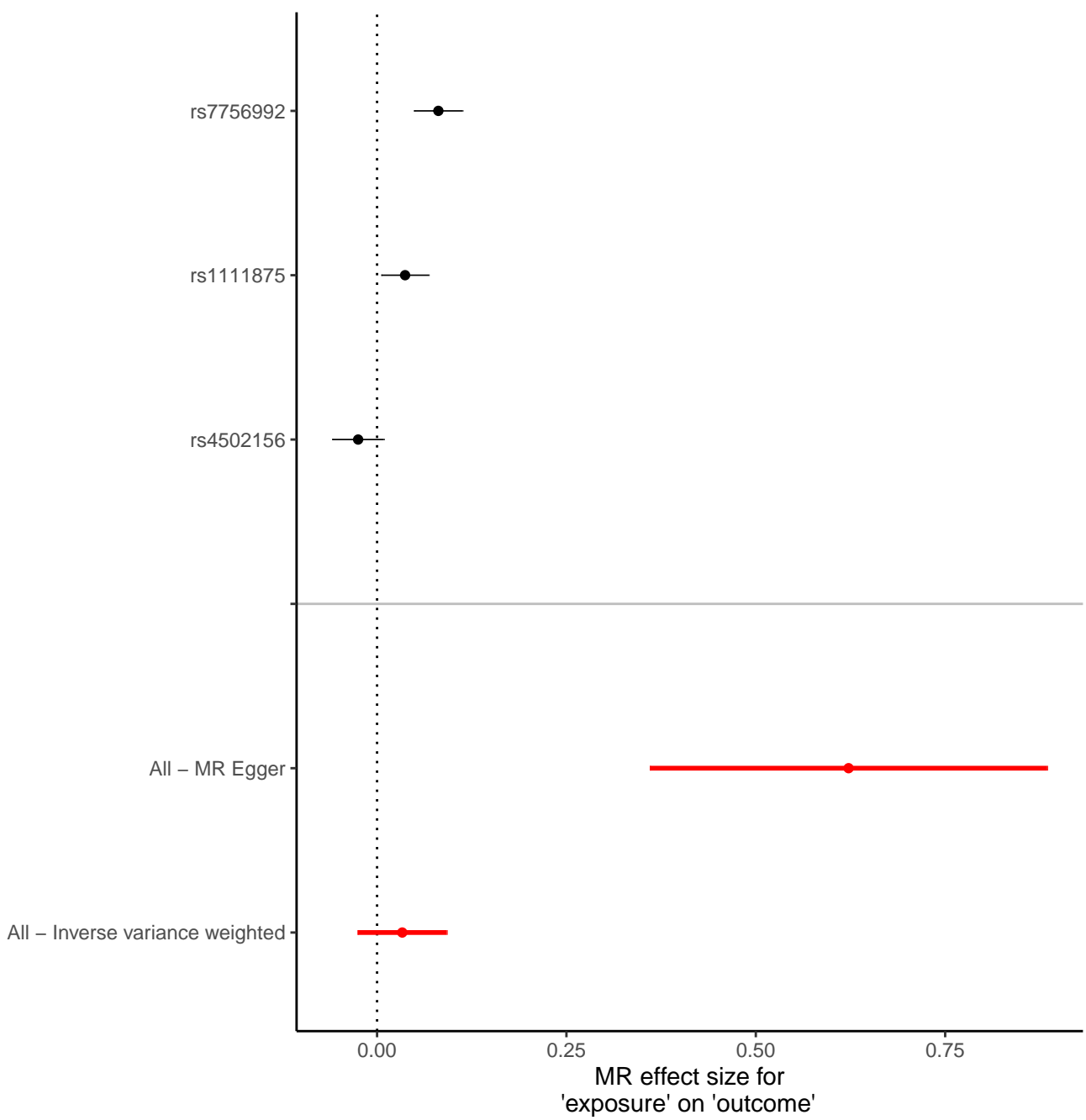


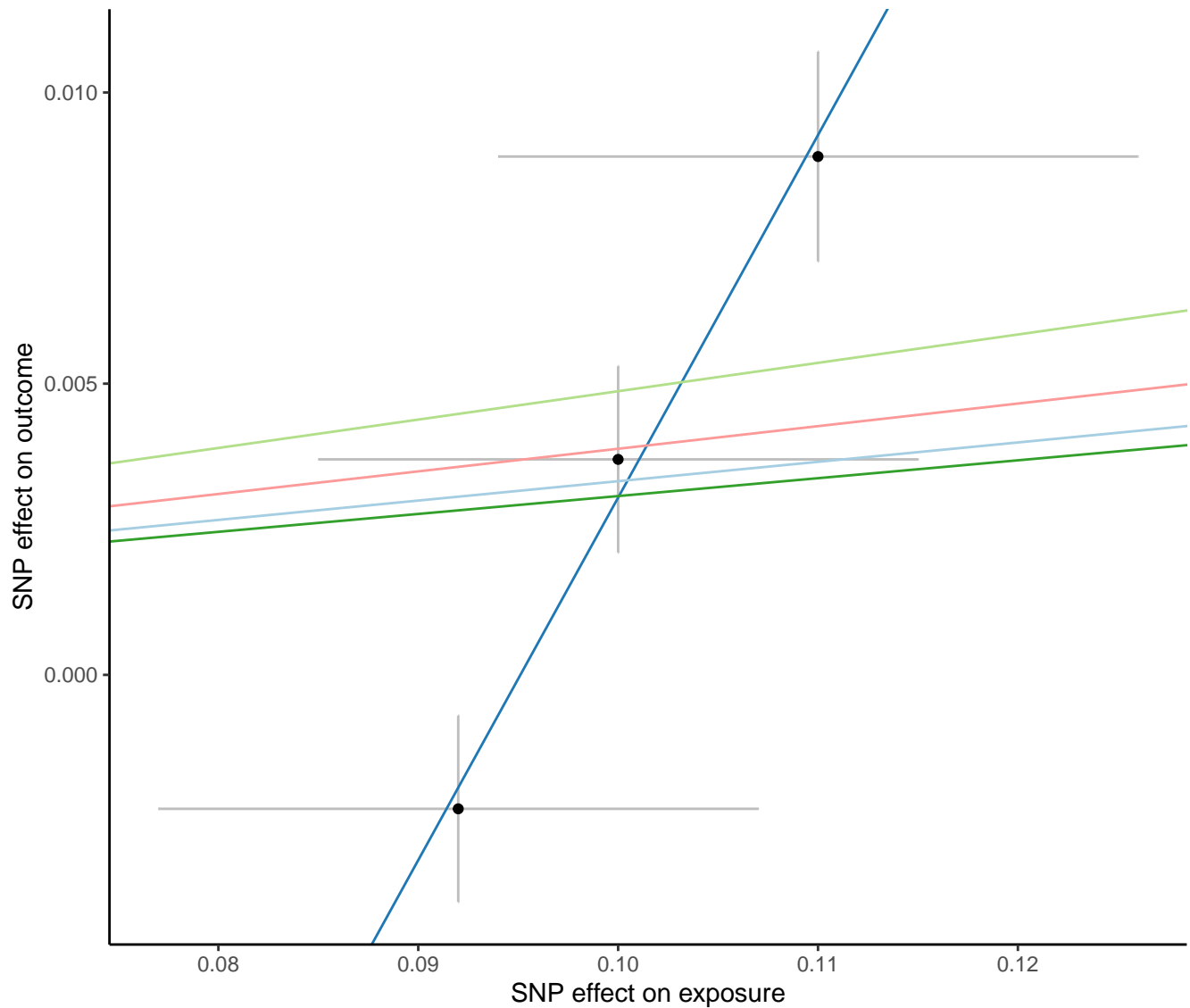
Supplementary File 3 — Exposure: CIRadjISI, Outcome: Anthropometric and Metabolic Traits

**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Body Mass Index**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**



MR Test

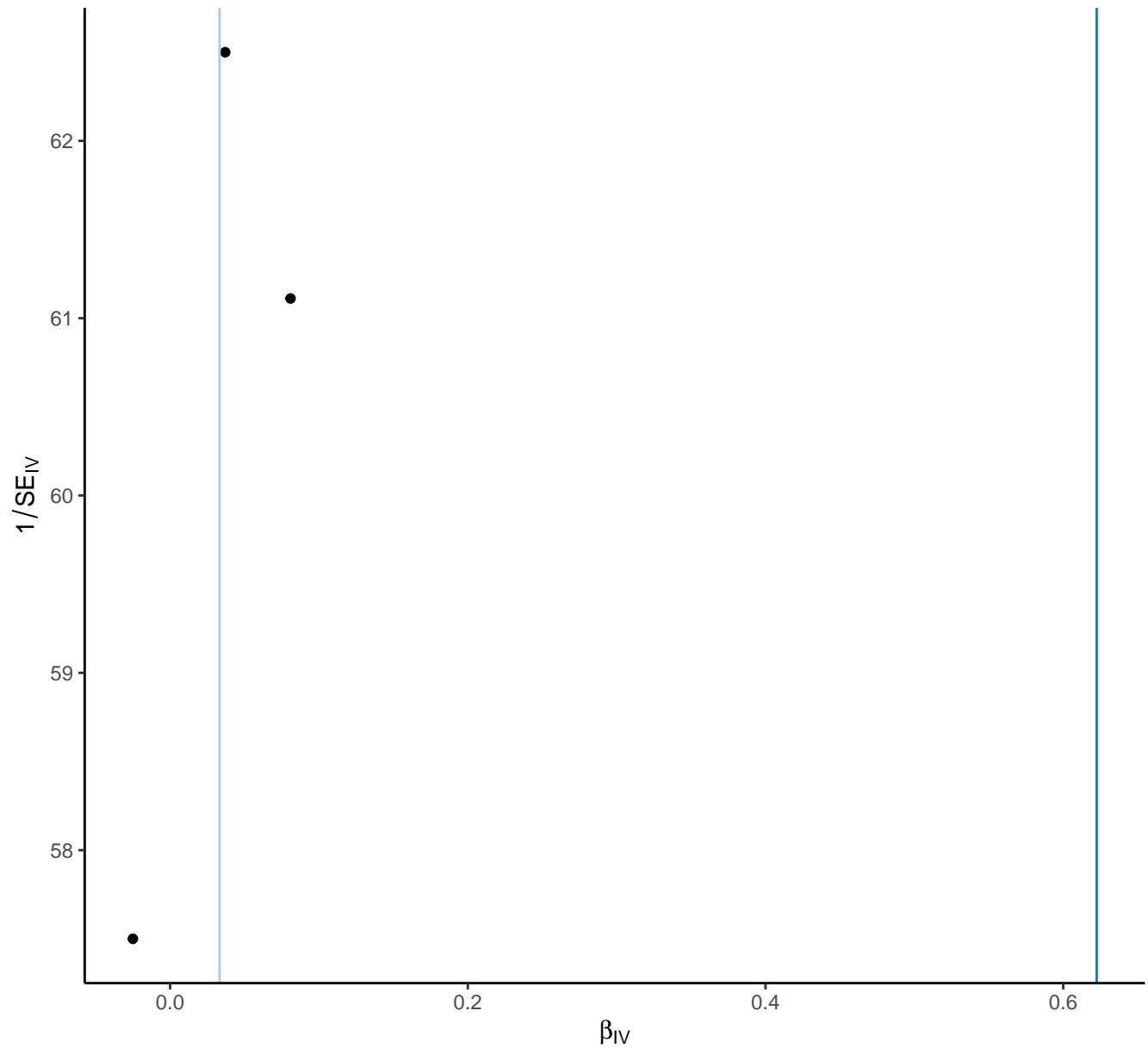
- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode



MR Method

Inverse variance weighted

MR Egger



rs4502156

rs1111875

rs7756992

All

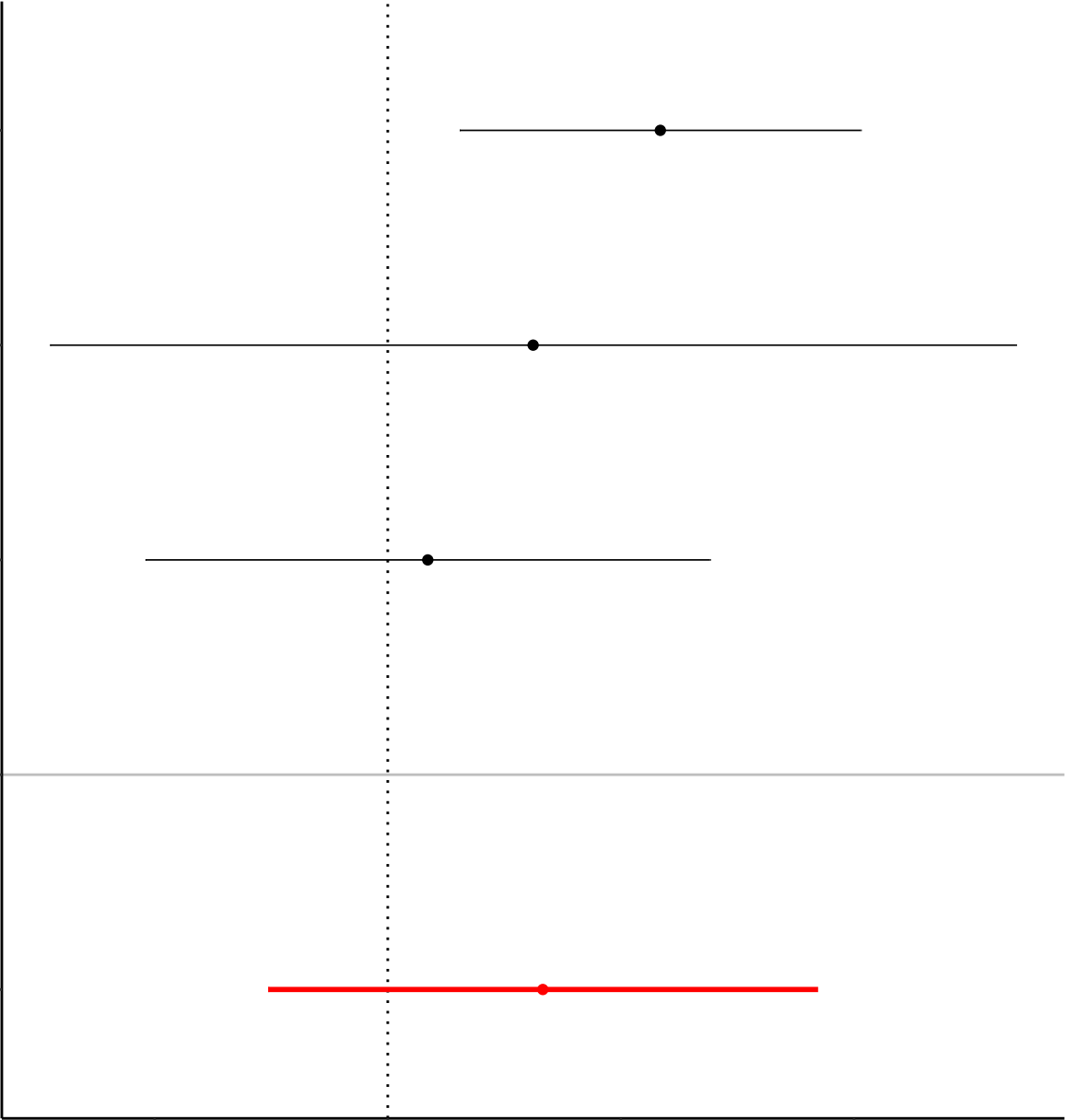
-0.05

0.00

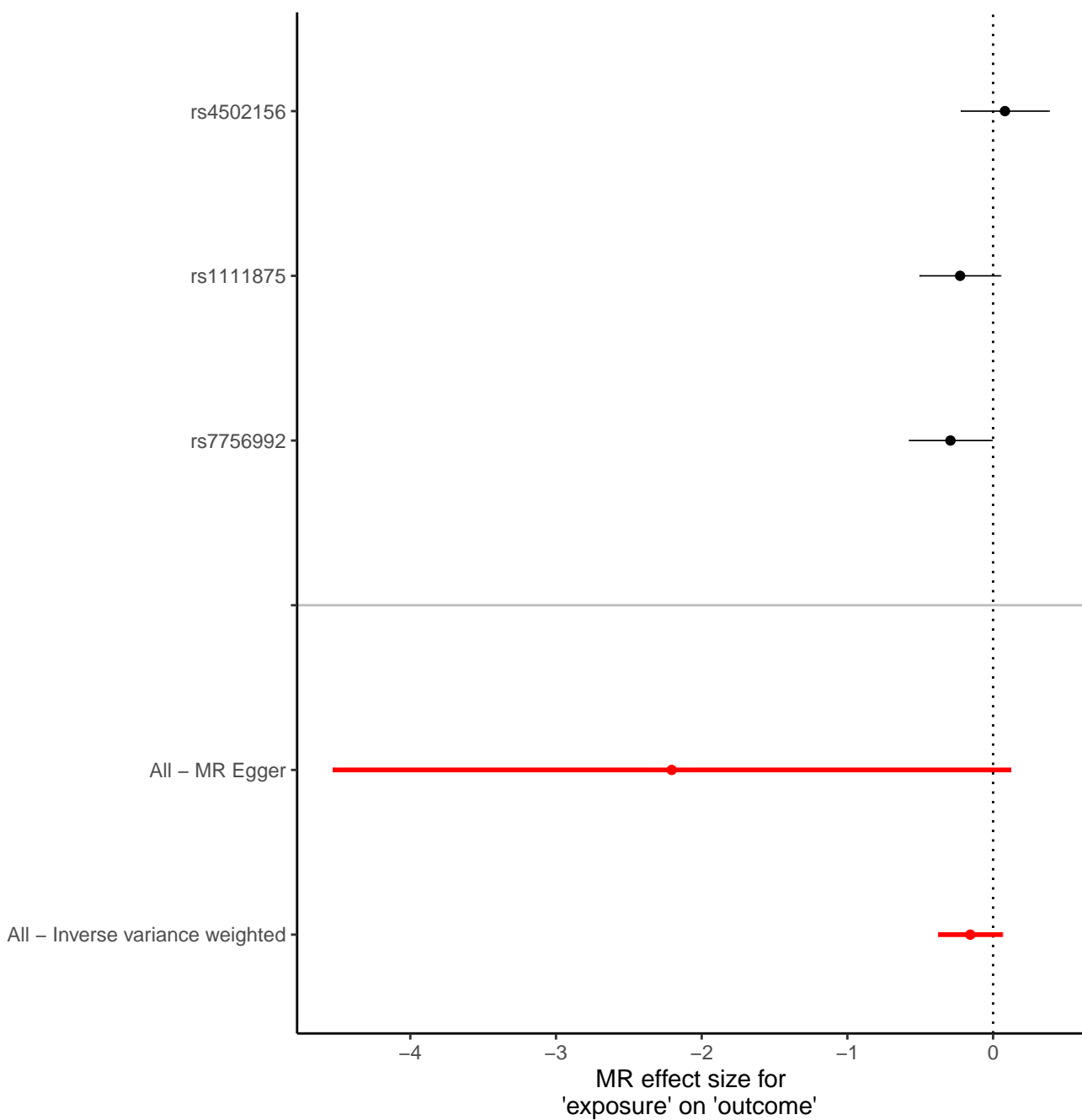
0.05

0.10

MR leave-one-out sensitivity analysis for  
'exposure' on 'outcome'



