ESSENTIAL EIGHT<br>BP & Lipid Control, DM Management, Exercise, Sleep, Smoking Cessation, Weight Management,<br>Diet & Nutrition Counseling |  |                                       |                                    |   |

Lala A, et al. *J Card Fail*. Published online August 13, 2025.

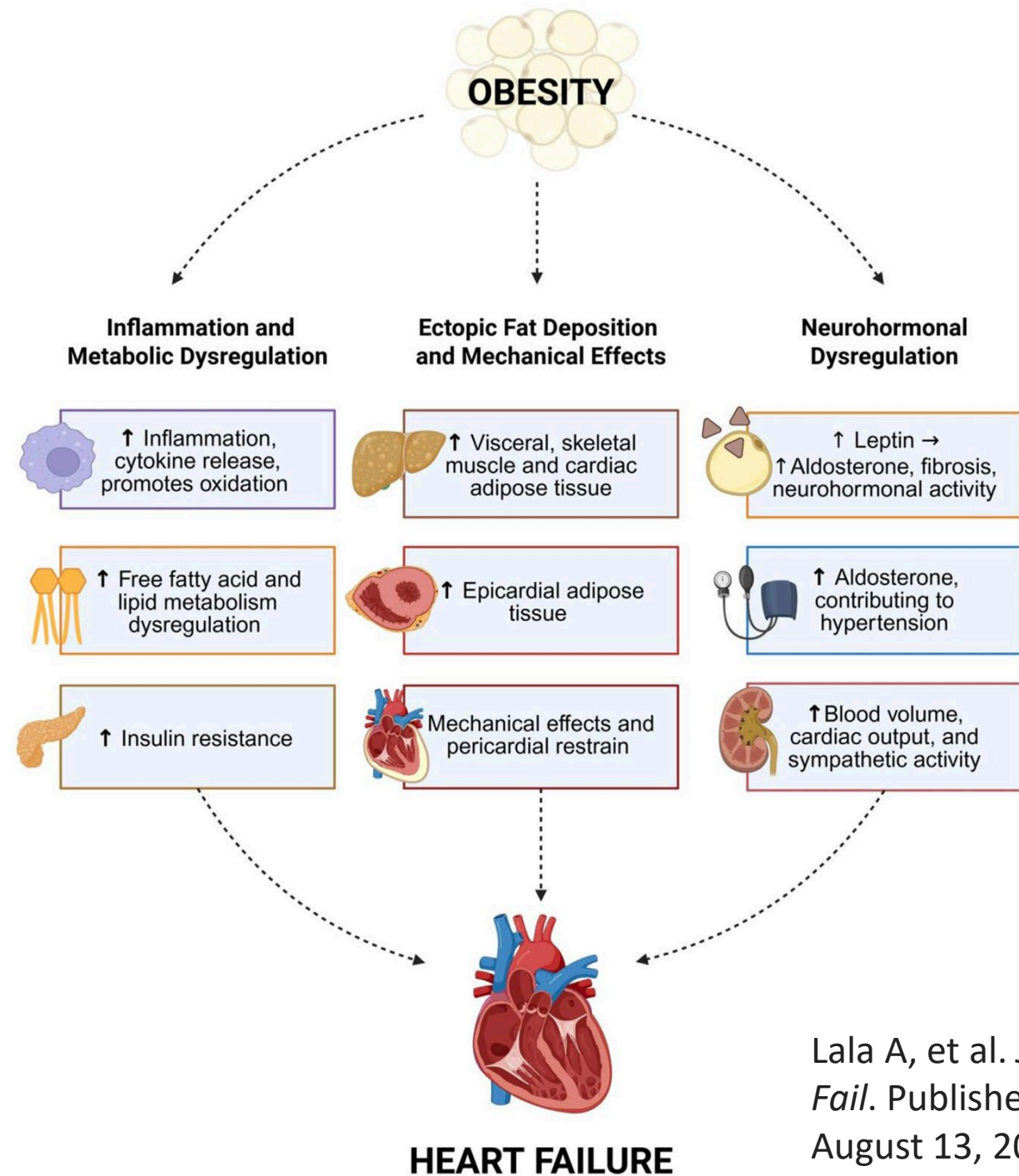


# RISK FACTORS FOR HEART FAILURE



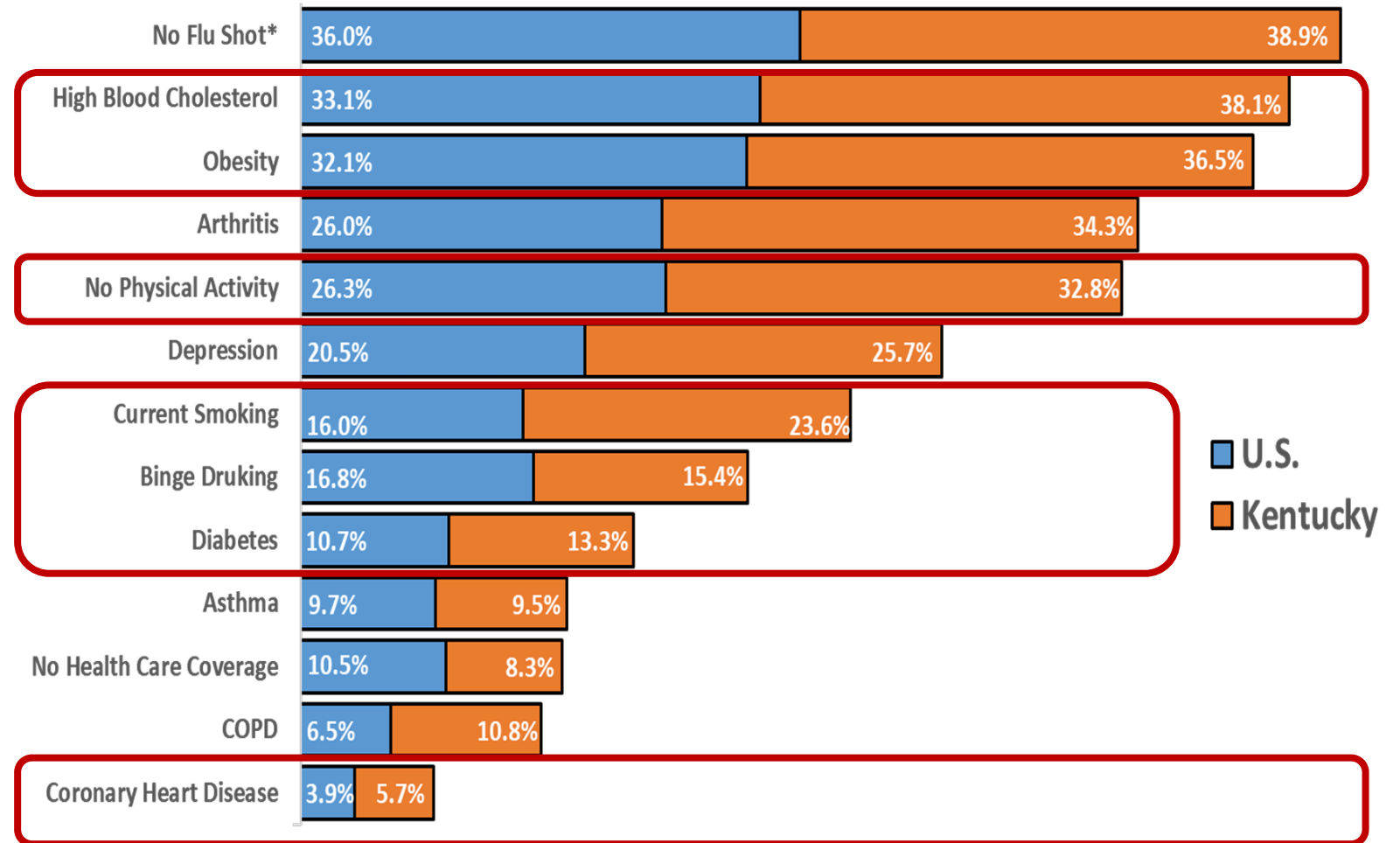
Lala A, et al. *J Card Fail*. Published online August 13, 2025.

# LINK BETWEEN OBESITY AND HEART FAILURE



Lala A, et al. *J Card Fail*. Published online August 13, 2025.

# HEART FAILURE RISK FACTORS IN KENTUCKY



2021 KyBRFS Annual Data Report



## FLU SHOTS AND YOUR HEART



### GET A FLU SHOT TO PROTECT YOUR HEART



1 out of 2 adults hospitalized with the flu also have heart disease

If you have **HEART DISEASE**, you're more likely to have **SERIOUS COMPLICATIONS** from the **FLU**.

These include:

- Pneumonia
- Heart attack
- Hospitalization
- Stroke
- Death

COVID-19 and the flu are a double threat.

### THE FLU VACCINE Can Help You:

- Lower the risk of a heart attack, stroke, or heart failure
- Avoid dangerous complications
- Stay healthy

### WHAT YOU CAN DO

Add a yearly flu shot to the steps you take to keep your heart healthy:



Exercise



Don't Smoke



Eat Heart-Healthy



Take Your Medication



GET VACCINATED!

### WHERE TO GET A FLU SHOT

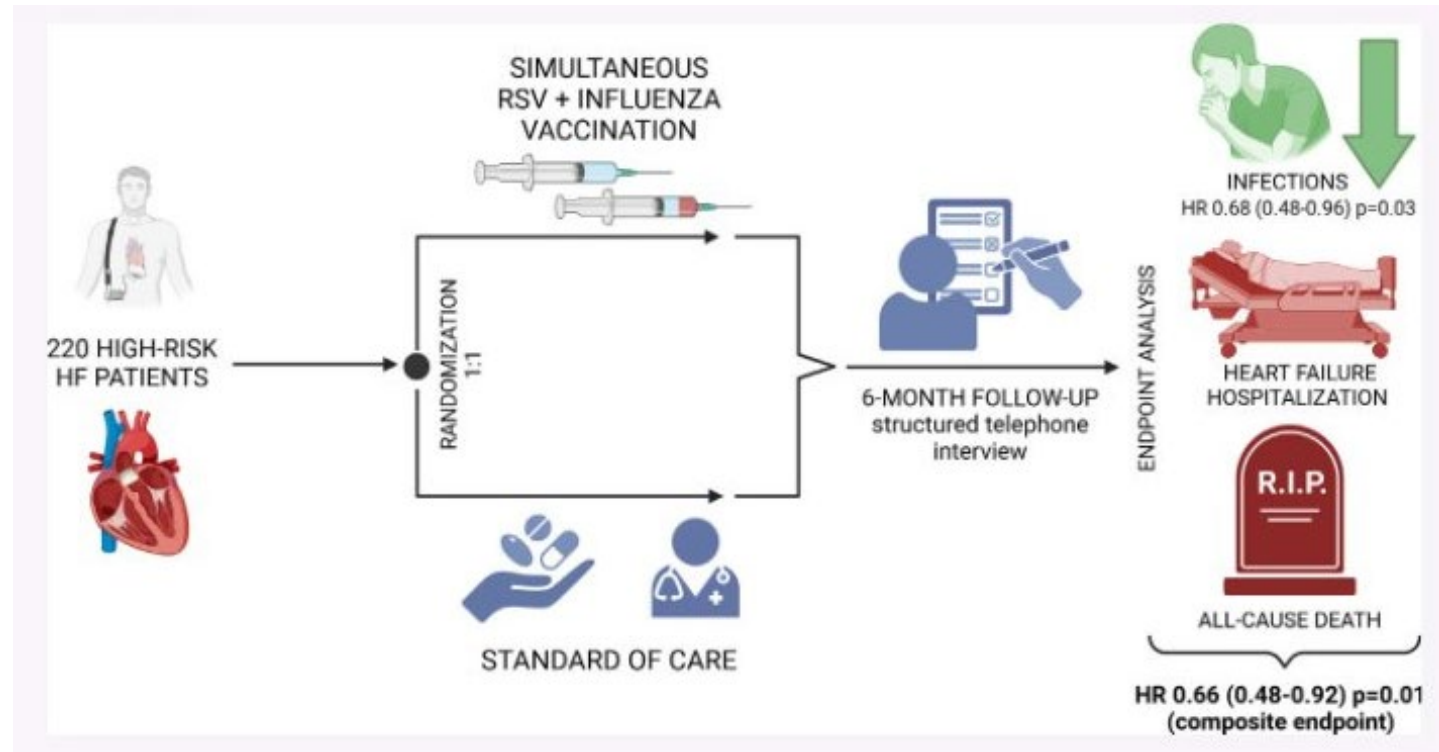
- Your doctor's office
- Your pharmacy
- Your community – go to [VaccineFinder.org](https://www.vaccinefinder.org)

For more information, visit [CardioSmart.org/Flu](https://CardioSmart.org/Flu)

@CardioSmart

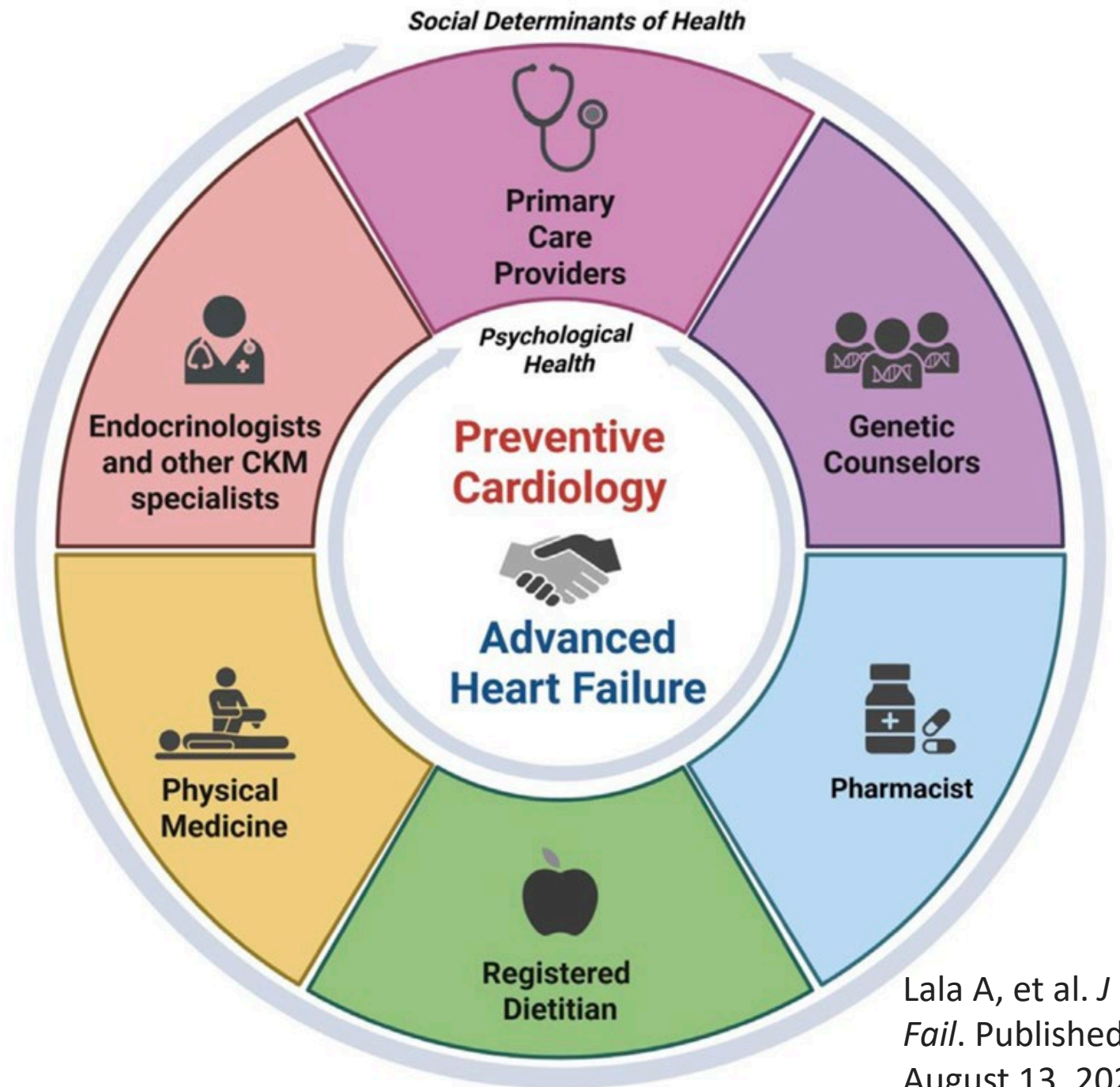
Information provided for educational purposes only. Please talk to your health care professional about your specific health needs. To download or order posters or other topics, visit [CardioSmart.org/Posters](https://CardioSmart.org/Posters)

# FLU VACCINATION



Biegus J, et al. *ESC Heart Fail*. Published online September 25, 2025.

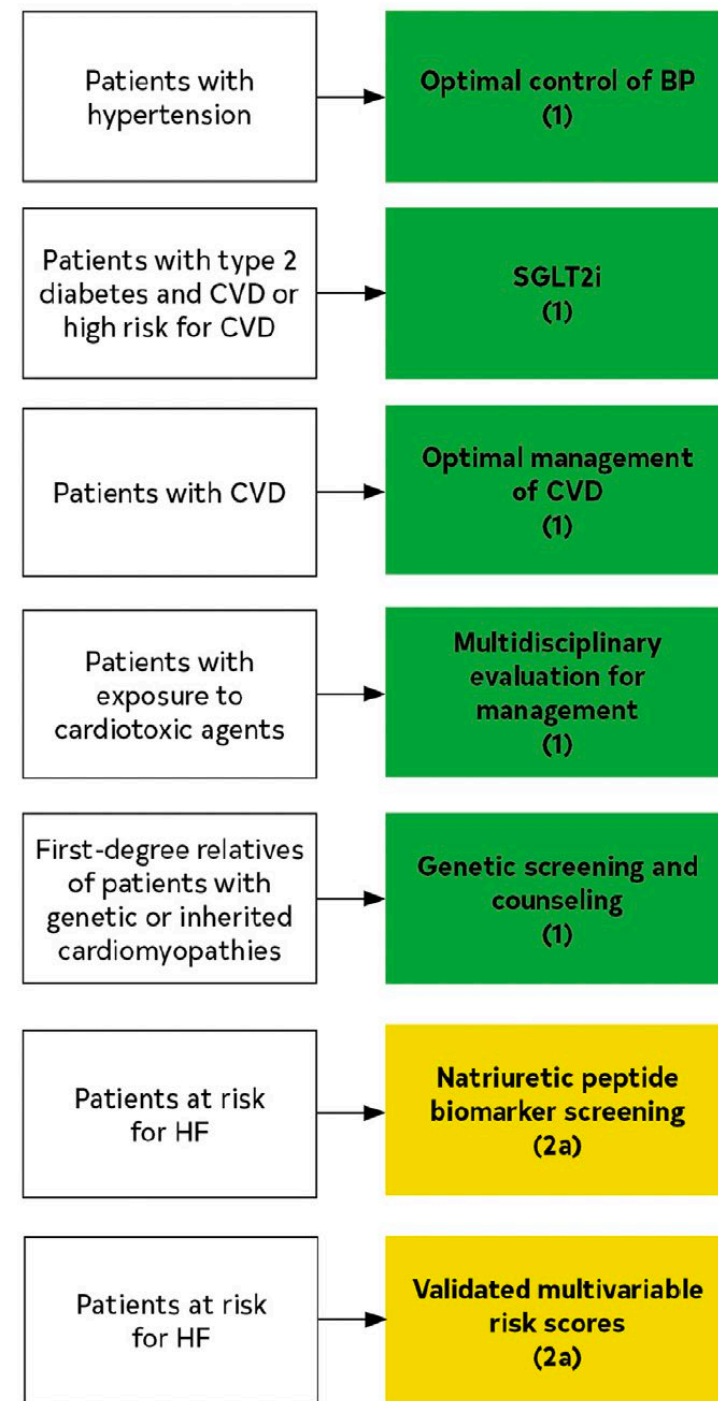
# MULTI-DISCIPLINARY PARTNERSHIP



Lala A, et al. *J Card Fail*. Published online August 13, 2025.



