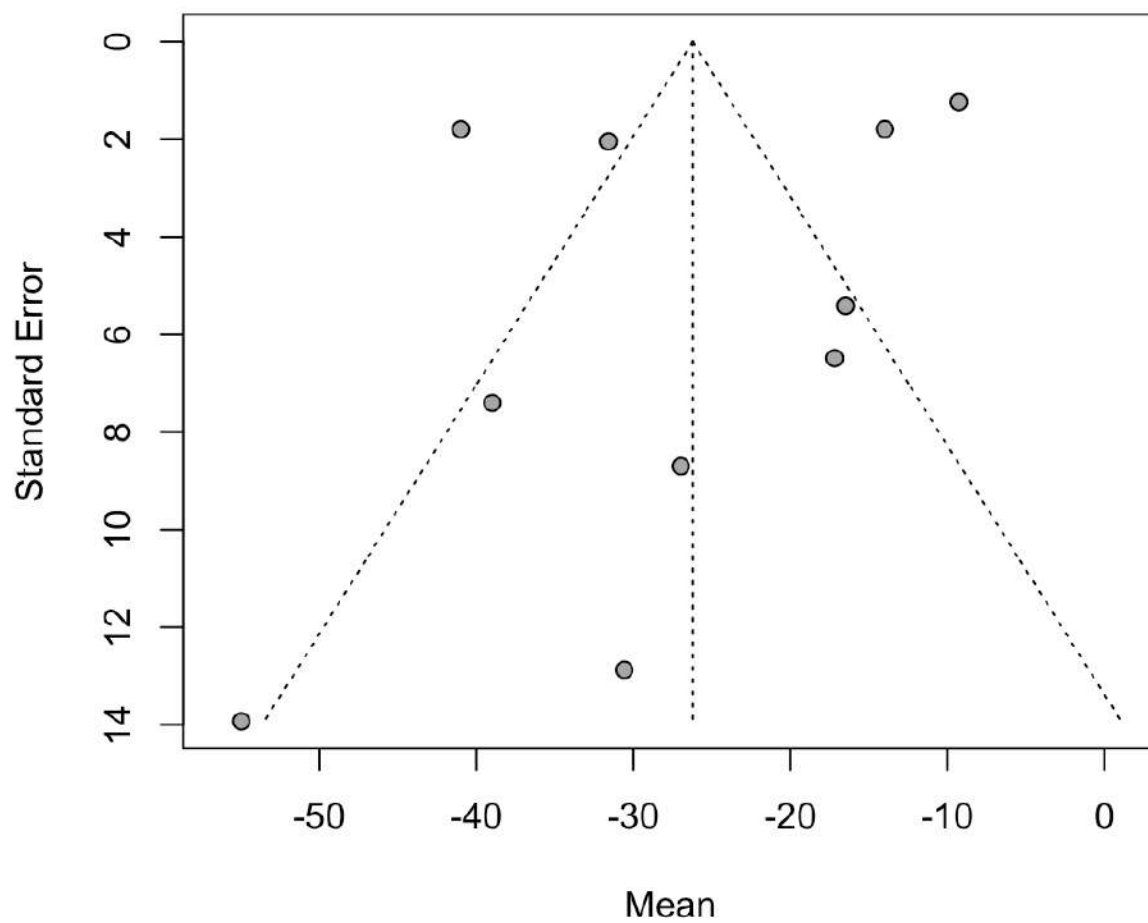
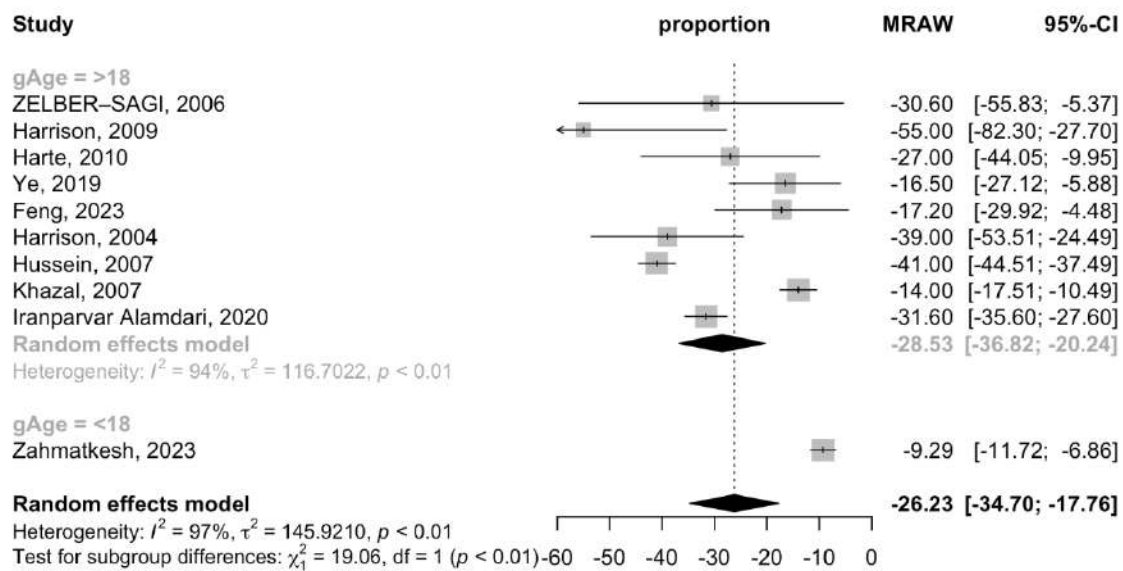


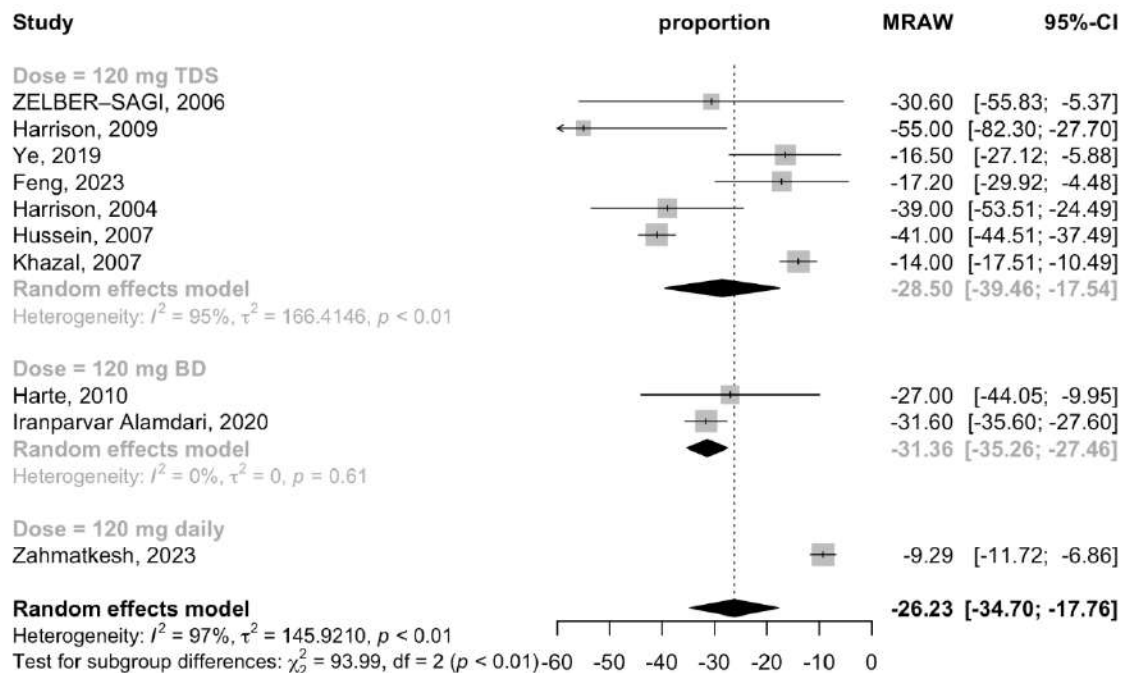
**Figure S1.** Forest plot describing the effect of orlistat administration on ALT levels (in RCTs)



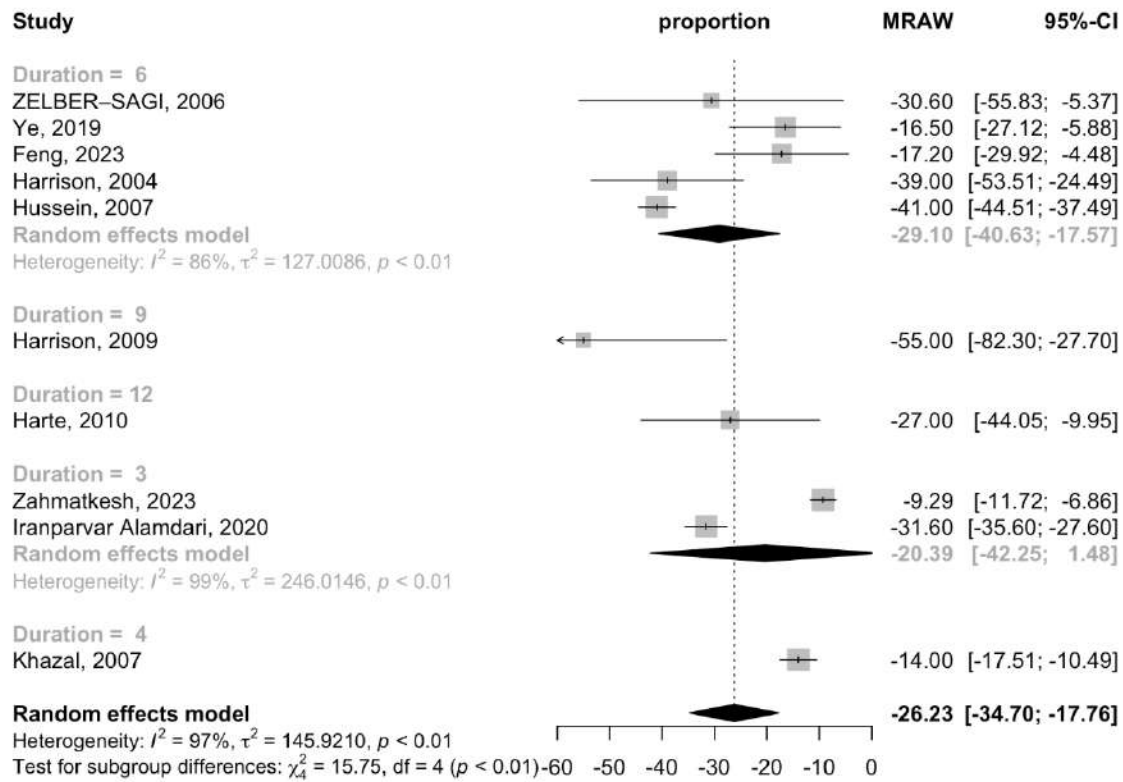
**Figure S2.** Funnel plot for the studies describing the effect of orlistat administration on ALT levels (in RCTs and single-arm trials)