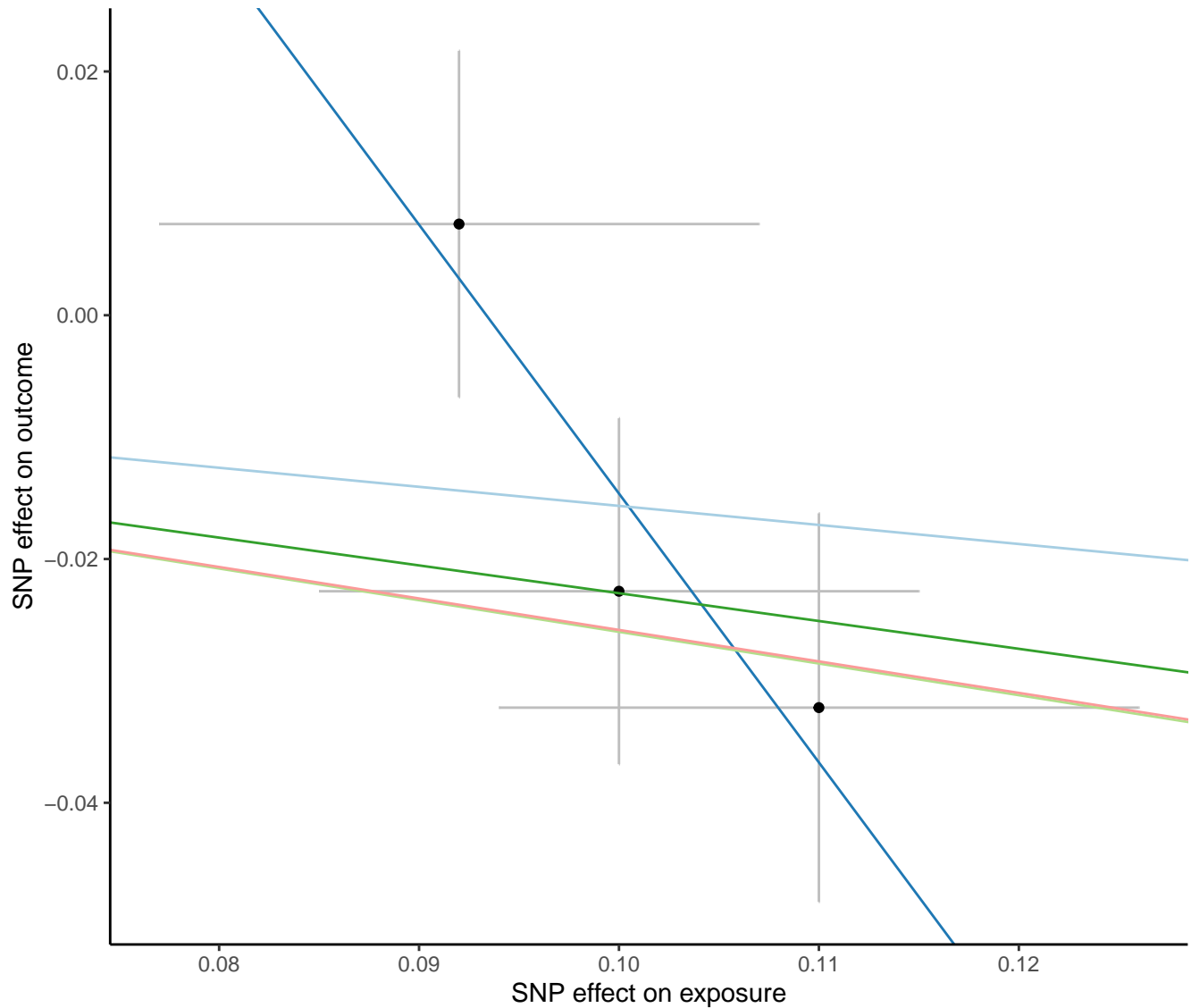
**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Coronary Artery Disease**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**





MR Test

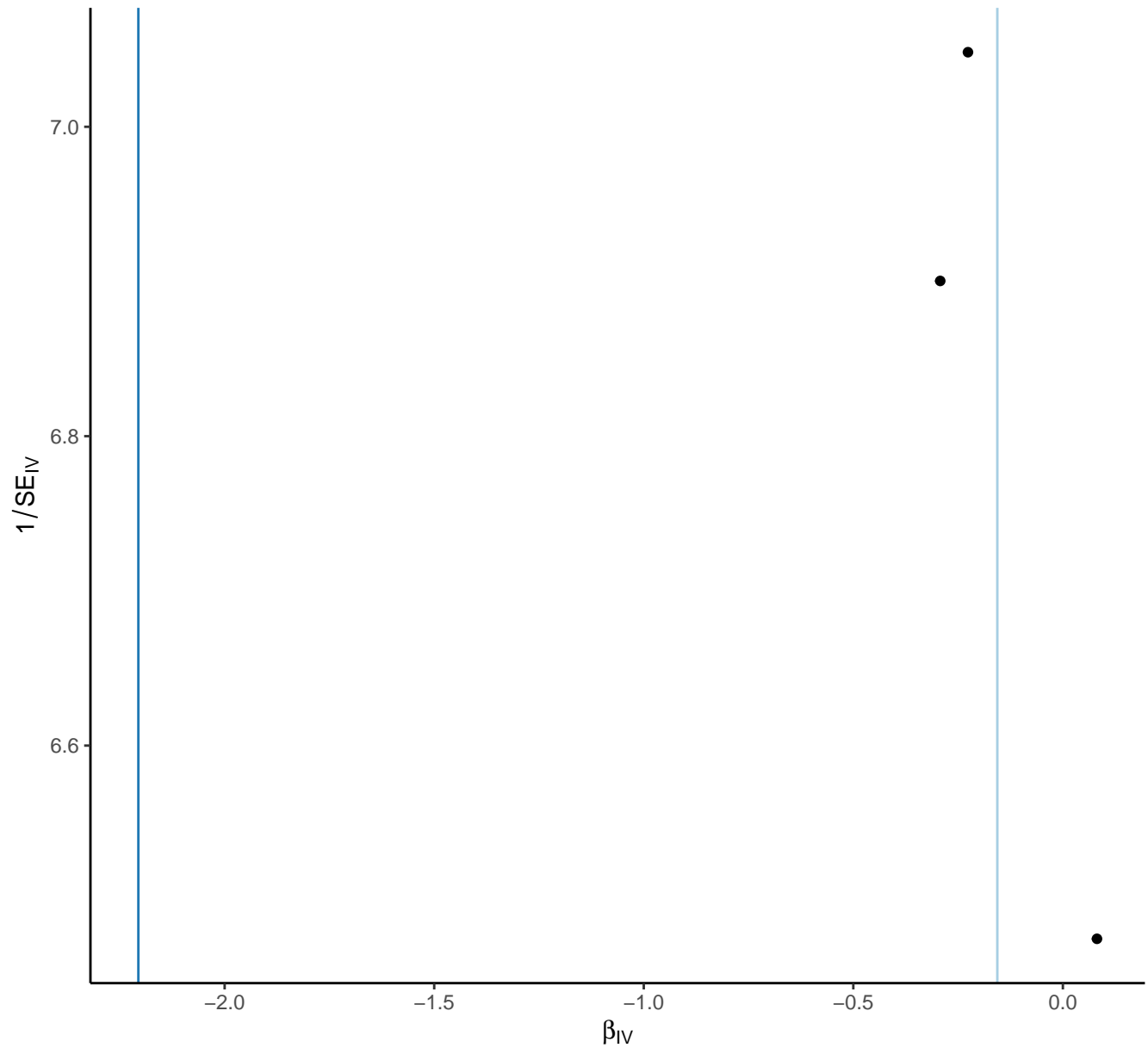
- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode