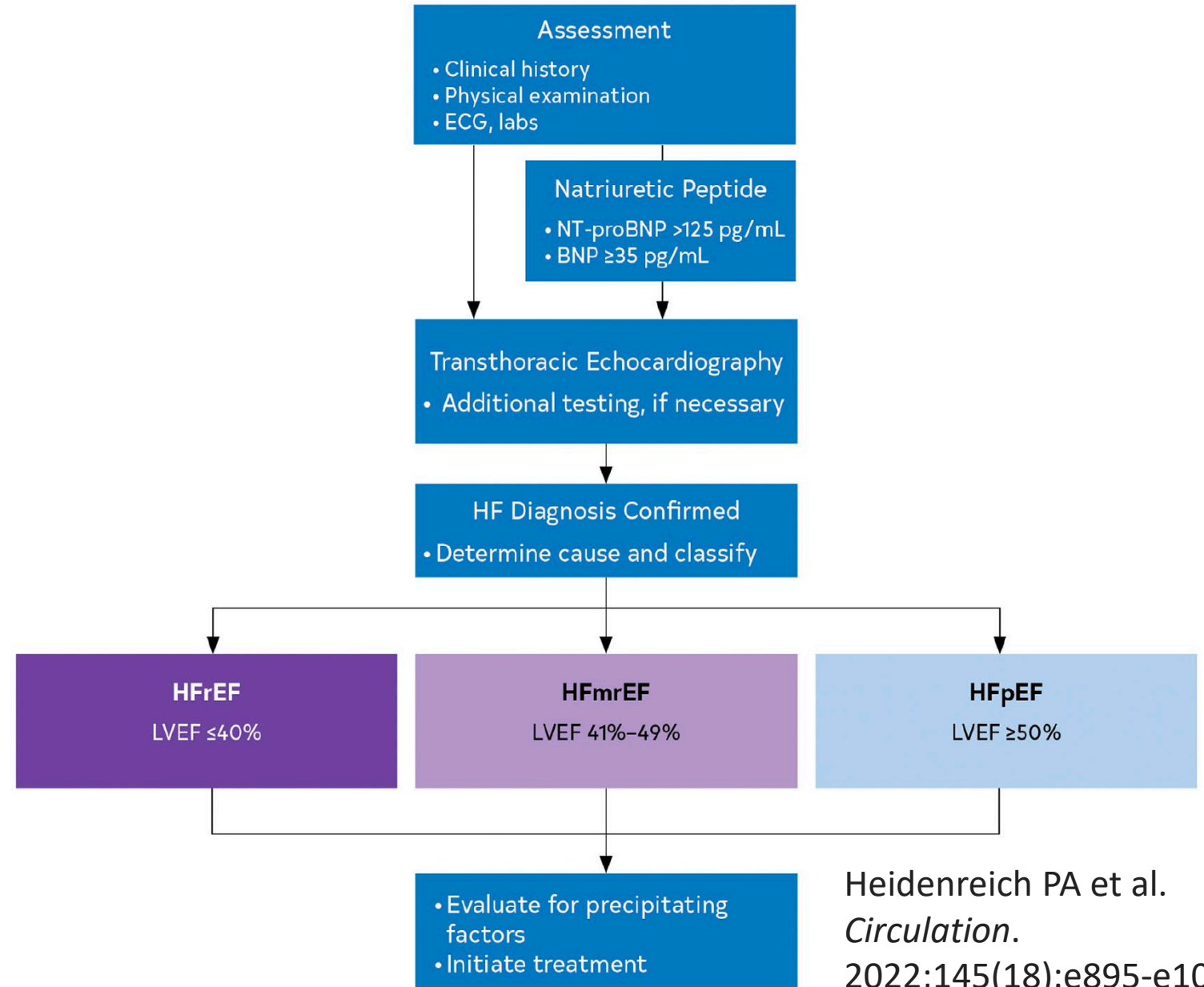
# STAGE A HEART FAILURE



Heidenreich PA et al.  
*Circulation*.  
2022;145(18):e895-  
e1032.

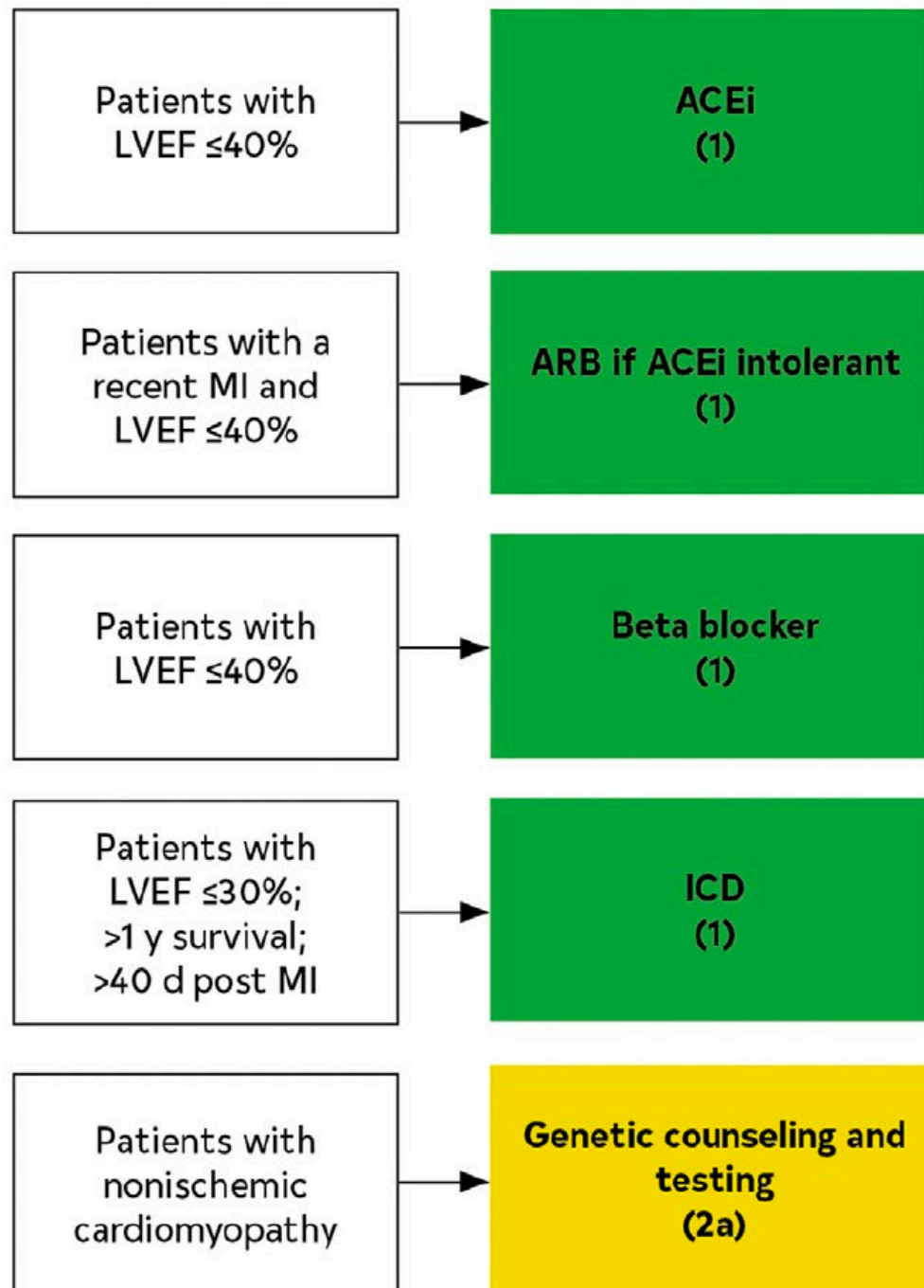


# DIAGNOSTIC ALGORITHM



Heidenreich PA et al.  
*Circulation*.  
2022;145(18):e895-e1032.

# STAGE B HEART FAILURE



Heidenreich PA et al.  
*Circulation*.  
2022;145(18):e895-e1032.

# Trajectory of Stage C Heart Failure

| New Onset/De Novo HF:  | Resolution of Symptoms:  | Persistent HF:  | Worsening HF:   |  |   |
|--|--|---|---|--|---|
| <ul style="list-style-type: none"><li>• Newly diagnosed HF</li><li>• No previous history of HF</li></ul> | <ul style="list-style-type: none"><li>• Resolution of symptoms/ signs of HF</li></ul> <table><tr><td>Stage C with previous symptoms of HF with persistent LV dysfunction</td><td>HF in remission with resolution of previous structural and/or functional heart disease*</td></tr></table> | Stage C with previous symptoms of HF with persistent LV dysfunction | HF in remission with resolution of previous structural and/or functional heart disease* | <ul style="list-style-type: none"><li>• Persistent HF with ongoing symptoms/signs and/or limited functional capacity</li></ul> | <ul style="list-style-type: none"><li>• Worsening symptoms/ signs/functional capacity</li></ul> |
| Stage C with previous symptoms of HF with persistent LV dysfunction                                      | HF in remission with resolution of previous structural and/or functional heart disease*  |   |   |  |   |

# HEART FAILURE CLASSIFICATION

**Table 4. Classification of HF by LVEF**

| Type of HF According to LVEF       | Criteria  |
|------------------------------------|---|
| HFrEF (HF with reduced EF)         | LVEF $\leq 40\%$  |
| HFimpEF (HF with improved EF)      | Previous LVEF $\leq 40\%$ and a follow-up measurement of LVEF $> 40\%$  |
| HFmrEF (HF with mildly reduced EF) | LVEF 41%–49%<br>Evidence of spontaneous or provokable increased LV filling pressures (eg, elevated natriuretic peptide, noninvasive and invasive hemodynamic measurement)     |
| HFpEF (HF with preserved EF)       | LVEF $\geq 50\%$<br>Evidence of spontaneous or provokable increased LV filling pressures (eg, elevated natriuretic peptide, noninvasive and invasive hemodynamic measurement) |

Heidenreich PA et al. *Circulation*. 2022;145(18):e895-e1032.



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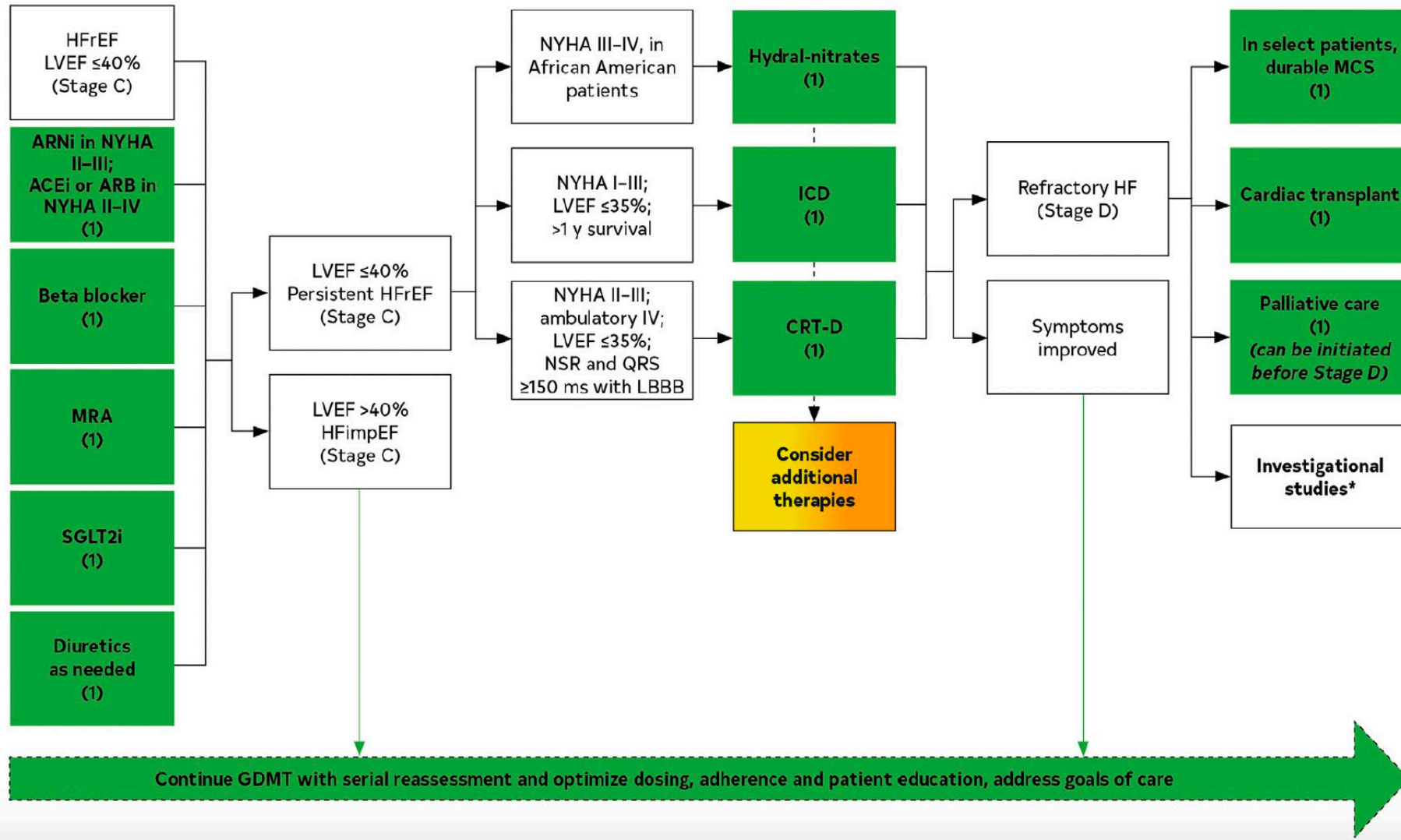
## Initiate and up-titrate GDMT



$$\text{NNT} = 4$$

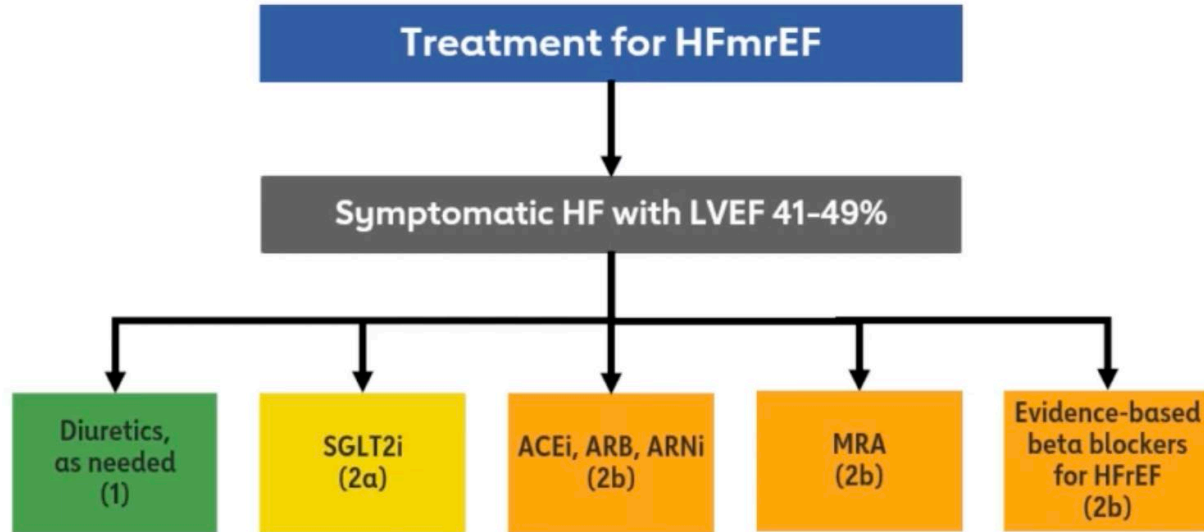


Heidenreich PA et al. *Circulation*. 2022;145(18):e895-e1032.





# Stage C Mildly Reduced Ejection Fraction

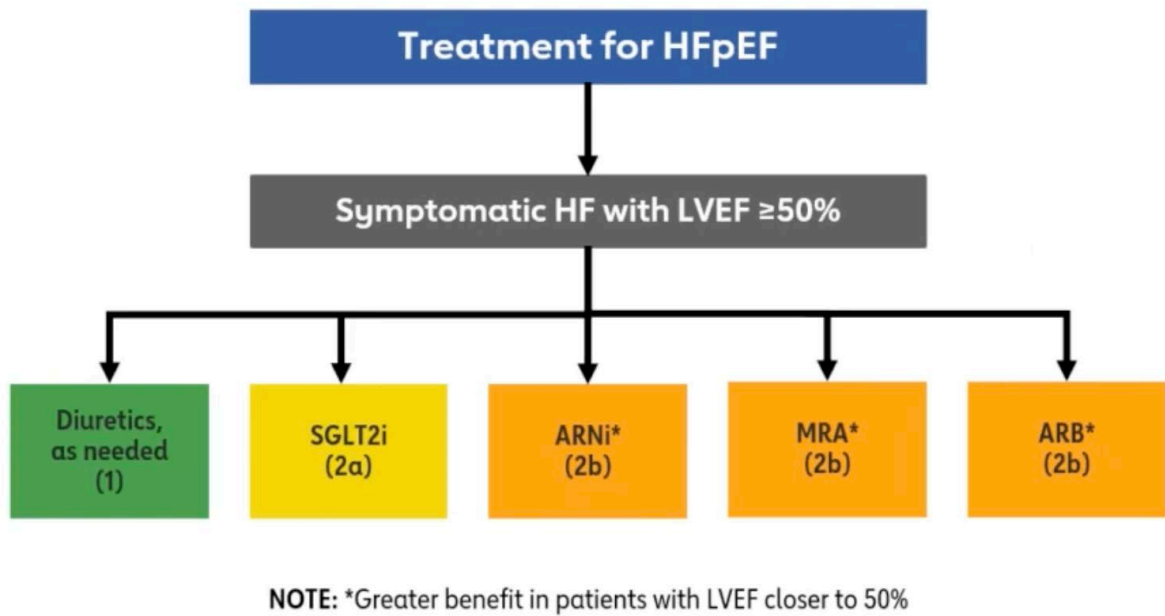


## Patients With HFimpEF

| COR | RECOMMENDATIONS  |
|-----|--|
| 1   | 1. In patients with HFimpEF after treatment, GDMT should be continued to prevent relapse of HF and LV dysfunction, even in patients who may become asymptomatic. (1) |

Heidenreich PA et al. *Circulation*. 2022;145(18):e895-e1032.

