

Summary of outcome trials of drugs with cardiorenal benefits

| Agent (outcome trial) | Population | Clinical outcomes (HR [95% CI] vs placebo) | | | | | | |
|--|---------------------------------------|--|-----------------------------|-----------------------------|----------------------|------------------------------|-----------------------------|--|
| | | MACE | CV mortality | All-cause mortality | Fatal/nonfatal MI | Fatal/nonfatal stroke | Hosp HF | Progression of CKD |
| GLP1-RA | | | | | | | | |
| Exenatide (EXSCEL) | CVD (73%) or CV risk factors | 0.91* (0.83-1.00) | 0.88 (0.76-1.02) | 0.86 (0.77-0.97) | 0.97 (0.85-1.10) | 0.85 (0.70-1.03) | - | - |
| Liraglutide (LEADER) | CVD (72%) or CV risk factors | 0.87* (0.78-0.97) | 0.78 (0.66-0.93) | 0.85 (0.74-0.97) | 0.86 (0.73-1.00) | 0.86 (0.71-1.06) | - | - |
| Semaglutide SC (SUSTAIN 6) | CVD (59%) or CV risk factors | 0.74* (0.58-0.95) | 0.98 (0.65-1.48) | 1.05 (0.74-1.50) | 0.74 (0.51-1.08)† | 0.61 (0.38-0.99)† | - | - |
| Semaglutide Oral (PIONEER 6) | CVD (85%) or CV risk factors | 0.79* (0.57-1.11) | 0.49 (0.27-0.92) | 0.50 (0.31-0.84) | 1.18 (0.73-1.90)† | 0.74 (0.35-1.57)† | - | - |
| Dulaglutide (REWIND) | CVD (31.5%) or CV risk factors | 0.88* (0.79-0.99) | 0.91 (0.78-1.06) | 0.90 (0.80-1.01) | 0.96 (0.79-1.16)† | 0.76 (0.61-0.95)† | - | - |
| Albiglutide (HARMONY) (withdrawn from market) | CVD or PVD | 0.78* (0.68-0.90) | 0.93 (0.73-1.19) | 0.95 (0.79-1.16) | 0.96 (0.79-1.15) | 0.76 (0.62-0.94) | - | - |
| SGLT2i | | | | | | | | |
| Empagliflozin (EMPA-REG) | CVD | 0.86* (0.74-0.99) | 0.62 (0.49-0.77) | 0.68 (0.57-0.82) | 0.87 (0.70-1.09) | 1.18 (0.89-1.56) | 0.65 (0.50-0.85) | 0.61 (0.53-0.70) |
| Canagliflozin (CANVAS PROGRAM) | CVD (66%) or CV risk factors | 0.86* (0.75-0.97) | 0.87 (0.72-1.06) | 0.87 (0.74-1.01) | 0.89 (0.73-1.09) | 0.87 (0.69-1.09) | 0.67 (0.52-0.87) | 0.73 (0.67-0.79) |
| Canagliflozin (CREDESCENCE) | CKD (eGFR 30-90 + proteinuria) | 0.80 (0.67-0.95) | 0.78 (0.61-1.00) | 0.83 (0.68-1.02) | - | - | 0.61 (0.47-0.80) | 0.70*² (0.59-0.82) |
| Dapagliflozin (DECLARE-TIMI) | CVD (41%) or CV risk factors | 0.93* (0.84-1.03) | 0.98 (0.82-1.17) | 0.93 (0.82-1.04) | 0.89 (0.77-1.01) | 1.01 (0.84-1.21) | 0.73 (0.61-0.88) | 0.76 (0.67-0.87) |
| Dapagliflozin (DAPA-HF) | CHF (reduced EF) ± DM (42%) | - ¹ | 0.82 (0.69-0.98) | 0.83 (0.71-0.97) | - | - | 0.70 (0.59-0.83) | 0.71 (0.44-1.16) |

* Primary outcome.

† Nonfatal events only.

Note: This table presents relative risk reduction versus placebo and NOT absolute risk reduction. Statistically significant results are shown in bold type, with cells outlined when this was the primary outcome. ¹DAPA-HF Primary Outcome = hospitalization for heart failure (Hosp HF) or cardiovascular (CV) death. Hazard ratio (HR) 0.74* (0.64-0.99), p<0.05. ²Primary outcome = chronic kidney disease (CKD) progression or CV death. ACS, acute coronary syndrome; CHF, congestive heart failure; CI, confidence interval; CVD, CV disease; DM, diabetes mellitus; EF, ejection fraction; eGFR, estimated glomerular filtration rate in mL/minute/1.73m²; GLP1-RA, glucagon-like peptide-1 receptor agonists; HR, hazard ratio; MACE, major cardiovascular events (CV death, nonfatal myocardial infarction [MI], nonfatal stroke); MI, myocardial infarction; PVD, peripheral vascular disease; SGLT2i, sodium-glucose cotransporter 2 inhibitors.