MR Method

Inverse variance weighted

MR Egger

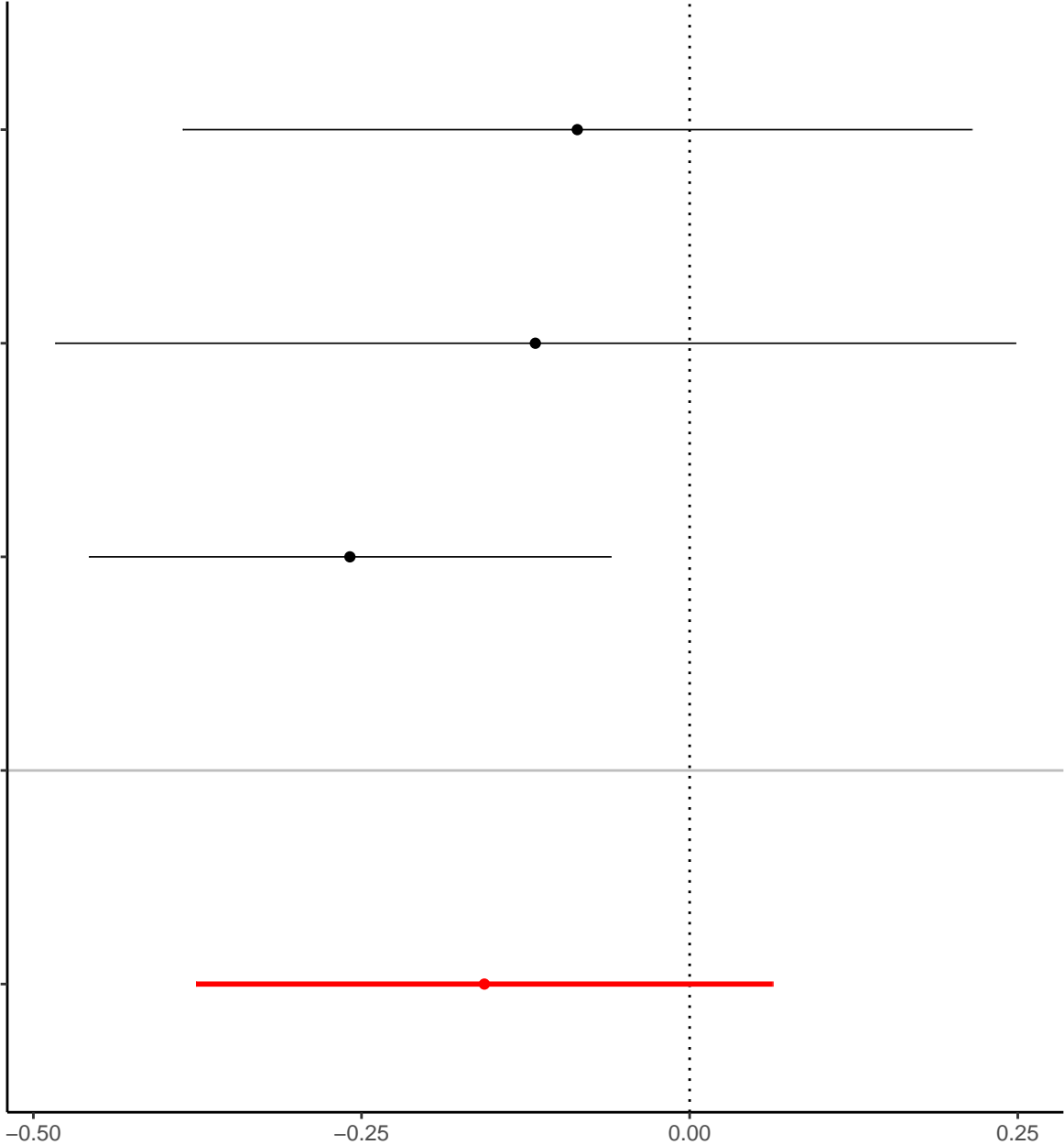


rs7756992

rs1111875

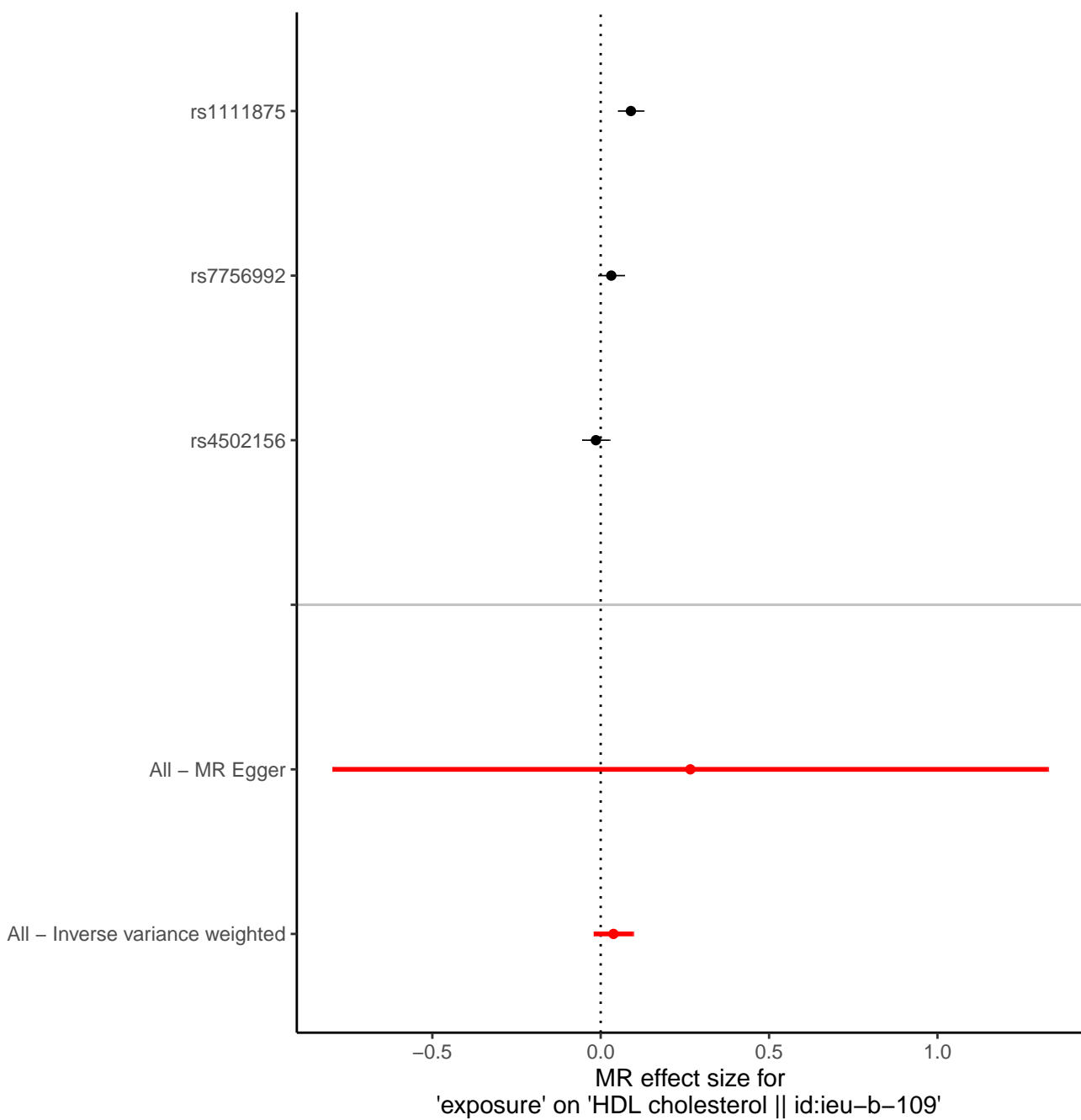
rs4502156

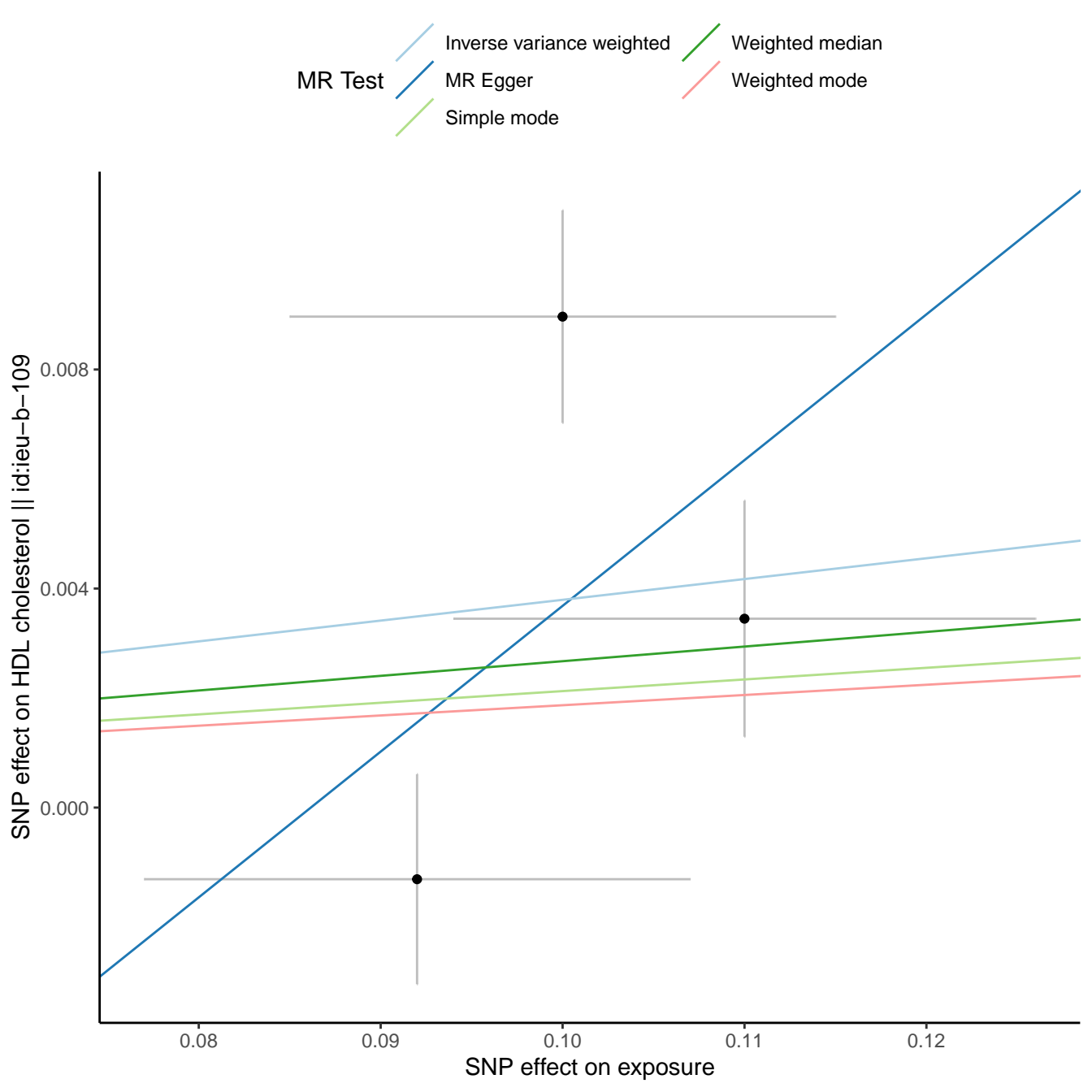
All

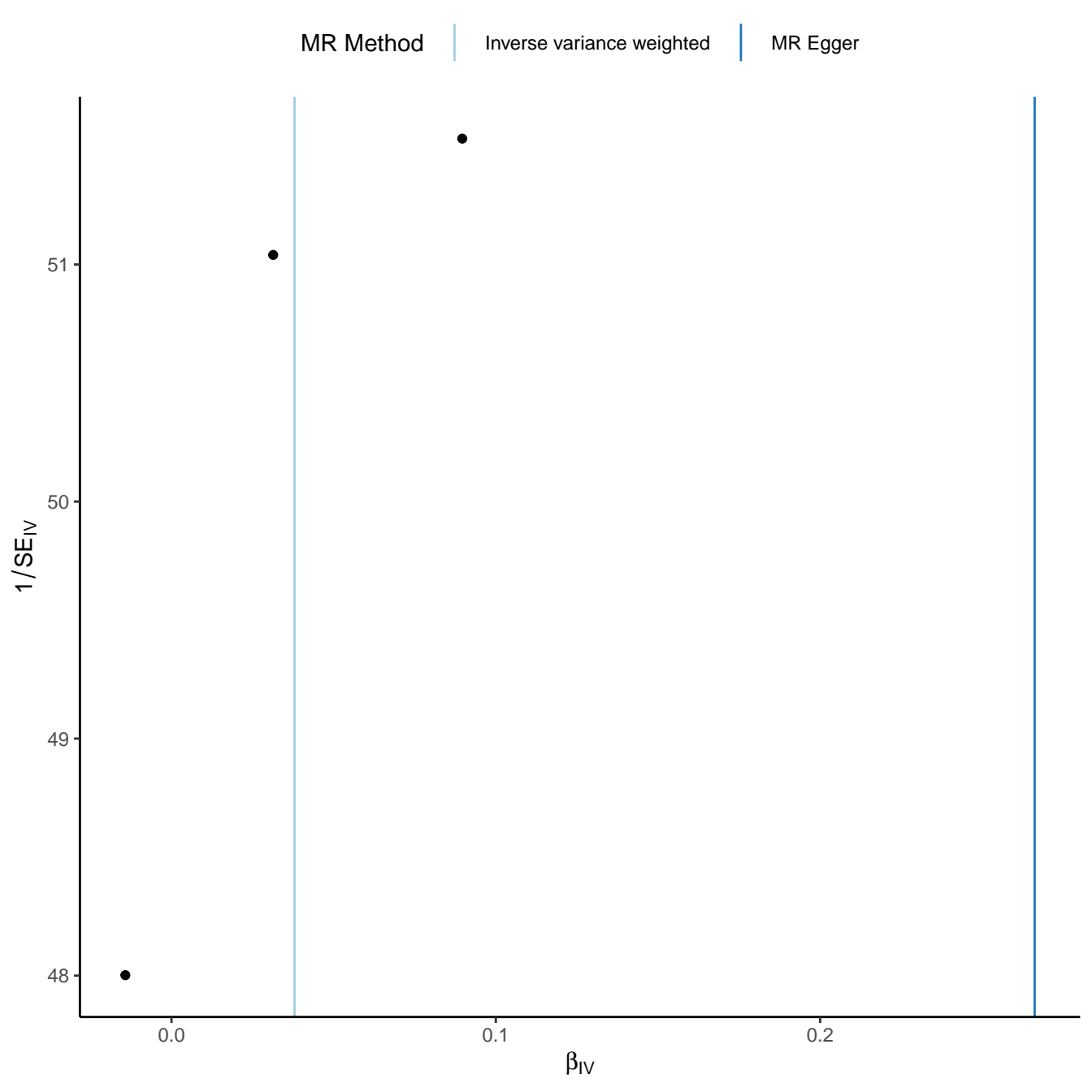


MR leave-one-out sensitivity analysis for  
'exposure' on 'outcome'

**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: High Density Lipoprotein**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**





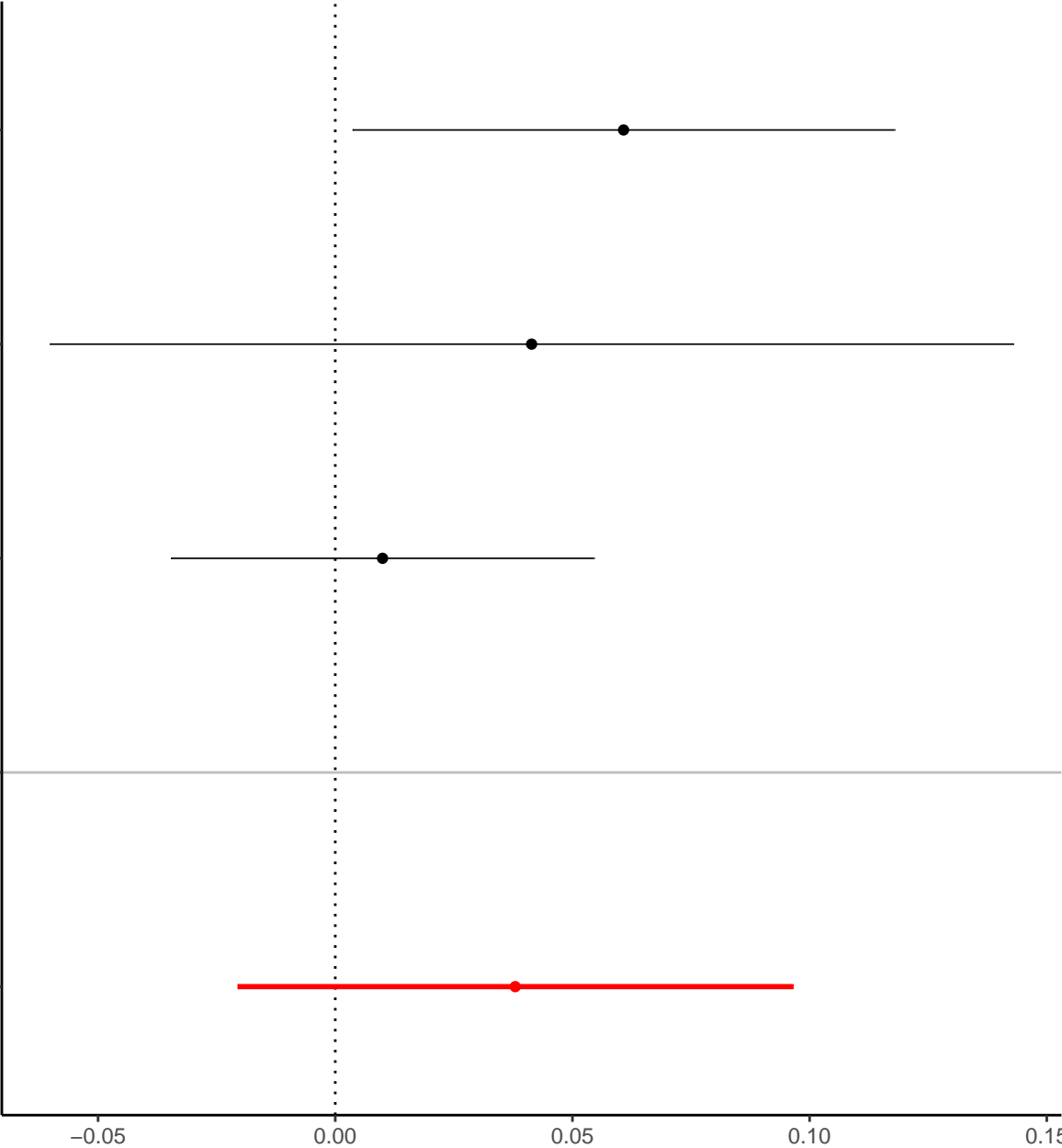


rs4502156

rs7756992

rs1111875

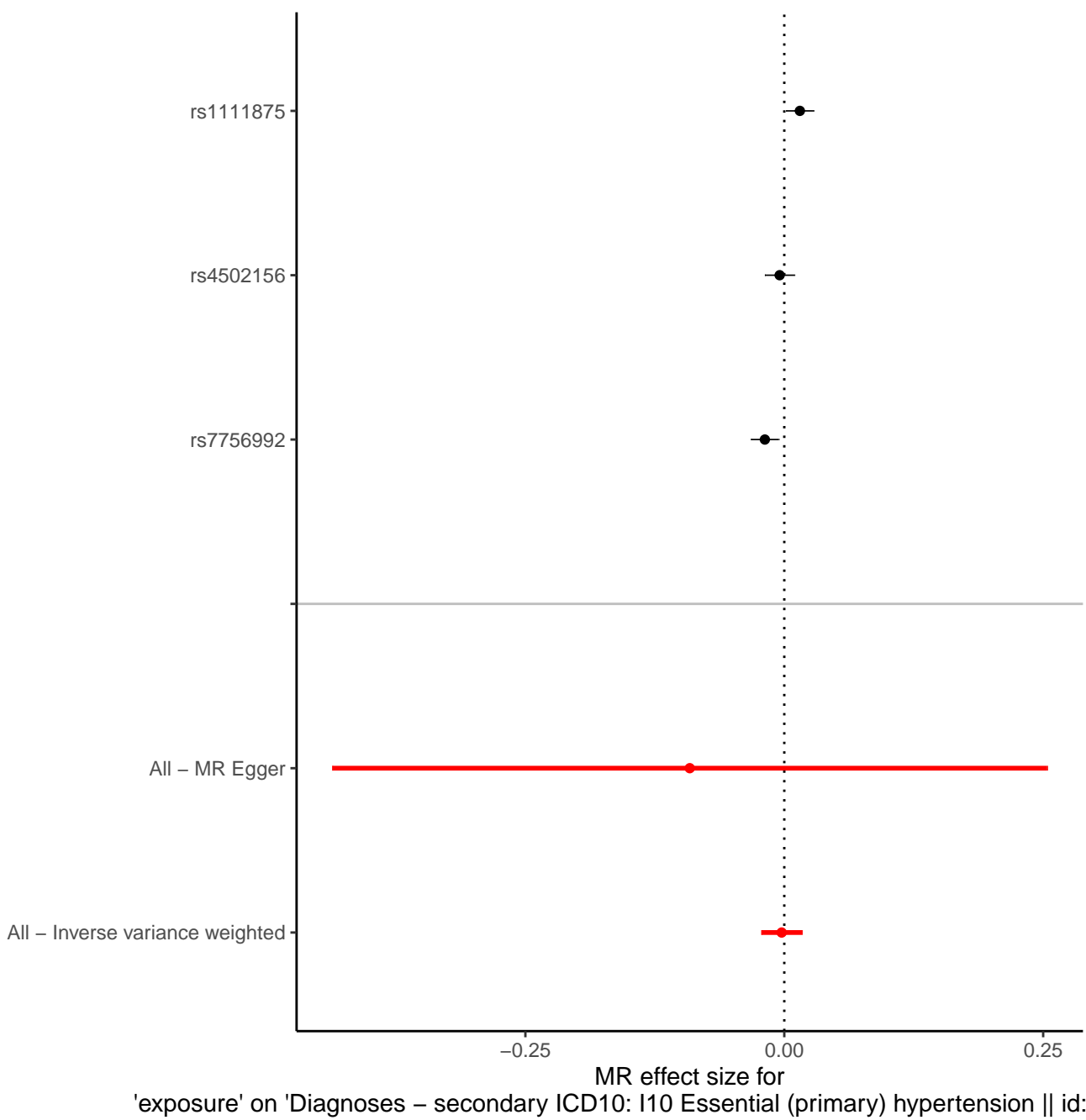
All



MR leave-one-out sensitivity analysis for 'exposure' on 'HDL cholesterol || id:ieu-b-109'

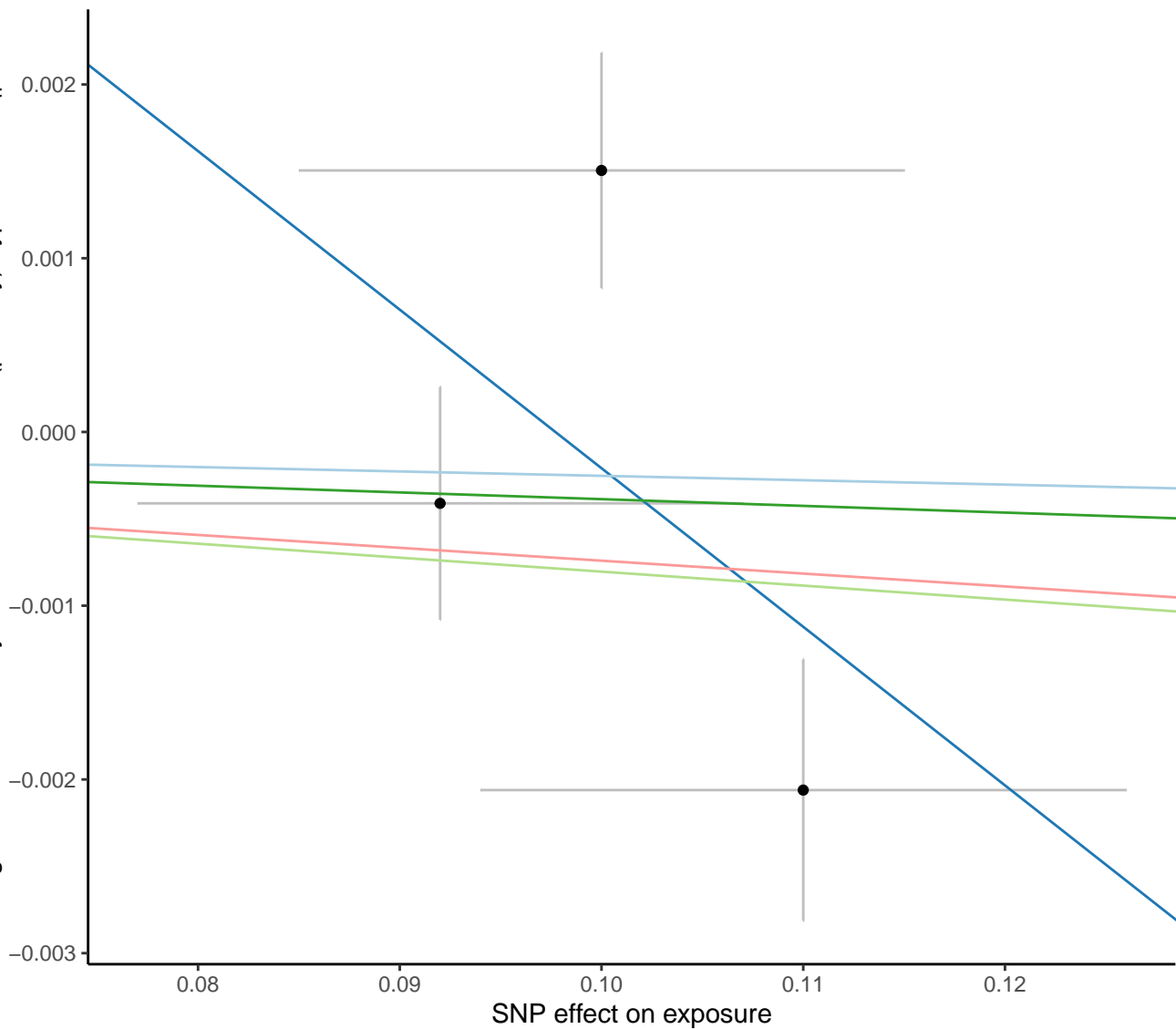


**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Hypertension**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**