# Stage C Heart Failure with Preserved EF



Heidenreich PA et al. *Circulation*. 2022;145(18):e895-e1032.

|               |      |   |
|---------------|------|---|
| 1             | C-LD | 1. Patients with HFpEF and hypertension should have medication titrated to attain blood pressure targets in accordance with published clinical practice guidelines to prevent morbidity. <sup>1-3</sup> |
| 2a            | B-R  | 2. In patients with HFpEF, SGLT2i can be beneficial in decreasing HF hospitalizations and cardiovascular mortality. <sup>4</sup>  |
| 2a            | C-EO | 3. In patients with HFpEF, management of AF can be useful to improve symptoms.  |
| 2b            | B-R  | 4. In selected patients with HFpEF, MRAs may be considered to decrease hospitalizations, particularly among patients with LVEF on the lower end of this spectrum. <sup>5-7</sup>                        |
| 2b            | B-R  | 5. In selected patients with HFpEF, the use of ARB may be considered to decrease hospitalizations, particularly among patients with LVEF on the lower end of this spectrum. <sup>8,9</sup>              |
| 2b            | B-R  | 6. In selected patients with HFpEF, ARNi may be considered to decrease hospitalizations, particularly among patients with LVEF on the lower end of this spectrum. <sup>10,11</sup>                      |
| 3: No-Benefit | B-R  | 7. In patients with HFpEF, routine use of nitrates or phosphodiesterase-5 inhibitors to increase activity or QOL is ineffective. <sup>12,13</sup>   |

## TREATMENT OF HFrEF

LVEF ≤40% (STAGE C)

ARNi in NYHA II-III; ACEi or ARB in NYHA II-IV  
(1)

Beta blocker  
(1)

MRA  
(1)

**SGLT2i**  
(1)

Diuretics, as needed  
(1)

## TREATMENT OF HFmrEF

SYMPTOMATIC HF WITH LVEF 41%-49%

Diuretics, as needed  
(1)

**SGLT2i**  
(2a)

ACEi, ARB, ARNi  
(2b)

MRA  
(2b)

Evidence-based beta blockers for HFrEF  
(2b)

## TREATMENT OF HFpEF

SYMPTOMATIC HF WITH LVEF ≥50%

Diuretics, as needed  
(1)

**SGLT2i**  
(2a)

ARNi<sup>†</sup>  
(2b)

MRA<sup>†</sup>  
(2b)

ARB<sup>†</sup>  
(2b)

## CLASS (STRENGTH) OF RECOMMENDATION

**CLASS 1 (STRONG)**  
Benefit >>> Risk

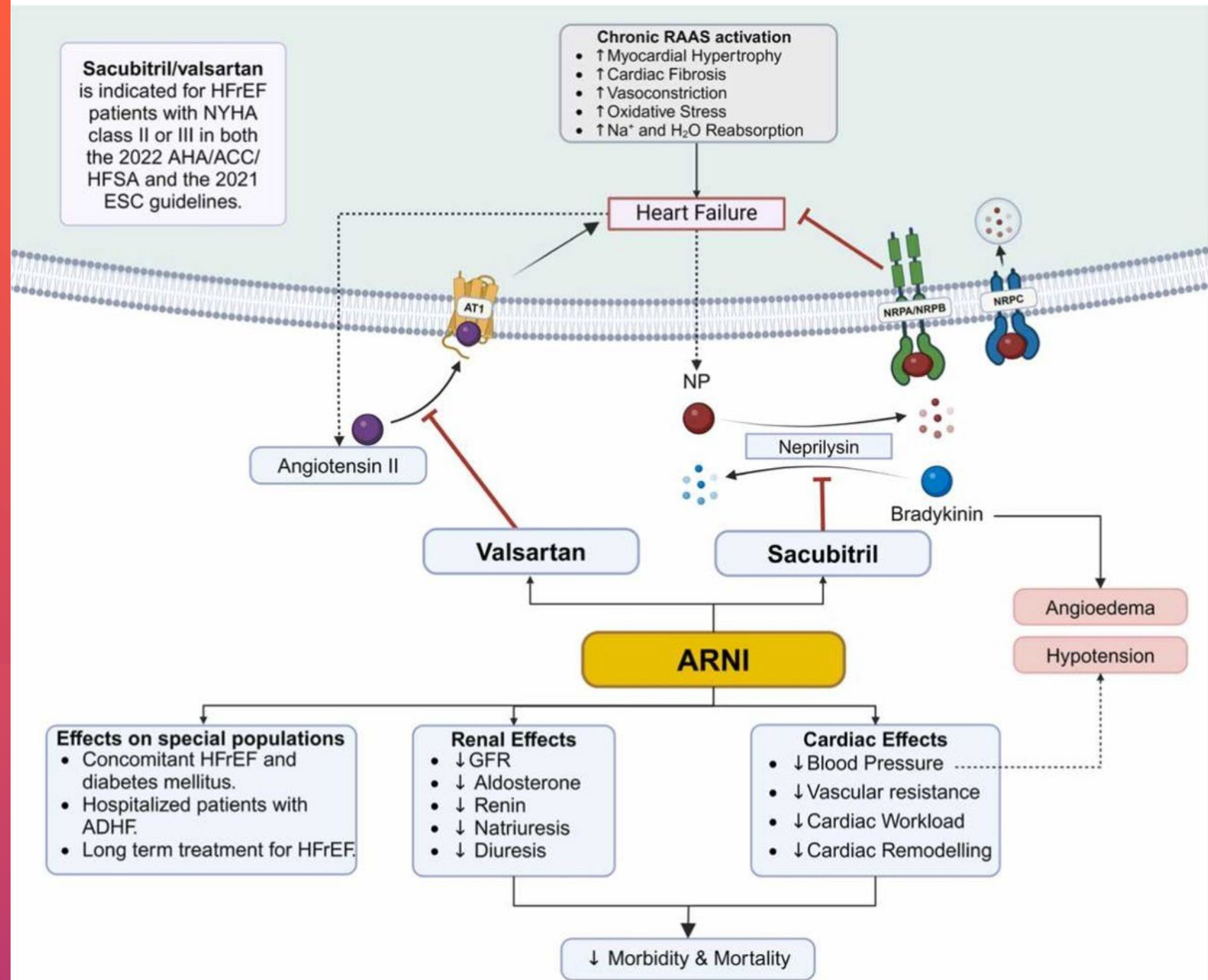
**CLASS 2a (MODERATE)**  
Benefit >> Risk

**CLASS 2b (WEAK)**  
Benefit ≥ Risk



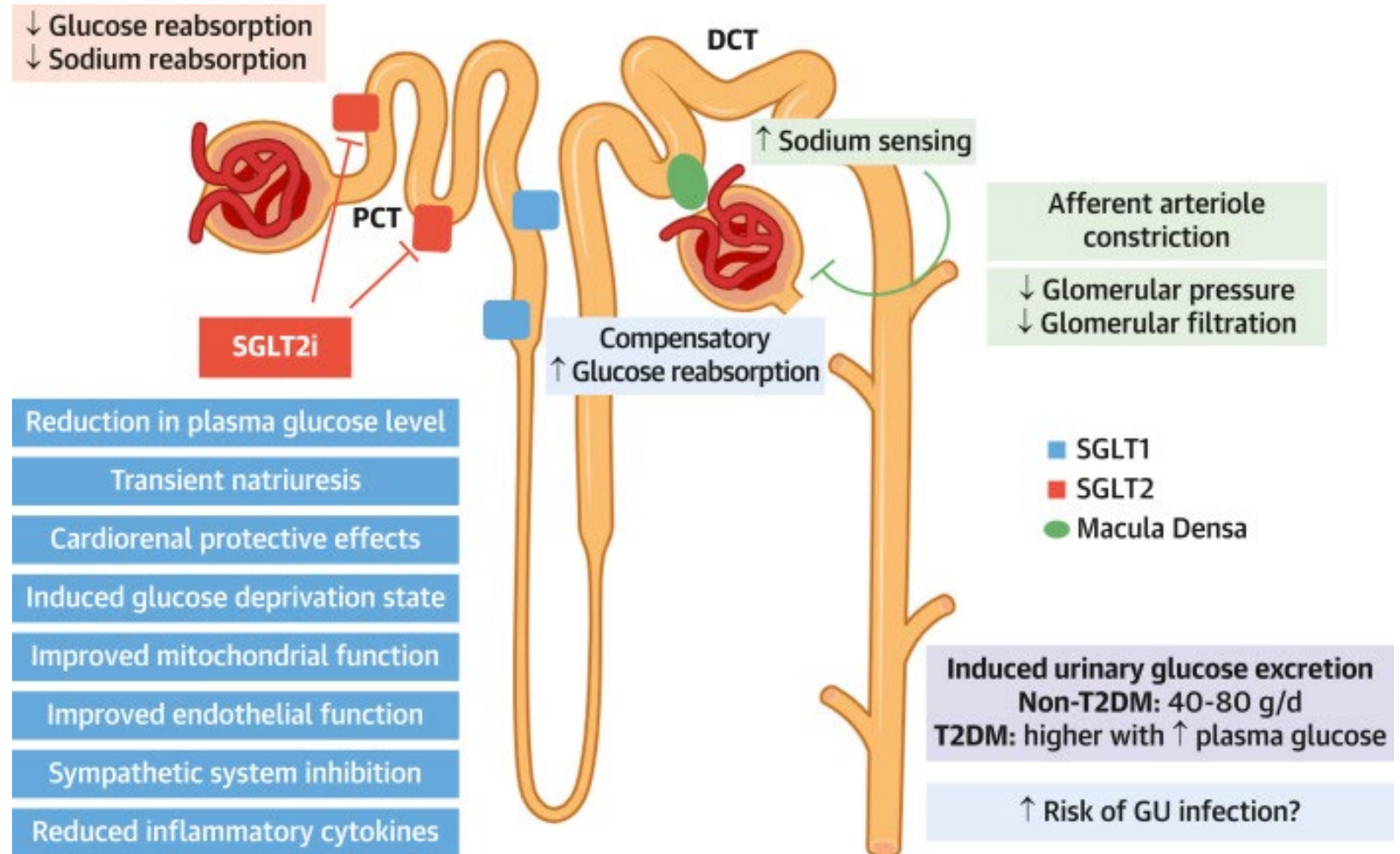
# SACUBITRIL- VALSARTAN

**Sacubitril/valsartan** is indicated for HFrEF patients with NYHA class II or III in both the 2022 AHA/ACC/HFSA and the 2021 ESC guidelines.



# SGLT2 inhibitors

- Diuresis
- Natriuresis
- Glucosuria
- Decreases plasma volume
- Reduces arterial stiffness
- Decreases blood pressure
- Renal protective





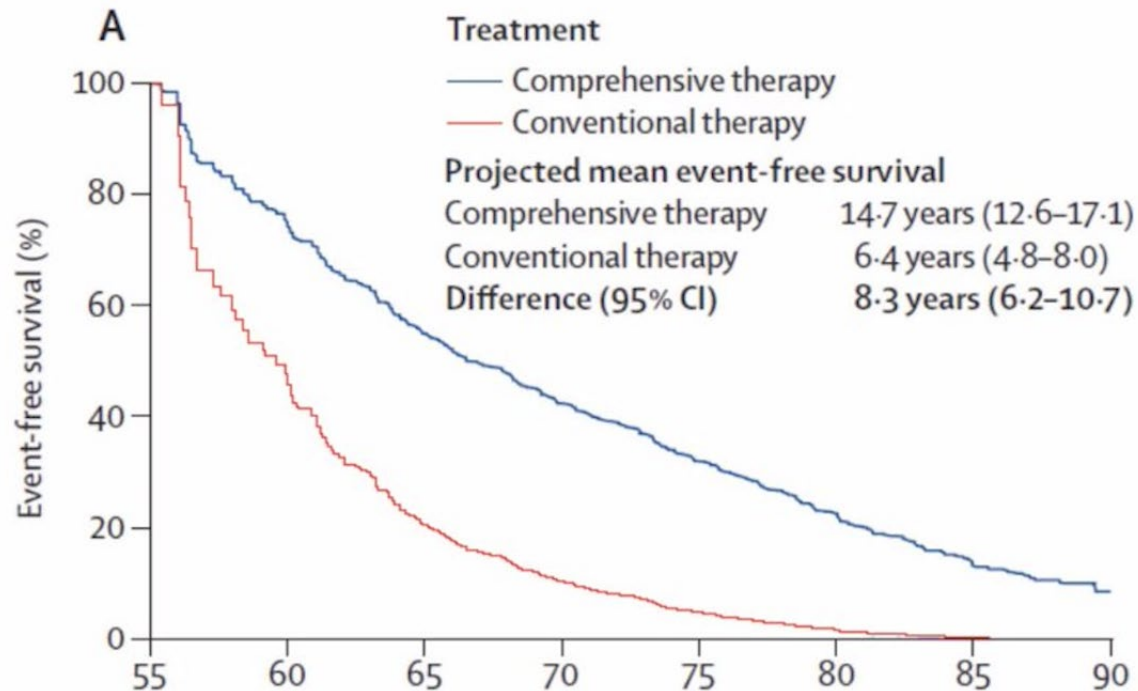
Mr. Johnson, a 54 year old male has non-ischemic cardiomyopathy, NYHA II symptoms, HFrEF (LVEF 30%) and sees you in clinic for the first time. BP 120/80 mm Hg, HR 97 bpm, Creatinine 1. Euvolemic. What would you do?



# Two vs Four Drugs ??

**Comprehensive Therapy (ARNI+BB+MRA+SGLT2i) vs. Conventional Therapy (ACEi/ARB + BB) in a 55-year-old patient with HFrEF**

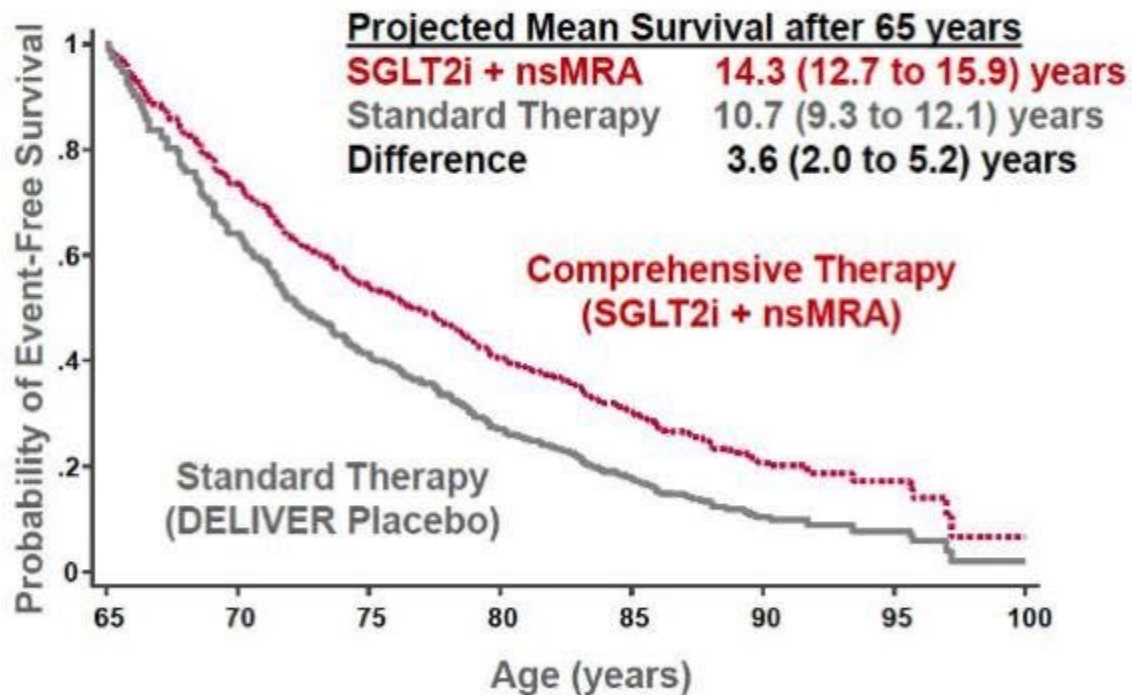
**8.3 additional years of event-free survival**



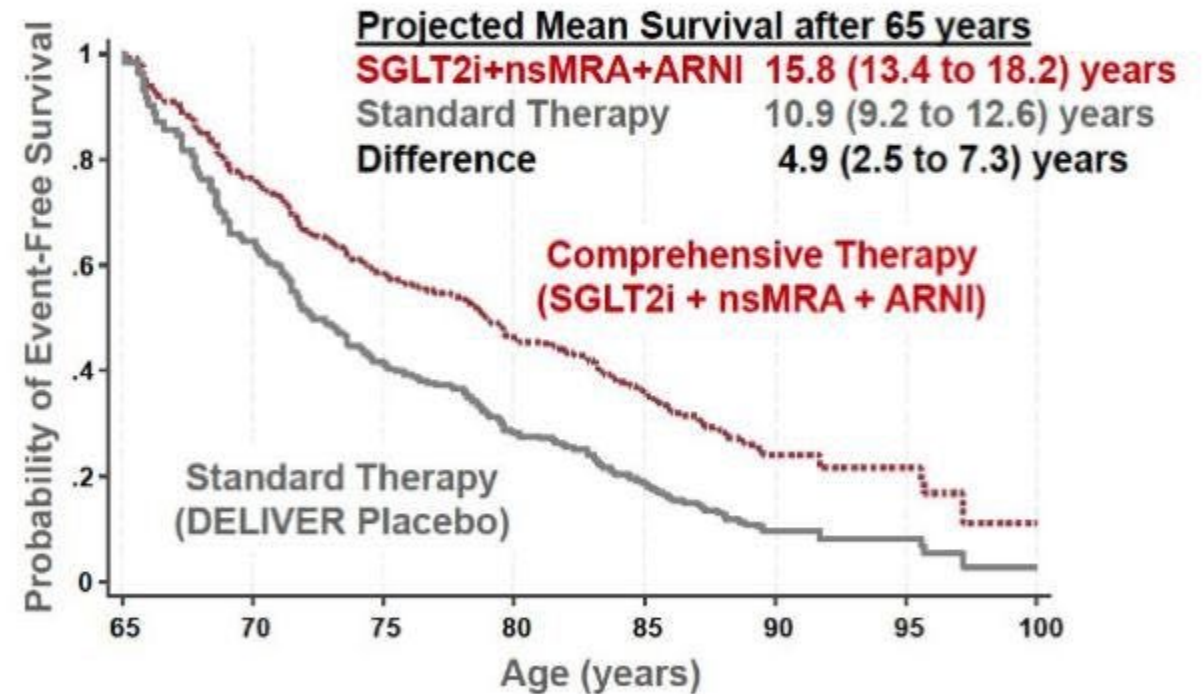
Use of comprehensive disease modifying medical therapy can further reduce cardiovascular mortality in HFrEF by 50%!!!!

