Confidence In Network Meta Analysis - CINeMA 2.0.0 - Project: **HFpEF network meta-analysis**

Incoherence

Define clinically important size of effect: Odds ratio

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Relative effect estimates below **1.000** and above **1.000** are considered clinically important.

Importance of Incoherence depends on the variability of direct and indirect effects in relation to a clinically important size of effect

Global test based on a random-effects design-by-treatment interaction model

 χ^2 statistic: 1.912 (2 degrees of freedom), P value: 0.384

Local tests: Separating indirect from direct evidence

Comparison Evidence: mixed	ACEI:ARB
NMA odds ratio: Direct odds ratio: Indirect odds ratio:	0.831(0.433,1.594) 0.443(0.018,11.167) 0.853(0.439,1.659)
Inconsistency measures Ratio of odds ratios: P value:	0.519(0.019,14.009) 0.697
Incoherence judgment	Some concerns 🗸

Comparison Evidence: mixed	ACEI:Placebo
NMA odds ratio:	0.845(0.448,1.595)
Direct odds ratio:	0.835(0.442,1.577)
Indirect odds ratio:	407.198(0.000,834213564.430)
Inconsistency measure Ratio of odds ratios: P value:	os 0.002(0.000,4261.174) 0.404
Incoherence judgment	Some concerns ✓

Comparison Evidence: direct	ARB:ARNI
Direct odds ratio: Inconsistency measures: Not	1.036(0.882,1.217) applicable
Incoherence judgment	No concerns 🗸

Comparison ARB:Placebo
Evidence: direct

Direct odds ratio: 1.016(0.876,1.179)
Inconsistency measures: Not applicable

Comparison Beta blockers:Placebo

No concerns

Evidence: direct

Incoherence judgment

Direct odds ratio: **0.588(0.363,0.953)**

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison Digoxin:Placebo

Evidence: direct

Direct odds ratio: 0.999(0.744,1.342)

Inconsistency measures: Not applicable

Incoherence judgment Major concerns 🗸

Comparison MRA:Placebo

Evidence: direct

Direct odds ratio: **0.910(0.756,1.096)**

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison Placebo:SGLT-2

Evidence: direct

Direct odds ratio: 1.063(0.918,1.232)

Inconsistency measures: Not applicable

Incoherence judgment No concerns 🗸

Comparison Placebo: Vericiguat 10mg

Evidence: direct

Direct odds ratio: **0.452(0.181,1.127)**

Inconsistency measures: Not applicable

Incoherence judgment No concerns 🗸

Comparison Placebo: Vericiguat 15mg

Evidence: direct

Direct odds ratio: **0.697(0.261,1.860)**

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison Vericiguat 10mg:Vericiguat 15mg

Evidence: direct

Direct odds ratio: 1.542(0.680,3.499)

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison **ACEI:ARNI Evidence: indirect** Indirect odds ratio: 0.861(0.440,1.684) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison **ACEI:Beta blockers Evidence: indirect** Indirect odds ratio: 1.437(0.647,3.193) **Inconsistency measures:** Not applicable Incoherence judgment Some concerns **✓** Comparison **ACEI:Digoxin Evidence: indirect** Indirect odds ratio: 0.846(0.420,1.703) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ACEI:MRA Evidence: indirect** Indirect odds ratio: 0.929(0.479,1.799) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ACEI:SGLT-2 Evidence: indirect** Indirect odds ratio: 0.899(0.468, 1.725) Inconsistency measures: Not applicable Incoherence judgment No concerns **ACEI:Vericiguat 10mg** Comparison **Evidence: indirect** Indirect odds ratio: 0.382(0.126,1.163) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison ACEI:Vericiguat 15mg **Evidence: indirect** Indirect odds ratio: 0.589(0.183, 1.897) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison **ARB:Beta blockers Evidence: indirect** Indirect odds ratio: 1.731(1.044,2.869)