MR Test

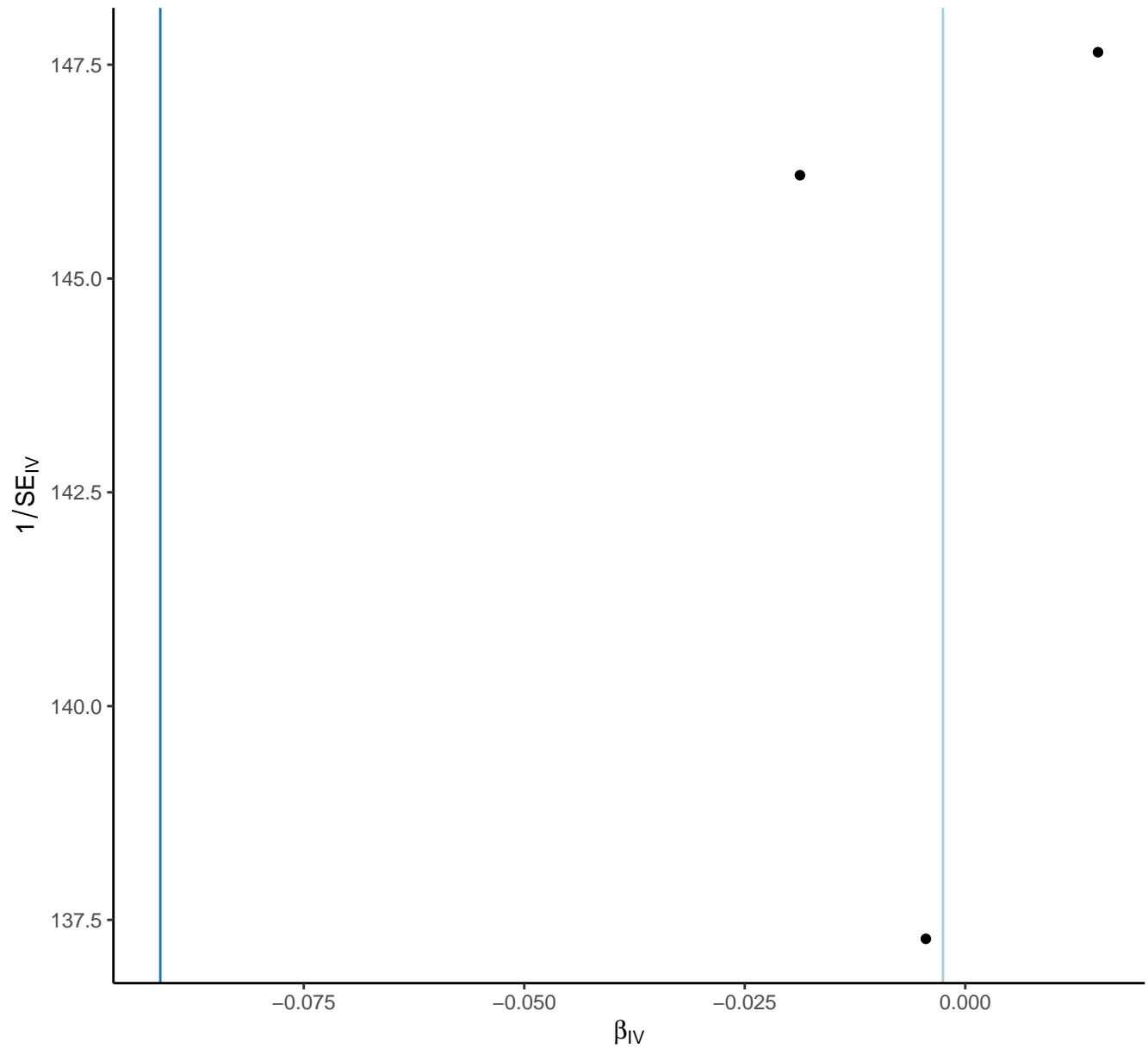
- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode



MR Method

Inverse variance weighted

MR Egger



rs7756992

rs4502156

rs1111875

All

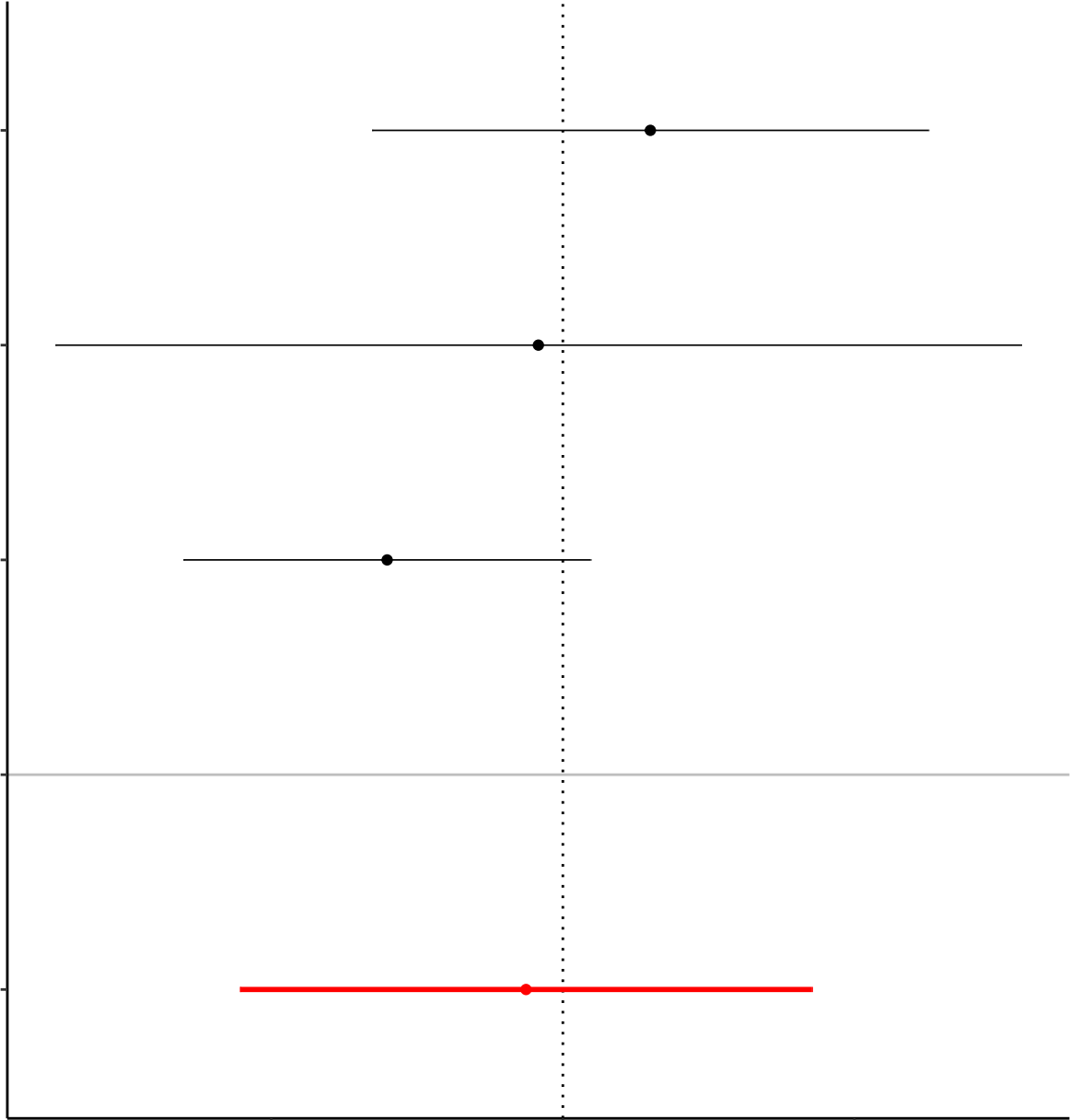
-0.02

0.00

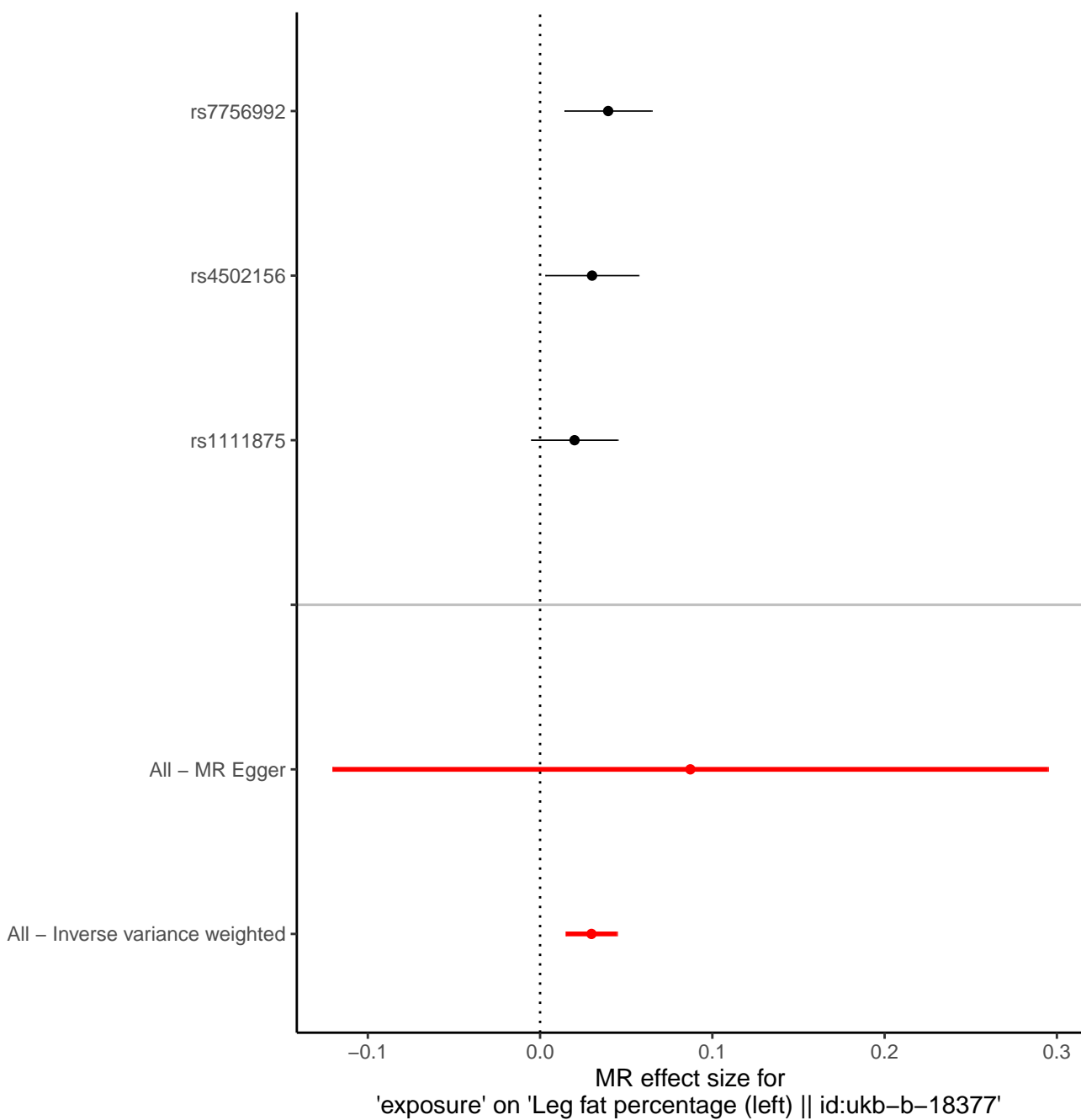
0.02

MR leave-one-out sensitivity analysis for

'exposure' on 'Diagnoses – secondary ICD10: I10 Essential (primary) hypertension || id:ukb-b-1

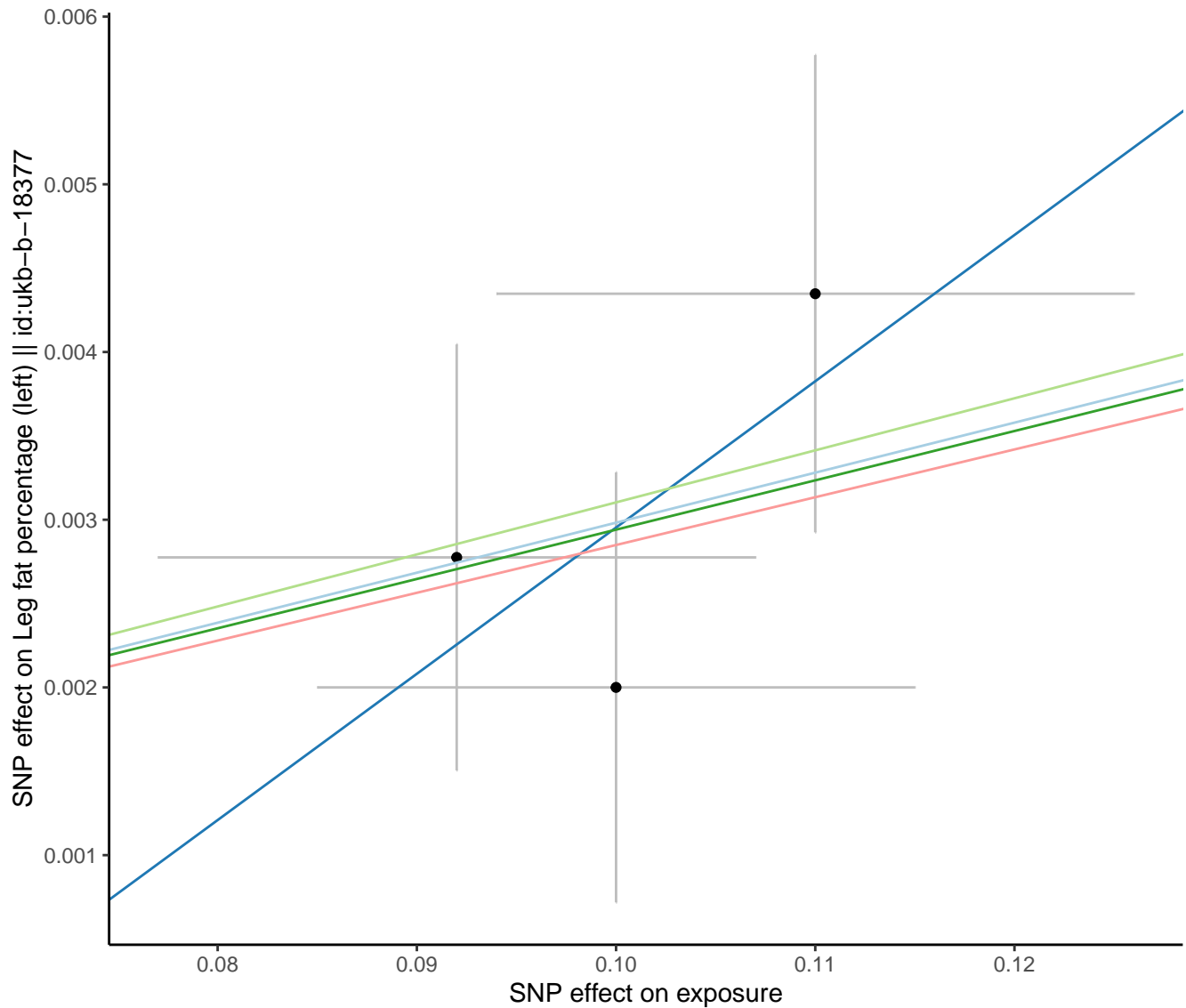


**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Leg Fat Percent - Left**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**