# Lifetime Benefits of Comprehensive Medical Therapy in Heart Failure with Mildly Reduced or Preserved Ejection Fraction

Overall Population with HFmrEF/HFpEF



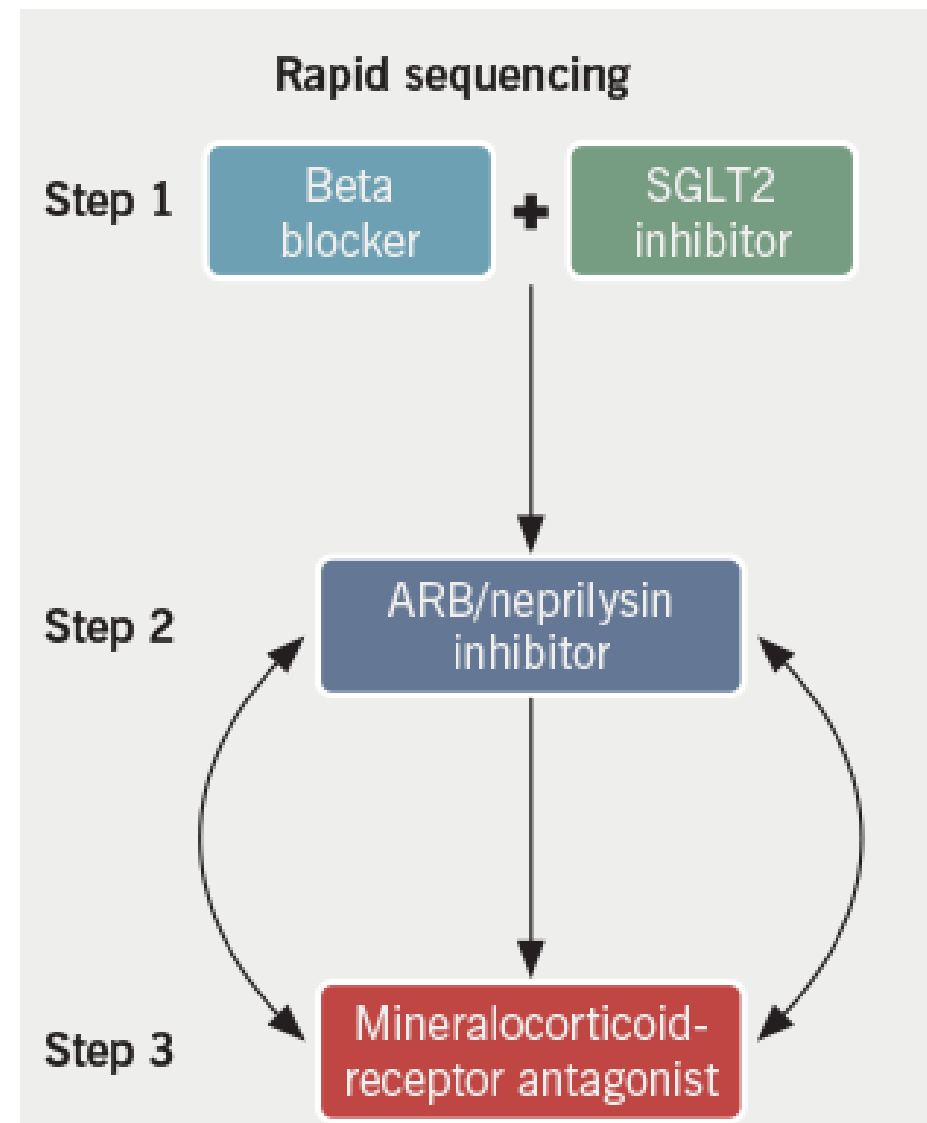
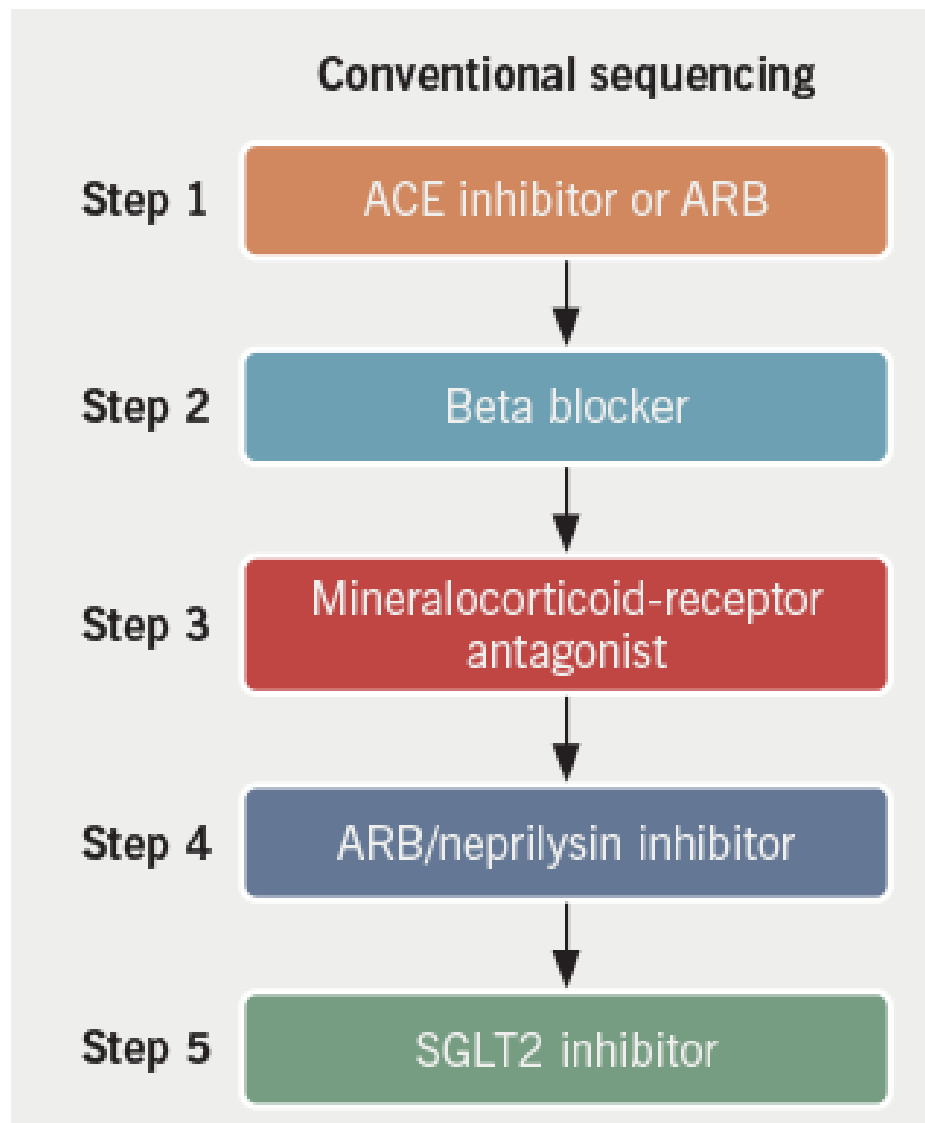
Patients with HFmrEF/HFpEF and LVEF Below Normal (<60%)



Vaduganathan et al. Nature Medicine (2025)



# How would you start GDMT?



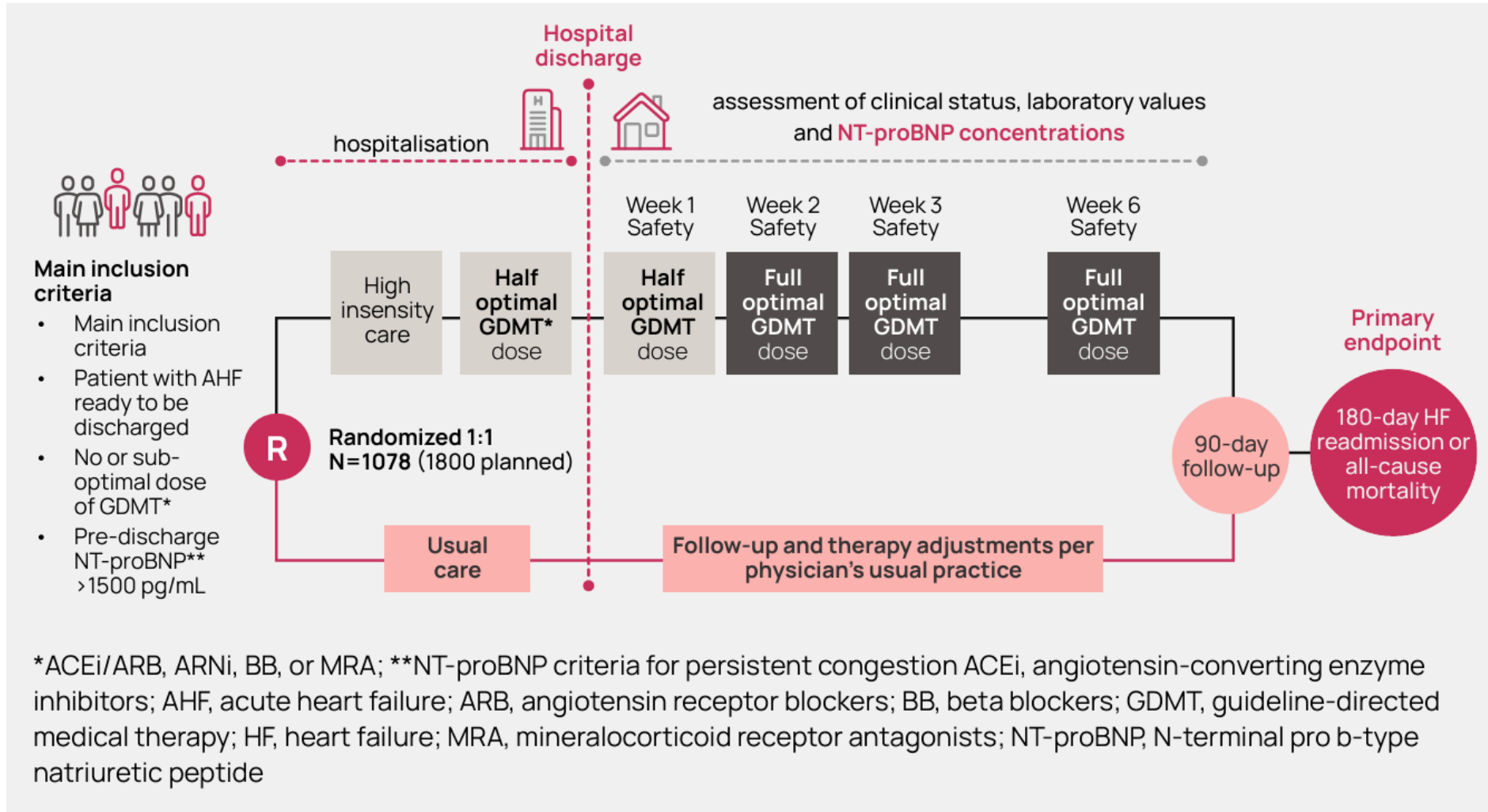


Rapid  
Sequencing

VS

Simultaneous  
Initiation

# STRONG-HF TRIAL



# STRONG-HF TRIAL

## Results

The high intensity care group: **34% relative** and **8.1% absolute risk reduction (ARR)** in the combination of death or heart failure readmission.<sup>14</sup>



CV (cardiovascular) death

**26% lower**

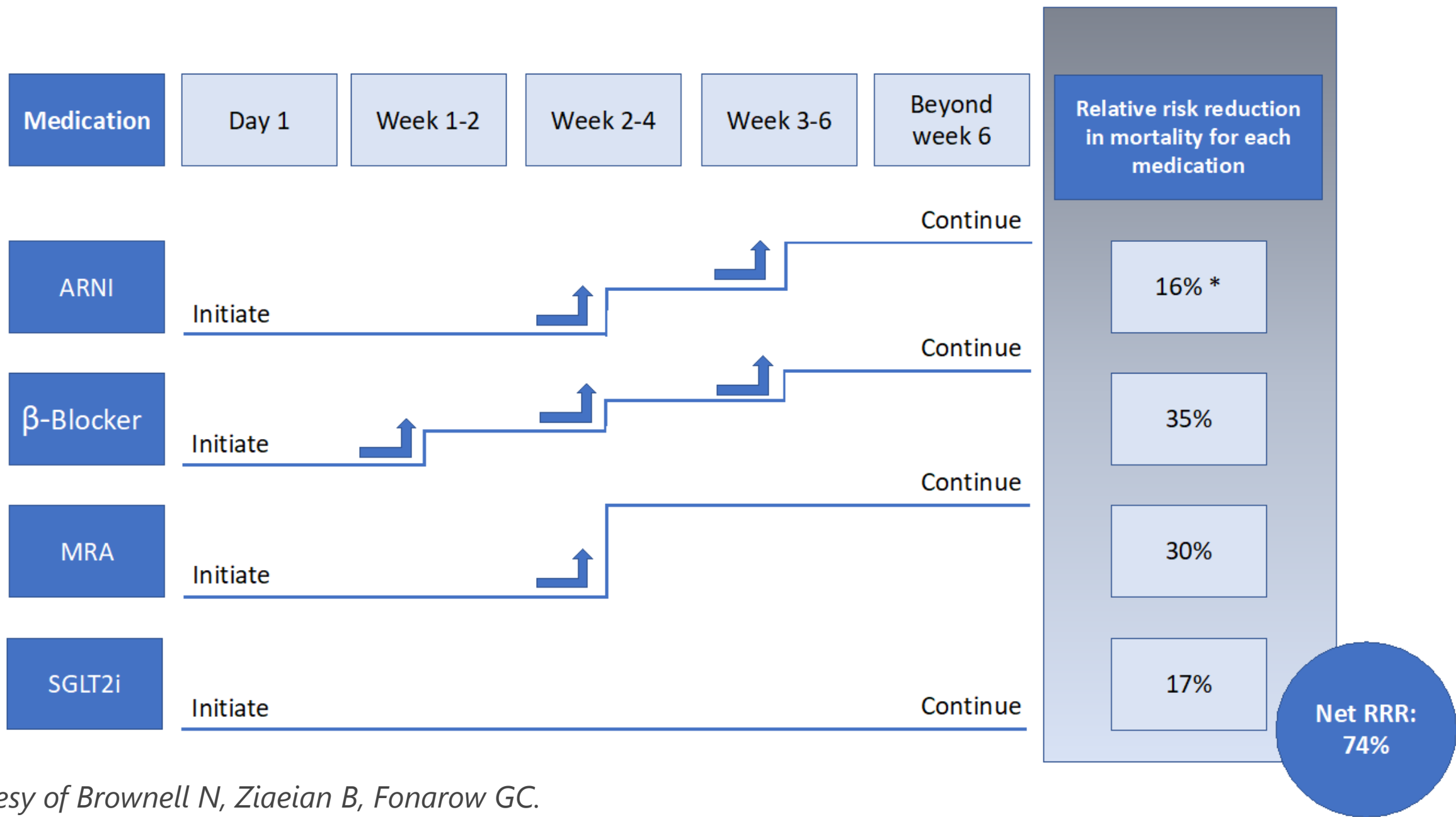
HF readmission

**44% lower**

All-cause death

**16% lower**

**STRONG-HF study results demonstrated clear benefits for acute heart failure patients by adapting the strategy of care.**



Courtesy of Brownell N, Ziaieian B, Fonarow GC.



# 3

## Simultaneous/Rapid Sequence Initiation of ALL 4 Medications – within 3 months or sooner



Frequent follow-up visits for titration and lab monitoring



For hospitalized patients, all 4 pillars should be initiated before discharge.

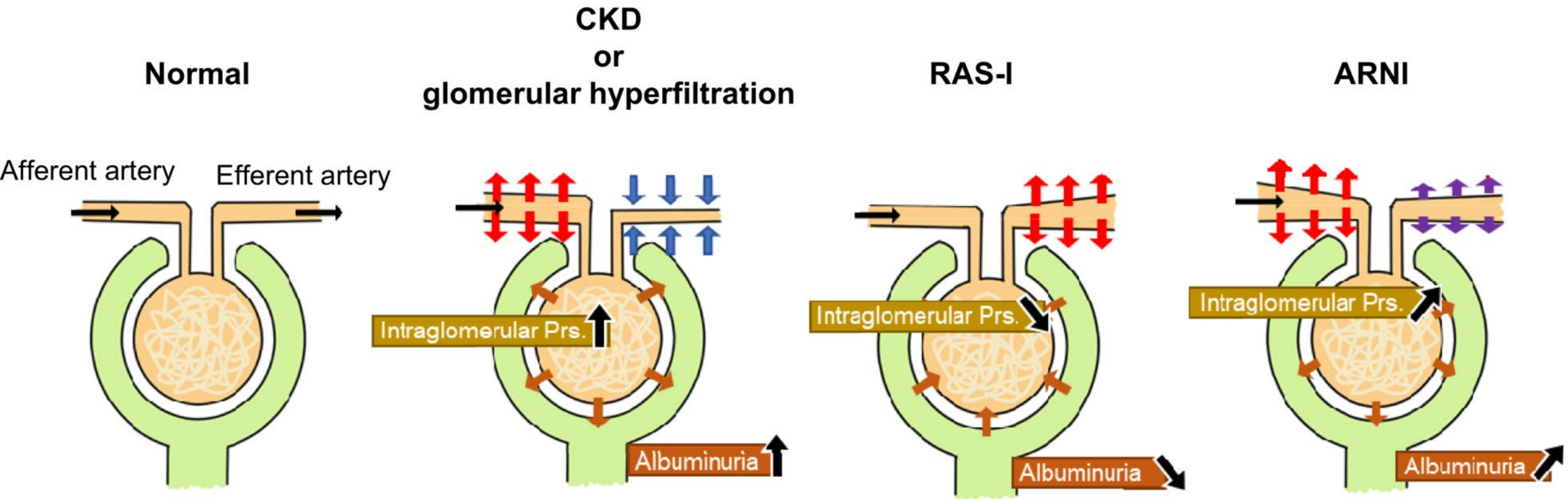


SGLT2 inhibitors can be started earlier or concomitantly with other agents due to their safety profile



Mr. Johnson got started on sacubitril-valsartan 24/26 mg BID. Creatinine was 1 at the time of initiation of the drug. In 2 weeks, his BP is 110/67 mm Hg, HR is 90 BPM. BMP done in 2 weeks showed a creatinine of 1.3. What would you do?

# RAS-I and ARNI and Kidney function



Beldhuis IE, et al. *Circulation*. 2022;145(9):693-712.

# ARNI

## Effect on Renal Function

The effect of ARNI on renal function is not entirely clear, but is attributed to higher circulating natriuretic peptide levels, the improved clinical status, an effect on renal podocyte function and the need for less loop diuretics.

|   |   |   |
|---|---|---|
| Early decline in eGFR after initiation (0.5-1.0 mL/min/1.73m <sup>2</sup> ) <sup>80</sup> | Long term slope in eGFR less with ARNI vs ACEi: -1.61 vs. -2.04 mL/min/1.73m <sup>2</sup> /year <sup>80</sup> | Change in serum creatinine/eGFR similar between ARNI/ACEi in PARADIGM-HF and PIONEER <sup>80,81</sup> |
|---|---|---|

## Management of substantial increase in serum creatinine/drop in eGFR during initiation/uptitration

In the context of uptitration of ARNI some increase in serum creatinine / drop in eGFR is expected and acceptable. The survival benefit seen with this class of drugs far outweigh the risks associated with this perceived worsening of renal function (WRF)

| $\Delta$ serum creatinine (%) | Max serum creatinine (mg/dL) | Min eGFR mL/min/1.73m <sup>2</sup> | Max serum potassium (mmol/L) | Action advised   |
|-------------------------------|------------------------------|------------------------------------|------------------------------|--|
| < 50                          | 2.5 mg/dL                    | 30                                 | 5.0                          | None, uptitrate and evaluate renal function and electrolytes                             |
| 50-100                        | 3.5 mg/dL                    | 20                                 | 5.5                          | Evaluate clinical status and other causes of WRF. Consider halving ARNI and re-evaluate  |
| > 100                         | > 3.5 mg/dL                  | < 20                               | > 5.5                        | Evaluate clinical status and other causes of WRF. Consider stopping ARNI and re-evaluate |

Rechallenge after 2-4 weeks (if possible at lower dose) when dosing reduced or stopped all together if renal function has improved

Beldhuis IE, et al. *Circulation*. 2022;145(9):693-712.



