Adiposity and TG

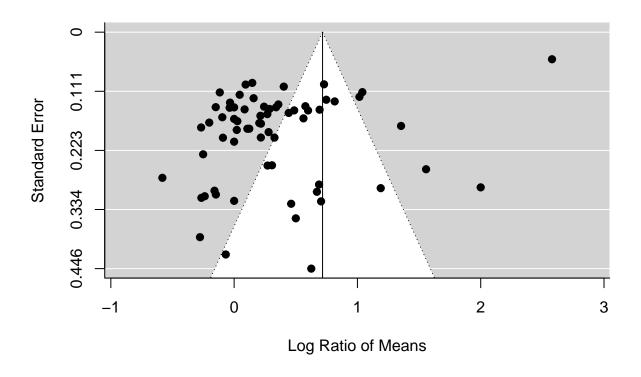
Hamza

6 October 2019

```
load(file="Effect_Size_All_Traits.RData")
load(file="Effect_Size_All_Traits_lnCVR.RData")
Adiposity_lnRR <- subset(Effect_Size_All_Traits, Effect_Size_All_Traits$Trait == "Adiposity")
Adiposity_lnRR_MG <- subset(Adiposity_lnRR, Adiposity_lnRR$Exposure_Type == "Multigenerational")
Adiposity_lnRR_OF <- subset(Adiposity_lnRR, Adiposity_lnRR$Exposure_Type == "One off")</pre>
```

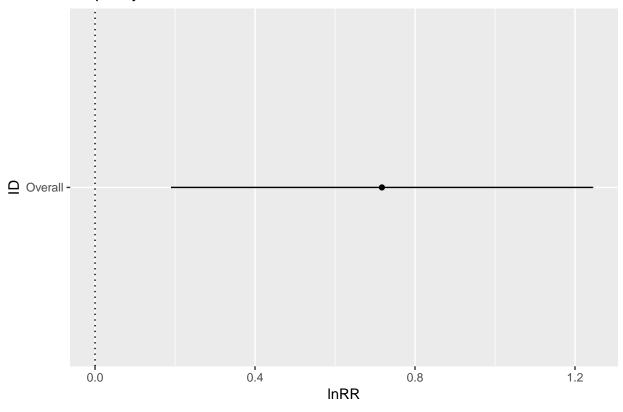
Adiposity

```
#Overall analysis, not split
Adiposity_overall_lnRR_Omods <- rma.mv(yi, vi,random = list(~1 | Paper_ID,~1 | Cohort_ID,~1 | ES_ID), data=Ad
summary(Adiposity_overall_lnRR_Omods)
## Multivariate Meta-Analysis Model (k = 65; method: REML)
##
    logLik Deviance
                           AIC
                                     BIC
                                              AICc
## -21.9638
             43.9275 51.9275
                                 60.5631
                                           52.6055
## Variance Components:
##
                       sqrt nlvls fixed
                                              factor
##
              estim
## sigma^2.1 0.6663 0.8163
                                            Paper_ID
                                10
## sigma^2.2 0.0547 0.2339
                                25
                                       no Cohort_ID
## sigma^2.3 0.0212 0.1457
                                65
                                               ES_ID
## Test for Heterogeneity:
## Q(df = 64) = 2075.5744, p-val < .0001
## Model Results:
##
## estimate
                                     ci.lb
                se
                      zval
                              pval
                                             ci.ub
   0.7172 0.2690 2.6659 0.0077 0.1899 1.2445 **
##
##
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
funnel(Adiposity_overall_lnRR_Omods)
```



```
\#Tibble\ of\ overall\ results
Adiposity_overall_lnRR <- tibble(
  ID = "Overall",
  lnRR = 0.7172,
  ci.lb = 0.1899,
  ci.ub = 1.2445
## Warning: `list_len()` is deprecated as of rlang 0.2.0.
## Please use `new_list()` instead.
## This warning is displayed once per session.
plot_adiposity_overall <- ggplot(Adiposity_overall_lnRR, aes(x=ID, y=lnRR)) +</pre>
  geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
  geom_point(aes(x = ID, y = lnRR), position = position_dodge(0.3))+
  geom_hline(yintercept = 0, lty = "dotted") +
  labs(title ="Adiposity differences, overall", x = "ID", y = "lnRR") +
  coord_flip()
plot_adiposity_overall
```