MR Test

- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode

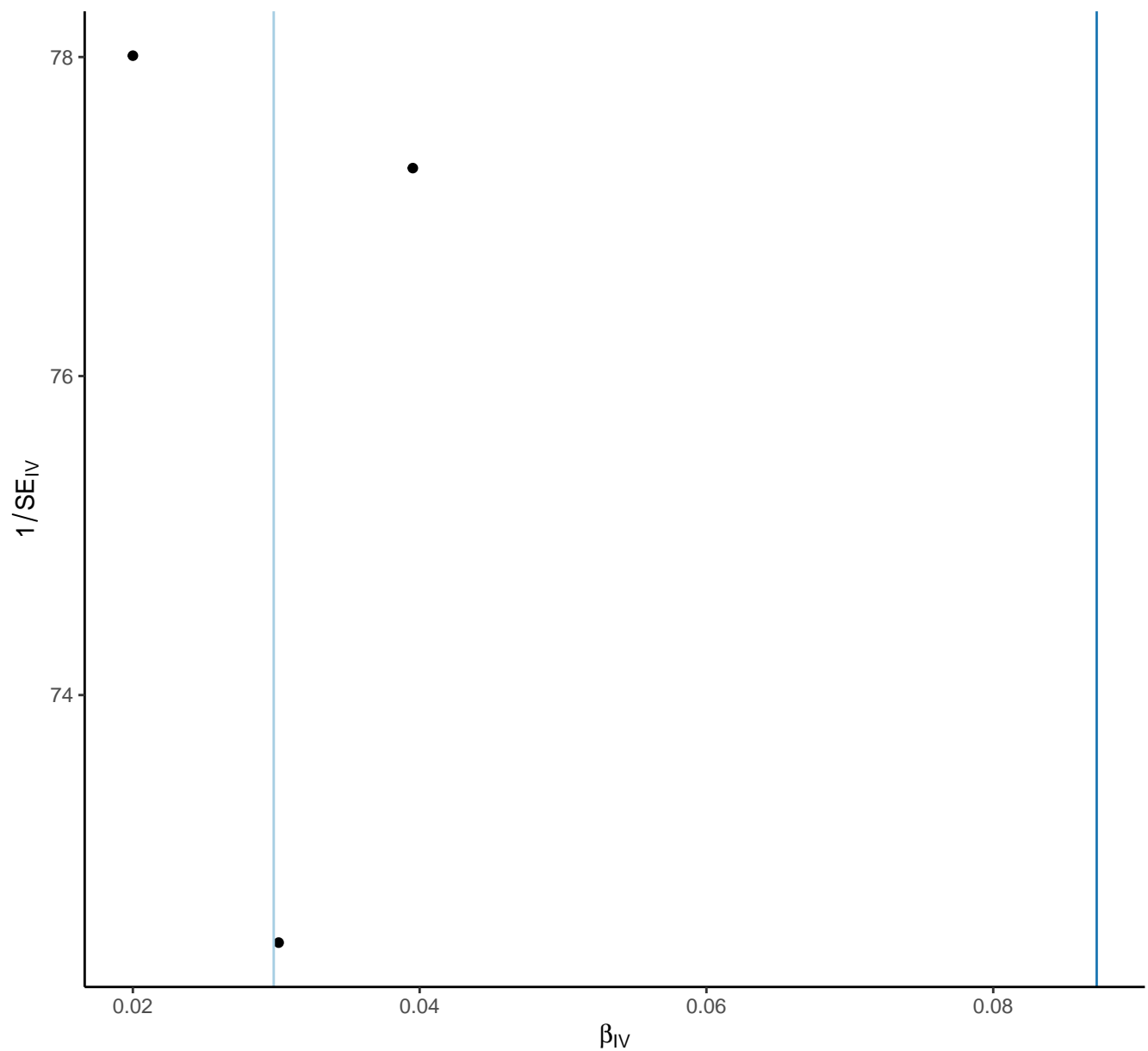




MR Method

Inverse variance weighted

MR Egger



rs1111875

rs4502156

rs7756992

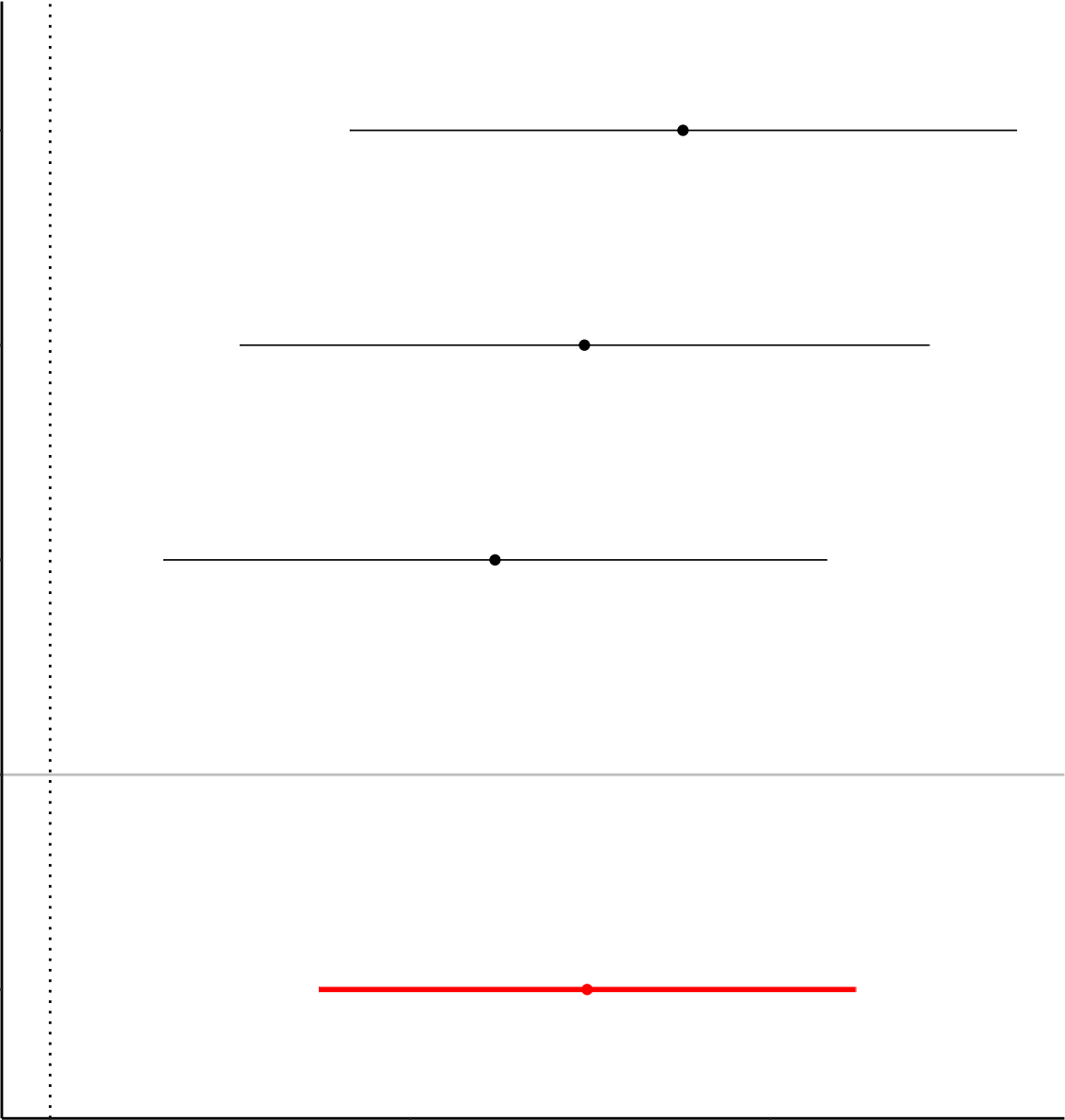
All

0.00

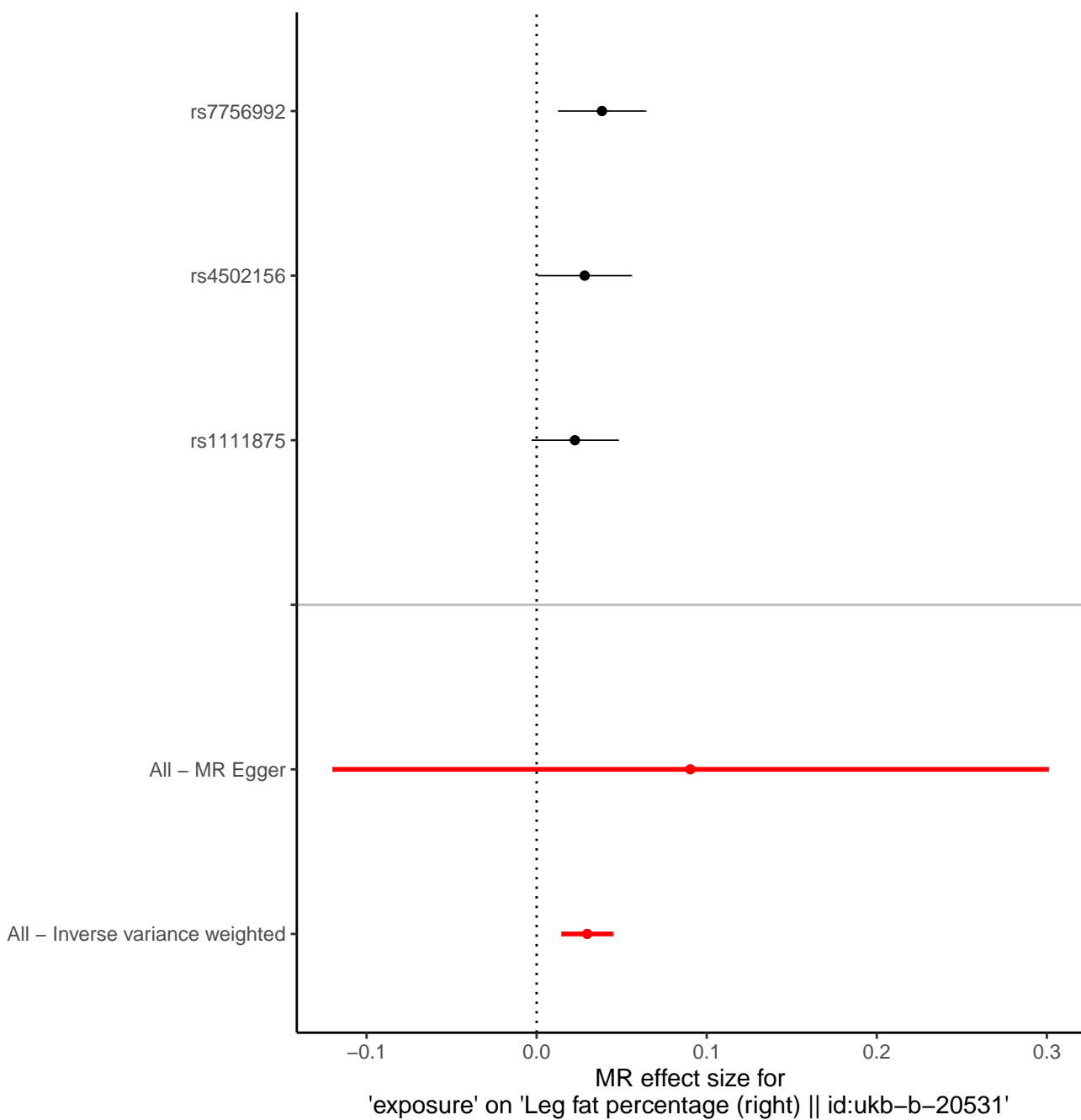
0.02

0.04

MR leave-one-out sensitivity analysis for  
'exposure' on 'Leg fat percentage (left) || id:ukb-b-18377'

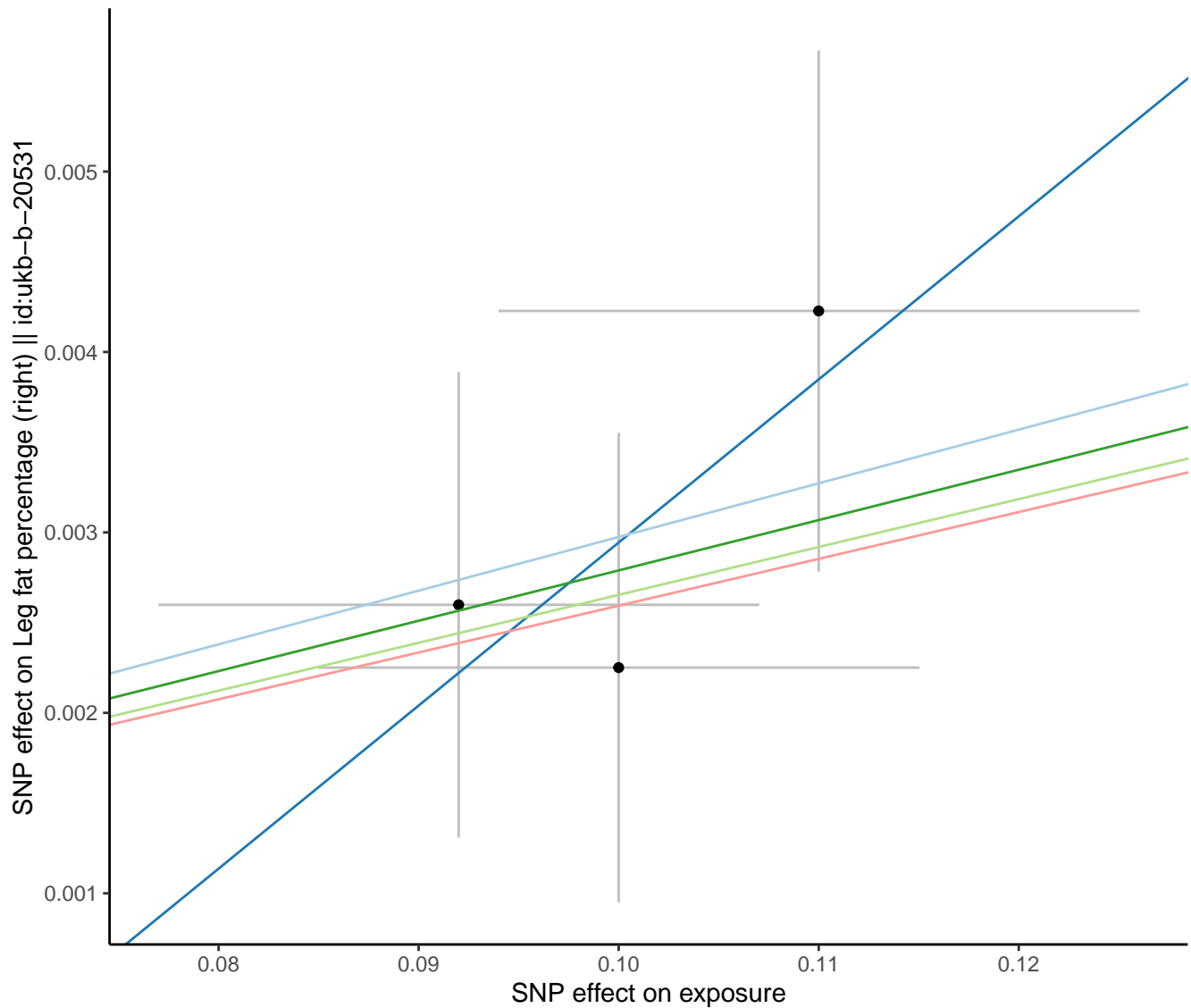


**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Leg Fat Percent - Right**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**



MR Test

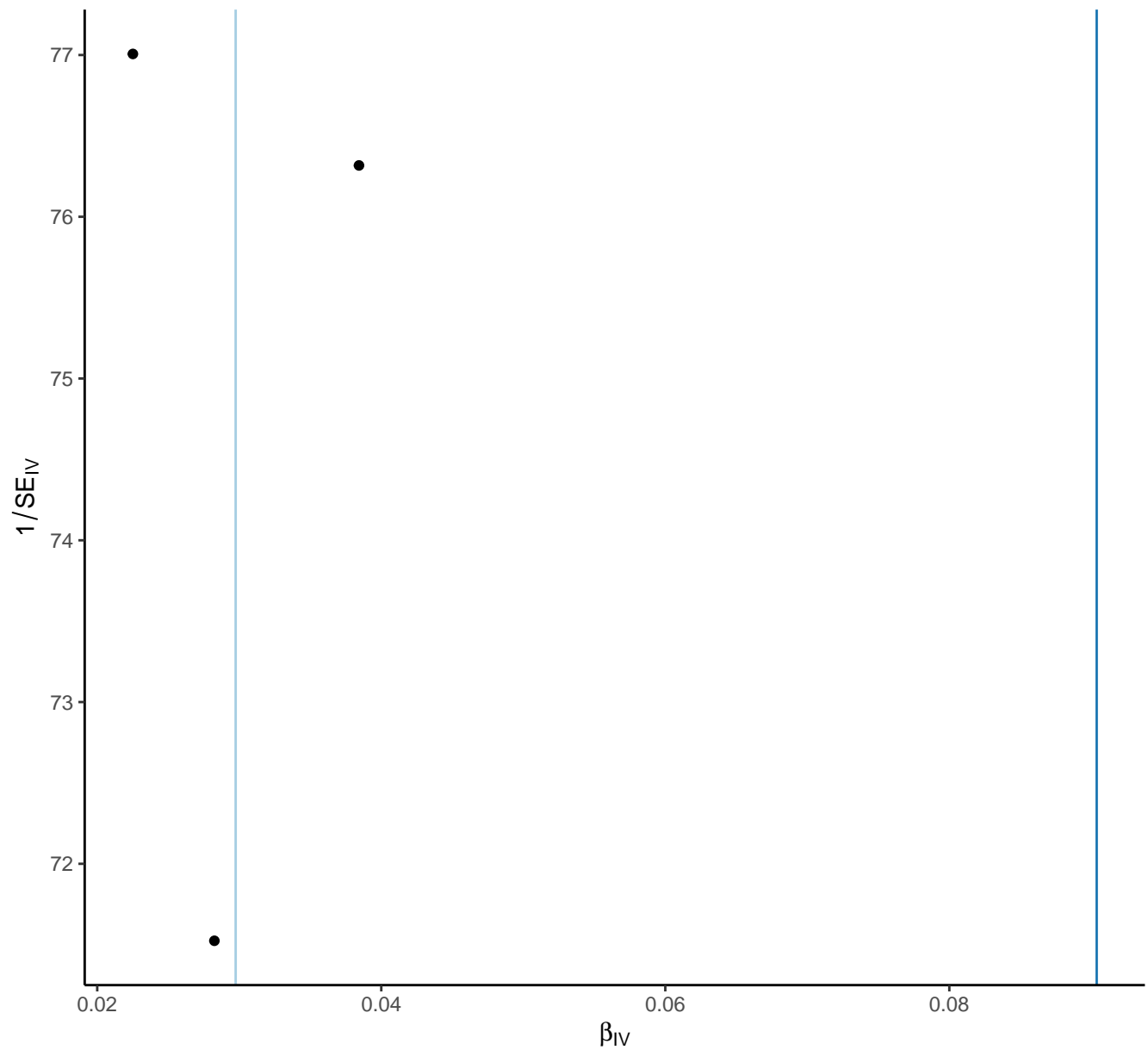
- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode



MR Method

Inverse variance weighted

MR Egger



rs1111875

rs4502156

rs7756992

All

0.00

0.01

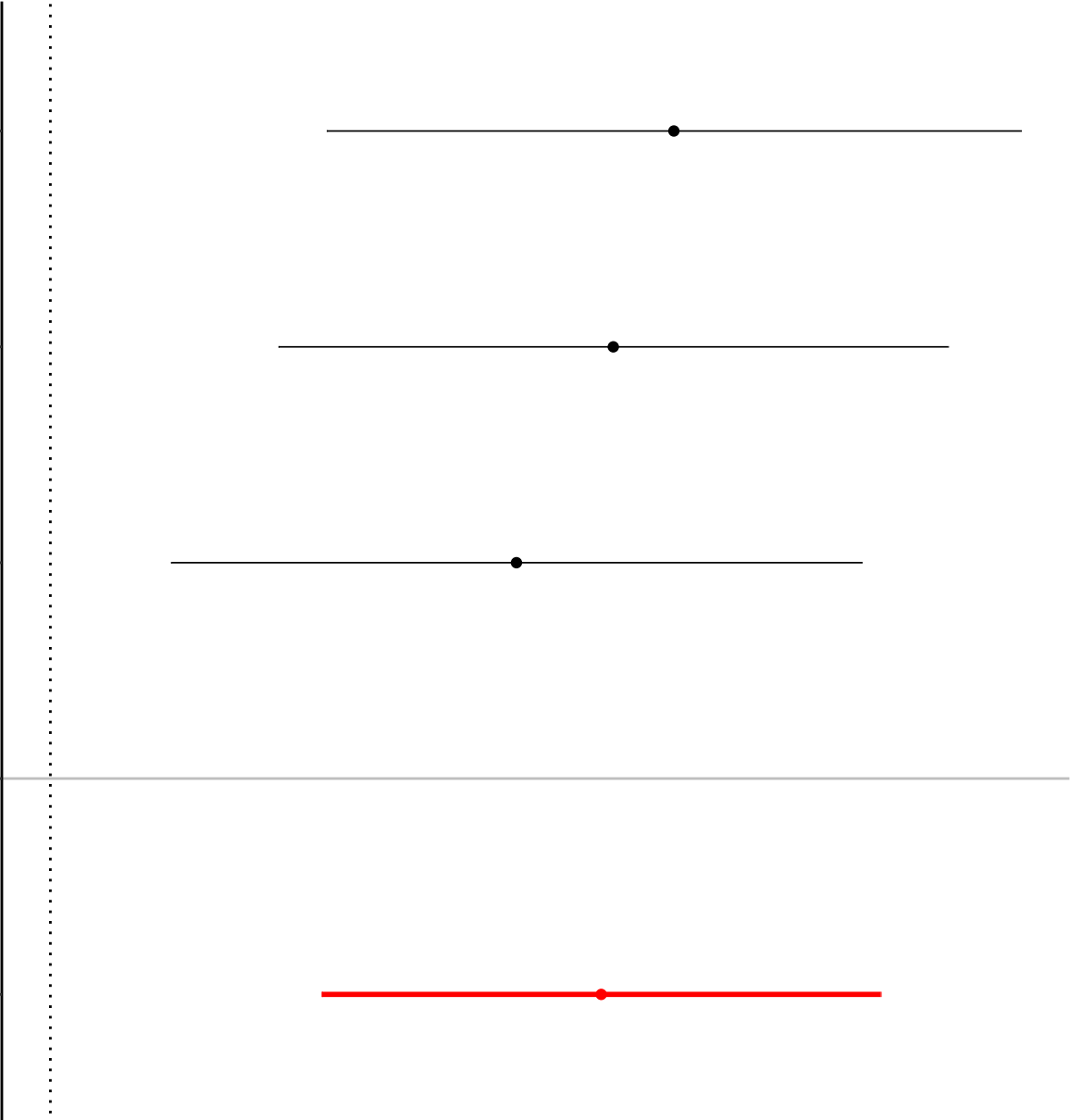
0.02

0.03

0.04

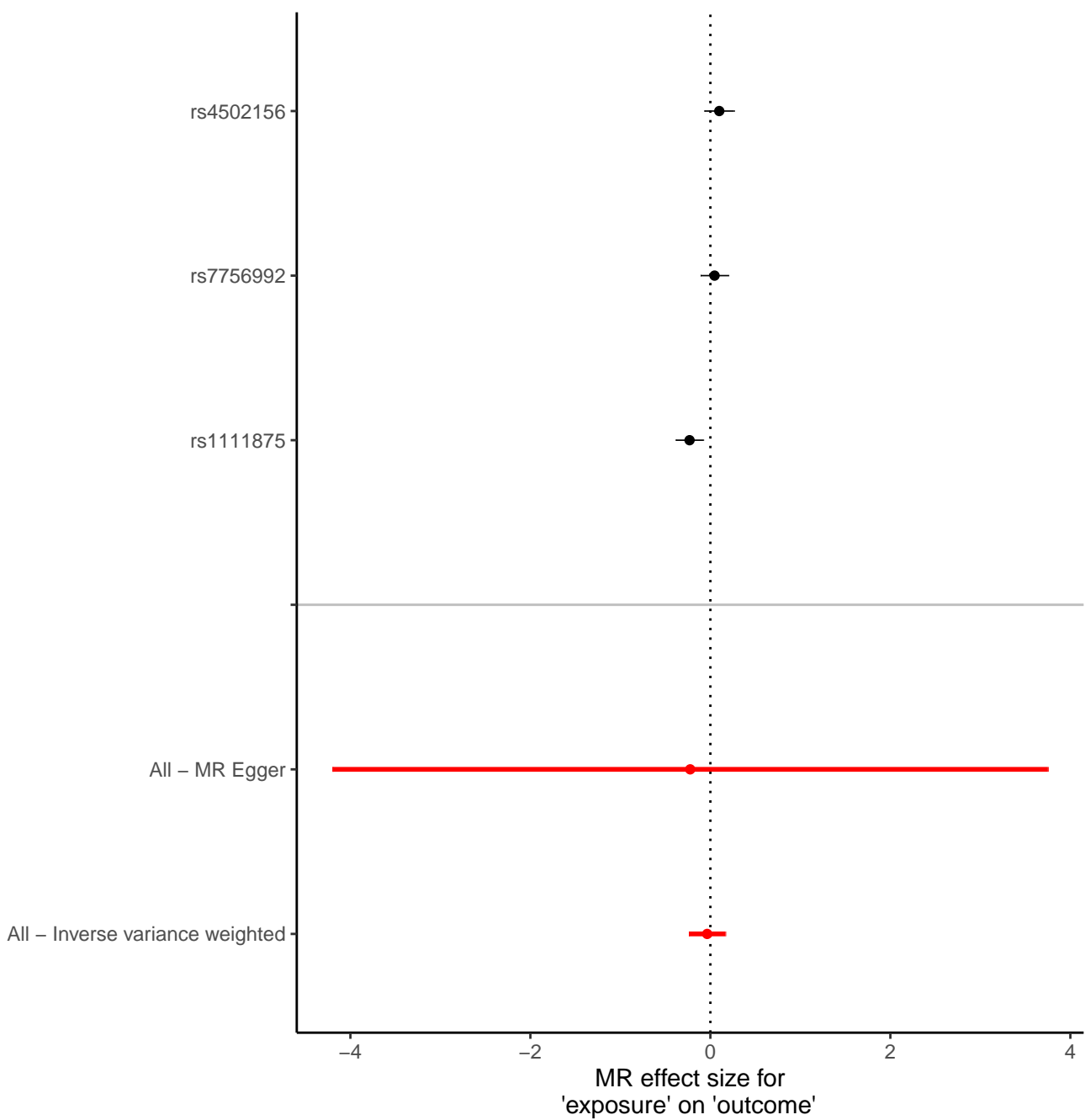
0.05

MR leave-one-out sensitivity analysis for  
'exposure' on 'Leg fat percentage (right) || id:ukb-b-20531'



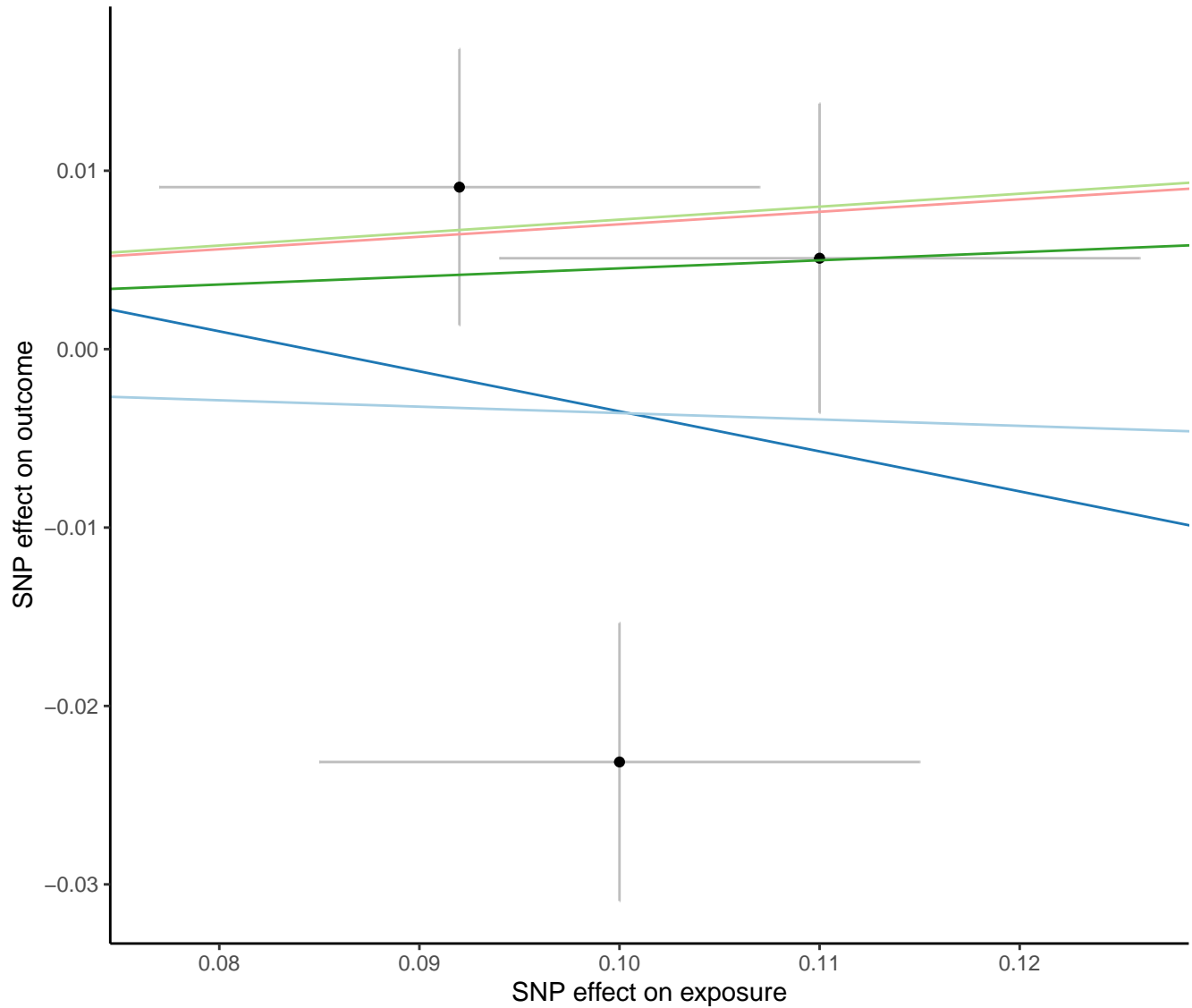
**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Liver Fat**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**





MR Test

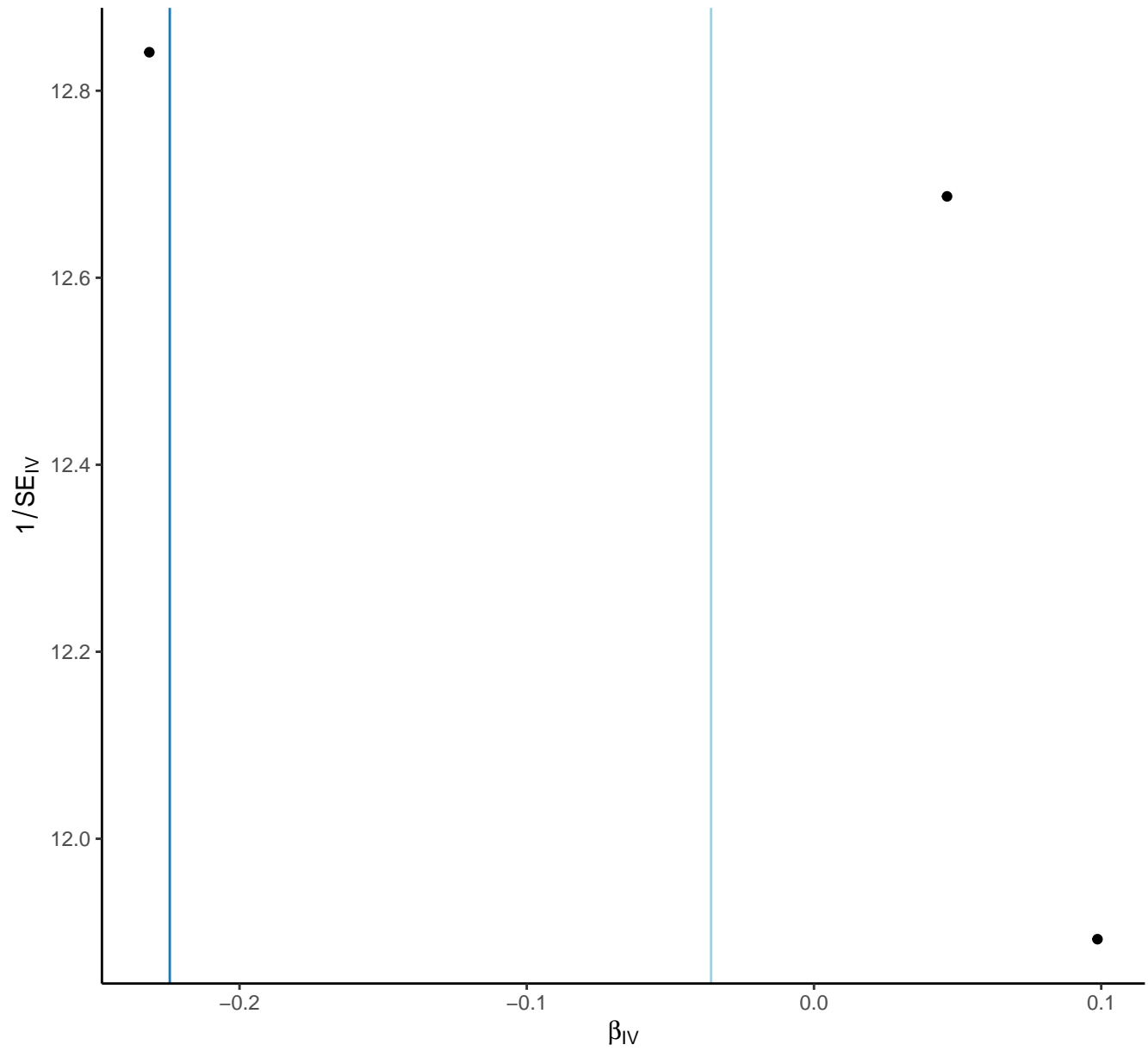
- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode