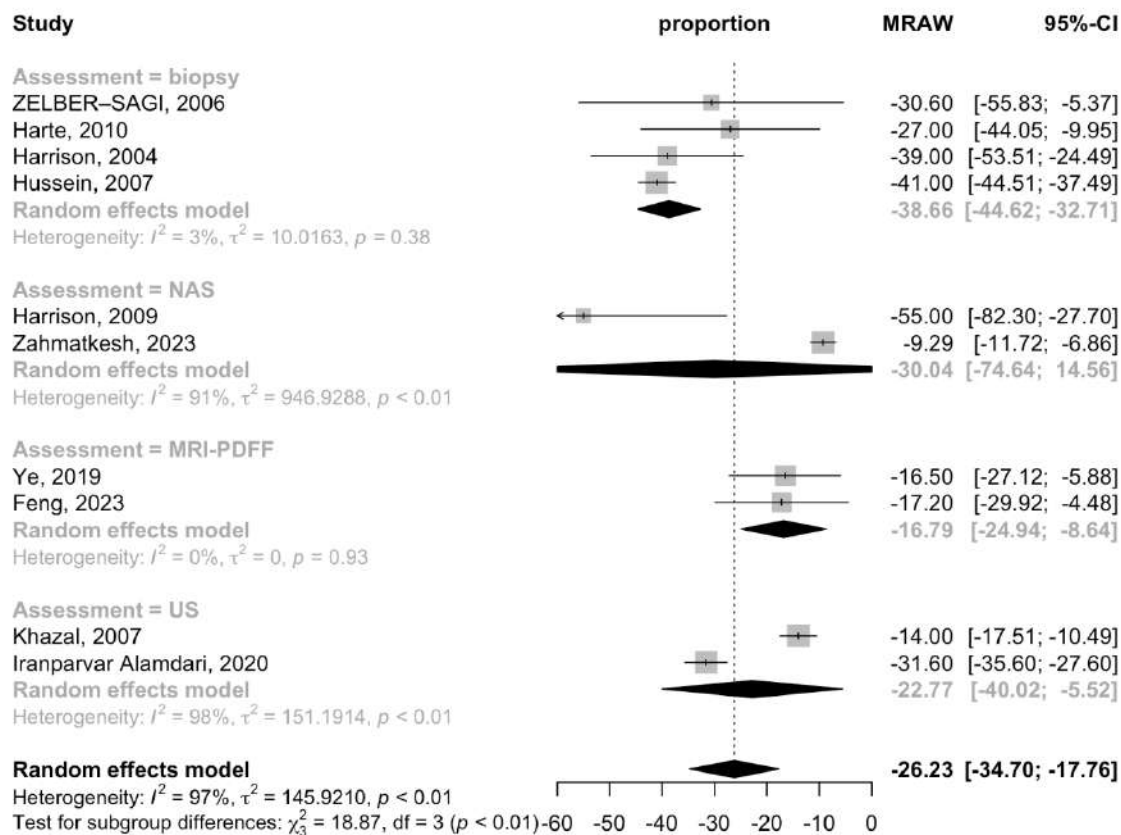
**Figure S3.** Forest plot describing the effect of orlistat administration on ALT levels (in RCTs and single-arm trials) (Subgroup analysis based on age of participants)



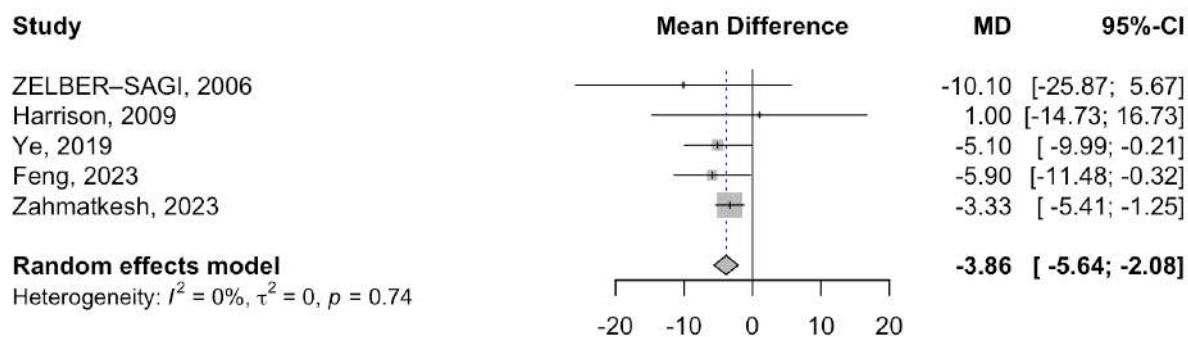
**Figure S4.** Forest plot describing the effect of orlistat administration on ALT levels (in RCTs and single-arm trials) (Subgroup analysis based on intake dose)



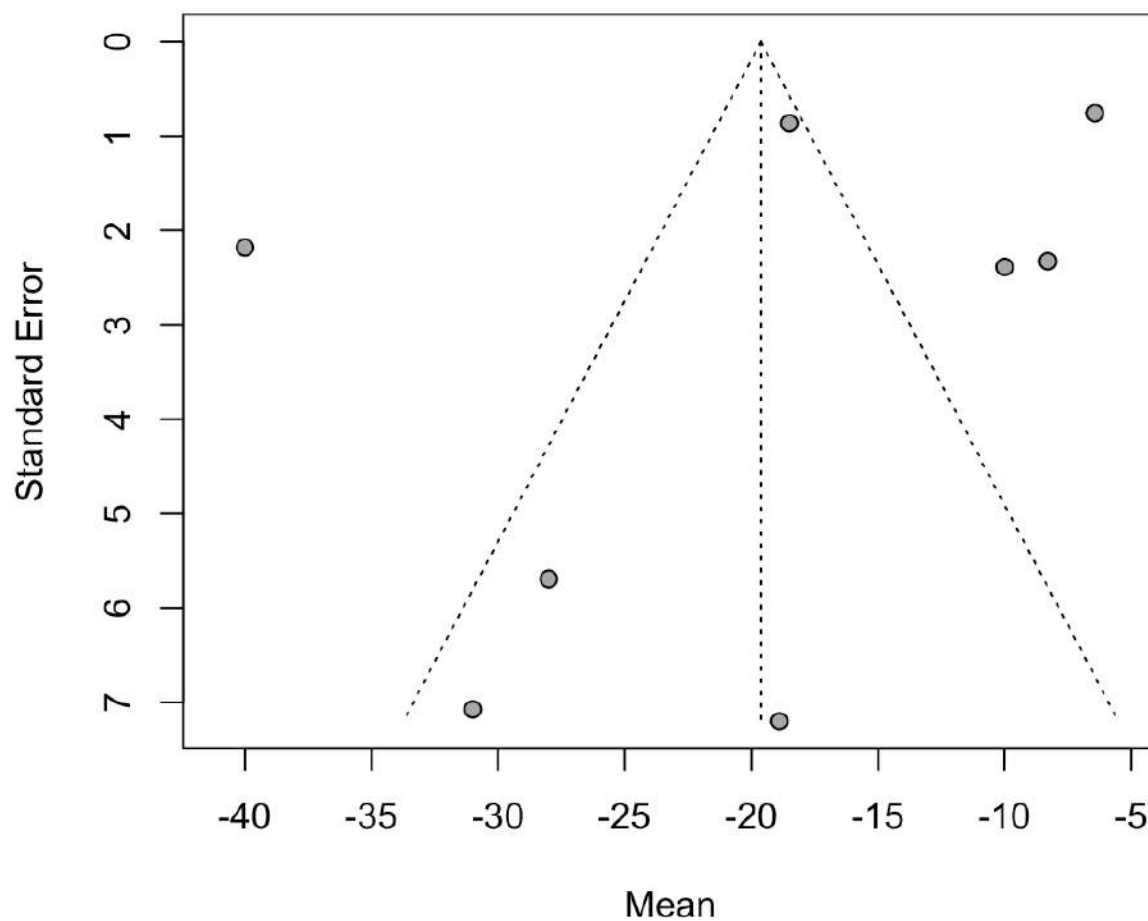
**Figure S5.** Forest plot describing the effect of orlistat administration on ALT levels (in RCTs and single-arm trials) (Subgroup analysis based on study duration)



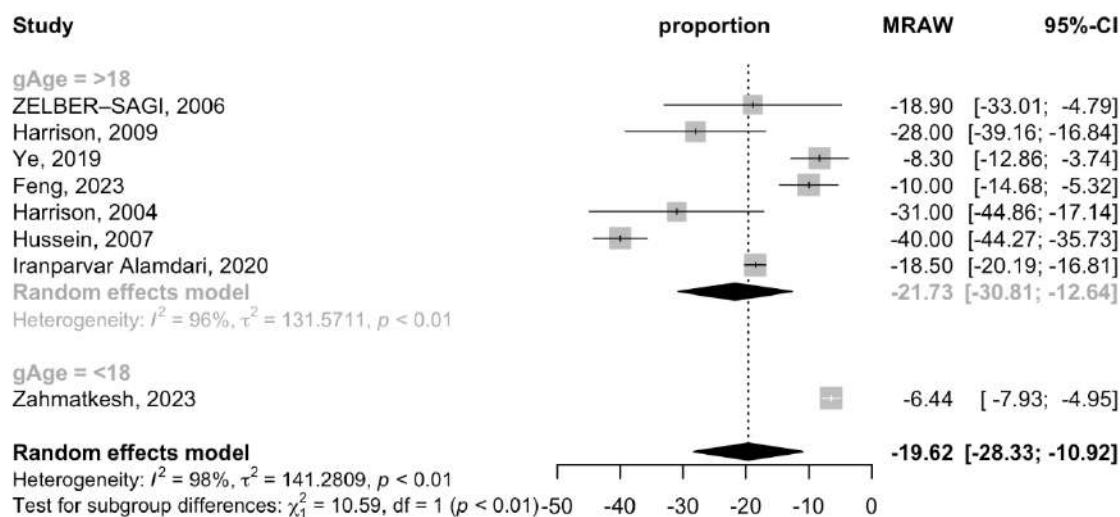
**Figure S6.** Forest plot describing the effect of orlistat administration on ALT levels (in RCTs and single-arm trials) (Subgroup analysis based on NAFLD detection method)