Adiposity differences, overall



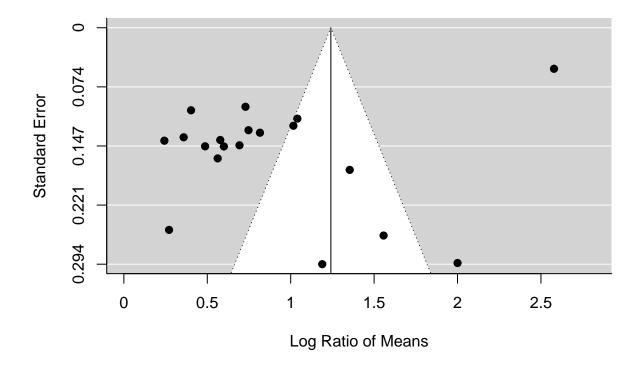
Splitting by exposure type

```
Adiposity_overall_lnRR_MG <- rma.mv(yi, vi,random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID), data=Adiposity_overall_lnRR_MG)
```

```
## Multivariate Meta-Analysis Model (k = 19; method: REML)
##
##
    logLik Deviance
                            AIC
                                     BIC
                                               AICc
## -10.6610
              21.3221
                       29.3221
                                 32.8836
                                            32.3990
## Variance Components:
##
##
                        sqrt nlvls fixed
                                              factor
              estim
## sigma^2.1 0.6268 0.7917
                                 6
                                       no
                                            Paper_ID
## sigma^2.2 0.0000 0.0000
                                 9
                                       no Cohort_ID
## sigma^2.3 0.0654 0.2557
                                19
                                               ES_ID
## Test for Heterogeneity:
## Q(df = 18) = 1024.7727, p-val < .0001
## Model Results:
```

```
##
## estimate se zval pval ci.lb ci.ub
## 1.2407 0.3375 3.6762 0.0002 0.5792 1.9022 ***
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

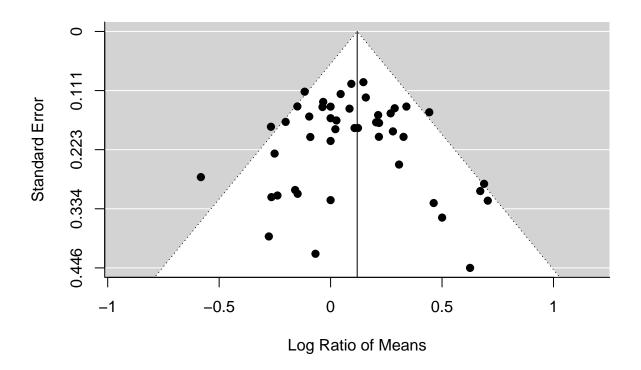
funnel (Adiposity_overall_lnRR_MG)
```



```
Adiposity_overall_lnRR_OF <- rma.mv(yi, vi,random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID), data=Adiposity_overall_lnRR_OF)
```

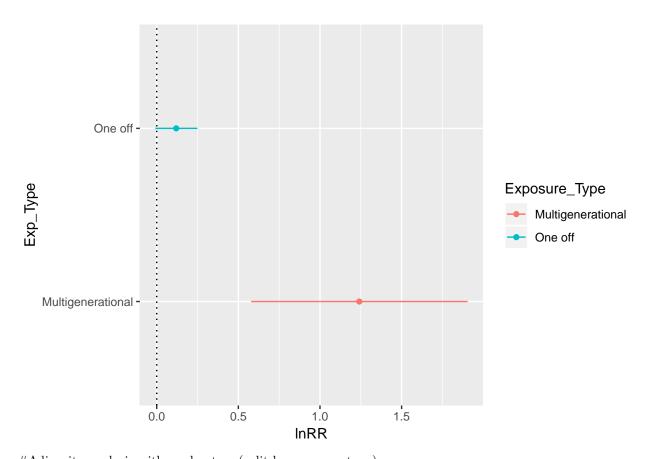
```
##
## Multivariate Meta-Analysis Model (k = 46; method: REML)
##
##
     logLik Deviance
                             AIC
                                        {\tt BIC}
                                                 AICc
                         -4.7588
##
     6.3794 -12.7588
                                    2.4678
                                              -3.7588
##
## Variance Components:
##
##
               {\tt estim}
                         sqrt nlvls fixed
                                                 factor
## sigma^2.1 0.0040 0.0629
                                   6
                                               Paper_ID
                                          no
## sigma^2.2 0.0304 0.1743
                                  16
                                              Cohort_ID
                                         no
                                                  ES_ID
## sigma^2.3 0.0000 0.0000
                                  46
                                         no
```

```
##
## Test for Heterogeneity:
## Q(df = 45) = 64.6623, p-val = 0.0288
## Model Results:
##
## estimate
                      zval
                              pval
                                      ci.lb
                                              ci.ub
                se
    0.1194 0.0649 1.8383 0.0660 -0.0079 0.2466 .
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
funnel (Adiposity_overall_lnRR_OF)
```



```
#Plotting when split by exp type

plot_Adiposity_exp_type <- ggplot(Adiposity_Exp_lnRR, aes(x=Exposure_Type, y=lnRR, colour = Exposure_Type
    geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
    geom_point(aes(x = Exposure_Type, y = lnRR), position = position_dodge(0.3))+
    geom_hline(yintercept = 0, lty = "dotted") +
    labs(x = "Exp_Type", y = "lnRR") +
    coord_flip()
plot_Adiposity_exp_type</pre>
```