# SGLT2 inhibitors

## Effect on Renal Function

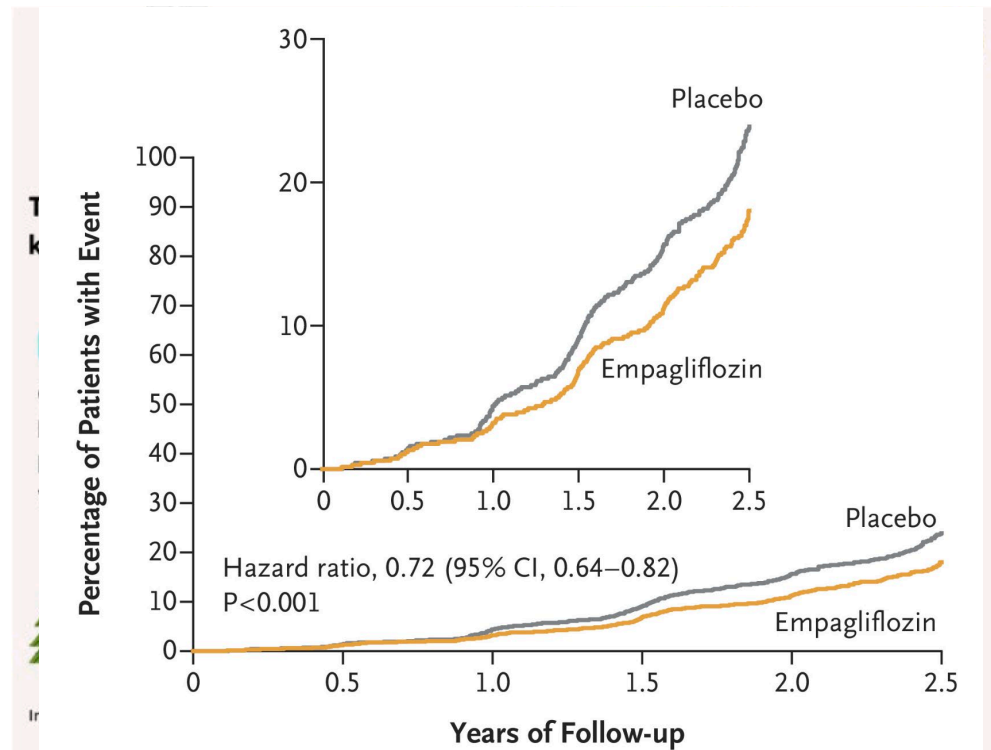
It is hypothesized that SGLT2i cause afferent arteriolar vasoconstriction (and possibly some efferent vasodilation) due to activated tubuloglomerular feedback caused by more distal sodium delivery to macula densa.

|   |  |   |
|---|--|---|
| Early decline in eGFR after initiation<br>(0.3-4.0 mL/min/1.73m <sup>2</sup> ) <sup>85,86</sup> | Long term slope in eGFR less with SGLT2i vs Placebo: -0.6 to 1.09 vs. -2.3 to 2.9 mL/min/1.73m <sup>2</sup> /year <sup>85,86</sup> | Drop in eGFR with SGLT2i no reason to discontinue |
|---|--|---|

## Management of substantial increase in serum creatinine/drop in eGFR during initiation/uptitration

In the context of initiation of SGLT2i some increase in serum creatinine / drop in eGFR is expected and acceptable.

| Δ serum creatinine (%) | Max serum creatinine (mg/dL) | Min eGFR mL/min/1.73m <sup>2</sup> | Action advised   |
|------------------------|------------------------------|------------------------------------|--|
| < 50                   | 2.5 mg/dL                    | 30                                 | None, continue SGLT2i and reevaluate renal function regularly  |
| 50-100                 | 3.5 mg/dL                    | 20                                 | Continue SGLT2i if eGFR/or serum creatinine are acceptable. Evaluate other causes in parallel. SGLT2i do not cause hyperkalemia. Evaluate potassium if creatinine rises steeply  |
| > 100                  | > 3.5 mg/dL                  | < 20                               | Such large increases in serum creatinine are unexpected with SGLT2i and should prompt further evaluation. SGLT2i do not cause hyperkalemia. Evaluate potassium if creatinine rises steeply.<br><br>If deemed clinically appropriate, continue SGLT2i with close monitoring; if no other option, stop SGLT2i.<br><br>Rechallenge after 2-4 weeks (if possible at lower dose) when dosing reduced or stopped all together if renal function has improved |



Packer M, et al. *N Engl J Med*. 2020;383(15):1413-1424.

Beldhuis IE, et al. *Circulation*. 2022;145(9):693-712.

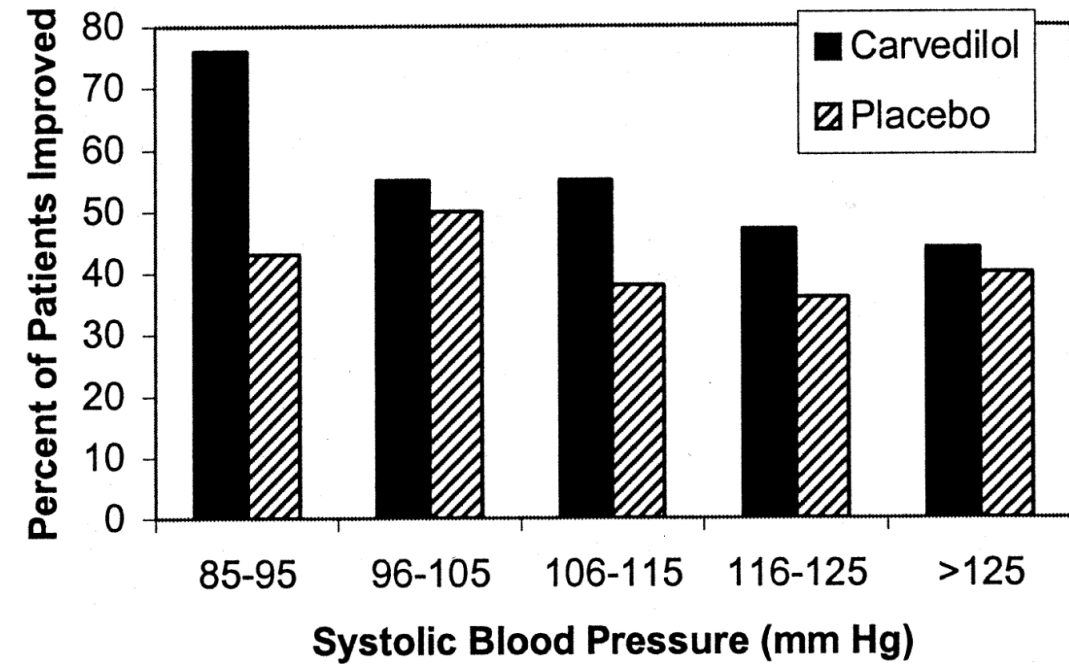
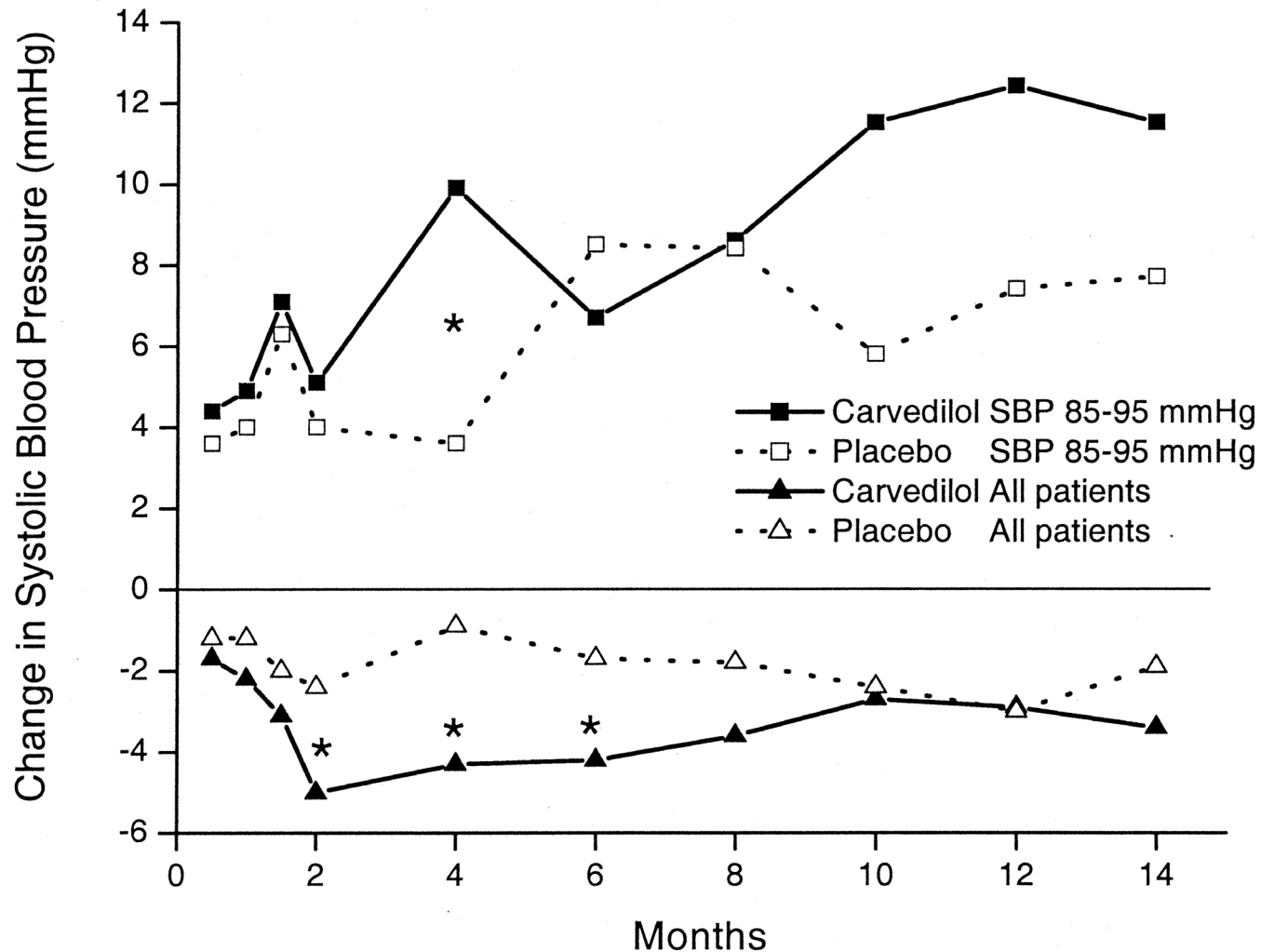


Mr. Johnson has now tolerated sacubitril-valsartan 49/51 mg BID, spironolactone 25 mg daily, metoprolol succinate 50 mg daily and on the 5-week BMP check creatinine is 1.1, potassium 4. His blood pressure is 95/62 mm Hg, HR 70 bpm. JVP 7 cm H<sub>2</sub>O. He denies any new symptoms. What is your next step?

# How to handle blood pressure before and after ACEI/ARB/ARNI/BB initiation?

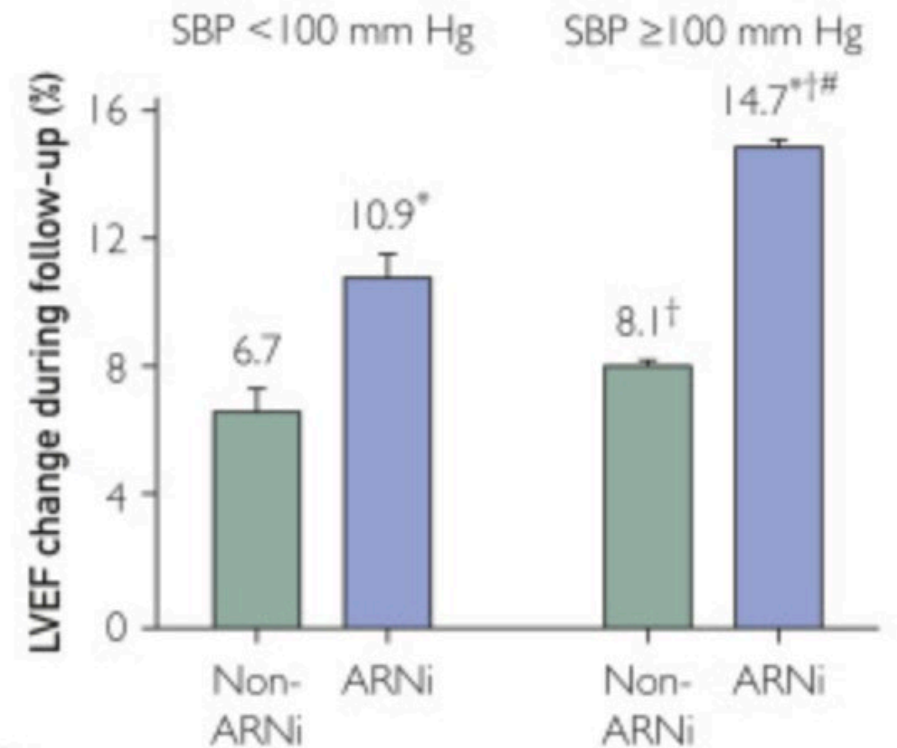
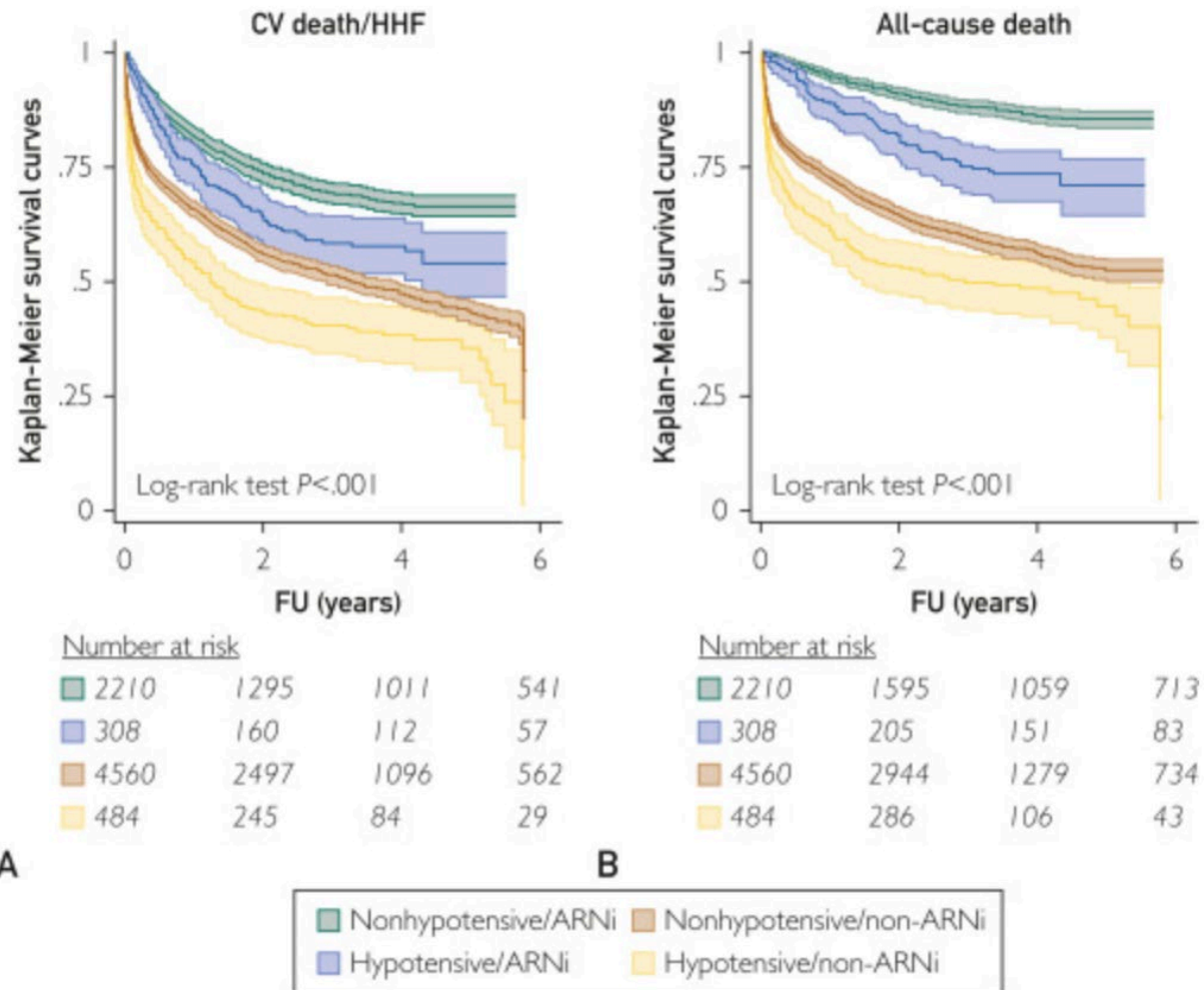
- Before initiation, BP contraindication:
  - Symptomatic hypotension (<100-110 SBP) (consistently dizzy, weak, oliguric)
  - Asymptomatic hypotension (<90 SBP). For ARNI, asymptomatic hypotension <100 SBP (PARADIGM, LIFE trials)
- After initiation
  - Asymptomatic hypotension <90 SBP is accepted. DO NOT stop the medication.
  - Mild occasional dizziness or weakness after initiation – stop nitrates, reduce diuretics especially if patients are not congested
  - Severe symptoms – can consider decreasing dose or stopping if patient does not tolerate the medication (!!! RED FLAG SIGN !!!)

