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

### Heterogeneity

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 heterogeneity depends on the variability of effects in relation to a clinically important size of effect

The estimated value of between-study variance for the network meta-analysis is **0.000** 

Comparison ACEI:ARB Evidence: mixed

NMA estimate: **0.831** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.433,1.594) interval: (0.353,1.952)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ARB:ARNI Evidence: mixed

NMA estimate: 1.036

95% intervals for NMA estimate

Confidence interval:

Prediction (0.882,1.217) interval: (0.839,1.279)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ~

Comparison ACEI:Placebo Evidence: mixed

NMA estimate: **0.845** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.448,1.595) interval: (0.368,1.944)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ARB:Placebo Evidence: mixed

NMA estimate: 1.018

95% intervals for NMA estimate

Confidence interval:

Prediction (0.877,1.180) interval: (0.837,1.236)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison

Beta blockers:Placebo

**Evidence:** mixed

NMA estimate: **0.588** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.363,0.953) interval: (0.312,1.108)

Prediction interval extends into clinically important effects in **both** directions

Heterogeneity judgment

Major concerns ✓

Comparison MRA:Placebo Evidence: mixed

NMA estimate: **0.910** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.756,1.096) interval: (0.714,1.161)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ~

Comparison Digoxin:Placebo Evidence: mixed

NMA estimate: **0.999** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.744,1.342) interval: (0.679,1.471)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison Placebo:SGLT-2

**Evidence:** mixed

NMA estimate: 1.063

95% intervals for NMA estimate

Confidence interval:

Prediction (0.918,1.232) interval: (0.877,1.289)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison

Placebo: Vericiguat 10mg

**Evidence:** mixed

NMA estimate: **0.452** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.181,1.127) interval: (0.136,1.499)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns >

Comparison

Placebo:Vericiguat 15mg

**Evidence:** mixed

NMA estimate: 0.697

95% intervals for NMA estimate

Confidence interval:

Prediction (0.261,1.860) interval: (0.192,2.526)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison

Vericiguat 10mg:Vericiguat 15mg

**Evidence:** mixed

NMA estimate: 1.542

95% intervals for NMA estimate

Confidence interval:

Prediction (0.680,3.499) interval: (0.527,4.516)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns 🗸

Comparison ACEI:ARNI

**Evidence: indirect** 

NMA estimate: **0.861** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.440,1.684) interval: (0.357,2.075)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison

**Evidence: ACEI:Beta blockers** 

indirect

NMA estimate: 1.437

95% intervals for NMA estimate

Confidence interval:

Prediction (0.647,3.193) interval: (0.505,4.094)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ~

Comparison ACEI:MRA Evidence: indirect

NMA estimate: **0.929** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.479,1.799) interval: (0.390,2.211)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ACEI:Digoxin Evidence: indirect

NMA estimate: **0.846** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.420,1.703) interval: (0.338,2.118)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ACEI:SGLT-2

**Evidence: indirect** 

NMA estimate: **0.899** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.468,1.725) interval: (0.382,2.113)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison

**ACEI:Vericiguat 10mg** 

**Evidence: indirect** 

NMA estimate: **0.382** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.126,1.163) interval: (0.089,1.645)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ~

Comparison

**Evidence: ARB:Beta blockers** 

indirect

NMA estimate: 1.731

95% intervals for NMA estimate

Confidence interval:

Prediction (1.044,2.869) interval: (0.892,3.359)

Prediction interval extends into clinically important effects in

**both** directions

Heterogeneity judgment

Major concerns 🗸

Comparison

**ACEI:Vericiguat 15mg** 

**Evidence: indirect** 

NMA estimate: **0.589** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.183,1.897) interval: (0.127,2.730)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ARB:Digoxin Evidence: indirect

NMA estimate: 1.018

95% intervals for NMA estimate

Confidence interval:

Prediction (0.732,1.416) interval: (0.661,1.570)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison ARB:MRA Evidence: indirect

NMA estimate: 1.118

95% intervals for NMA estimate

Confidence interval:

Prediction (0.882,1.418) interval: (0.819,1.527)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ✓

#### Comparison

**ARB:Vericiguat 10mg** 

**Evidence: indirect** 

NMA estimate: **0.460** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.182,1.161) interval: (0.137,1.549)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ~

Comparison ARB:SGLT-2

**Evidence: indirect** 

NMA estimate: 1.082

95% intervals for NMA estimate

Confidence interval:

Prediction (0.878,1.333) interval: (0.823,1.423)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

#### Comparison

ARB:Vericiguat 15mg

**Evidence: indirect** 

NMA estimate: 0.709

95% intervals for NMA estimate

Confidence interval:

Prediction (0.263,1.914) interval: (0.193,2.608)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison

**Evidence: ARNI:Beta blockers** 

indirect

NMA estimate: 1.670

95% intervals for NMA estimate

Confidence interval:

Prediction (0.983,2.840) interval: (0.833,3.350)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ARNI:MRA Evidence: indirect

NMA estimate: 1.079

95% intervals for NMA estimate

Confidence interval:

Prediction (0.810,1.438) interval: (0.741,1.572)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ✓

Comparison ARNI:Digoxin Evidence: indirect

NMA estimate: 0.983

95% intervals for NMA estimate

Confidence interval:

Prediction (0.681,1.419) interval: (0.607,1.591)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison ARNI:Placebo Evidence: indirect

NMA estimate: 0.982

95% intervals for NMA estimate

Confidence interval:

Prediction (0.789,1.223) interval: (0.737,1.309)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison ARNI:SGLT-2 Evidence: indirect

NMA estimate: 1.044

### 95% intervals for NMA estimate

Confidence interval:

Prediction (0.802,1.359) interval: (0.739,1.476)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

#### Comparison

**ARNI:Vericiguat 15mg** 

**Evidence: indirect** 

NMA estimate: **0.685** 

### 95% intervals for NMA estimate

Confidence interval:

Prediction (0.251,1.872) interval: (0.183,2.560)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ✓

#### **Comparison**

**ARNI:Vericiguat 10mg** 

**Evidence: indirect** 

NMA estimate: **0.444** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.173,1.136) interval: (0.129,1.523)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

#### Comparison

Beta blockers:Digoxin

**Evidence: indirect** 

NMA estimate: **0.588** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.334,1.036) interval: (0.280,1.236)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison

**Evidence: Beta blockers:MRA** 

indirect

NMA estimate: **0.646** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.385,1.084) interval: (0.328,1.274)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison

Beta blockers:Vericiguat

10mg

**Evidence: indirect** 

NMA estimate: **0.266** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.095,0.747) interval: (0.068,1.031)

Prediction interval extends into clinically important effects in **both** directions

Heterogeneity judgment

Major concerns 🗸

Comparison

**Beta blockers:SGLT-2** 

**Evidence: indirect** 

NMA estimate: **0.625** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.377,1.036) interval: (0.322,1.213)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

Comparison

**Beta blockers:Vericiguat** 

15mg

**Evidence: indirect** 

NMA estimate: **0.410** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.137,1.224) interval: (0.098,1.721)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison Digoxin:MRA Evidence: indirect

NMA estimate: 1.098

### 95% intervals for NMA estimate

Confidence interval:

Prediction (0.775,1.555) interval: (0.695,1.733)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ✓

#### Comparison

Digoxin:Vericiguat 10mg
Evidence: indirect

NMA estimate: **0.452** 

## 95% intervals for NMA estimate

Confidence interval:

Prediction (0.173,1.180) interval: (0.128,1.592)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns 🗸

### Comparison Digoxin:SGLT-2 Evidence: indirect

NMA estimate: 1.062

95% intervals for NMA estimate

Confidence interval:

Prediction (0.764,1.477) interval: (0.690,1.636)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

#### Comparison

Digoxin:Vericiguat 15mg
Evidence: indirect

NMA estimate: 0.697

95% intervals for NMA estimate

Confidence interval:

Prediction (0.250,1.941) interval: (0.182,2.671)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

Comparison MRA:SGLT-2 Evidence: indirect

NMA estimate: **0.968** 

### 95% intervals for NMA estimate

Confidence interval:

Prediction (0.764,1.226) interval: (0.709,1.320)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

#### Comparison

MRA: Vericiguat 15mg

**Evidence: indirect** 

NMA estimate: 0.635

### 95% intervals for NMA estimate

Confidence interval:

Prediction (0.234,1.723) interval: (0.171,2.352)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns ✓

#### **Comparison**

MRA: Vericiguat 10mg

**Evidence: indirect** 

NMA estimate: **0.411** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.162,1.046) interval: (0.121,1.398)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

No concerns

#### Comparison

SGLT-2:Vericiguat 10mg

**Evidence: indirect** 

NMA estimate: **0.425** 

95% intervals for NMA estimate

Confidence interval:

Prediction (0.168,1.073) interval: (0.126,1.432)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment

#### Comparison

SGLT-2:Vericiguat 15mg

**Evidence: indirect** 

NMA estimate: **0.656** 

95% intervals for NMA

estimate

Confidence interval:

Prediction (0.243,1.769) interval: (0.178,2.410)

Confidence and prediction intervals **agree** in relation to clinically important effect

Heterogeneity judgment