A diposity analysis with moderators (split by exposure type)

```
Adiposity_MG_FO <- rma.mv(yi, vi, mods = ~FO_Parent_Exposed-1, random = list(~1 Paper_ID,~1 Cohort_ID,~

## Warning in rma.mv(yi, vi, mods = ~FO_Parent_Exposed - 1, random = list(~1

## | : Redundant predictors dropped from the model.

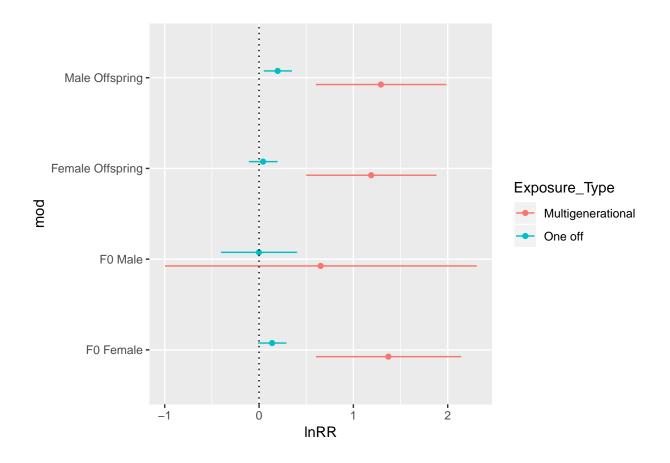
Adiposity_MG_FO

## ## Multivariate Meta-Analysis Model (k = 19; method: REML)
```

```
##
## Variance Components:
##
##
                        sqrt nlvls fixed
                                               factor
               {\tt estim}
## sigma^2.1 0.6981 0.8355
                                  6
                                        no
                                             Paper_ID
## sigma^2.2 0.0000 0.0000
                                  9
                                           Cohort ID
                                       no
## sigma^2.3 0.0653 0.2555
                                                ES ID
                                 19
                                       no
## Test for Residual Heterogeneity:
## QE(df = 17) = 818.7222, p-val < .0001
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 12.8877, p-val = 0.0016
## Model Results:
##
##
                                                                ci.lb
                                                                        ci.ub
                            estimate
                                                zval
                                                        pval
                                          se
## FO_Parent_ExposedFemale
                              1.3709 0.3911
                                             3.5051
                                                      0.0005
                                                               0.6043 2.1375
## F0_Parent_ExposedMale
                              0.6531 0.8420 0.7757 0.4379 -0.9971 2.3034
## FO_Parent_ExposedFemale ***
## F0_Parent_ExposedMale
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_OF_FO <- rma.mv(yi, vi, mods = ~FO_Parent_Exposed-1, random = list(~1 Paper_ID,~1 Cohort_ID,~
## Warning in rma.mv(yi, vi, mods = ~FO_Parent_Exposed - 1, random = list(~1
## | : Redundant predictors dropped from the model.
Adiposity_OF_F0
##
## Multivariate Meta-Analysis Model (k = 46; method: REML)
##
## Variance Components:
##
                        sqrt nlvls fixed
##
                                               factor
               estim
## sigma^2.1 0.0063 0.0791
                                  6
                                             Paper_ID
## sigma^2.2 0.0324 0.1799
                                 16
                                            Cohort_ID
                                        no
## sigma^2.3 0.0000 0.0000
                                                ES_ID
                                 46
                                        no
##
## Test for Residual Heterogeneity:
## QE(df = 44) = 61.4679, p-val = 0.0418
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 3.4316, p-val = 0.1798
##
## Model Results:
##
##
                            estimate
                                          se
                                                 zval
                                                         pval
                                                                 ci.lb
                                               1.8524 0.0640 -0.0080
## F0_Parent_ExposedFemale
                              0.1384 0.0747
```

```
## FO_Parent_ExposedMale
                            -0.0011 0.2041 -0.0055 0.9956 -0.4012
##
                            ci.ub
## FO Parent ExposedFemale 0.2849
## F0_Parent_ExposedMale
                           0.3990
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_MG_Sex <- rma.mv(yi, vi, mods = ~Sex-1, random = list(~1 Paper_ID,~1 Cohort_ID,~1 ES_ID), dat
## Warning in rma.mv(yi, vi, mods = ~Sex - 1, random = list(~1 | Paper_ID, :
## Redundant predictors dropped from the model.
Adiposity_MG_Sex
##
## Multivariate Meta-Analysis Model (k = 19; method: REML)
##
## Variance Components:
##
                       sqrt nlvls fixed
##
                                              factor
              estim
                                           Paper_ID
## sigma^2.1 0.6373 0.7983
                                6
## sigma^2.2 0.0000 0.0000
                                 9
                                       no Cohort_ID
## sigma^2.3 0.0701 0.2647
                                19
                                               ES_ID
##
## Test for Residual Heterogeneity:
## QE(df = 17) = 845.9961, p-val < .0001
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 13.6125, p-val = 0.0011
##
## Model Results:
##
##
             estimate
                           se
                                 zval
                                         pval
                                               ci.lb
                                                       ci.ub
             1.1890 0.3512 3.3850 0.0007 0.5005 1.8774
## SexFemale
## SexMale
               1.2919 0.3513 3.6771 0.0002 0.6033 1.9805
##
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_OF_Sex <- rma.mv(yi, vi, mods = ~Sex-1, random = list(~1 Paper_ID,~1 Cohort_ID,~1 ES_ID), dat
## Warning in rma.mv(yi, vi, mods = ~Sex - 1, random = list(~1 | Paper_ID, :
## Redundant predictors dropped from the model.
Adiposity_OF_Sex
## Multivariate Meta-Analysis Model (k = 46; method: REML)
## Variance Components:
```

```
##
                        sqrt nlvls fixed
##
                                               factor
               estim
## sigma^2.1 0.0080 0.0892
                                6
                                        no
                                             Paper ID
                                        no Cohort_ID
## sigma^2.2 0.0193 0.1388
                                 16
## sigma^2.3 0.0000 0.0000
                                 46
                                        no
                                                ES ID
##
## Test for Residual Heterogeneity:
## QE(df = 44) = 54.1902, p-val = 0.1396
##
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 8.1136, p-val = 0.0173
## Model Results:
##
##
              estimate
                                  zval
                                          pval
                                                  ci.lb
                                                         ci.ub
                            se
## SexFemale
               0.0423 0.0761 0.5554 0.5786
                                               -0.1069 0.1914
## SexMale
                0.1976 0.0746 2.6471 0.0081
                                                 0.0513 0.3438 **
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_Exp_mods_lnRR <- tibble(
  Exposure_Type = c("Multigenerational", "Multigenerational", "One off", "One off", "Multigenerational",
  mod = c("F0 Female", "F0 Male", "F0 Female", "F0 Male", "Female Offspring", "Male Offspring", "Female Of
  lnRR = c(1.3709, 0.6531, 0.1384, -0.0011, 1.1890, 1.2919, 0.0423, 0.1976),
  ci.lb = c(0.6043, -0.9971, -0.0080, -0.4012, 0.5005, 0.6033, -0.1069, 0.0513),
  ci.ub = c(2.1375, 2.3034, 0.2849, 0.3990, 1.8774, 1.9805, 0.1914, 0.3438)
)
plot_Adiposity_lnRR_mods_exp <- ggplot(Adiposity_Exp_mods_lnRR, aes(x=mod, y=lnRR, colour=Exposure_Type
  geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
  geom_point(aes(x = mod, y = lnRR), position = position_dodge(0.3))+
  geom_hline(yintercept = 0, lty = "dotted") +
  labs(x = "mod", y = "lnRR") +
  coord_flip()
plot_Adiposity_lnRR_mods_exp
```