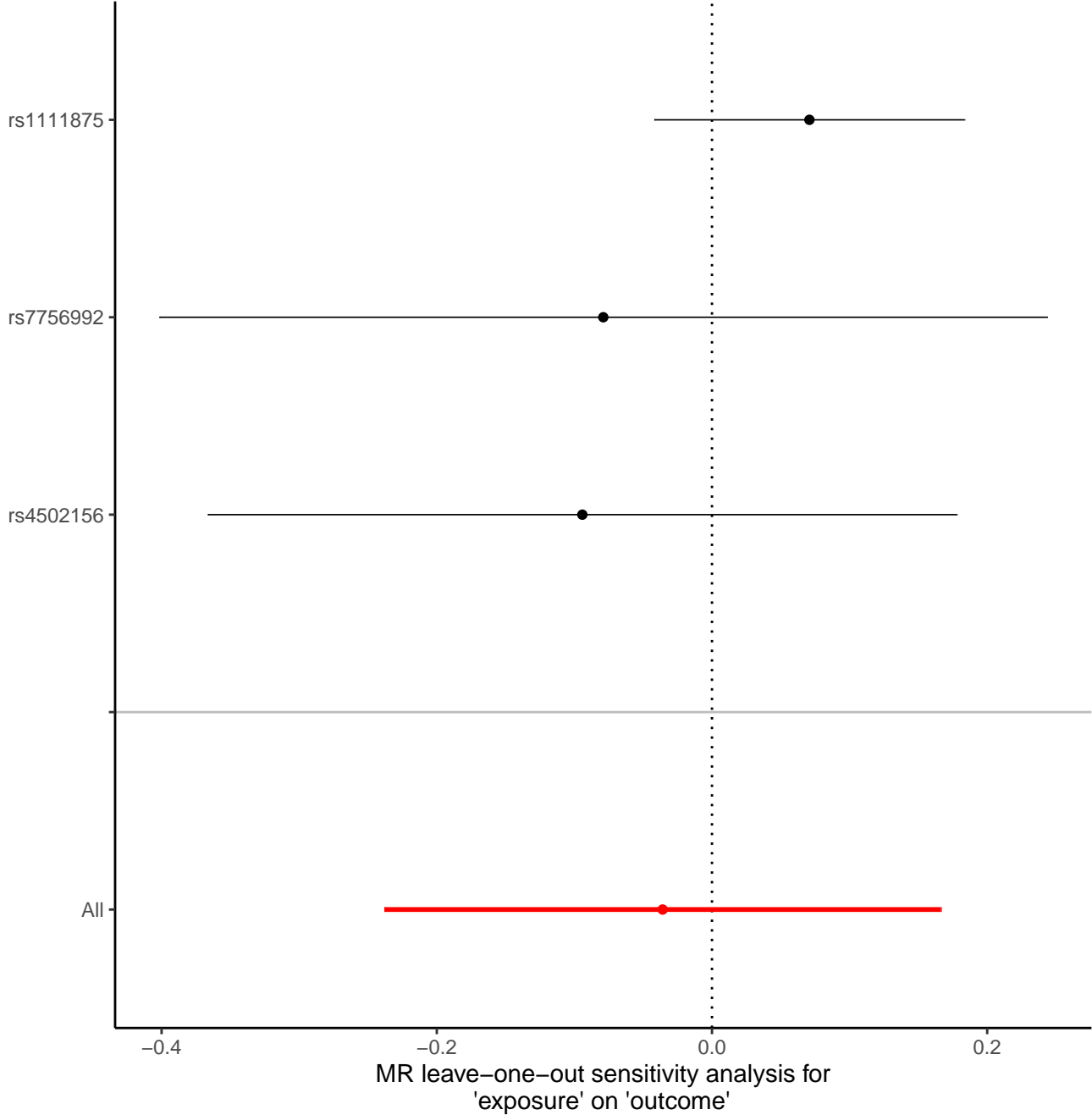


MR Method

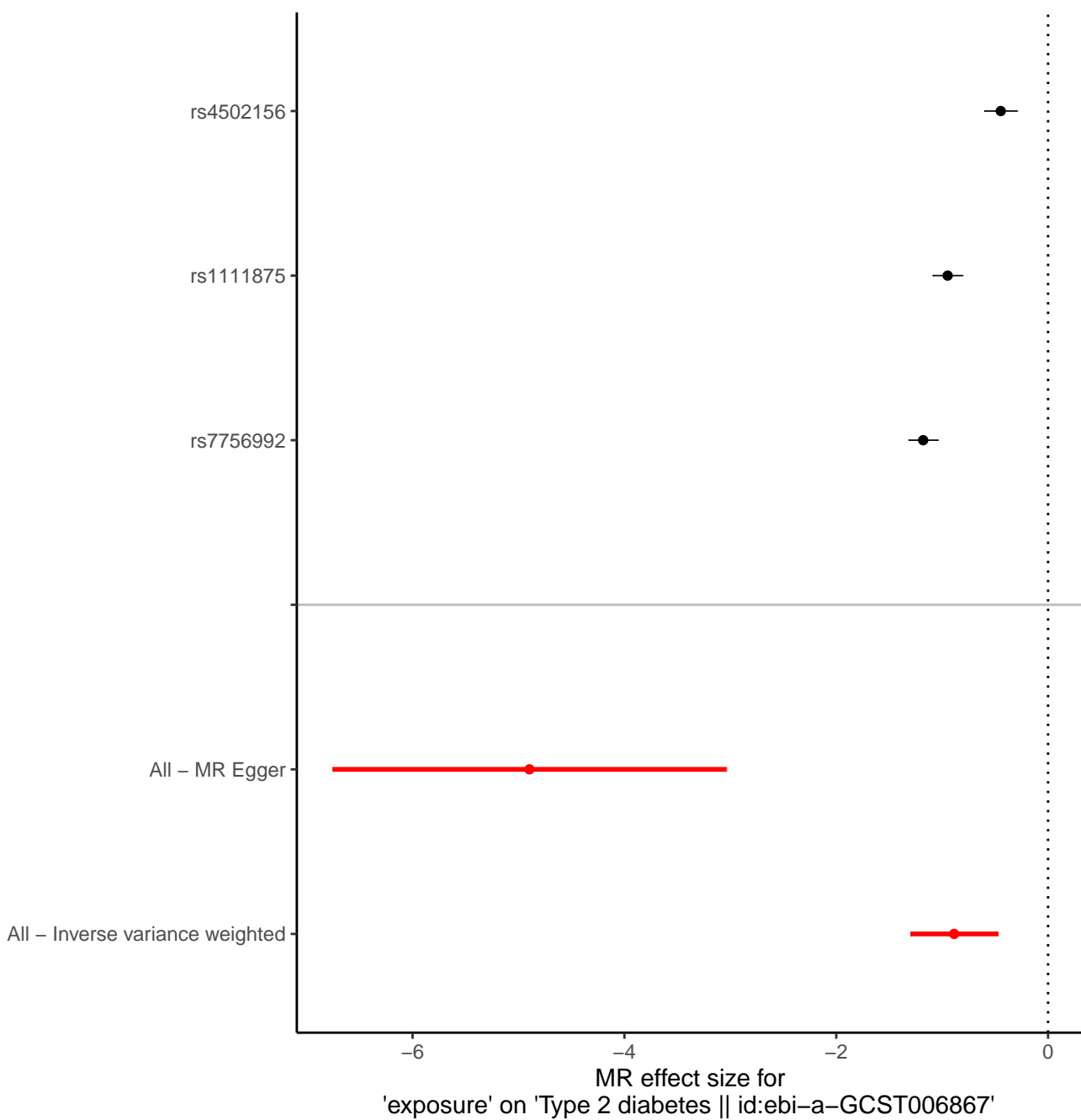
Inverse variance weighted

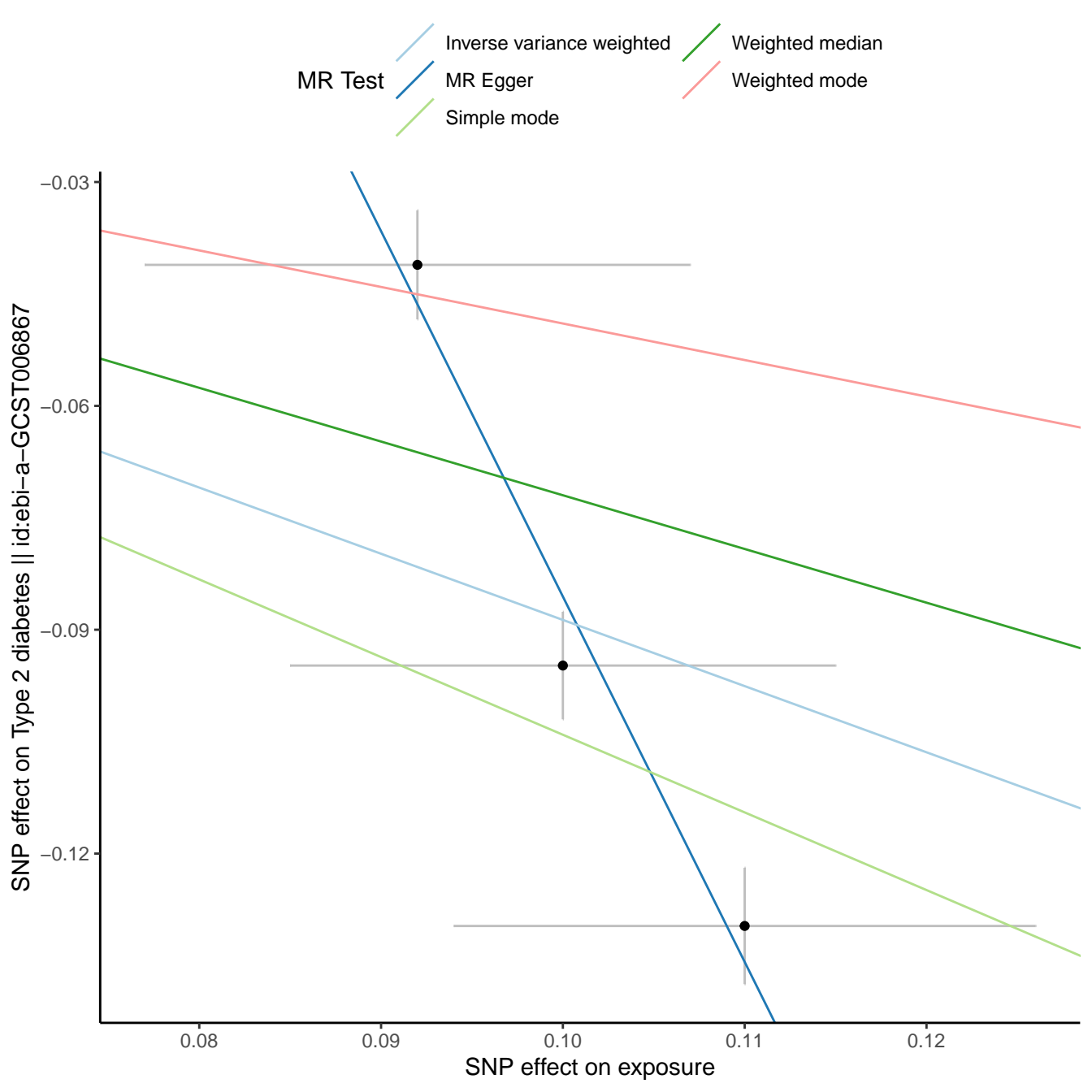
MR Egger





**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Type 2 Diabetes**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**

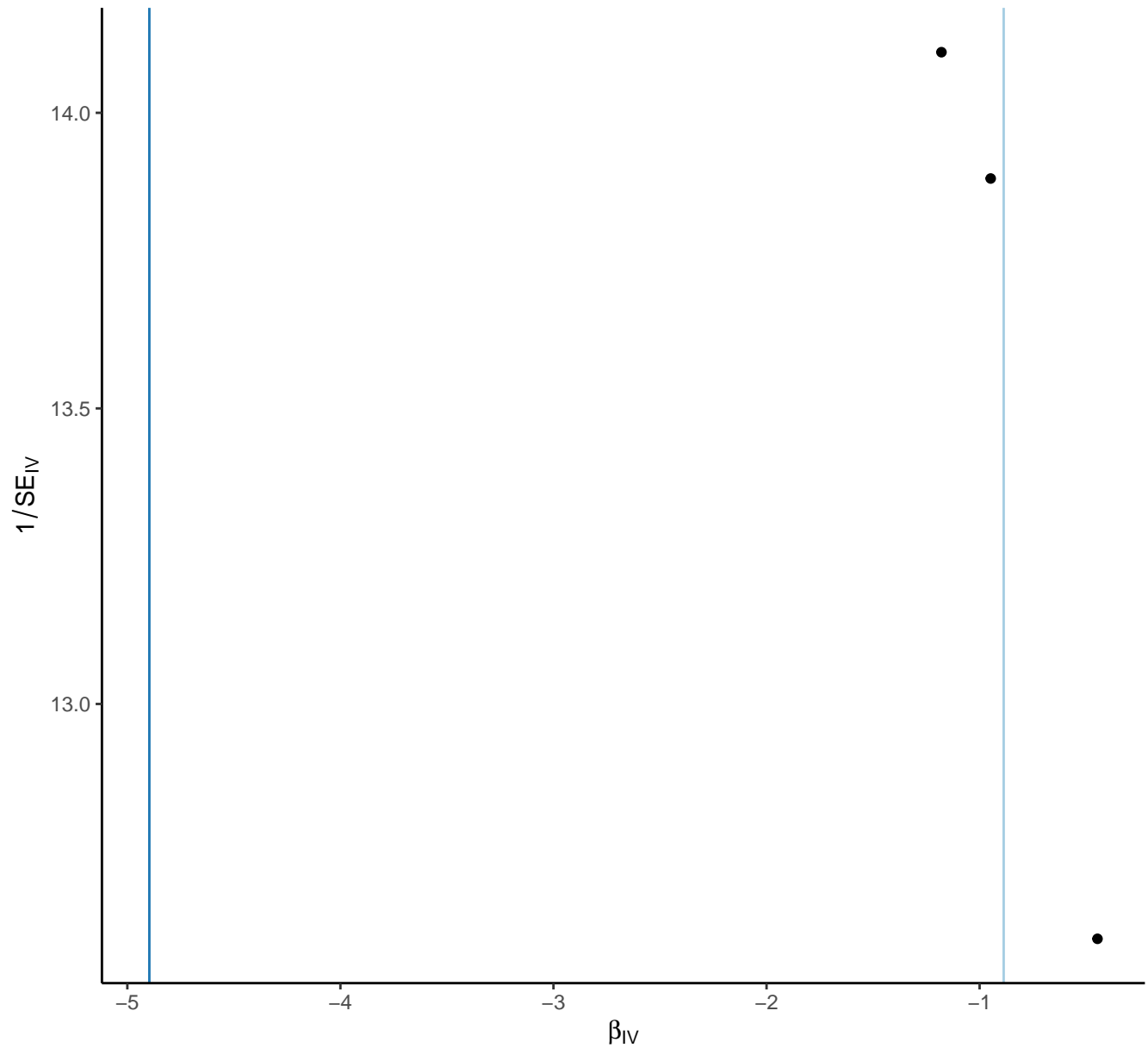




MR Method

Inverse variance weighted

MR Egger





rs7756992

rs1111875

rs4502156

All

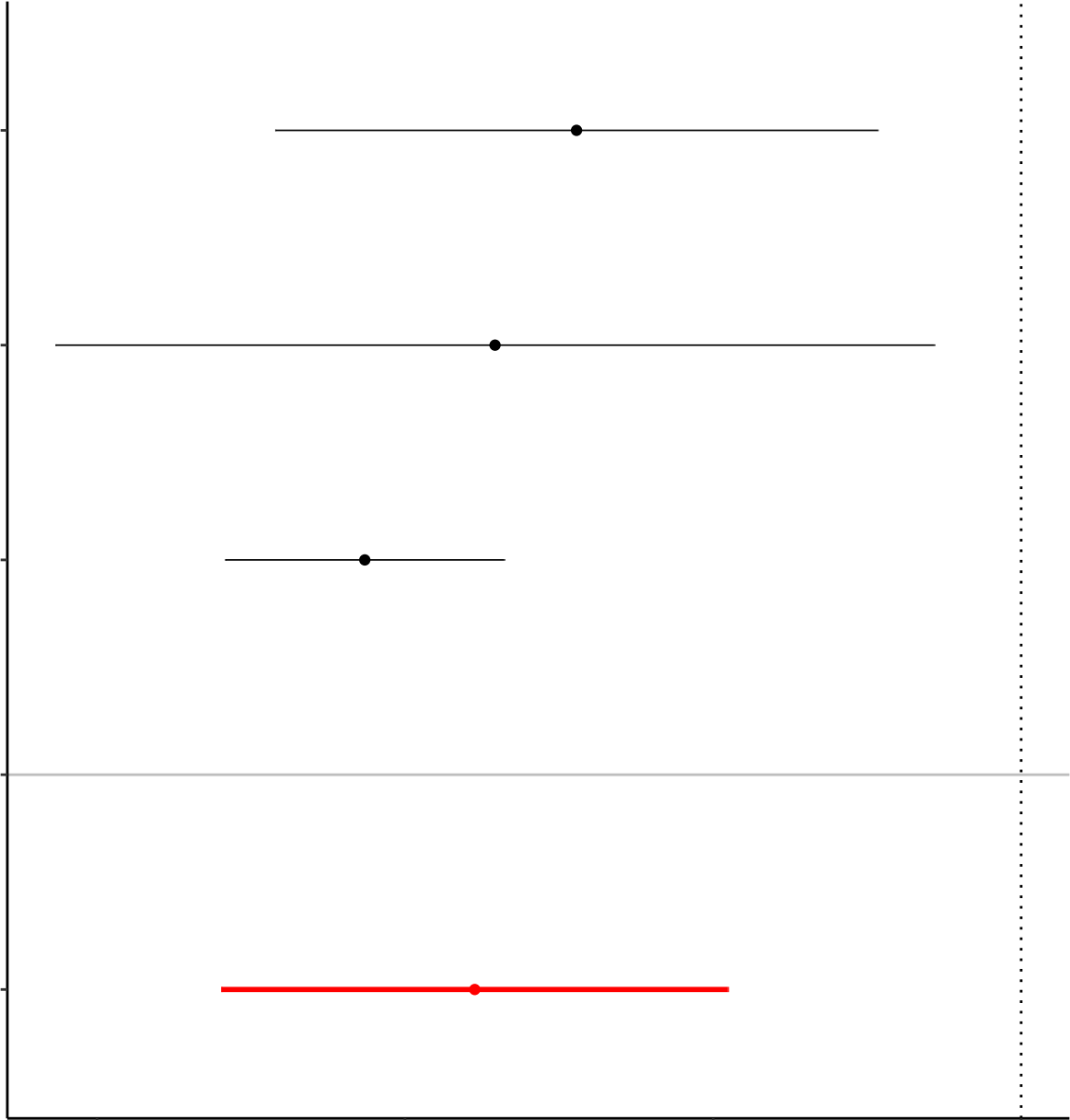
-1.5

-1.0

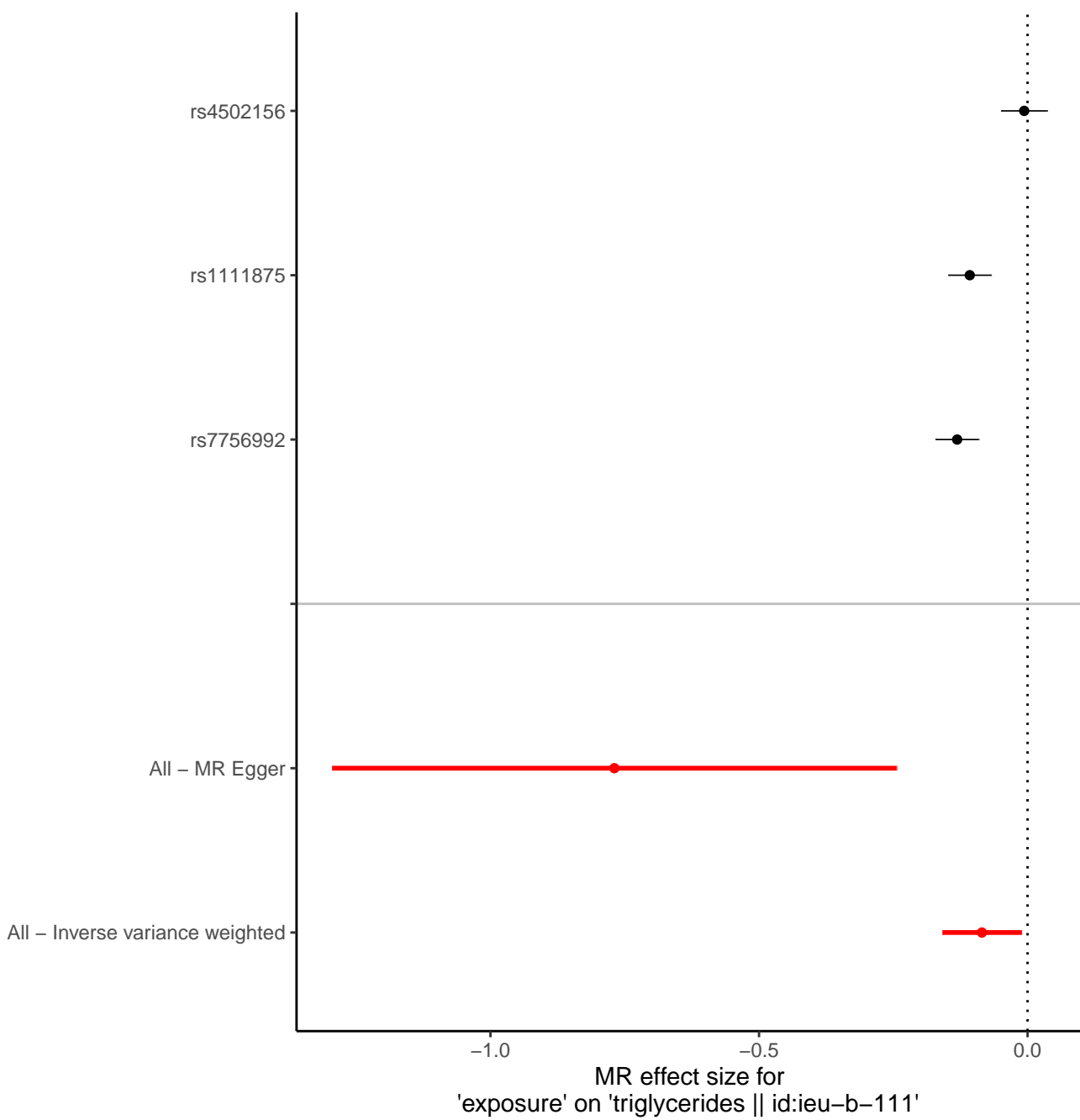
-0.5

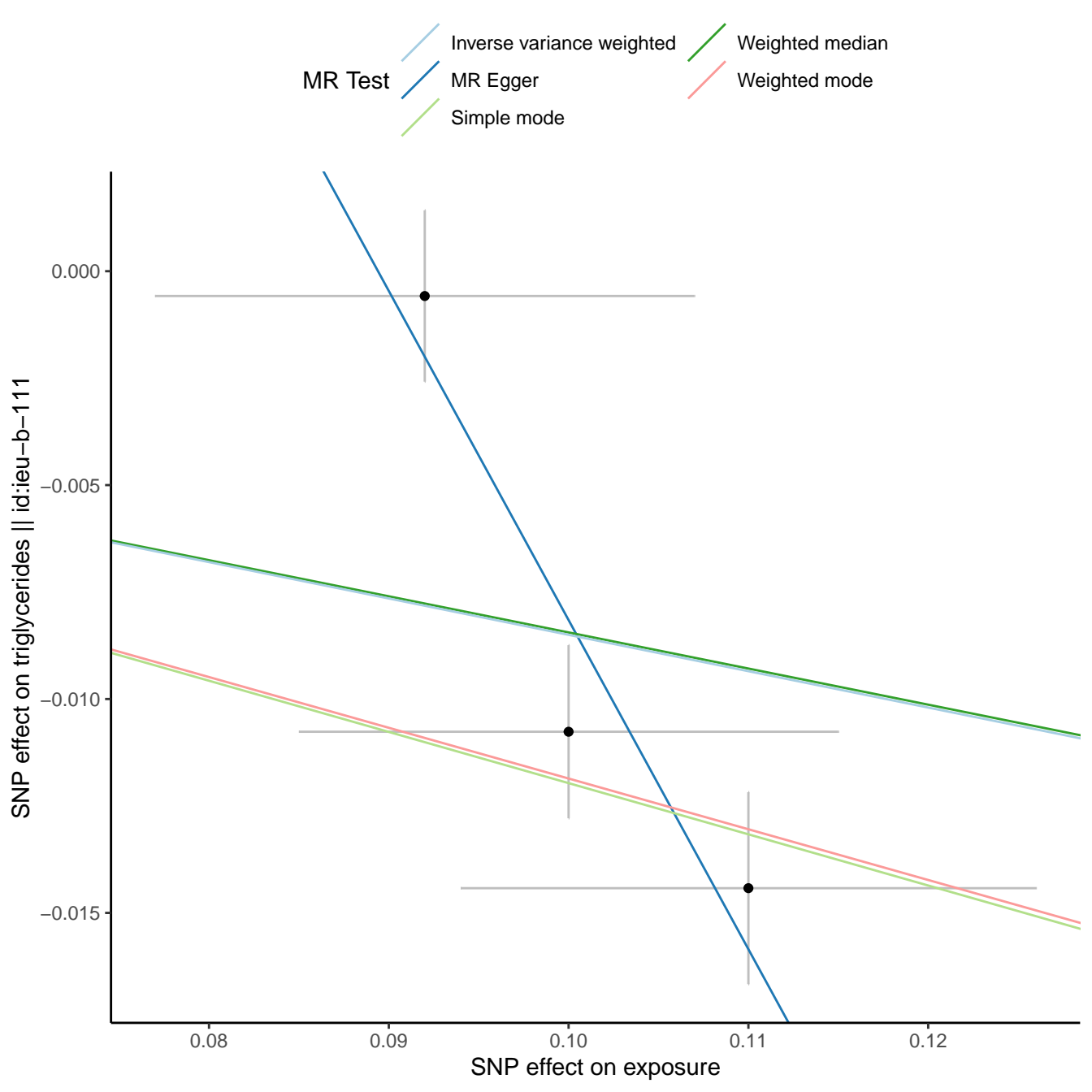
0.0

MR leave-one-out sensitivity analysis for  
'exposure' on 'Type 2 diabetes || id:ebi-a-GCST006867'