Inconsistency measures: Not applicable

No concerns

Incoherence judgment

Comparison **ARB:Digoxin Evidence: indirect** Indirect odds ratio: 1.018(0.732,1.416) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ARB:MRA Evidence: indirect** Indirect odds ratio: 1.118(0.882,1.418) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ARB:SGLT-2 Evidence: indirect** Indirect odds ratio: 1.082(0.878, 1.333) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ARB:Vericiguat 10mg Evidence: indirect** Indirect odds ratio: 0.460(0.182,1.161) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison ARB: Vericiguat 15mg **Evidence: indirect** Indirect odds ratio: 0.709(0.263,1.914) Inconsistency measures: Not applicable Incoherence judgment No concerns **ARNI:Beta blockers** Comparison **Evidence: indirect** Indirect odds ratio: 1.670(0.983, 2.840) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ARNI:Digoxin Evidence: indirect** Indirect odds ratio: 0.983(0.681,1.419) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison ARNI:MRA **Evidence: indirect** 1.079(0.810,1.438) Indirect odds ratio:

Inconsistency measures: Not applicable

No concerns

Incoherence judgment

Comparison ARNI:Placebo **Evidence: indirect** Indirect odds ratio: 0.982(0.789,1.223) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison **ARNI:SGLT-2 Evidence: indirect** Indirect odds ratio: 1.044(0.802, 1.359) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison **ARNI:Vericiguat 10mg Evidence: indirect** Indirect odds ratio: 0.444(0.173,1.136) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **ARNI:Vericiguat 15mg Evidence: indirect** Indirect odds ratio: 0.685(0.251,1.872) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison Beta blockers:Digoxin **Evidence: indirect** Indirect odds ratio: 0.588(0.334,1.036) Inconsistency measures: Not applicable Incoherence judgment No concerns Beta blockers: MRA Comparison **Evidence: indirect** Indirect odds ratio: 0.646(0.385,1.084) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **Beta blockers:SGLT-2 Evidence: indirect** Indirect odds ratio: 0.625(0.377,1.036)

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison Beta blockers: Vericiguat 10mg **Evidence: indirect**

Indirect odds ratio: 0.266(0.095, 0.747)

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison Beta blockers: Vericiguat 15mg **Evidence: indirect** Indirect odds ratio: 0.410(0.137,1.224) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison Digoxin:MRA **Evidence: indirect** Indirect odds ratio: 1.098(0.775, 1.555) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison **Digoxin:SGLT-2 Evidence: indirect** 1.062(0.764,1.477) Indirect odds ratio: **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **Digoxin:Vericiguat 10mg Evidence: indirect** Indirect odds ratio: 0.452(0.173,1.180) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison **Digoxin:Vericiguat 15mg Evidence: indirect** Indirect odds ratio: 0.697(0.250,1.941) Inconsistency measures: Not applicable Incoherence judgment No concerns MRA:SGLT-2 Comparison **Evidence: indirect** Indirect odds ratio: 0.968(0.764,1.226) **Inconsistency measures:** Not applicable Incoherence judgment No concerns Comparison MRA: Vericiguat 10mg **Evidence: indirect** Indirect odds ratio: 0.411(0.162,1.046) Inconsistency measures: Not applicable Incoherence judgment No concerns Comparison

MRA: Vericiguat 15mg **Evidence: indirect** Indirect odds ratio: 0.635(0.234,1.723) Inconsistency measures: Not applicable Incoherence judgment No concerns

Comparison SGLT-2:Vericiguat 10mg

Evidence: indirect

Indirect odds ratio: **0.425(0.168,1.073)**

Inconsistency measures: Not applicable

Incoherence judgment No concerns

Comparison SGLT-2:Vericiguat 15mg

Evidence: indirect

Indirect odds ratio: **0.656(0.243,1.769)**

Inconsistency measures: Not applicable

Incoherence judgment No concerns