ADiposity lnCVR

```
#Subset Adiposity lnCVR Data
Adiposity_lnCVR <- subset(Effect_Size_All_Traits_lnCVR, Effect_Size_All_Traits_lnCVR$Trait == "Adiposit
Adiposity_lnCVR_MG <- subset(Adiposity_lnCVR, Adiposity_lnCVR$Exposure_Type == "Multigenerational")
Adiposity_lnCVR_OF <- subset(Adiposity_lnCVR, Adiposity_lnCVR$Exposure_Type == "One off")
Adiposity_overall_lnCVR_Omods <- rma.mv(yi, vi, random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID), data=
summary(Adiposity_overall_lnCVR_Omods)
##
## Multivariate Meta-Analysis Model (k = 65; method: REML)
##
##
    logLik Deviance
                            AIC
                                      BIC
                                               AICc
## -44.7927
              89.5854
                        97.5854 106.2210
                                            98.2634
##
## Variance Components:
##
##
                        sqrt nlvls fixed
               estim
                                               factor
```

```
## sigma^2.1 0.0650 0.2549
                                10
                                            Paper_ID
                                       no
## sigma^2.2 0.0348 0.1867
                                25
                                       no Cohort_ID
## sigma^2.3 0.0000 0.0000
                                65
                                       no
                                               ES ID
##
## Test for Heterogeneity:
## Q(df = 64) = 84.9840, p-val = 0.0408
## Model Results:
##
## estimate
                       zval
                               pval
                                       ci.lb
## -0.0529 0.1167 -0.4535
                            0.6502
                                    -0.2818 0.1759
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
#Tibble of overall results
Insulin overall lnCVR <- tibble(</pre>
 ID = "Overall",
 lnCVR = -0.0529,
 ci.lb = -0.2818,
 ci.ub = 0.175
)
```