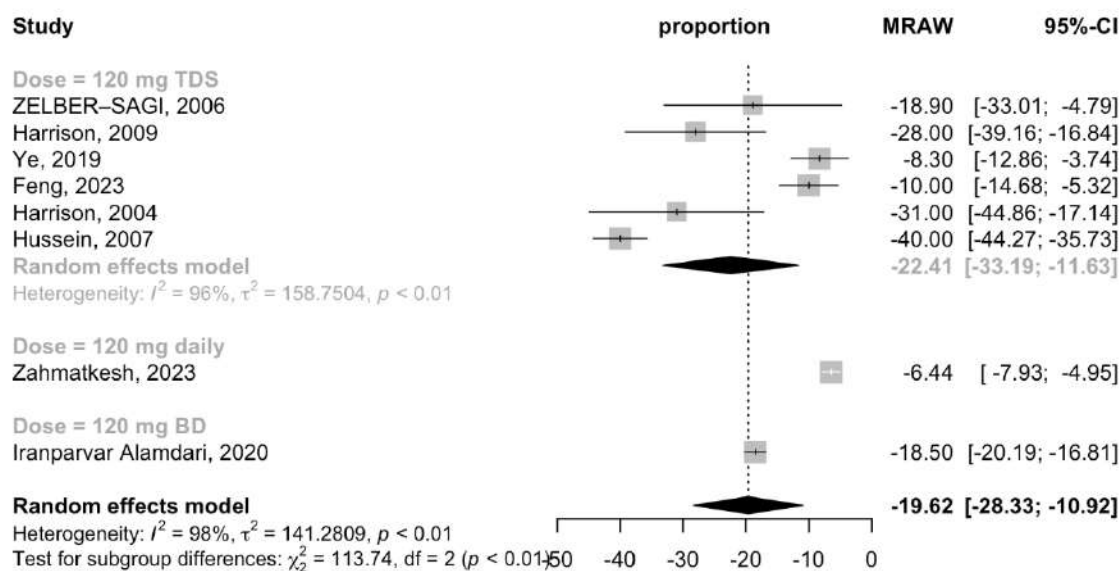
**Figure S7.** Forest plot describing the effect of orlistat administration on AST levels (in RCTs)



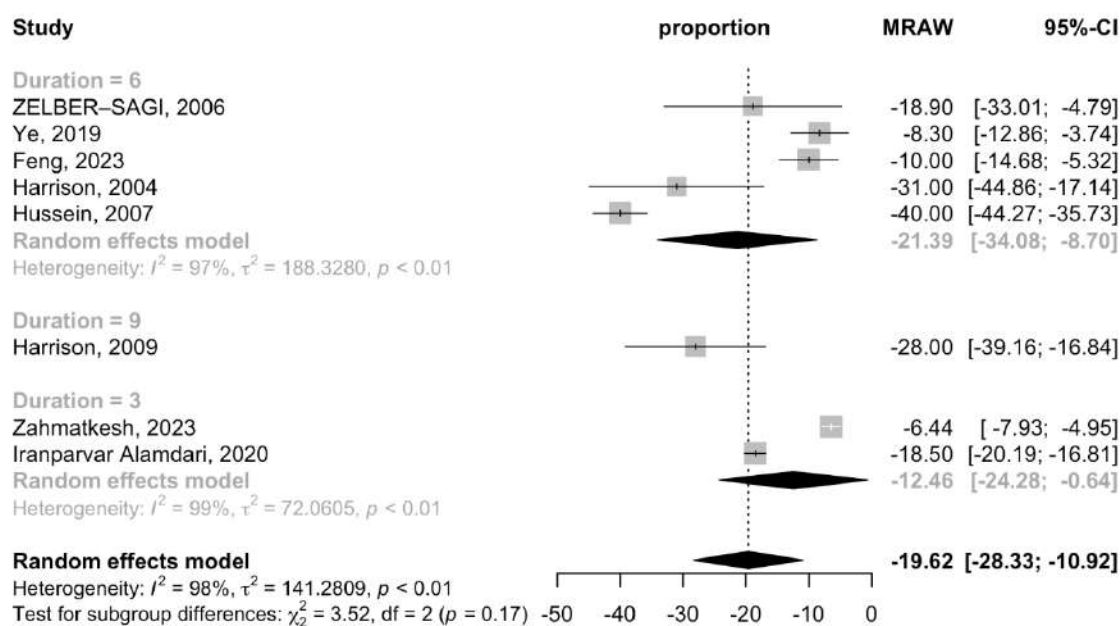
**Figure S8.** Funnel plot for the studies describing the effect of orlistat administration on AST levels (in RCTs and single-arm trials)



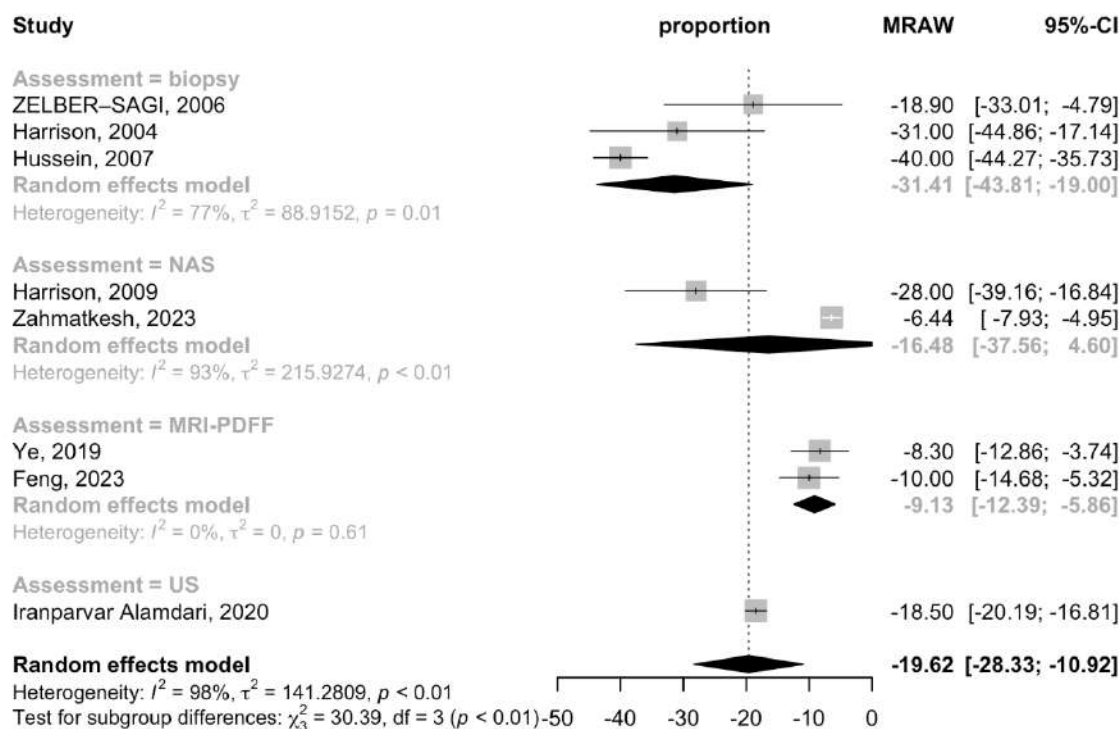
**Figure S9.** Forest plot describing the effect of orlistat administration on AST levels (in RCTs and single-arm trials) (Subgroup analysis based on age of participants)



**Figure S10.** Forest plot describing the effect of orlistat administration on AST levels (in RCTs and single-arm trials) (Subgroup analysis based on intake dose)

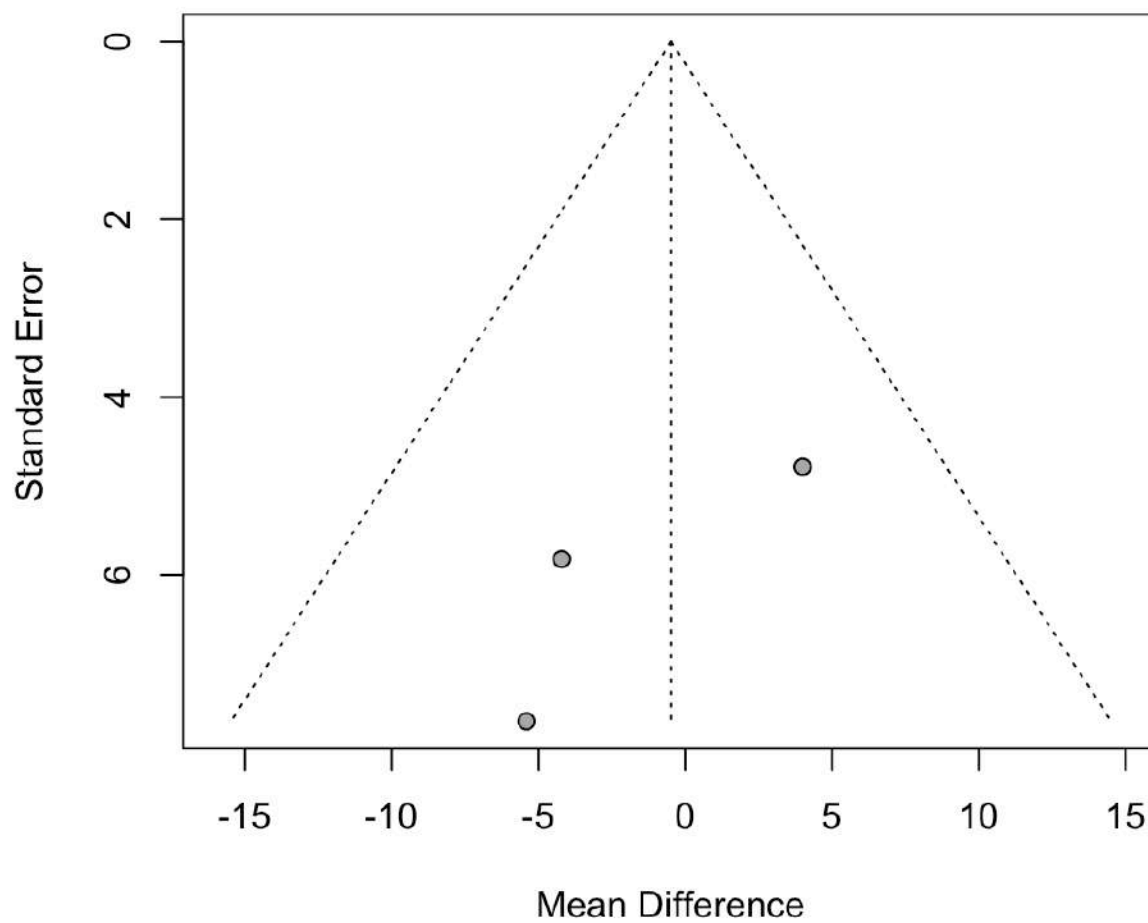


**Figure S11.** Forest plot describing the effect of orlistat administration on AST levels (in RCTs and single-arm trials) (Subgroup analysis based on study duration)