# CARVEDILOL - COPENICUS trial





# ARNI

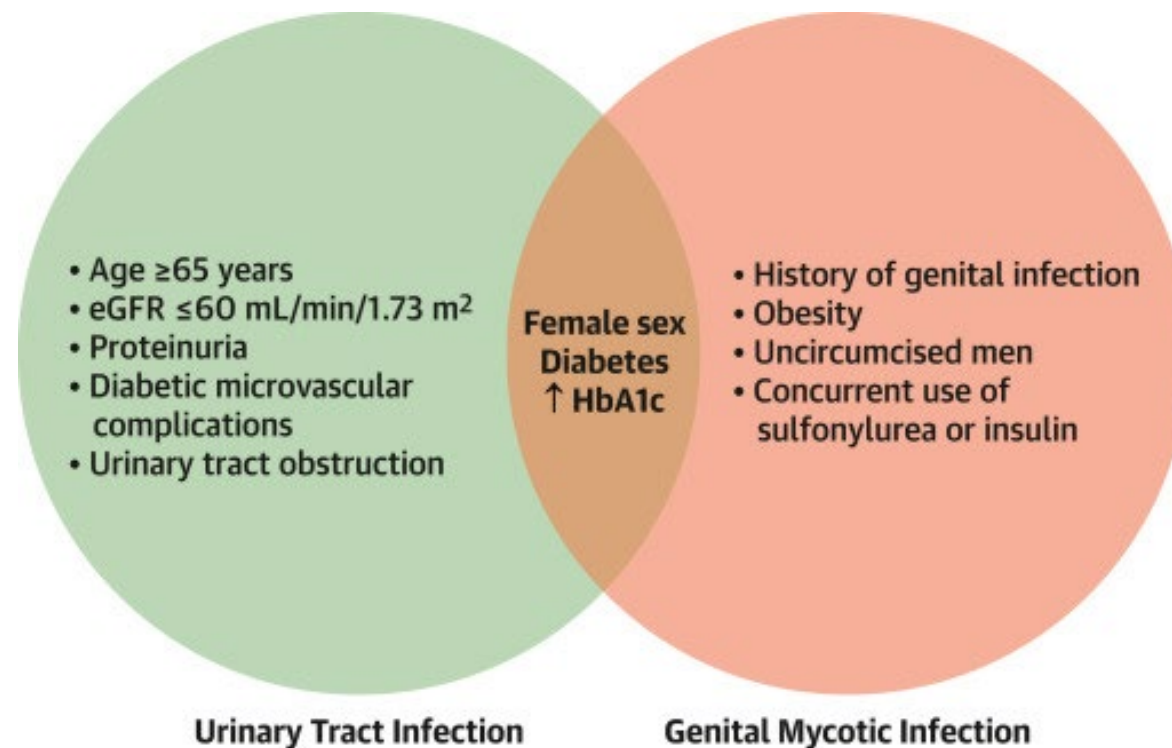
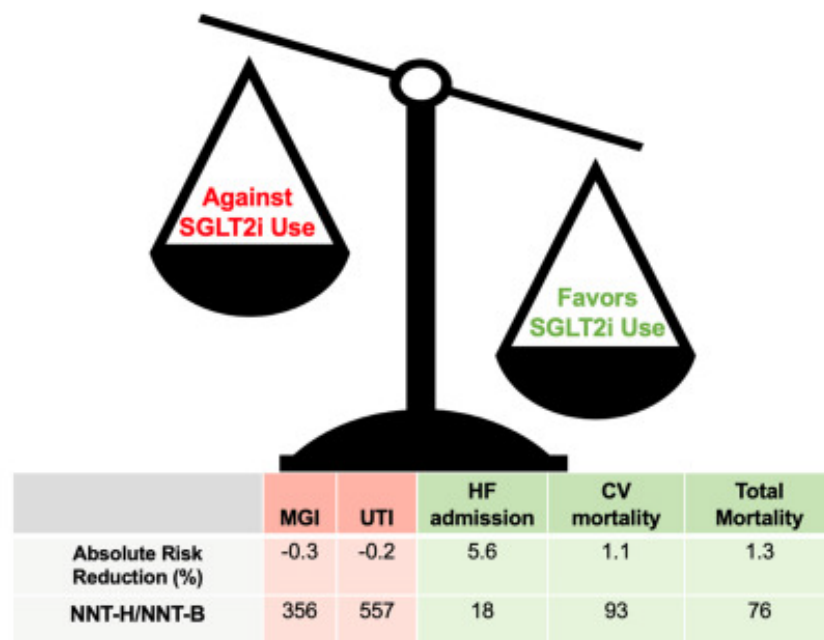




Mr. Johnson was started on dapagliflozin 10 mg daily. 3 weeks later he calls you saying he is having urinary urgency and itching in his genital area. What would you do next?

# SGLT2 inhibitors and UTI

Benefit:Risk Assessment of SGLT2i Use from MGIs and UTIs in Individuals with HF



Risk Factors for GU infections

# SGLT2I AND UTI

## Assessing Risk of MGI and UTI Before Starting SGLT2 Inhibitors

| Initiate:†                         | Do Not Initiate:                                  |
|------------------------------------|---|
| Demographics                       | Current UTI/MGI infection                         |
| - Female*                          | ADPKD   |
| - Uncircumcised male*              | Recurrent MGI or UTI (special attention to MDRO)Δ |
| - Obesity                          | ΔRefer to specialist for guidance                 |
| - Older Individuals                |   |
| - Non-white                        |   |
| Increased Estrogen Levels          |   |
| - Postmenopausal Estrogen Therapy  |   |
| Medications                        |   |
| - Antimicrobials                   |   |
| - Corticosteroids                  |   |
| Immunosuppression                  |   |
| - HIV                              |   |
| - Chemotherapy/Steroids            |   |
| Urologic Procedures                |   |
| - Orthotopic Bladder Substitution* |   |
| - Urinary Outlet Obstruction*      |   |
| - Artificial Urinary Sphincter     |   |
| - Implantable Penile Prosthesis    |   |
| Type 2 Diabetes Mellitus*          |   |
| History of Fournier's Gangrene     |   |

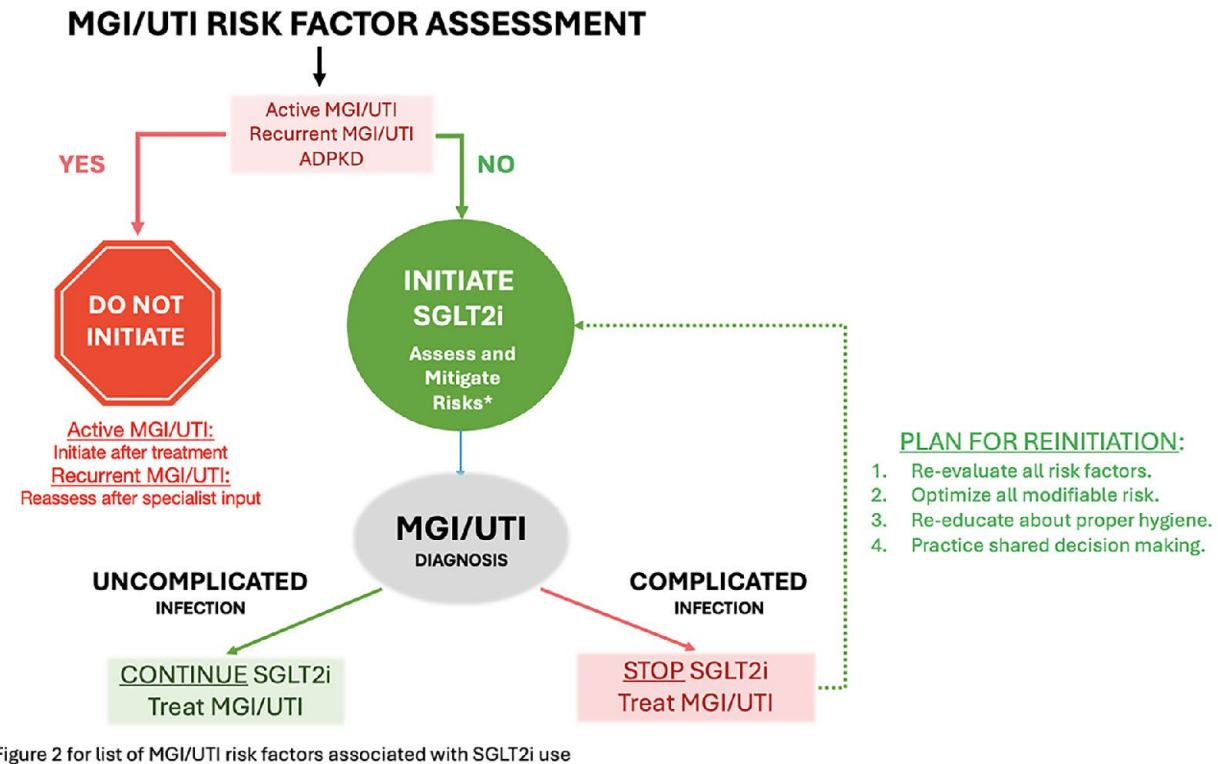
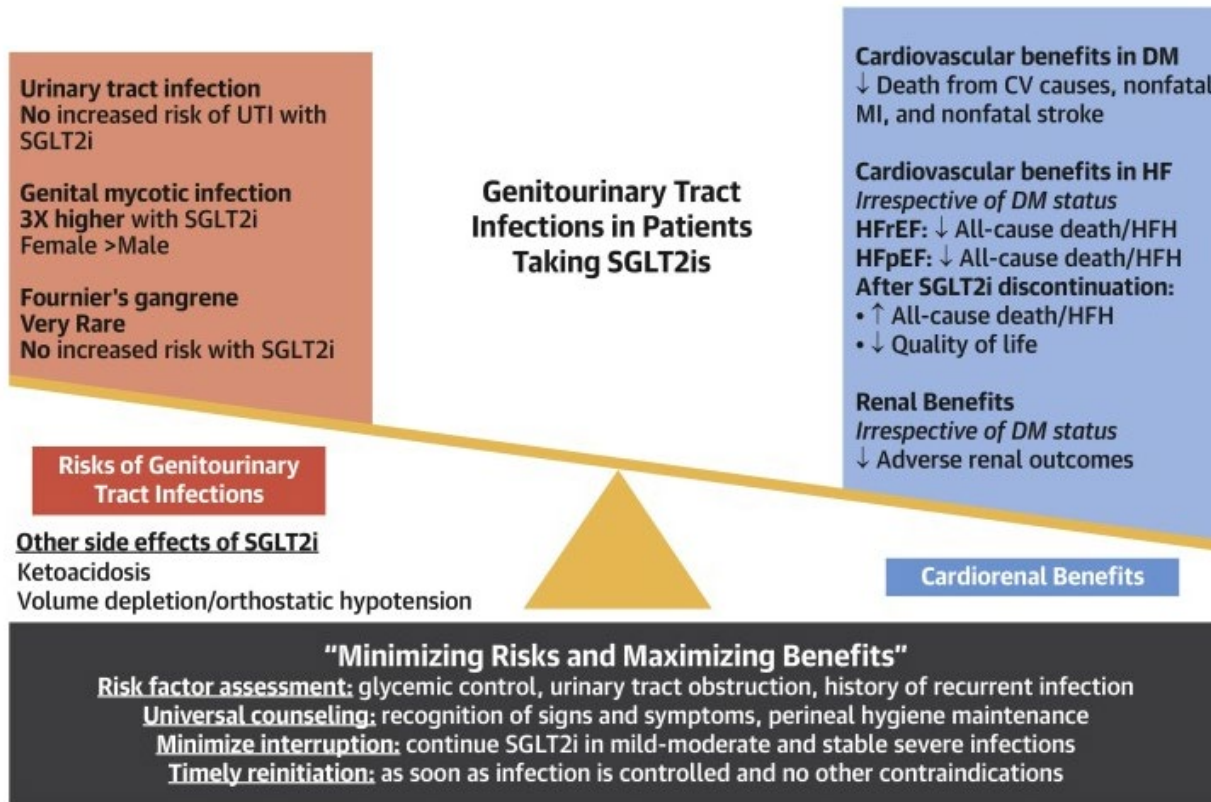
\*Risk Factors for UTI/MGI from SGLT2 inhibitor Use

†Practice Shared Decision Making

Duvalyan A, et al. *J Card Fail.* 2024;30(8):1031-1040.



# SGLT2 inhibitors and UTI



# SGLT2 INHIBITORS AND UTI

## Key Question

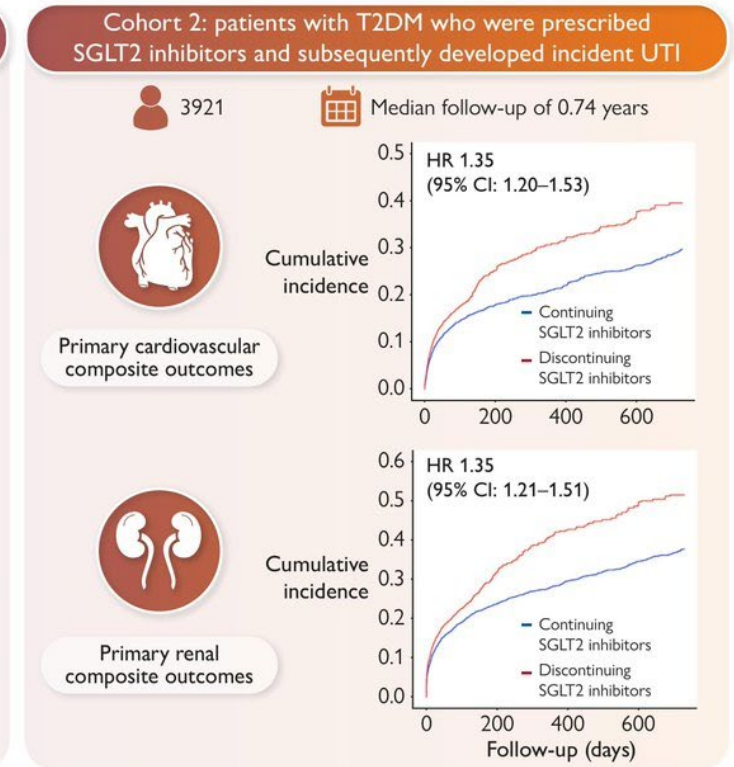
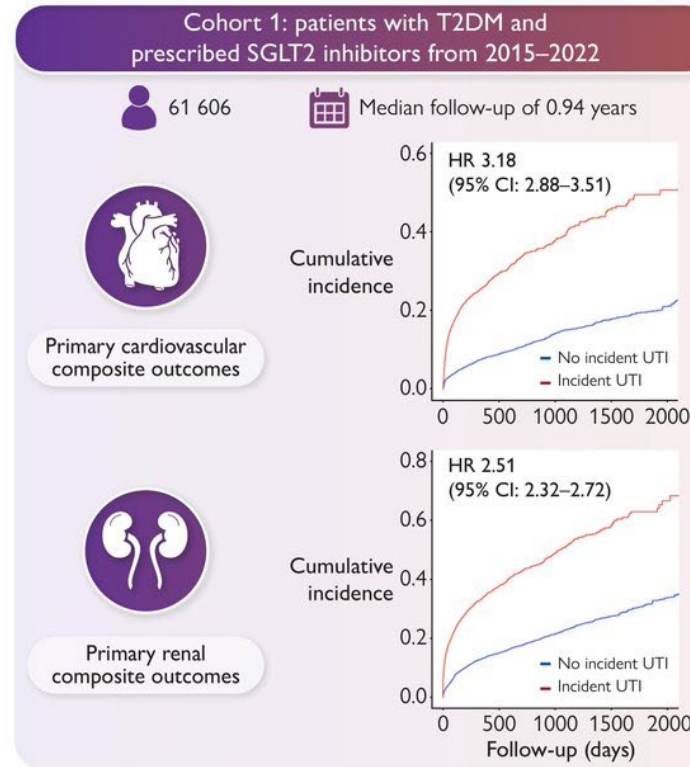
What is the prognosis of new-onset UTI and discontinuing SGLT2 inhibitors after UTI in patients with T2DM prescribed SGLT2 inhibitors?

## Key Finding

Incident UTI was associated with a significantly higher risk of cardiovascular and renal events than no incident UTI. Discontinuing SGLT2 inhibitors following UTI was associated with a higher risk of cardiovascular and renal events, compared to continued therapy.

## Take Home Message

Although new-onset UTI may serve as a marker of subsequent adverse events, continued use of SGLT2 inhibitors after UTI may effectively mitigate the complications.



SGLT2, sodium-glucose cotransporter-2; T2DM, type 2 diabetes mellitus; UTI, urinary tract infection

Wu MZ, et al. *European Heart Journal*.

Mei-Zhen Wu et al, *European Heart Journal*, 2025



# INPATIENT GDMT INITIATION GUIDE

COURTESY OF  
MAYA  
IGNASZEWSKI, MD  
AND JESSI CLARK,  
PHARMD, BCCP



1

**Establish** diagnosis of stage C HFrEF

- Stage C: current or previous heart failure signs or symptoms
- HFrEF: heart failure with LVEF  $\leq 40\%$

**Address** congestion with diuretics

2

**Initiate** guideline directed medical therapy (GDMT)

- Goal is to get patients on all 4 categories below
- Order doesn't matter, use patient factors
- Start 1-2 agents at a time, some patients can tolerate starting low doses of all 4 groups within one week

3

**Titrate** GDMT to target doses as tolerated

- Double doses every 2-4 weeks as vitals/labs allow, consider more often inpatient
- Starting doses listed below, target doses in [brackets]
- **Epic** tip: use SmartText UKHC HFrEF in notes to document GDMT progress

**Schedule** follow up with cardiology 2 weeks post-discharge & consider cardiac rehab

Afterload reduction: choose one

| ARNI<br>angiotensin receptor -<br>neprilysin inhibitor  | ACEi<br>Angiotensin - converting<br>enzyme inhibitor                                     | ARB<br>Angiotensin receptor<br>blocker   | Alternative/additional<br>afterload reduction  |
|---|--|--|--|
| Sacubitril-valsartan (Entresto)<br>24-26 [97-103] mg BID<br><i>*start at 49-51 mg BID if on<br/>moderate dose vasodilator</i>                               | Lisinopril 2.5-5 [40] mg QD<br>Enalapril 2.5 [10] mg BID<br>Captopril 6.25 [50] mg TID   | Valsartan 40 [160] mg BID<br>Losartan 25 [150] mg QD<br>Candesartan 4 [32] mg QD   | Hydralazine +<br>nitrate<br>Hydralazine 25 [100] mg TID<br>Isosorbide dinitrate (Isordil)<br>20 [40] mg TID<br><i>*outpatient combination product<br/>Bidil 37.5-20 mg TID</i>   |
| <ul style="list-style-type: none"> <li>• Use: first line if SBP &gt;100 mmHg, requires a 36 hour washout from last ACEi, <b>cost check first</b></li> </ul> | <ul style="list-style-type: none"> <li>• Use: first line if not using an ARNI</li> </ul> | <ul style="list-style-type: none"> <li>• Use: if plan to switch to ARNI eventually or if cannot tolerate ACEi</li> </ul> | <ul style="list-style-type: none"> <li>• Use: reserve for contraindications to other vasodilators &amp; additional therapy for Black patients on all 4 pillars</li> <li>• <b>Contraindicated:</b> lupus</li> <li>• <b>Monitor:</b> BP</li> </ul> |
| <p><b>Contraindicated:</b> AKI, K&gt;5.5, angioedema, pregnancy, bilateral renal artery stenosis<br/><b>Monitor:</b> BP, K, SCr (~30% bump expected)</p>    |  |  |  |

## Beta Blocker (BB)

metoprolol succ 25 [200] mg daily, carvedilol 6.25 [25, 50 if >85 kg] mg BID, bisoprolol 2.5 [10] mg daily

- Use: consider 1st if tachycardic or hypertensive, carvedilol offers more BP control
- **Contraindicated:** decompensated, symptomatic bradycardia, severe lung disease
- **Monitor:** HR, BP



## Mineralocorticoid Receptor Antagonist (MRA)

spironolactone 12.5-25 [25] mg daily, eplerenone 25 [50] mg daily

- Use: consider 1st if volume overloaded or hypokalemic; use eplerenone if gynecomastia
- **Contraindicated:** K > 5.0, SCr > 2.5 for males or > 2.0 for females
- **Monitor:** BP, SCr, K (stop if K > 5.5)

## Sodium-Glucose Cotransporter-2 Inhibitor (SGLT2i)

dapagliflozin (Farxiga) 10 mg daily, empagliflozin (Jardiance) 10 mg daily [flat dose, no titration]

- Use: consider 1st if hyperglycemic, volume overloaded, or have less BP room; **cost check first**
- **Contraindicated:** recurrent genital mycotic infection, T1DM, eGFR < 25 mL/min
- **Monitor:** BP, SCr (~30% bump expected), BG, signs/symptoms of hypoglycemia & DKA