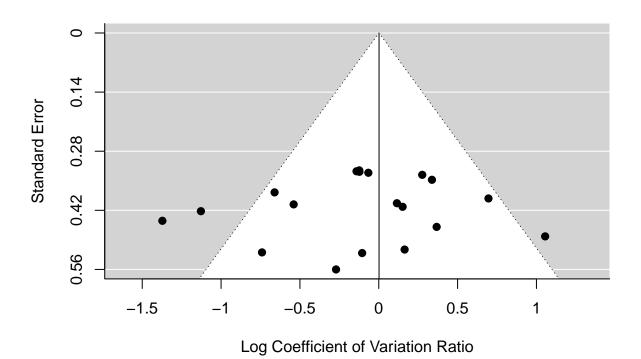
Adiposity lnCVR (split by exposure type)

##

```
Adiposity_overall_lnCVR_MG <- rma.mv(yi, vi,random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID), data=Adip
summary(Adiposity_overall_lnCVR_MG)
## Multivariate Meta-Analysis Model (k = 19; method: REML)
##
    logLik Deviance
                            AIC
                                      BIC
                                               AICc
##
## -15.0157
              30.0314
                        38.0314
                                  41.5929
                                            41.1083
##
## Variance Components:
##
##
                        sqrt nlvls fixed
                                               factor
               estim
## sigma^2.1 0.0000 0.0000
                                             Paper_ID
                                  6
## sigma^2.2 0.1139 0.3376
                                  9
                                            Cohort_ID
                                        no
## sigma^2.3 0.0630 0.2510
                                 19
                                                ES_ID
##
## Test for Heterogeneity:
## Q(df = 18) = 33.8343, p-val = 0.0132
## Model Results:
##
## estimate
                               pval
                                       ci.lb
                                               ci.ub
                 se
                       zval
    0.0018 0.1733 0.0101 0.9919 -0.3380 0.3415
```

```
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

funnel (Adiposity_overall_lnCVR_MG)
```