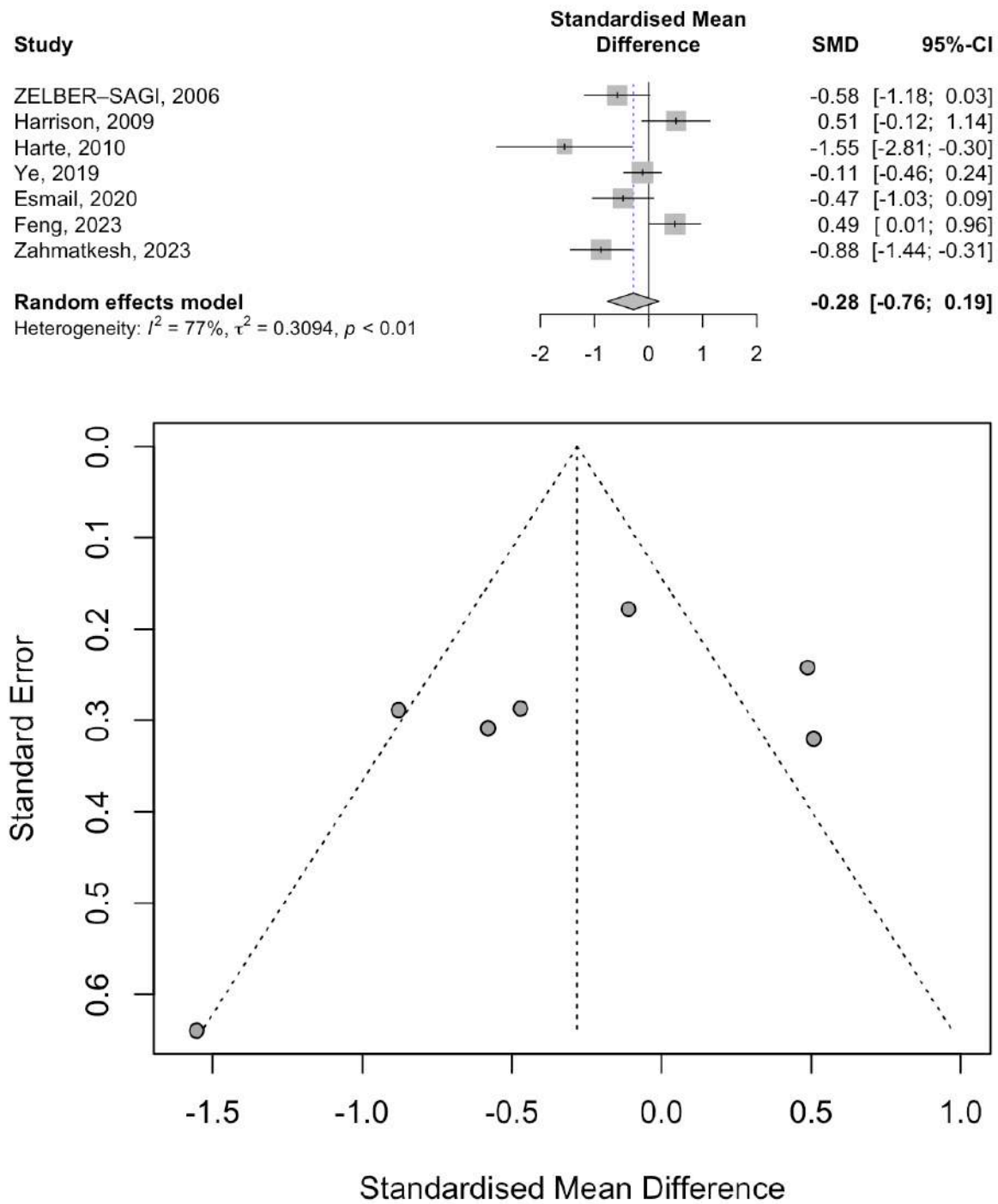


**Figure S12.** Forest plot describing the effect of orlistat administration on AST levels (in RCTs and single-arm trials) (Subgroup analysis based on NAFLD detection method)

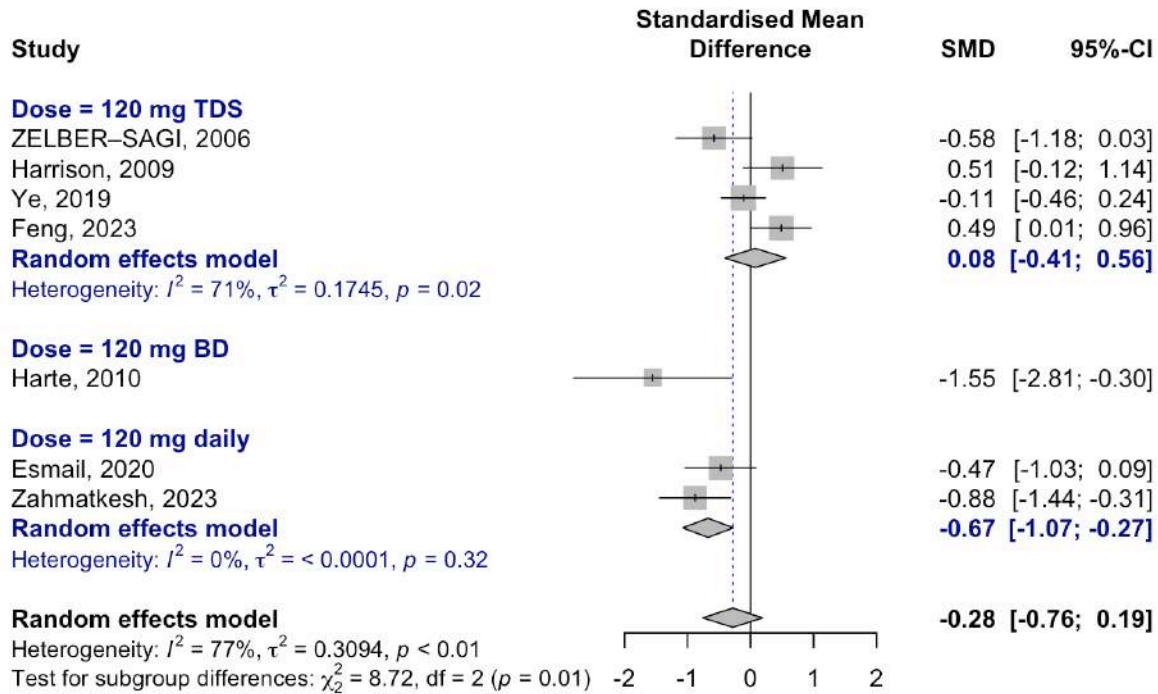




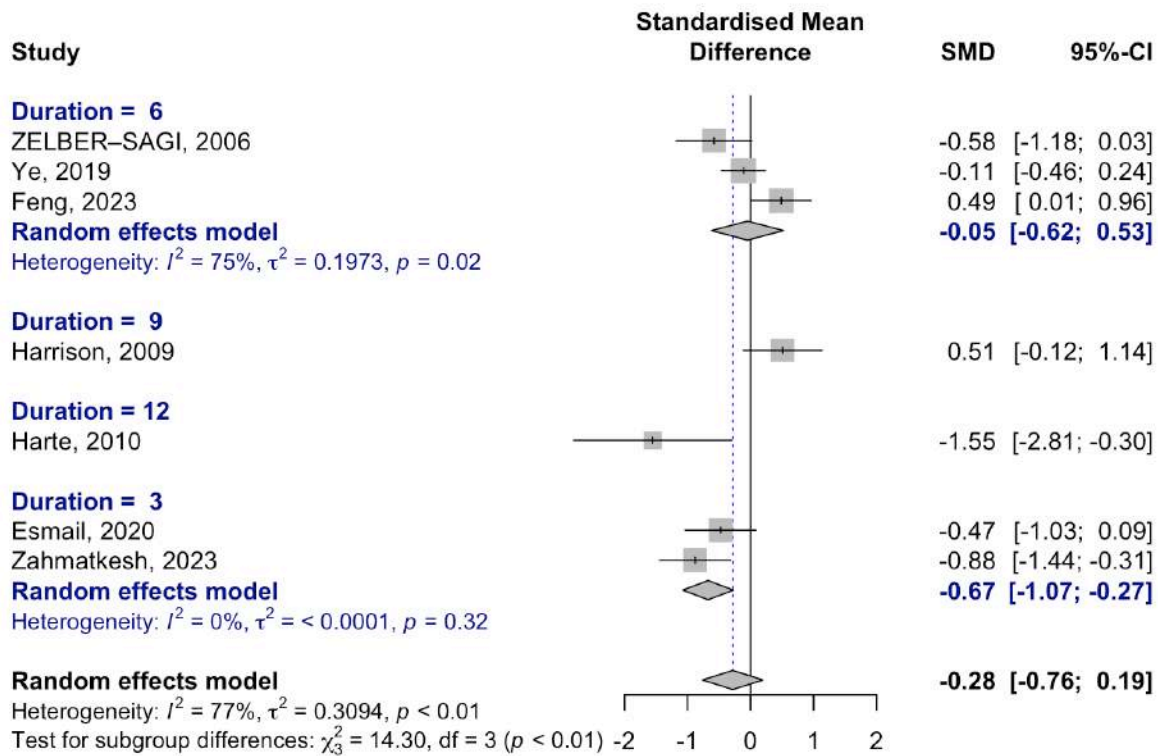
**Figure S13.** Funnel plot for the studies describing the effect of orlistat administration on ALP levels