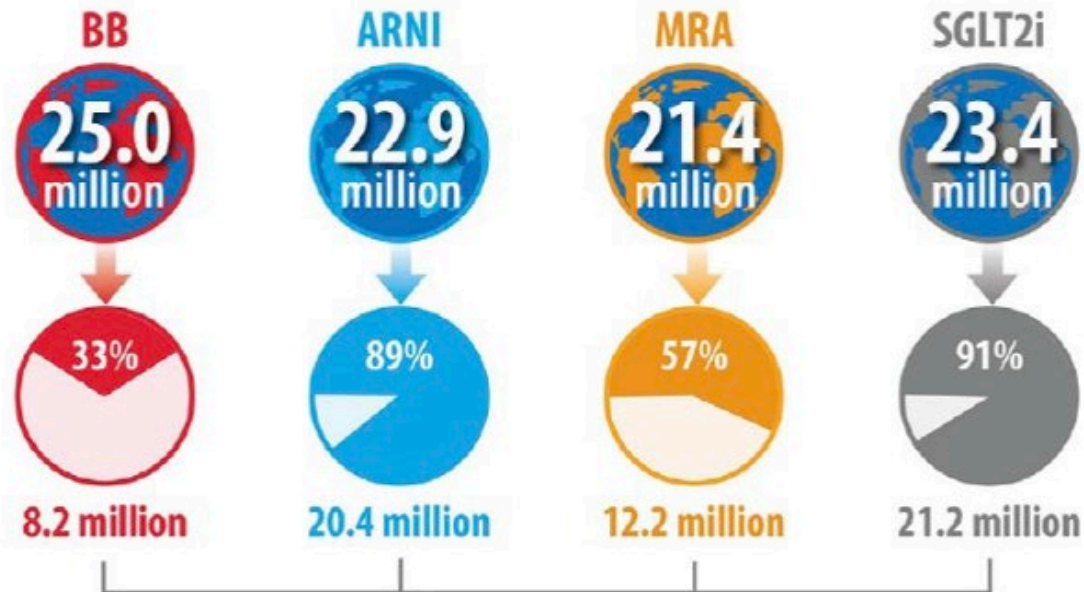
# Medication Uptitration Clinic





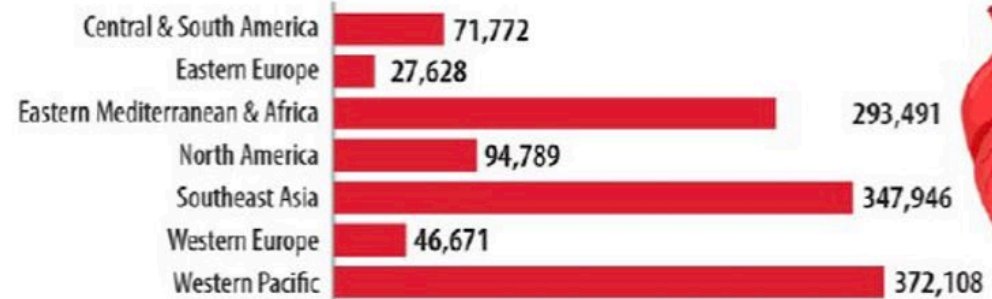
# HFrEF Prevalence is Estimated at **29 million worldwide**

Estimated patients worldwide with HFrEF eligible for GDMT



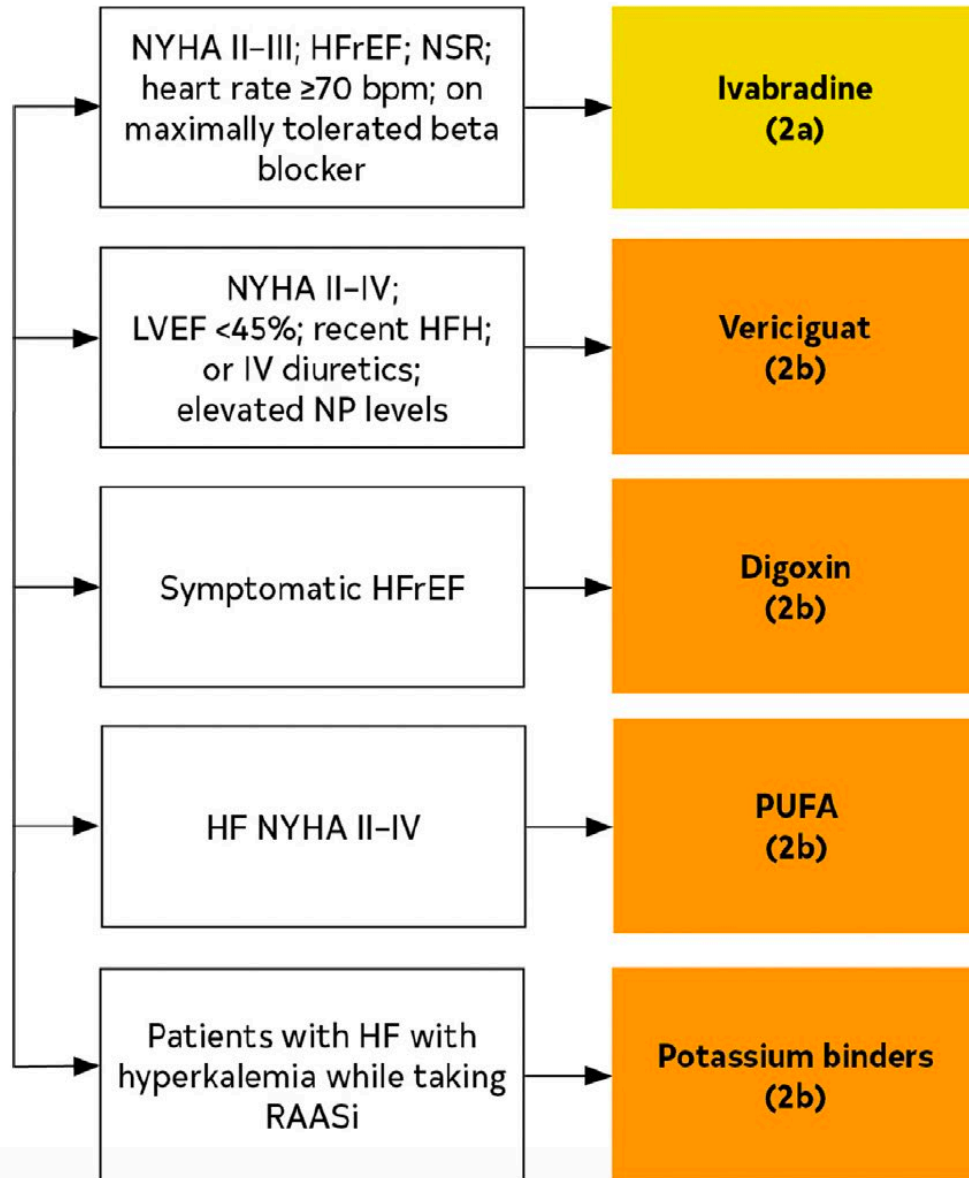
Estimated patients worldwide not on GDMT

Potential lives saved globally on optimal GDMT quadruple therapy

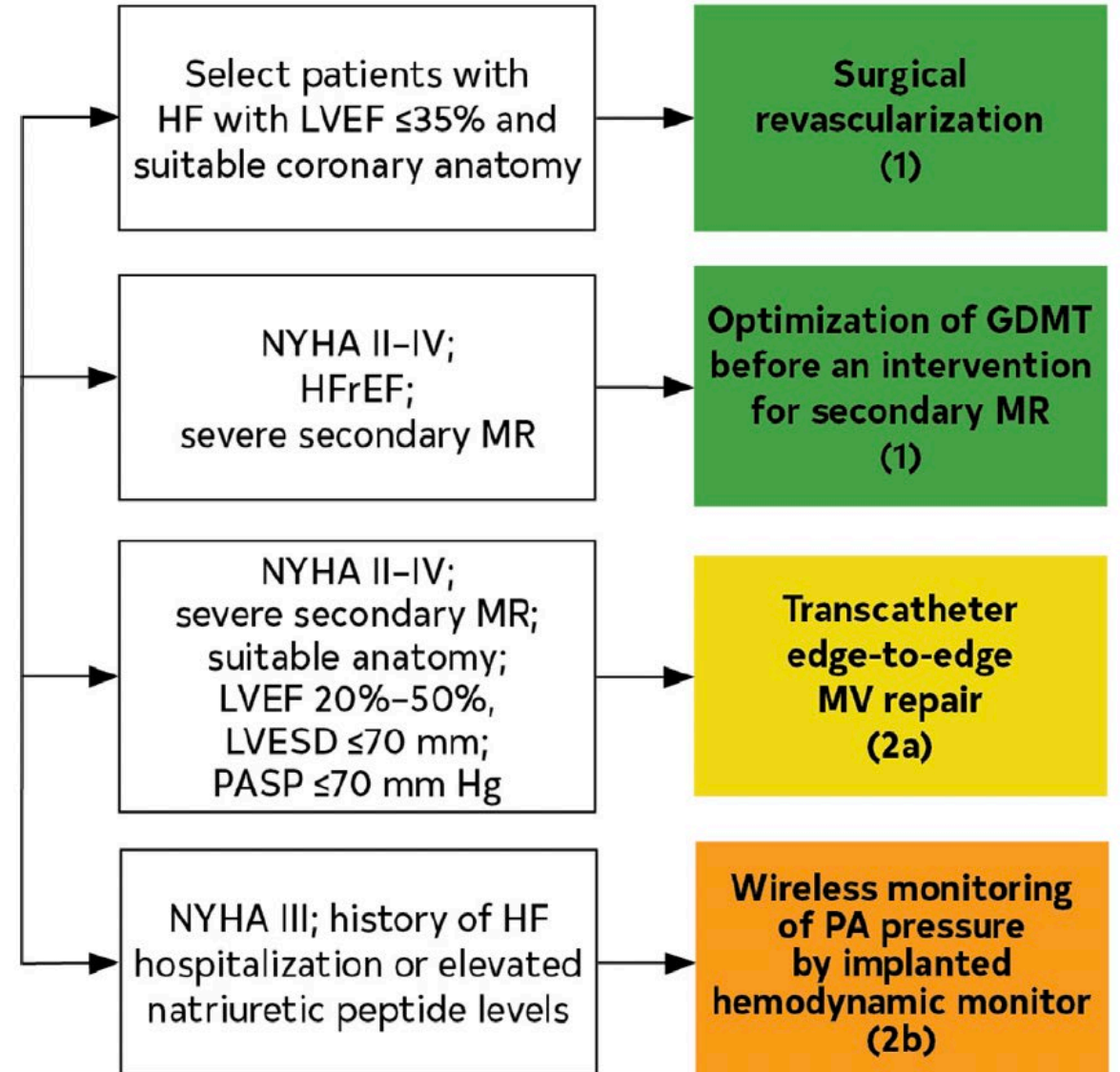


**1.2 million**  
potential lives saved per year

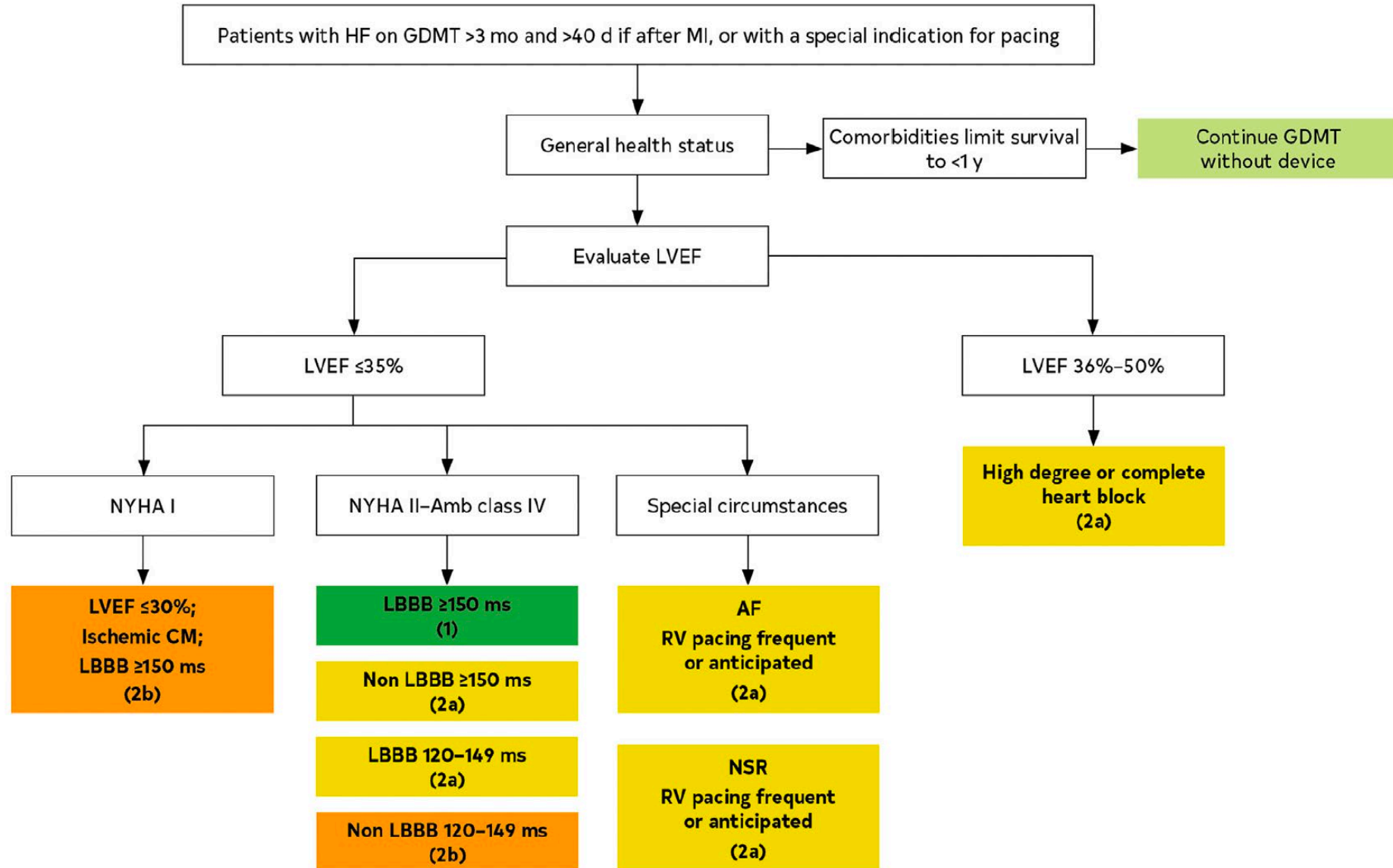
### Consider Additional Therapies Once GDMT Optimized



### Consider Additional Therapies Once GDMT Optimized

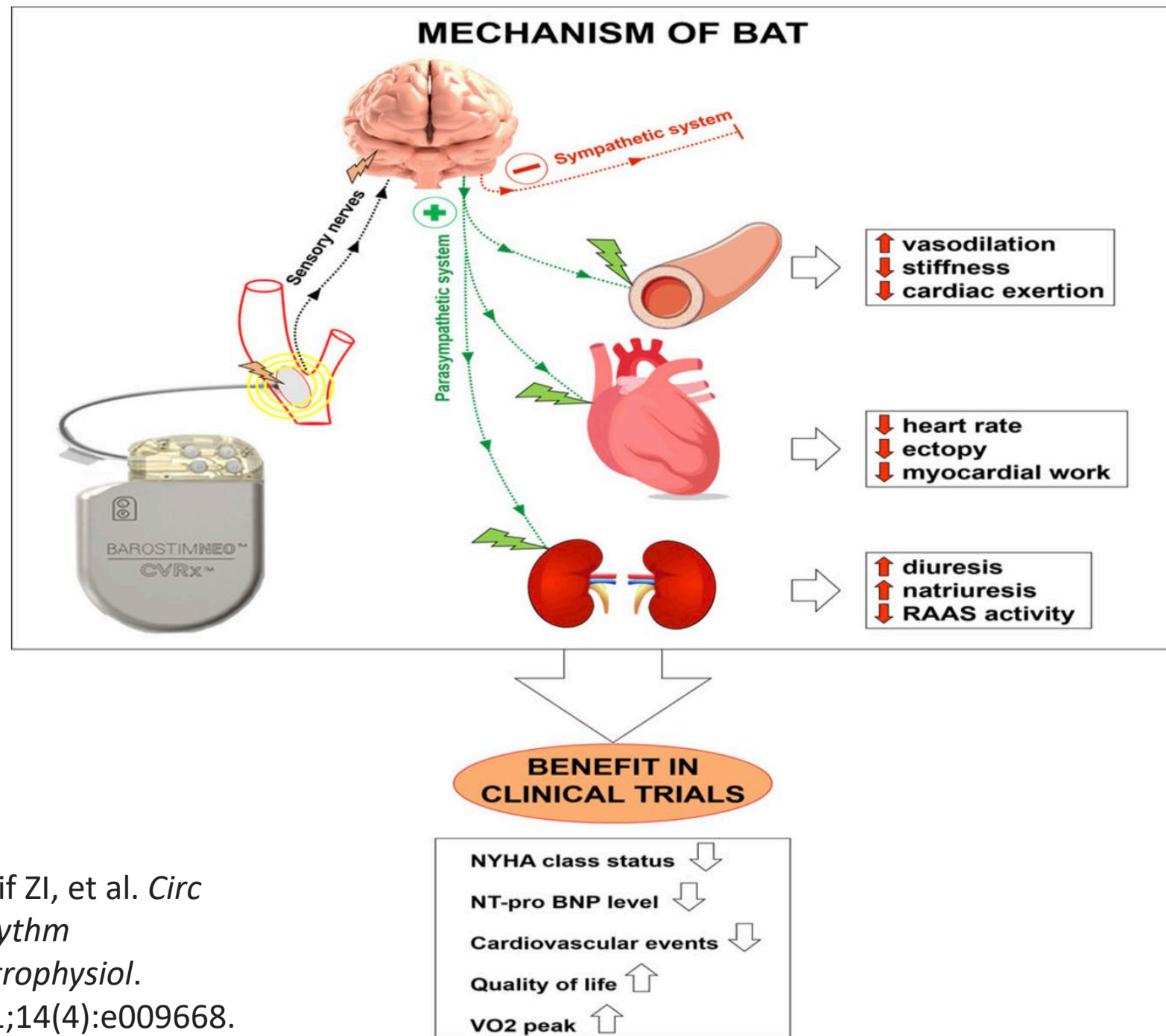


## CRT Recommendations





# BARORECEPTOR ACTIVATION THERAPY



Sharif ZI, et al. *Circ Arrhythm Electrophysiol.*  
2021;14(4):e009668.