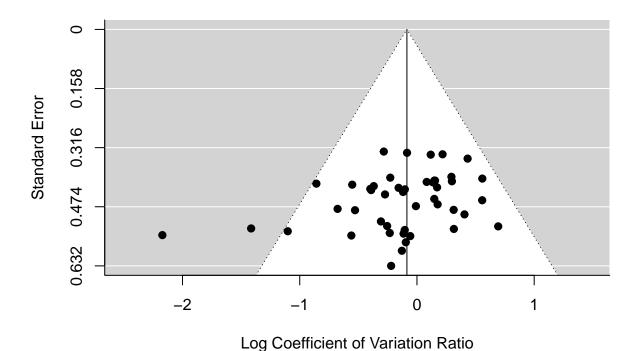


Adiposity_overall_lnCVR_OF <- rma.mv(yi, vi,random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID), data=Adip summary(Adiposity_overall_lnCVR_OF)

```
##
## Multivariate Meta-Analysis Model (k = 46; method: REML)
##
##
     logLik Deviance
                             AIC
                                       BIC
                                                AICc
## -28.5396
              57.0791
                         65.0791
                                   72.3058
                                             66.0791
##
## Variance Components:
##
##
               estim
                         sqrt nlvls
                                     fixed
                                                factor
## sigma^2.1 0.0577
                      0.2403
                                   6
                                              Paper_ID
## sigma^2.2
              0.0368
                      0.1917
                                  16
                                             Cohort_ID
                                         no
## sigma^2.3 0.0000 0.0000
                                  46
                                                 ES_ID
                                         no
##
## Test for Heterogeneity:
## Q(df = 45) = 51.1494, p-val = 0.2450
##
```

```
## Model Results:
##
## estimate se zval pval ci.lb ci.ub
## -0.0857 0.1430 -0.5992 0.5490 -0.3660 0.1946
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

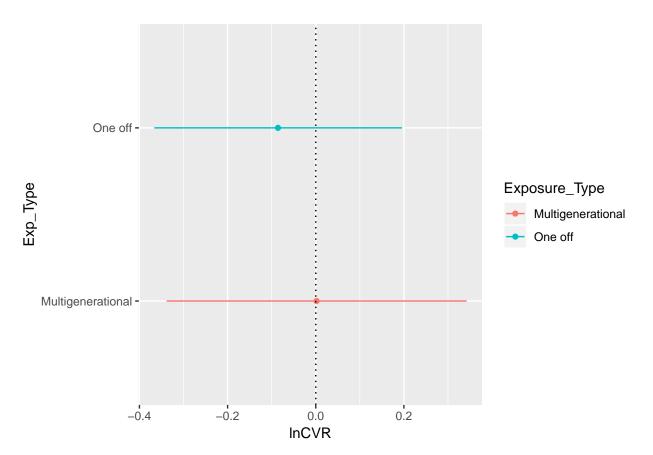
funnel (Adiposity_overall_lnCVR_OF)
```



Adiposity_Exp_lnCVR <- tibble(
 Exposure_Type = c("Multigenerational", "One off"),
 lnCVR = c(0.0018,-0.0857),
 ci.lb = c(-0.3380,-0.3660),
 ci.ub = c(0.3415,0.1946)
)</pre>
Adiposity_Exp_lnCVR

```
#Plotting when split by exp type

plot_Adiposity_exp_type_lnCVR <- ggplot(Adiposity_Exp_lnCVR, aes(x=Exposure_Type, y=lnCVR, colour = Exp
    geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
    geom_point(aes(x = Exposure_Type, y = lnCVR), position = position_dodge(0.3))+
    geom_hline(yintercept = 0, lty = "dotted") +
    labs(x = "Exp_Type", y = "lnCVR") +
    coord_flip()
plot_Adiposity_exp_type_lnCVR</pre>
```



Adiposity analysis with moderators (split by exposure type)

