

Supplementary Table 6. Certainty of evidence assessment with GRADE approach

Participants (studies)	Certainty assessment					Other consideration	Certainty of evidence	Summary of findings
	Risk of bias	Inconsistency	Indirectness	Imprecision	Impact			
eGFR slope								
28,038 (6 obs. studies)	Not serious	Not serious	Not serious	Not serious	None	⊕⊕○○ Low	MD, 0.64; 95% CI: 0.437–0.843; <i>P</i> <0.01	
Major adverse kidney events								
15,610 (3 obs. studies)	Not serious	Not serious	Not serious	Not serious	None	⊕⊕○○ Low	HR, 1.01; 95% CI: 0.869–1.173; <i>P</i> =0.87	
Hospitalized heart failure and cardiovascular death								
3,841 (2 obs. studies)	Not serious	Not serious	Not serious	Serious ^a	None	⊕○○○ Very low	HR, 0.824; 95% CI: 0.633–1.074; <i>P</i> =0.10	
Hospitalized heart failure								
12,030 (2 obs. studies)	Not serious	Not serious	Not serious	Very serious ^b	None	⊕○○○ Very low	HR, 1.059; 95% CI: 0.574–1.952; <i>P</i> =0.79	
All-cause mortality								
3,857 (2 obs. studies)	Not serious	Not serious	Not serious	Serious ^a	None	⊕○○○ Very low	HR, 0.83; 95% CI: 0.589–1.170; <i>P</i> =0.18	
Discontinuation of SGLT2i due to adverse events								
11,826 (4 obs. studies)	Not serious	Not serious	Not serious	Not serious	None	⊕⊕○○ Low	OR, 1.023; 95% CI: 0.832–1.257; <i>P</i> =0.4	
Serious adverse events								
9,518 (3 obs. studies)	Not serious	Not serious	Not serious	Not serious	None	⊕⊕○○ Low	OR, 0.991; 95% CI: 0.918–1.070; <i>P</i> =0.87	
Kidney-related adverse events								
11,826 (4 obs. studies)	Not serious	Not serious	Not serious	Serious ^c	None	⊕○○○ Very low	OR, 1.216; 95% CI: 0.873–1.695; <i>P</i> =0.21	
Acute kidney injury								
9,222 (3 obs. studies)	Not serious	Not serious	Not serious	Very serious ^b	None	⊕○○○ Very low	OR, 0.949; 95% CI: 0.510–1.767; <i>P</i> =0.84	
Volume depletion								
4,383 (2 obs. studies)	Not serious	Not serious	Not serious	Serious ^c	None	⊕○○○ Very low	OR, 1.190; 95% CI: 0.869–1.630; <i>P</i> =0.18	
Hyperkalemia								
7,443 (2 obs. studies)	Not serious	Not serious	Not serious	Serious ^c	None	⊕○○○ Very low	OR, 1.144; 95% CI: 0.954–1.371; <i>P</i> =0.10	

GRADE, Grading of Recommendations Assessment, Development and Evaluation; eGFR, estimated glomerular filtration rate; obs., observational; MD, mean difference; CI, confidence interval; HR, hazard ratio; SGLT2i, sodium-glucose cotransporter-2 inhibitor; OR, odds ratio.

^aThe confidence interval includes no difference and potential benefit, ^bThe confidence interval includes potential benefit and harm, ^cThe confidence interval includes no difference and potential harm.