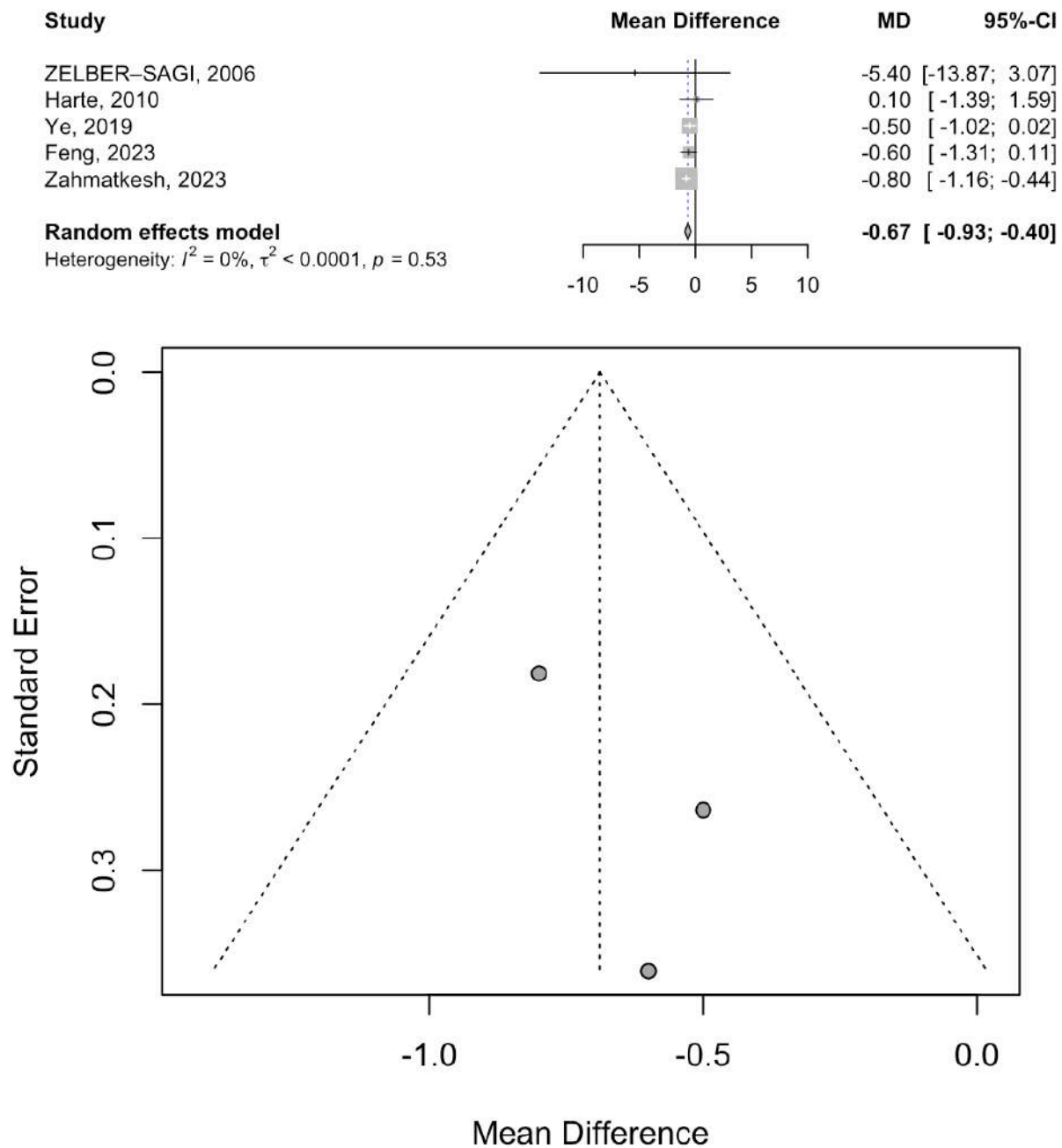
**Figure S14.** Forest and funnel plot for the studies describing the effect of orlistat administration on FBS



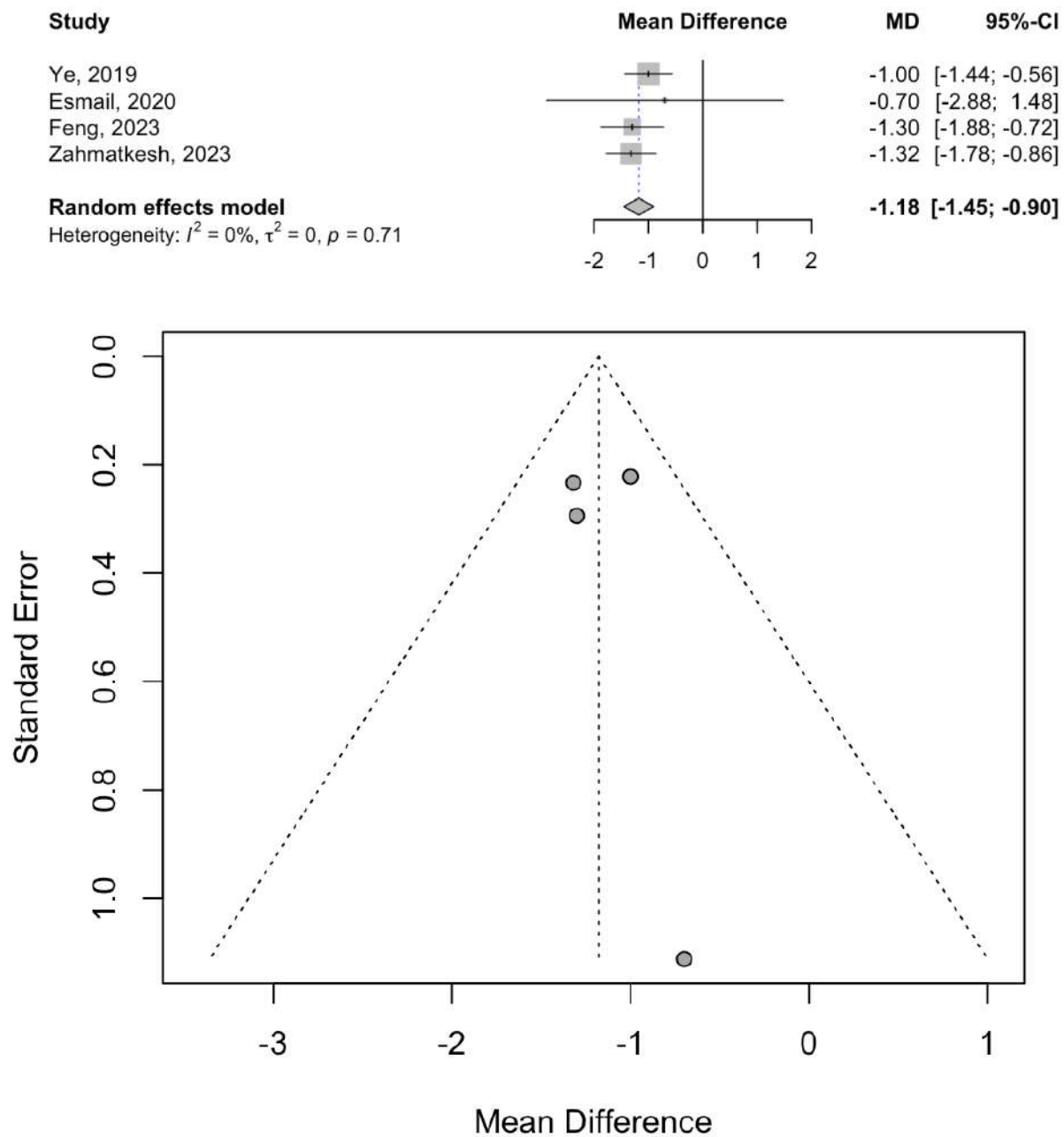
**Figure S15.** Forest plot for the studies describing the effect of orlistat administration on FBS (Subgroup analysis based on intake dose)



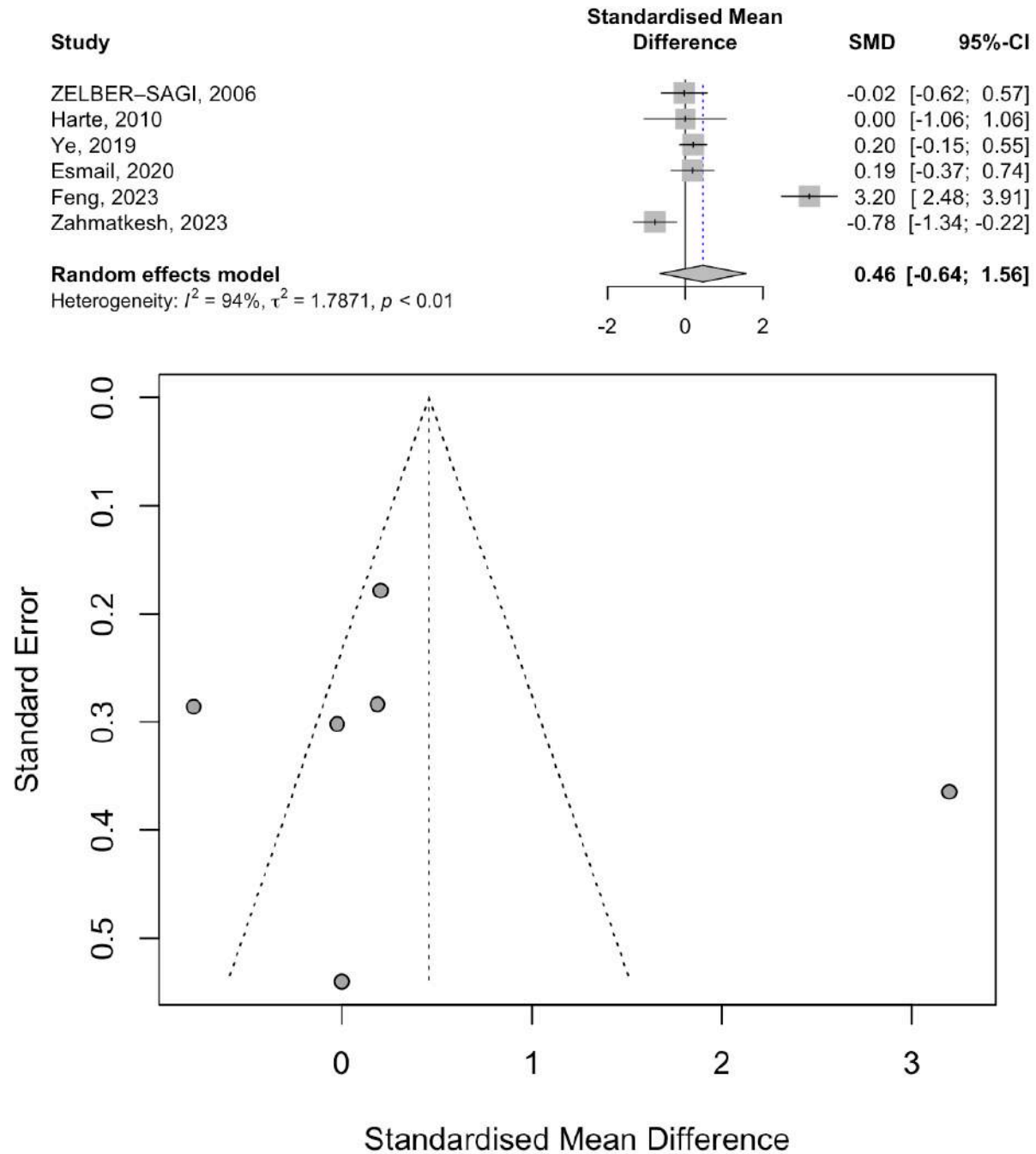
**Figure S16.** Forest plot for the studies describing the effect of orlistat administration on FBS (Subgroup analysis based on study duration)