**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Triglycerides**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**

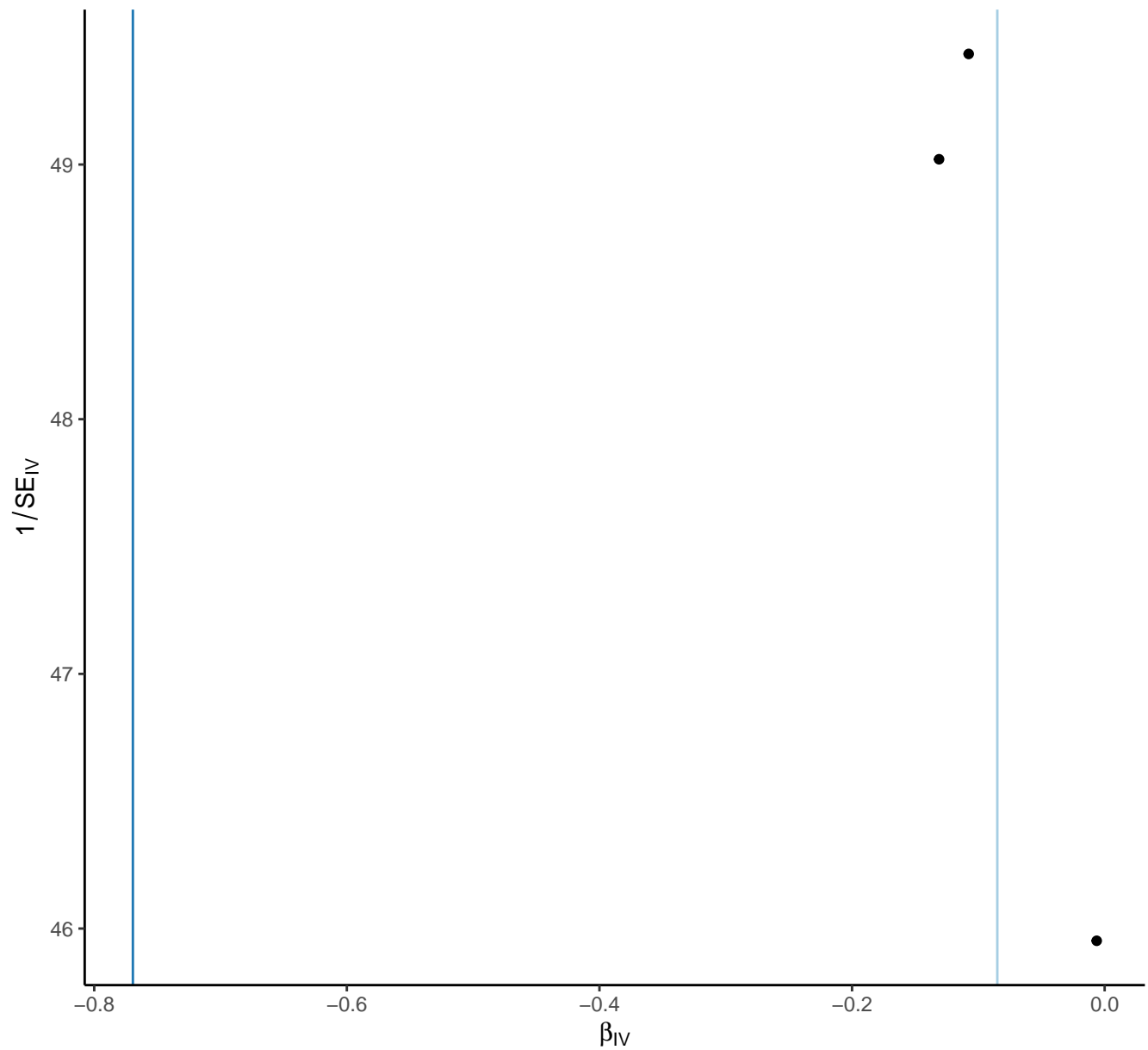




MR Method

Inverse variance weighted

MR Egger

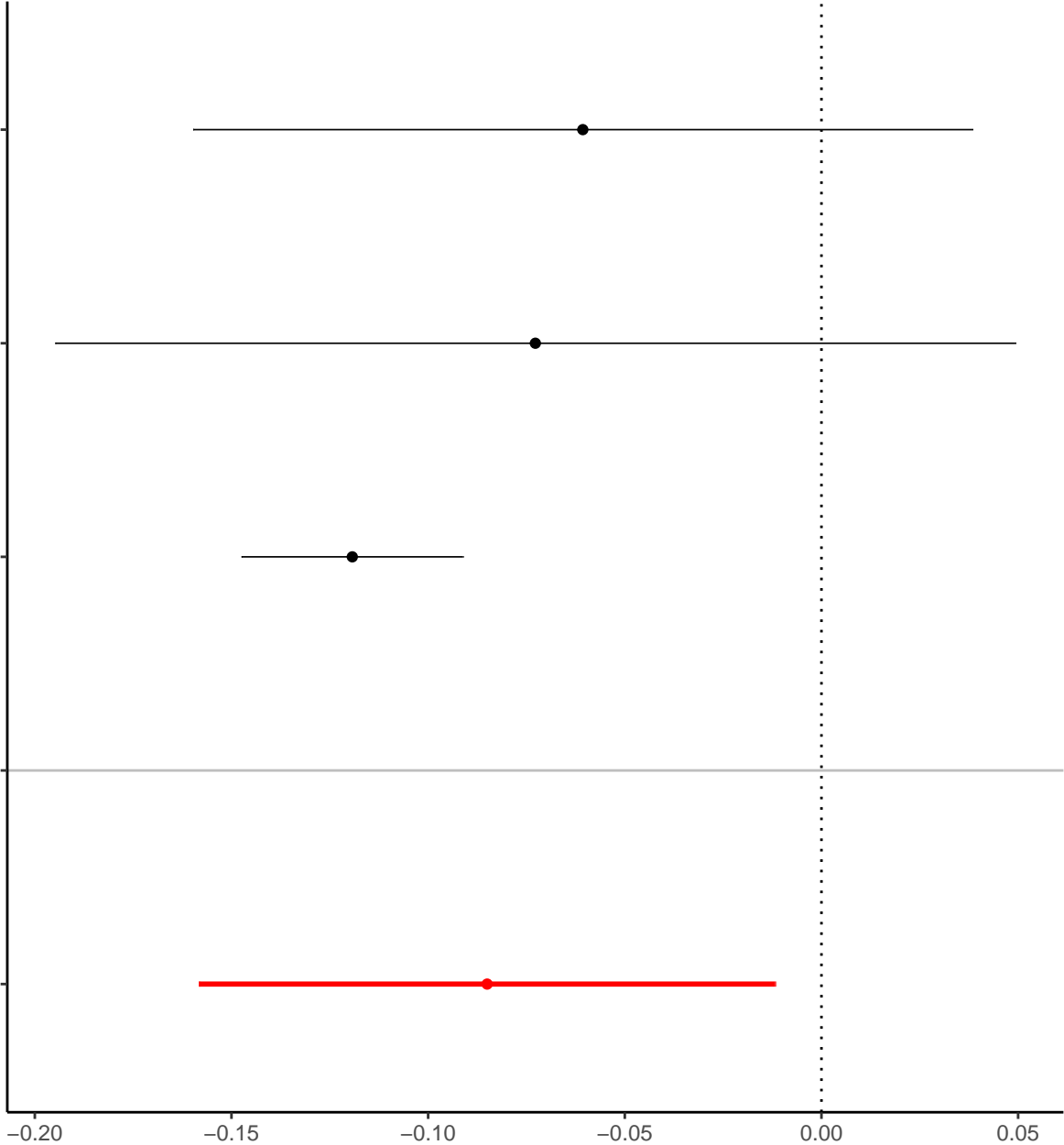


rs7756992

rs1111875

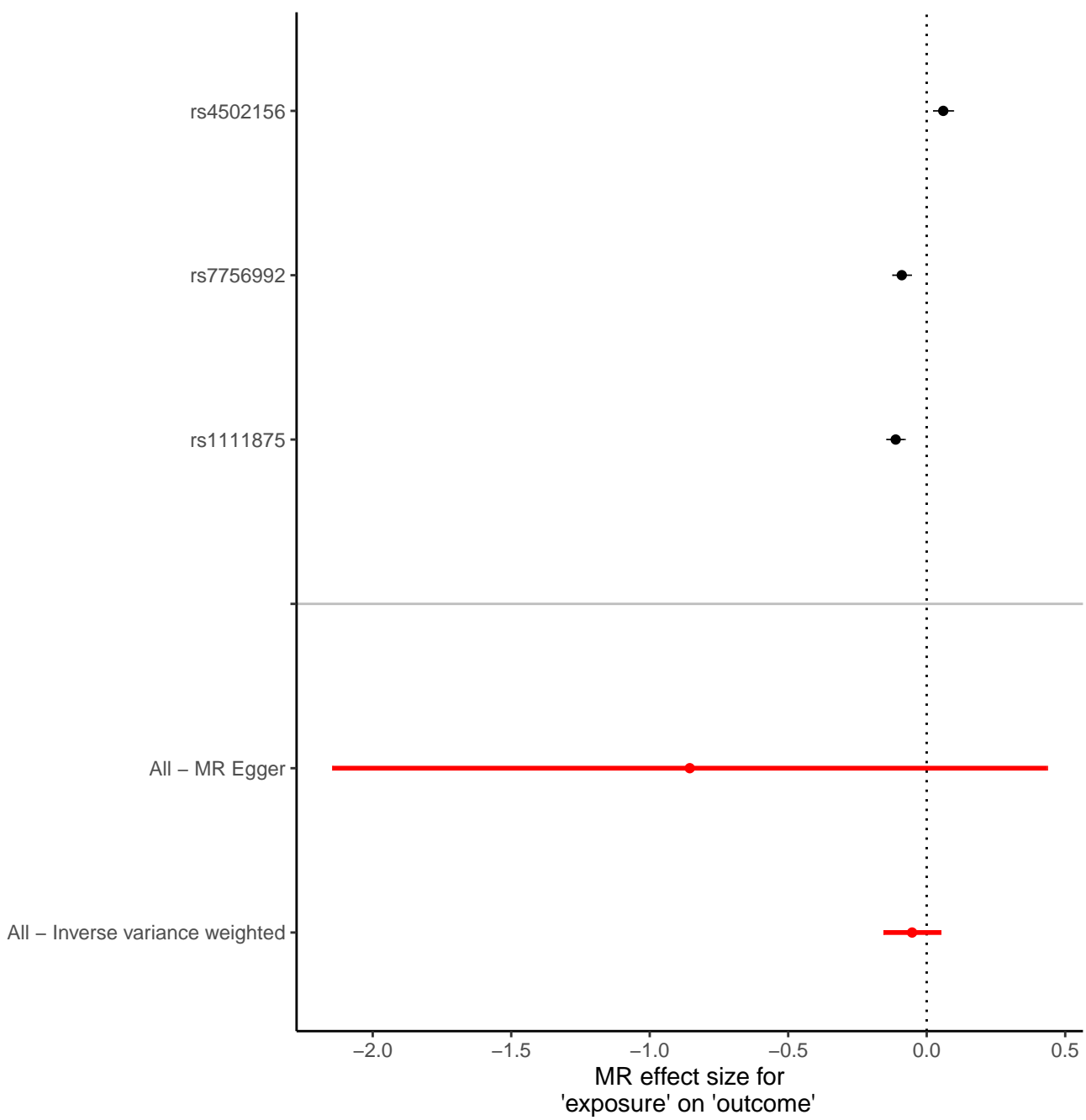
rs4502156

All



MR leave-one-out sensitivity analysis for 'exposure' on 'triglycerides || id:ieu-b-111'

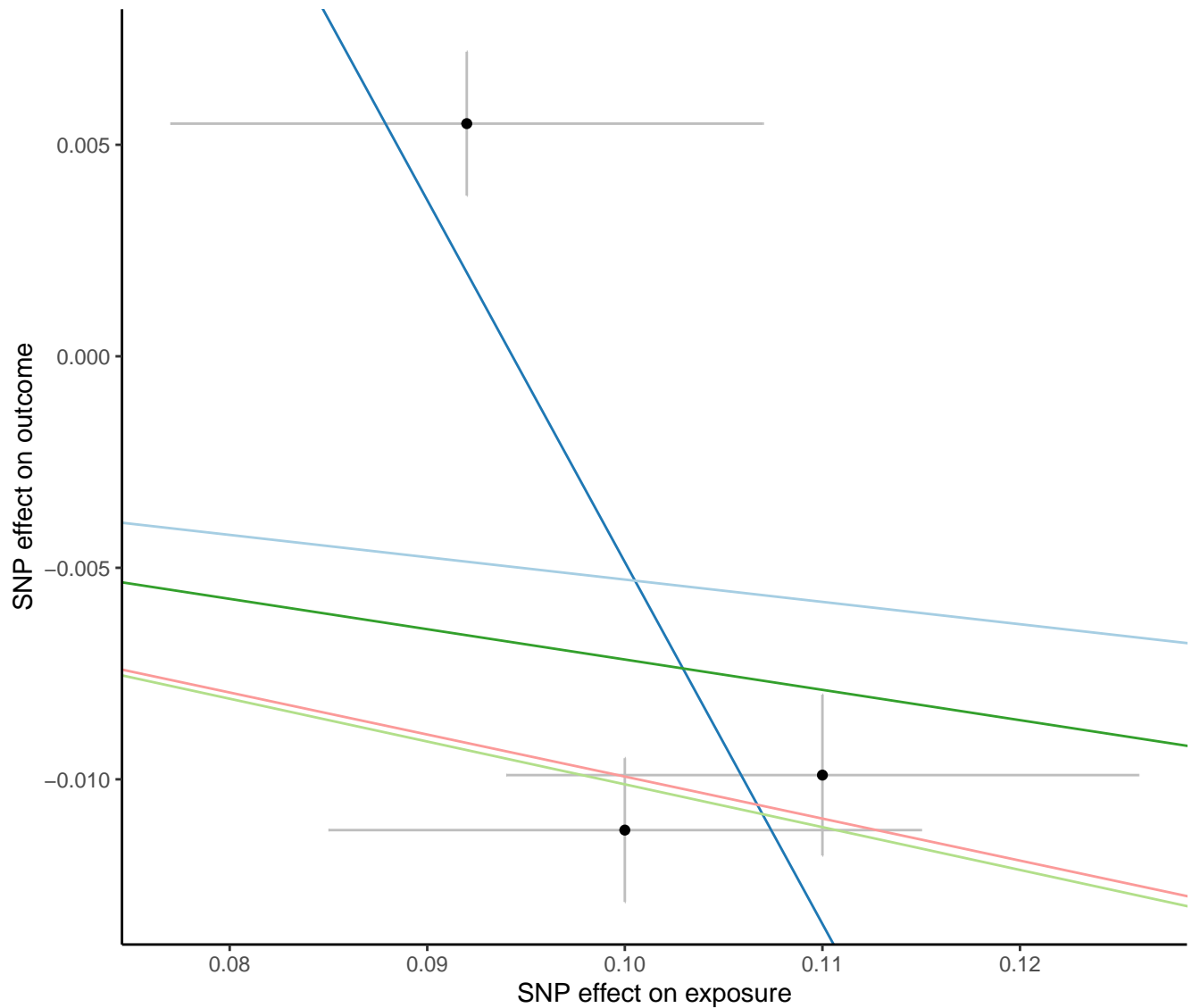
**Exposure: Corrected Insulin Response adjusted for ISI, Outcome: Waist-Hip Ratio adjusted for Body Mass Index**  
**Figures of Forest Plot, Comparison of Different MR Methods, Funnel Plot and Leave-one-out Sensitivity Analysis**





MR Test

- Inverse variance weighted
- MR Egger
- Simple mode
- Weighted median
- Weighted mode



MR Method

Inverse variance weighted

MR Egger