# Baroreceptor activation therapy

## Barostim Indications for Use

- NYHA III or NYHA II with recent history of NYHA III on GDMT\*
- LVEF  $\leq 35\%$
- NT-proBNP  $< 1600$  pg/mL
- Not indicated for CRT\*\*

\*Guideline-directed medical therapy (GDMT) according to 2022 AHA/ACC/ESC guidelines

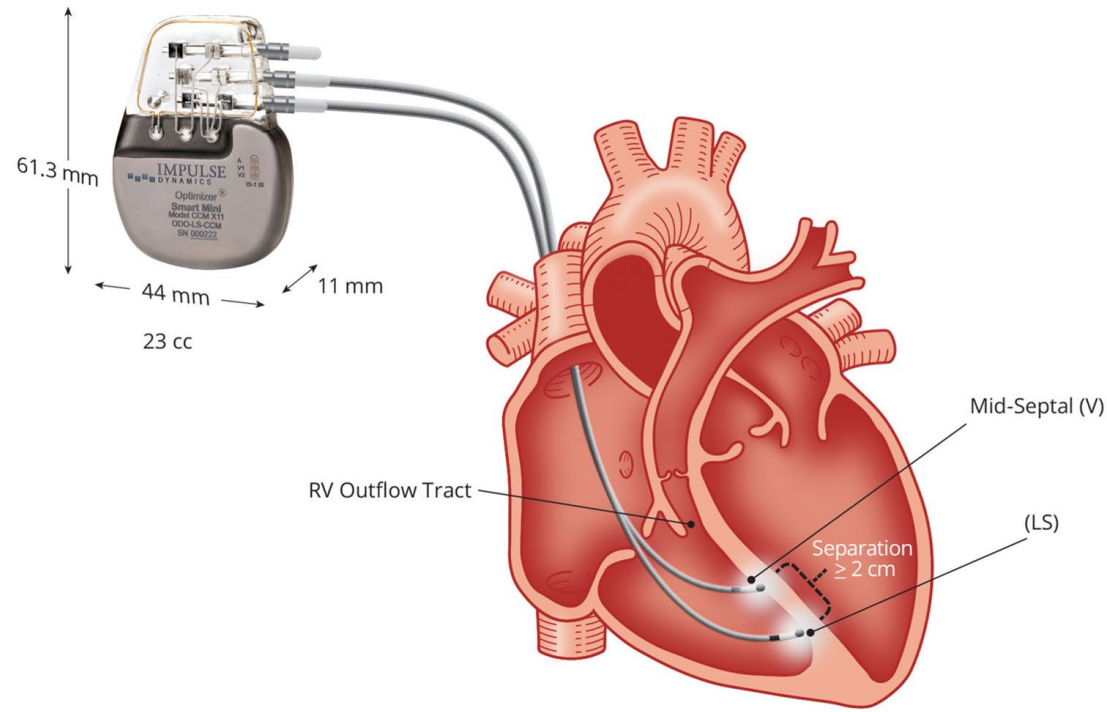
\*\*Or not receiving adequate response from existing CRT device

*NYHA = New York Heart Association; LVEF = left ventricular ejection fraction;  
CRT = cardiac resynchronization therapy; AHA/ACC/ESC = American Heart Association/American College of Cardiology/European Society of Cardiology*



# Cardiac Contractility Modulation

## CCM DEVICE AND ANATOMICAL LOCATION OF PACING WIRES



### Mechanism of action

Application of non-excitatory electric stimulation to the interventricular septum during the absolute refractory period

### Biomolecular changes

- Optimization of intra-cellular calcium homeostasis
  - ↑ titin phosphorylation
- Upregulation of pivotal cardioprotective genes
- Amplification of downstream proteomic signaling

### Alteration in myocardial properties

- Lusitropic effect with improved diastolic recoil
  - Increased left ventricular contractility

### Effect on functional and clinical outcomes

- ↑ ejection fraction reserve
- ↑ diastolic filling index
- ↑ exercise capacity
- ↑ functional status
- ↑ survival

# CCM - Indications

- Indicated to improve 6-minute hall walk distance, quality of life and functional status of NYHA Class III heart failure patients who remain symptomatic
  - Despite guideline directed medical therapy
  - Who are in normal sinus rhythm
  - Are not indicated for CRT
  - have an LVEF ranging from 25% to 45%.

# Decompensated Heart Failure



## Evaluation

| COR | RECOMMENDATIONS                 |
|-----|---------------------------------|
| 1   | Address precipitating factors   |
| 1   | Evaluate severity of congestion |
| 1   | Assess adequacy of perfusion    |



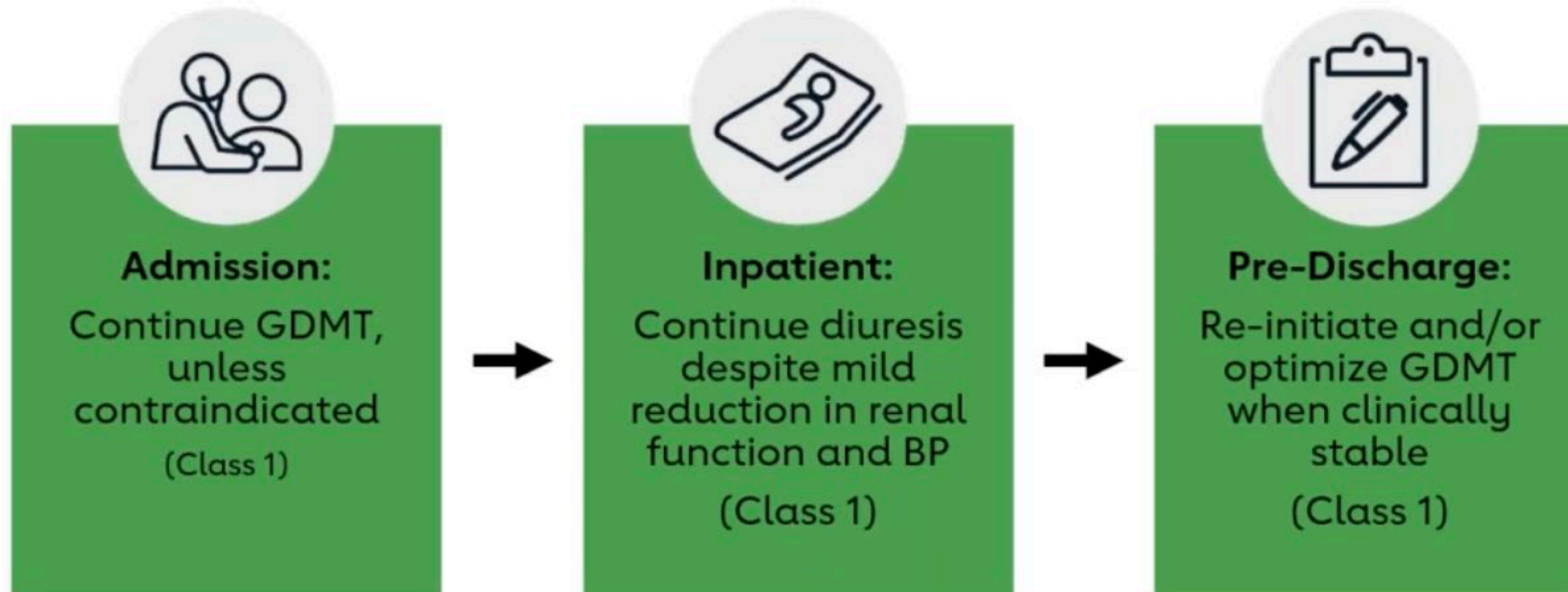
## Goals for GDMT

| COR | RECOMMENDATIONS            |
|-----|----------------------------|
| 1   | Optimize volume status     |
| 1   | Address reversible factors |
| 1   | Continue or initiate GDMT  |

## COMMON FACTORS PRECIPITATING HF HOSPITALIZATION

- |   |   |
|---|---|
| <ul style="list-style-type: none"> <li>• Acute coronary syndrome</li> <li>• Uncontrolled hypertension</li> <li>• Atrial fibrillation and arrhythmias</li> <li>• Additional cardiac disease</li> <li>• Acute infections</li> </ul> | <ul style="list-style-type: none"> <li>• Non-adherence to medications or diet</li> <li>• Anemia</li> <li>• Hypo-/Hyperthyroidism</li> <li>• Medications that increase sodium retention</li> <li>• Medications with negative inotrope</li> </ul> |
|---|---|

# Decompensated Heart Failure

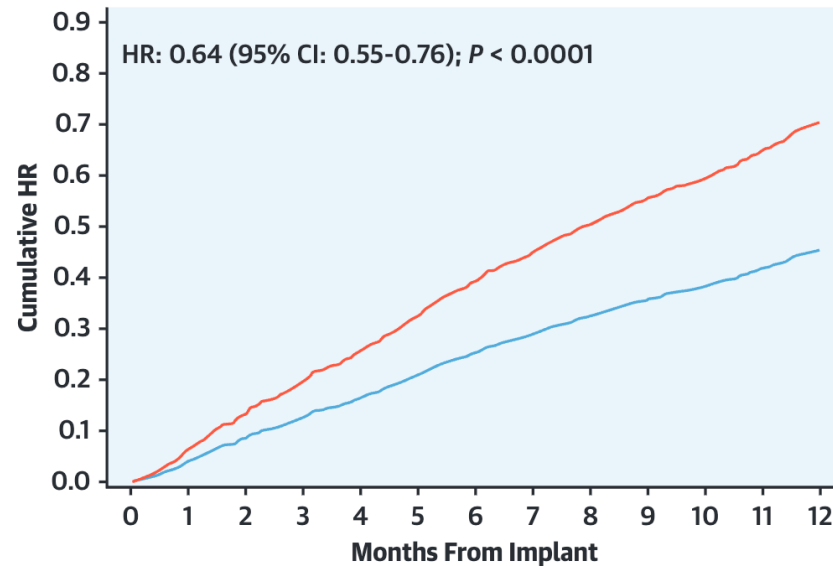
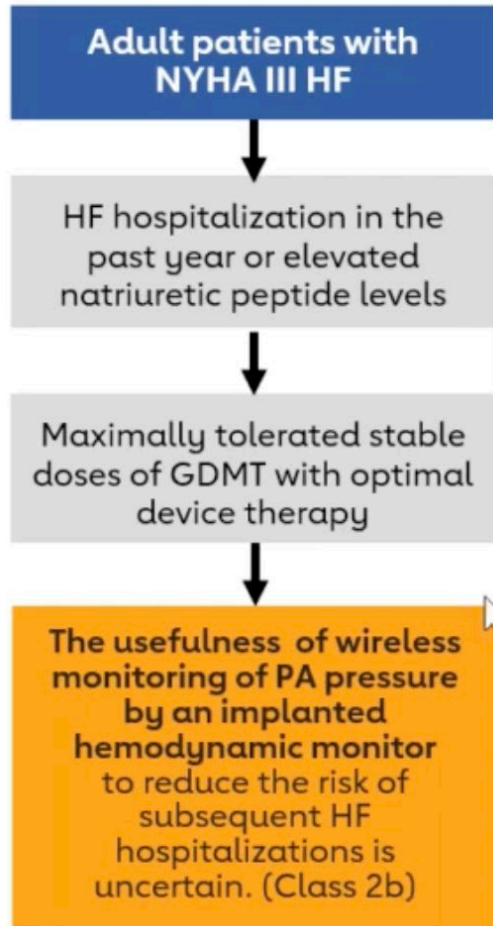


## Special considerations

- Consider discontinuation of beta blockers in patients with low cardiac output, severe volume overload, advanced AV block or ACEi/ARNi with angioedema
- VTE prophylaxis is recommended in all hospitalized patients



# Initial and Serial Evaluation



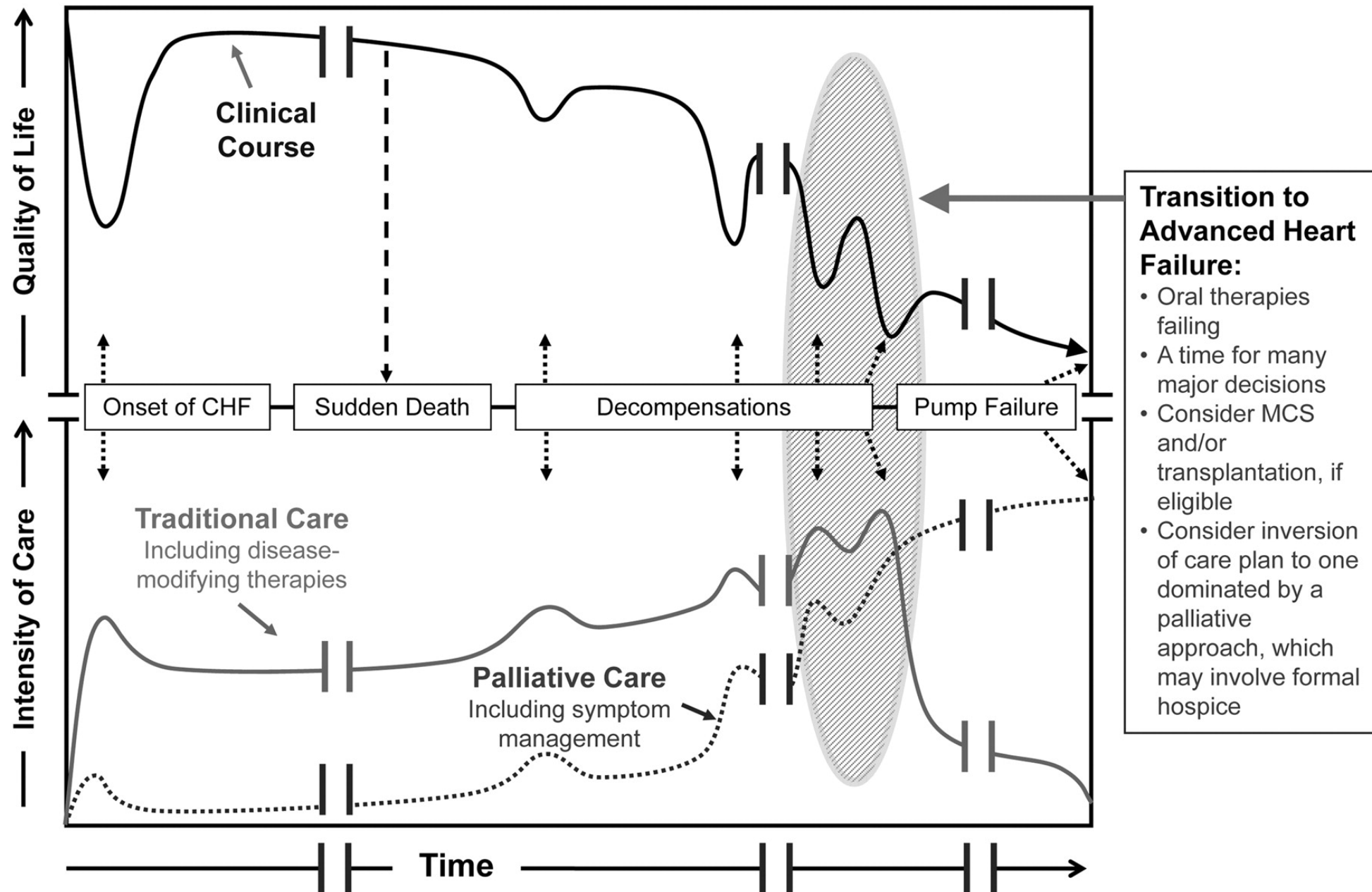
|             |     |     |     |     |     |     |     |     |     |     |     |     |     |
|-------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| No. At Risk |     |     |     |     |     |     |     |     |     |     |     |     |     |
| Treatment   | 666 | 662 | 655 | 635 | 601 | 569 | 539 | 511 | 485 | 468 | 438 | 408 | 342 |
| Control     | 684 | 674 | 664 | 635 | 607 | 575 | 554 | 532 | 514 | 484 | 456 | 429 | 352 |

— Treatment — Control

# Heart Failure Hospitalizations for Implantable Hemodynamic Monitoring and Medical Therapy in Pooled Population

| COR | RECOMMENDATIONS   |
|-----|---|
| 1   | 1. In patients with HF, assessment and documentation of NYHA functional                                 |
| 2a  | 4. In ambulatory patients with unexplained dyspnea, CPET is reasonable to evaluate the cause of dyspnea |





# IS IT STAGE D HEART FAILURE?

|          |   |
|----------|---|
| <b>I</b> | Need for <b>inotropes</b>                               |
| <b>N</b> | <b>New</b> York Heart Association Class IV              |
| <b>E</b> | Worsening <b>end-organ</b> dysfunction                  |
| <b>E</b> | <b>Ejection fraction</b> <20%                           |
| <b>D</b> | <b>Defibrillator</b> shocks for ventricular arrhythmias |
| <b>H</b> | Recurrent <b>HF</b> hospitalizations                    |
| <b>E</b> | <b>Escalating</b> diuretic dose                         |
| <b>L</b> | <b>Low blood pressure</b>                               |
| <b>P</b> | <b>Progressive</b> intolerance of GDMT                  |



How important is the family physician in the care of heart failure patients?

# Advanced Heart Failure, VAD & Transplant Program

## Medical Team



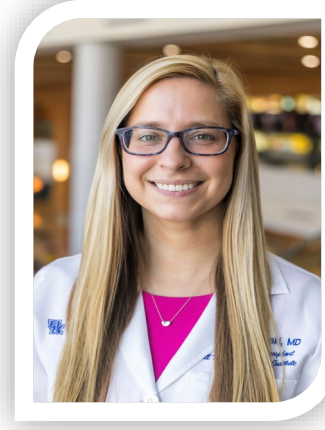
**Emma Birks, MD, PhD**  
Section Chief, Heart Failure



**Andrew Kolodziej, MD**  
Medical Director of Heart Transplant



**Navin Rajagopalan, MD**  
Director, Gill Affiliate Network



**Maya Ignaszewski, MD**



**Sonu Abraham, MBBS**

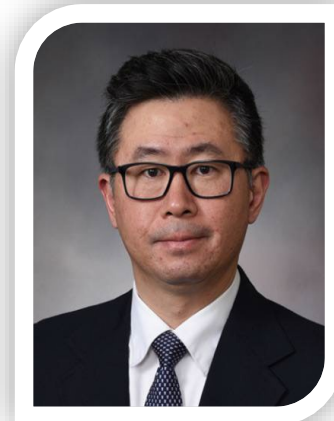
## Surgical Team



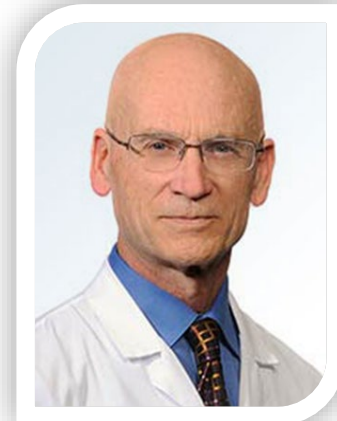
**Matthias Loebe, MD, PhD**  
Surgical Director, Heart  
Transplant



**Masashi Kawabori, MD**  
Surgical Director, MCS



**Jin Woo Chung, MD**



**Michael Sekela, MD**  
Chief, Cardiothoracic Surgery



# Specialized Care - Complex Cardiomyopathies

- Accepting referrals for diagnosis and/or management of cardiac amyloidosis (TTR and AL), sarcoidosis, and HCM.
- Comprehensive evaluation:
  - Heart biopsy
  - PYP nuclear scan
  - Cardiac MRI
  - Genetic screening
  - Multidisciplinary collaboration through **UK Amyloidosis Alliance**.
- Offering management of patients on long-term pharmacologic therapy (Tamafidis, Acoramidis, Amvuttra, Camzyos).



Sonu Abraham, MD, FACC

[sonu.abraham@uky.edu](mailto:sonu.abraham@uky.edu)



# Specialized Care - Cardio-Psychiatry

Holistic, Integrated patient care

- Allows for simultaneous treatment of cardiac and mental health needs

Improved patient outcomes

- Improved patient quality of life, medication adherence, and potential for reduced readmissions

Early detection and intervention of psychological impact

- Allows for prompt identification of issues following new diagnosis or after heart transplant/LVAD



Aaron Harris, DNP, APRN, PMHNP-BC





THANK YOU