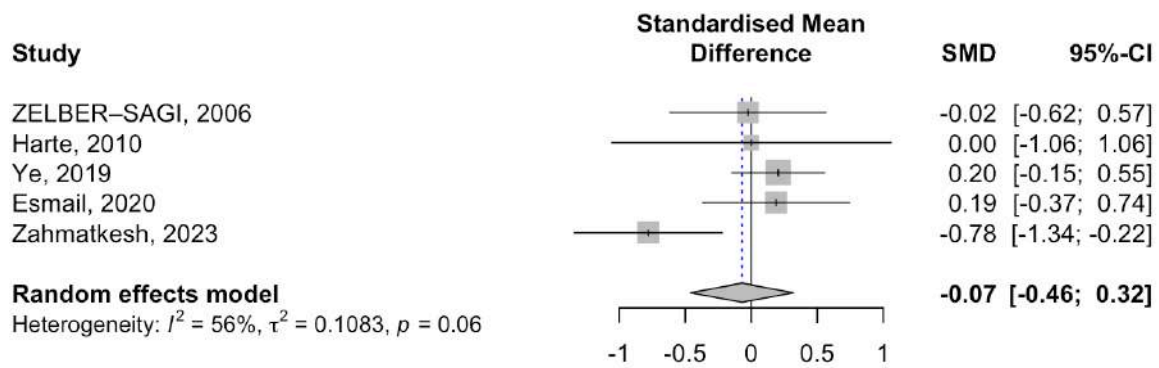
**Figure S17.** Forest and funnel plot for studies describing the effect of orlistat administration on HOMA



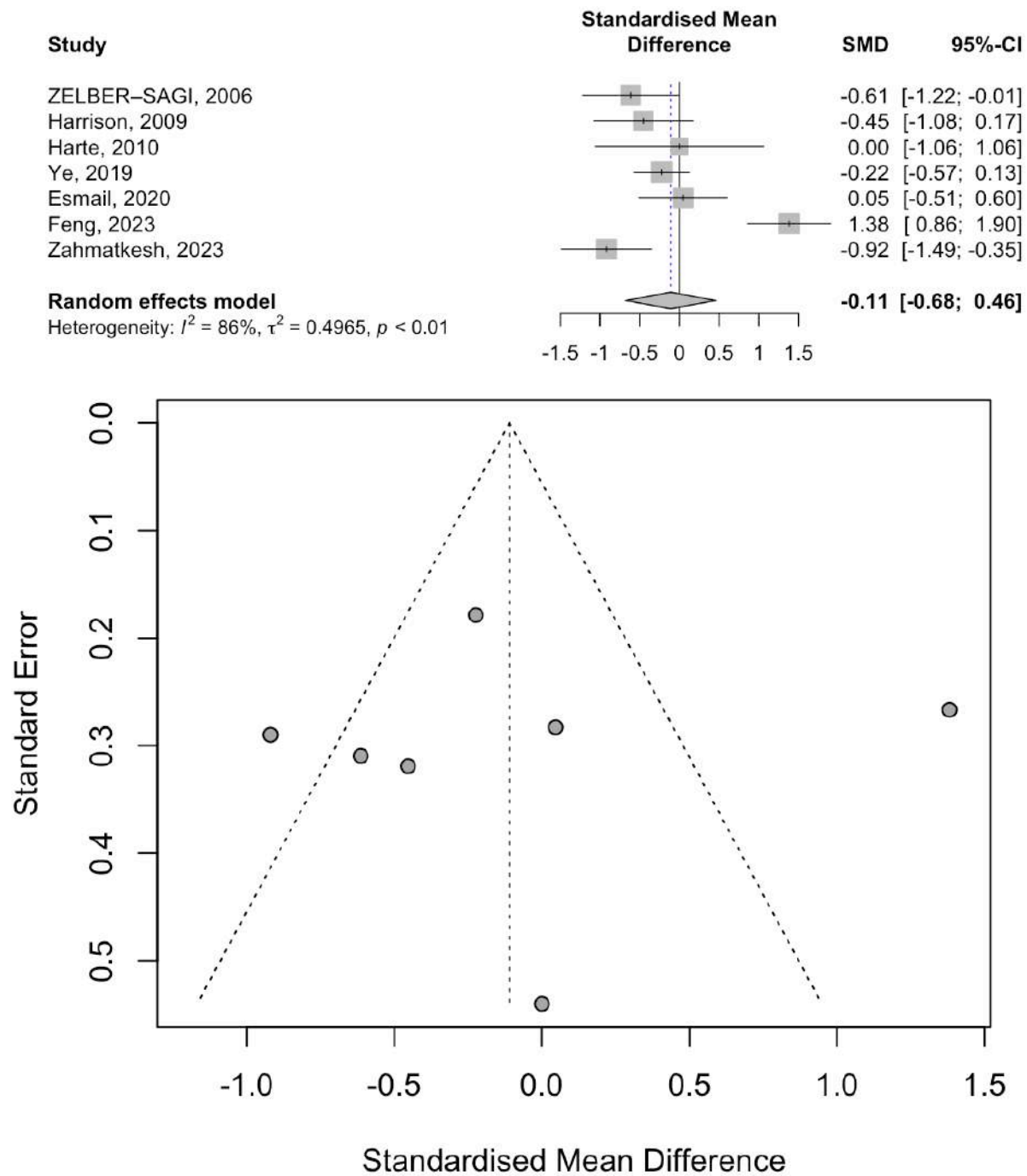
**Figure S18.** Forest and funnel plot for studies describing the effect of orlistat administration on BMI



**Figure S19.** Forest and funnel plot for studies describing the effect of orlistat administration on TG levels

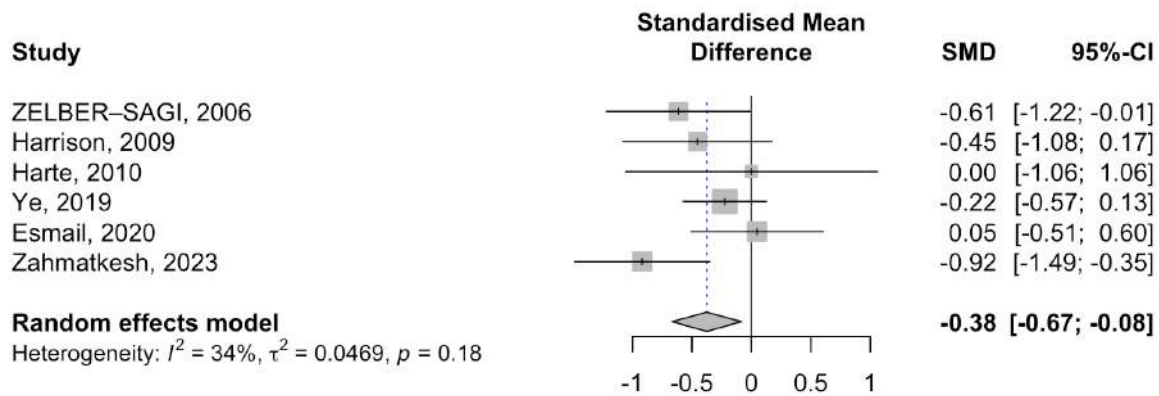


**Figure S20.** Forest plot for studies describing the effect of orlistat administration on TG levels (after removal of Feng's study)

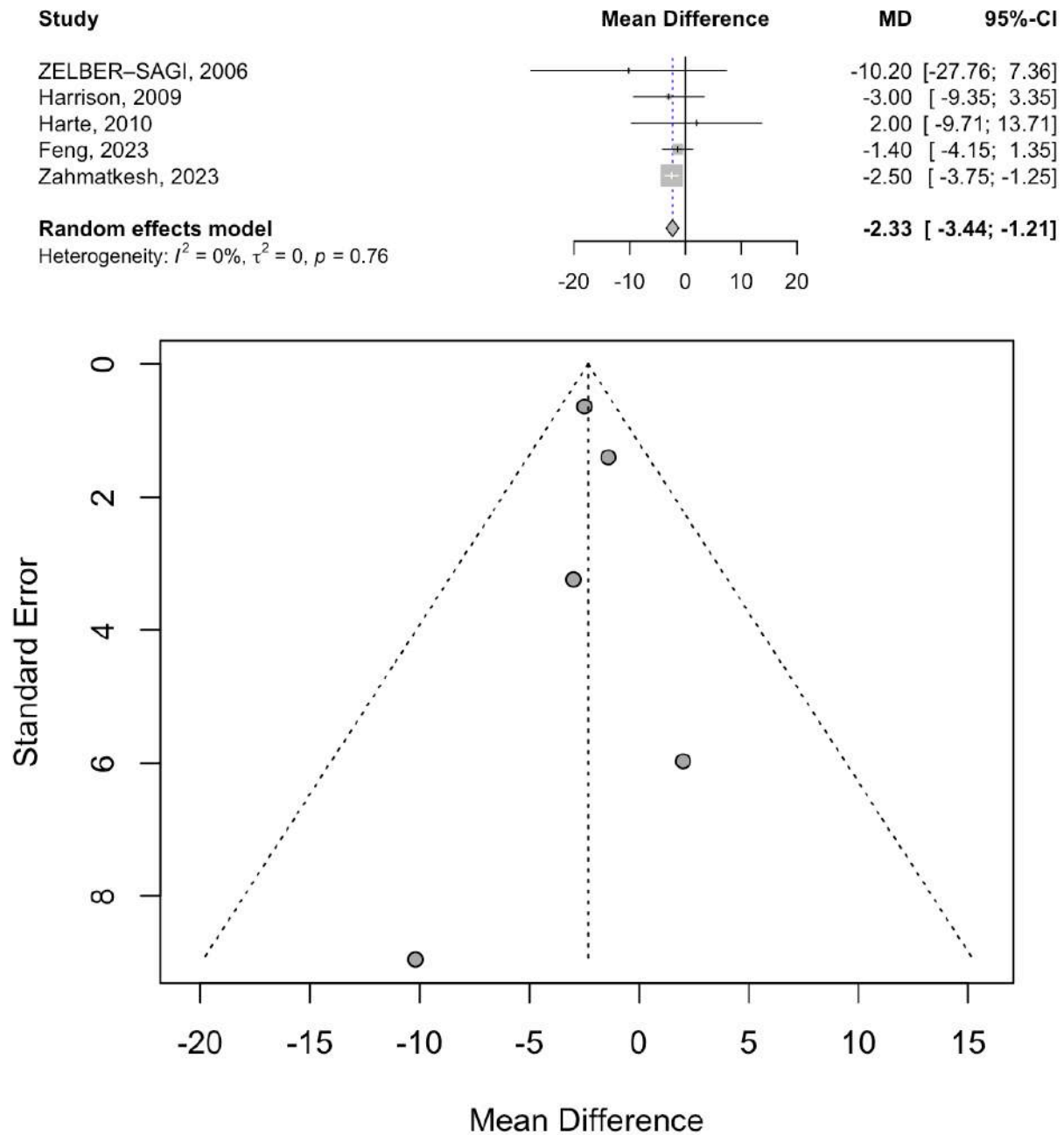


**Figure S21.** Forest and funnel plot for studies describing the effect of orlistat administration on cholesterol levels

