Supplementary Table 2. Risk of bias of included randomized controlled trial studies

| Study | Selection bias (random sequence generation) | Selection bias (allocation concealment) | Performance bias | Detection bias | Attrition bias | Reporting bias | Other bias |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| MET |  |  |  |  |  |  |  |
| Anderson et al. (2017) [1] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Codner et al. (2013) [2] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Hamilton et al. (2003) [3] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Jacobsen et al. (2009) [4] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Khan et al. (2006) [16] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Libman et al. (2015) [7] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Lund et al. (2008) [5] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Meyer et al. (2002) [8] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Nadeau et al. (2015) [9] | Unclear risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Nwosu et al. (2015) [10] | Unclear risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Petrie et al. (2017) [11] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Pitocco et al. (2013) [13] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Sarnblad et al. (2003) [14] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Ziaee et al. (2017) [15] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| AGI |  |  |  |  |  |  |  |
| Dimitriadis et al. (1988) [19] | Unclear risk | Unclear risk | Low risk | Low risk | Unclear risk | Low risk | Unclear risk |
| Hollander et al. (1997) [17] | Unclear risk | Unclear risk | Low risk | Low risk | Unclear risk | Low risk | Unclear risk |
| Kennedy et al. (1987) [23] | Unclear risk | Unclear risk | Low risk | Low risk | Unclear risk | Low risk | Unclear risk |
| Marena et al. (1991) [20] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Mcculloch et al. (1983) [21] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Riccardi et al. (1999) [18] | Unclear risk | Unclear risk | Low risk | Low risk | Unclear risk | Low risk | Unclear risk |
| Serrano-Rios et al. (1988) [22] | Unclear risk | Unclear risk | Low risk | Low risk | Unclear risk | Unclear risk | Unclear risk |
| TZD |  |  |  |  |  |  |  |
| Bhat et al. (2007) [25] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Stone et al. (2008) [29] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Strowig et al. (2005) [26] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Tafuri et al. (2013) [27] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Zdravkovic et al. (2006) [28] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| GLP-1 RA |  |  |  |  |  |  |  |
| Ahren et al. (2016) [30] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Dejgaard et al. (2016) [31] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Dube et al. (2018) [37] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Frandsen et al. (2015) [33] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Kuhadiya et al. (2016) [35] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Mathieu et al. (2016) [36] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |

(Continued to the next page)

Supplementary Table 2. Continued

| Study | Selection bias (random sequence generation) | Selection bias (allocation concealment) | Performance bias | Detection bias | Attrition bias | Reporting bias | Other bias |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| DPP-4i |  |  |  |  |  |  |  |
| Ellis et al. (2011) [39] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Farngren et al. (2012) [40] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Foley et al. (2008) [41] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Garg et al. (2013) [38] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| George et al. (2016) [42] | Unclear risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Schopman et al. (2015) [43] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| SGLT-2i |  |  |  |  |  |  |  |
| Biester et al. (2017) [56] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Dandona et al. (2017) [45] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Famulla et al. (2017) [47] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Garg et al. (2017) [49] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Henry et al. (2015) [50] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Kuhadiya et al. (2016) [53] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Pieber et al. (2015) [48] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Sands et al. (2015) [54] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Shimada et al. (2018) [55] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Buse et al. (2018) [44] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Danne et al. (2018) [46] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Mathieu et al. (2018) [65] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Rosenstock et al. (2018) [66] | Low risk | Low risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Pramlintide |  |  |  |  |  |  |  |
| Edelman et al. (2006) [57] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Herrmann et al. (2013) [60] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Ratner et al. (2004) [61] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Ratner et al. (2005) [62] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Riddle et al. (2015) [64] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |
| Whitehouse et al. (2002) [63] | Unclear risk | Unclear risk | Low risk | Low risk | Low risk | Low risk | Unclear risk |

MET, metformin; AGI, alpha glucosidase inhibitor; TZD, thiazolidinedione; GLP-1RA, glucagon-like peptide-1 receptor agonist; DPP-4i, dipeptidyl peptide 4 inhibitor; SGLT-2i, sodium glucose cotransporter 2 inhibitor.