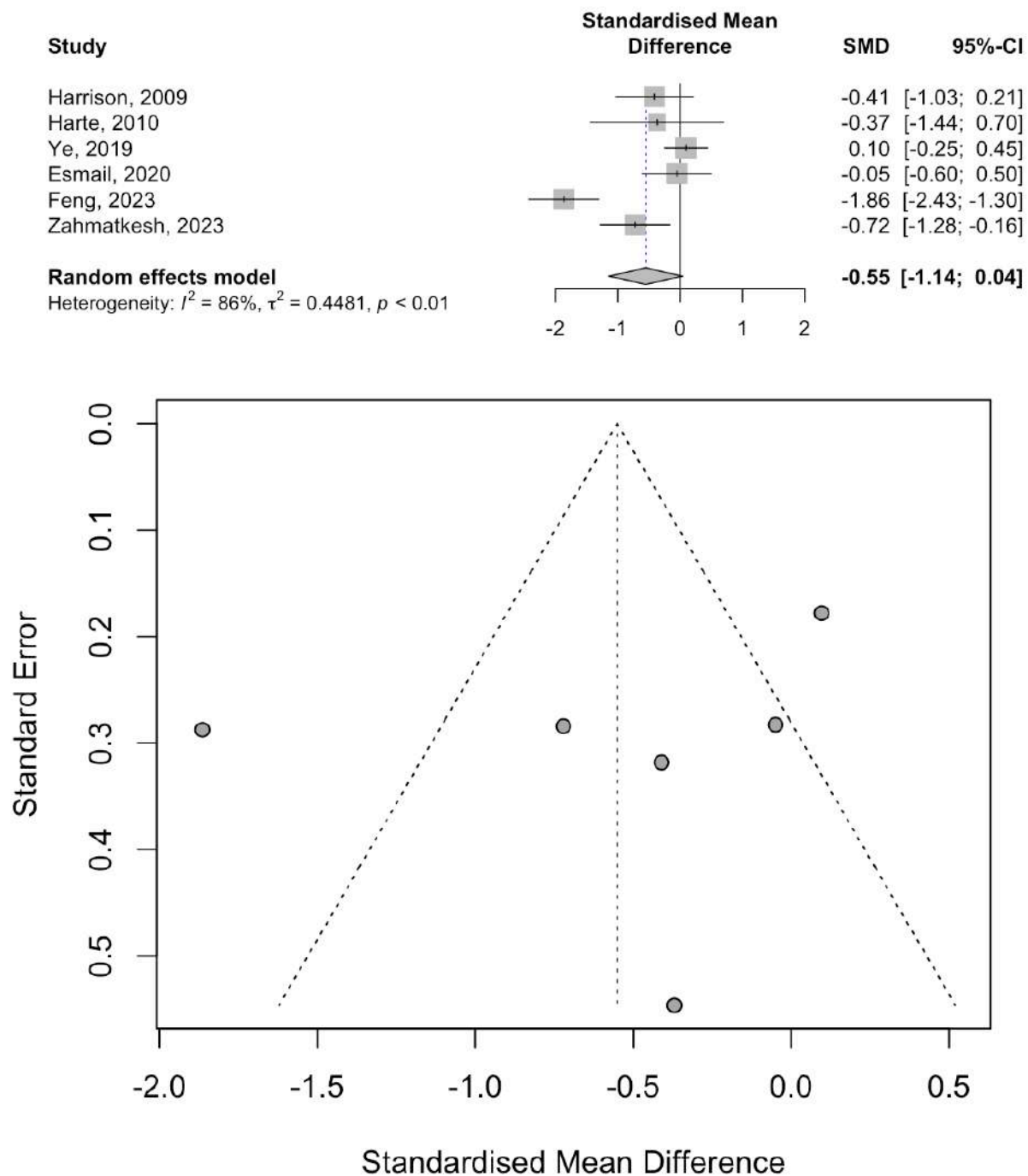




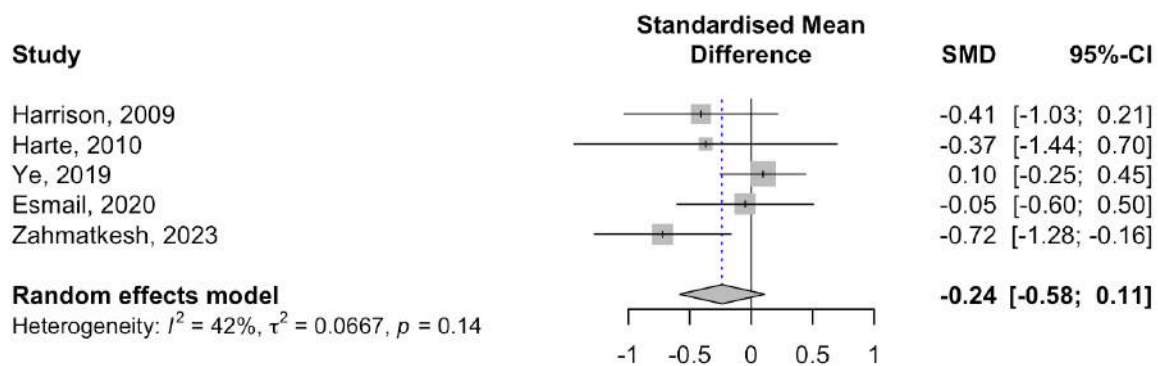
**Figure S22.** Forest plot for studies describing the effect of orlistat administration on cholesterol levels (after removal of Feng's study)



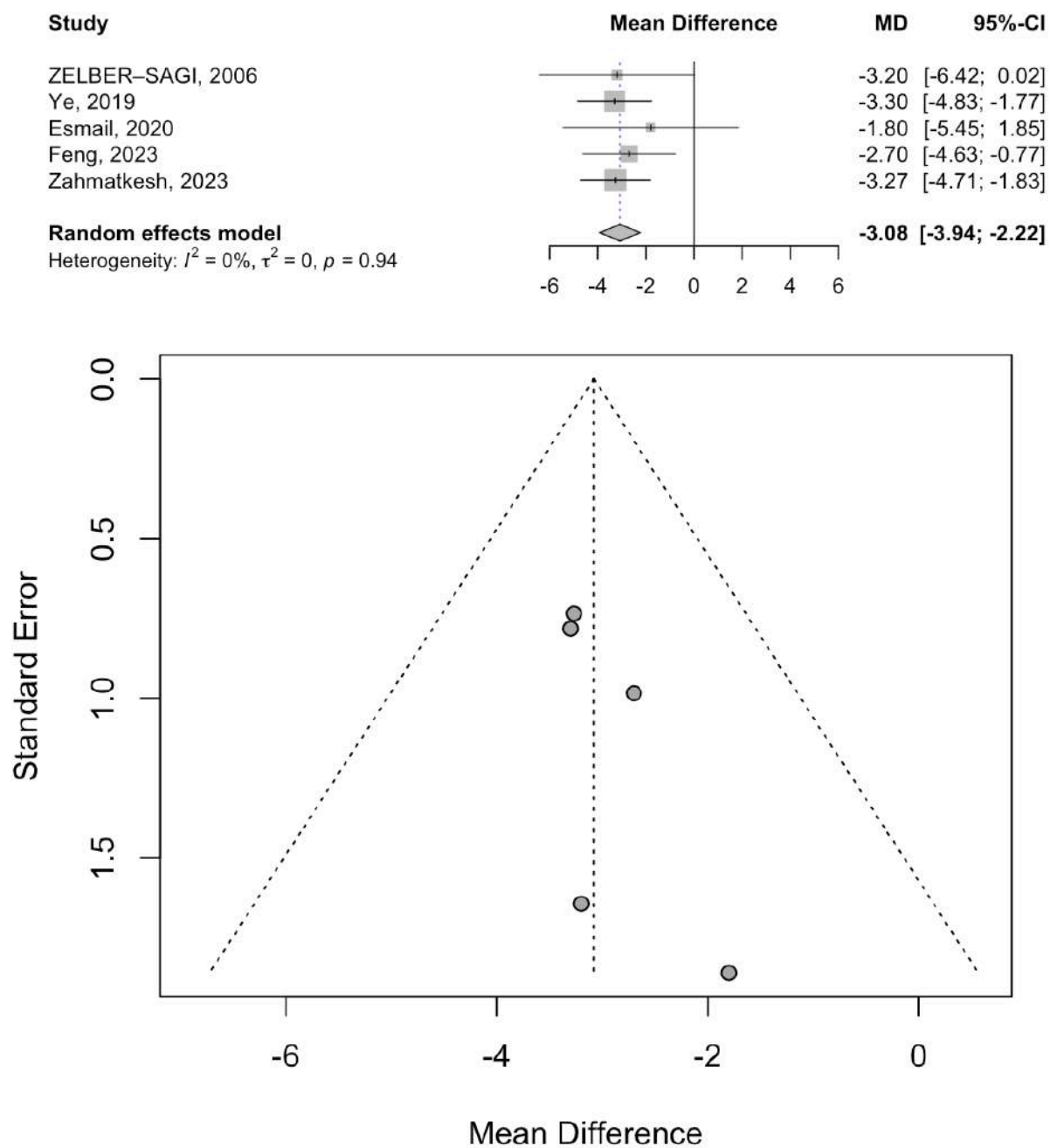
**Figure S23.** Forest and funnel plot for studies describing the effect of orlistat administration on insulin levels



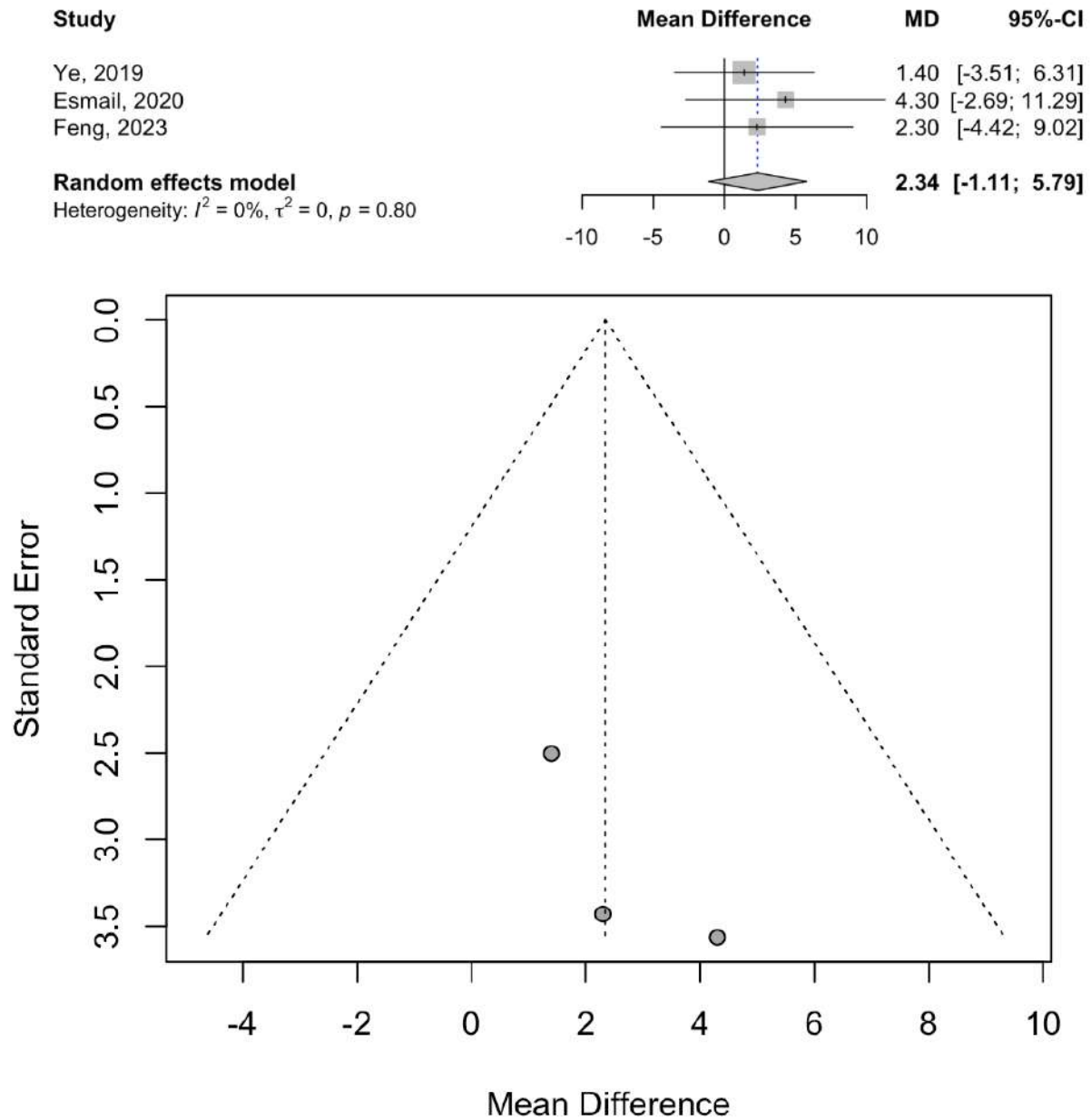
**Figure S24.** Forest and funnel plot for studies describing the effect of orlistat administration on LDL levels



**Figure S25.** Forest plot for studies describing the effect of orlistat administration on LDL levels (after removal of Feng's study)



**Figure S26.** Forest and funnel plot for studies describing the effect of orlistat administration on WC



**Figure S27.** Forest and funnel plot for studies describing the effect of orlistat administration on SBP

**Table S1.** JBI critical appraisal checklist for randomized controlled trials

<b>RCT studies included</b>	<b>1</b>	<b>2</b>	<b>3</b>	<b>4</b>	<b>5</b>	<b>6</b>	<b>7</b>	<b>8</b>	<b>9</b>	<b>10</b>	<b>11</b>	<b>12</b>	<b>13</b>
<b>Zahmatkesh et al.<sup>[15]</sup></b>	Yes	No	Yes	No	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
<b>Feng et al.<sup>[16]</sup></b>	Yes	Yes	Yes	Yes	Yes	Yes	n/a	Yes	Yes	Yes	Yes	Yes	Yes
<b>Wasta Esmail et al.<sup>[14]</sup></b>	Yes	No	Yes	No	No	Yes	n/a	Yes	Yes	Yes	Yes	Yes	Yes
<b>Ye et al.<sup>[17]</sup></b>	Yes	No	Yes	No	No	Yes	n/a	Yes	Yes	Yes	Yes	Yes	Yes
<b>Harte et al.<sup>[18]</sup></b>	No	Yes	Yes	Yes	Yes	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes
<b>Harrison et al.<sup>[19]</sup></b>	Yes	No	Yes	No	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
<b>Zelber–Sagi et al.<sup>[20]</sup></b>	Yes	Yes	Yes	Yes	Yes	Yes	n/a	Yes	Yes	Yes	Yes	Yes	Yes

- 1- Was true randomization used for assignment of participants to treatment groups?
- 2- Was allocation to treatment groups concealed?
- 3- Were treatment groups similar at the baseline?
- 4- Were participants blind to treatment assignment?
- 5- Were those delivering the treatment blind to treatment assignment?
- 6- Were treatment groups treated identically other than the intervention of interest?
- 7- Were outcome assessors blind to treatment assignment?
- 8- Were outcomes measured in the same way for treatment groups?
- 9- Were outcomes measured in a reliable way?
- 10- Was follow up complete and if not, were differences between groups in terms of their follow up adequately described and analyzed?
- 11- Were participants analyzed in the groups to which they were randomized?
- 12- Was appropriate statistical analysis used?
- 13- Was the trial design appropriate and any deviations from the standard RCT design (individual randomization, parallel groups) accounted for in the conduct and analysis of the trial?

**Table S2.** JBI critical appraisal checklist for quasi-experimental studies

<b>Single-arm trials included</b>	<b>1</b>	<b>2</b>	<b>3</b>	<b>4</b>	<b>5</b>	<b>6</b>	<b>7</b>	<b>8</b>	<b>9</b>
<b>Iranparvar Alamdari et al.<sup>[21]</sup></b>	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes
<b>Khazal et al.<sup>[13]</sup></b>	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes
<b>Hussein et al.<sup>[22]</sup></b>	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes
<b>Harrison et al.<sup>[23]</sup></b>	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	Yes

- 1- Is it clear in the study what is the “cause” and what is the “effect” (i.e. there is no confusion about which variable comes first)?
- 2- Was there a control group?
- 3- Were participants included in any comparisons similar?
- 4- Were the participants included in any comparisons receiving similar treatment/care, other than the exposure or intervention of interest?
- 5- Were there multiple measurements of the outcome, both pre and post the intervention/exposure?
- 6- Were the outcomes of participants included in any comparisons measured in the same way?
- 7- Were outcomes measured in a reliable way?
- 8- Was follow-up complete and if not, were differences between groups in terms of their follow-up adequately described and analyzed?
- 9- Was appropriate statistical analysis used?



**Table S3.** Sensitivity analysis for studies assessing TG levels

Study	Estimate	CI_lb	CI_ub	p_value	Tau2	I2
ZELBER–SAGI	0.5589	-0.77799	1.895794	0.41257	2.206851	96.29354
Harte	0.544935	-0.77555	1.86542	0.418611	2.187304	96.79489
Ye	0.514243	-0.84422	1.872707	0.458124	2.269922	95.36896
Esmail	0.516276	-0.83592	1.868477	0.454266	2.258144	96.26787
Feng	-0.07074	-0.46142	0.319943	0.722672	0.108736	57.7949
Zahmatkesh	0.712844	-0.50195	1.927641	0.2501	1.799965	95.37402

**Table S4.** Sensitivity analysis for studies assessing cholesterol levels

Study	Estimate	CI_lb	CI_ub	p_value	Tau2	I2
ZELBER–SAGI	-0.02513	-0.672	0.621734	0.939294	0.55273	87.65158
Harrison	-0.05281	-0.71398	0.608357	0.875598	0.582644	88.31157
Harte	-0.12379	-0.77349	0.525905	0.708812	0.582994	89.31295
Ye	-0.08874	-0.77257	0.595082	0.799222	0.616837	86.267
Esmail	-0.13712	-0.81169	0.537461	0.690345	0.606665	88.31534
Feng	-0.37592	-0.66872	-0.08311	0.01186	0.047308	36.49487
Zahmatkesh	0.030263	-0.56862	0.62915	0.921105	0.458542	85.20076

**Table S5.** Sensitivity analysis for studies assessing LDL levels

Study	Estimate	CI_lb	CI_ub	p_value	Tau2	I2
Harrison	-0.58087	-1.29994	0.138199	0.113358	0.569794	88.06166
Harte	-0.57903	-1.26635	0.108283	0.098702	0.540852	88.84267
Ye	-0.70468	-1.35436	-0.055	0.033513	0.431989	80.88303
Esmail	-0.65704	-1.34244	0.028372	0.060268	0.504801	86.14197
Feng	-0.23803	-0.58207	0.106012	0.175092	0.066837	45.24189
Zahmatkesh	-0.51718	-1.23872	0.204365	0.16007	0.570103	87.54376

**Table S6.** Sensitivity analysis for studies assessing FBS levels

Study	Estimate	CI_lb	CI_ub	p_value	Tau2	I2
ZELBER–SAGI	-0.24824	-0.80909	0.312607	0.38566	0.38823	83.57852
Harrison	-0.41123	-0.89844	0.075982	0.098066	0.272885	78.34587
Harte	-0.16758	-0.61696	0.281803	0.46485	0.24191	78.16519
Ye	-0.33762	-0.92707	0.251819	0.261591	0.42588	81.5384
Esmail	-0.26809	-0.84157	0.305382	0.359529	0.407805	83.91835
Feng	-0.42207	-0.89049	0.046342	0.077386	0.238131	73.96895
Zahmatkesh	-0.17993	-0.6824	0.322533	0.482762	0.292321	78.92562