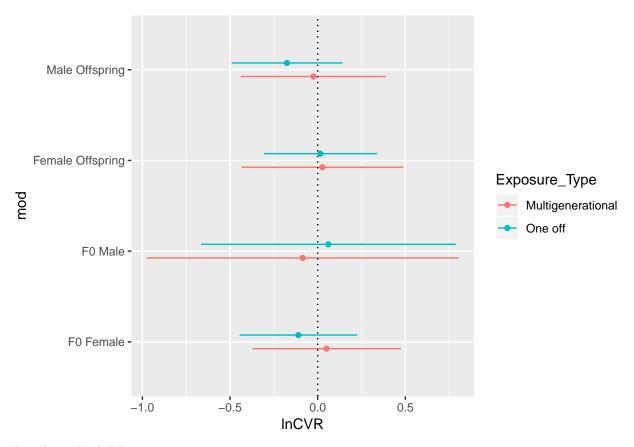
```
Adiposity_MG_FO_lnCVR <- rma.mv(yi, vi, mods = ~FO_Parent_Exposed-1, random = list(~1|Paper_ID,~1|Cohor ## Warning in rma.mv(yi, vi, mods = ~FO_Parent_Exposed - 1, random = list(~1 ## | : Redundant predictors dropped from the model.

Adiposity_MG_FO_lnCVR ## ## Multivariate Meta-Analysis Model (k = 19; method: REML)
```

```
##
## Variance Components:
##
##
                        sqrt nlvls fixed
                                               factor
               {\tt estim}
## sigma^2.1 0.0000 0.0000
                                  6
                                        no
                                             Paper_ID
## sigma^2.2 0.1770 0.4208
                                  9
                                            Cohort ID
                                        no
## sigma^2.3 0.0597 0.2443
                                                ES ID
                                 19
                                        no
## Test for Residual Heterogeneity:
## QE(df = 17) = 33.8068, p-val = 0.0089
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 0.0905, p-val = 0.9557
## Model Results:
##
##
                            estimate
                                                                 ci.lb
                                                 zval
                                                         pval
                                          se
## FO_Parent_ExposedFemale
                              0.0503 0.2150
                                               0.2338 0.8152
                                                               -0.3711
## F0_Parent_ExposedMale
                             -0.0858   0.4530   -0.1895   0.8497   -0.9736
                             ci.ub
## FO_Parent_ExposedFemale 0.4717
## F0_Parent_ExposedMale
                            0.8020
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_OF_FO_lnCVR <- rma.mv(yi, vi, mods = ~FO_Parent_Exposed-1, random = list(~1 Paper_ID,~1 Cohor
## Warning in rma.mv(yi, vi, mods = ~FO_Parent_Exposed - 1, random = list(~1
## | : Redundant predictors dropped from the model.
Adiposity_OF_FO_lnCVR
##
## Multivariate Meta-Analysis Model (k = 46; method: REML)
##
## Variance Components:
##
                        sqrt nlvls fixed
##
                                               factor
               estim
## sigma^2.1 0.0730 0.2702
                                  6
                                             Paper_ID
## sigma^2.2 0.0441 0.2100
                                 16
                                            Cohort_ID
                                        no
## sigma^2.3 0.0000 0.0000
                                                ES_ID
                                 46
                                        no
##
## Test for Residual Heterogeneity:
## QE(df = 44) = 49.6318, p-val = 0.2588
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 0.4459, p-val = 0.8002
##
## Model Results:
##
##
                            estimate
                                          se
                                                 zval
                                                         pval
## F0_Parent_ExposedFemale
                           -0.1104 0.1705 -0.6478 0.5171 -0.4445
```

```
## F0_Parent_ExposedMale
                             0.0600 0.3703 0.1619 0.8713 -0.6657
##
                            ci.ub
## FO Parent ExposedFemale
                           0.2237
## F0_Parent_ExposedMale
                           0.7857
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_MG_Sex_lnCVR <- rma.mv(yi, vi, mods = ~Sex-1, random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID
Adiposity_MG_Sex_lnCVR
##
## Multivariate Meta-Analysis Model (k = 19; method: REML)
##
## Variance Components:
##
##
                       sqrt nlvls fixed
                                              factor
              estim
## sigma^2.1 0.0000 0.0000
                                 6
                                            Paper_ID
                                       no
## sigma^2.2 0.1121 0.3347
                                 9
                                       no
                                           Cohort_ID
                                               ES ID
## sigma^2.3 0.0795 0.2820
                                19
                                       no
## Test for Residual Heterogeneity:
## QE(df = 17) = 32.9481, p-val = 0.0114
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 0.0371, p-val = 0.9816
##
## Model Results:
##
##
             estimate
                                                  ci.lb
                                                          ci.ub
                           se
                                  zval
                                          pval
                                                -0.4326 0.4860
## SexFemale
               0.0267
                       0.2343
                                0.1139 0.9093
## SexMale
              -0.0252 0.2105 -0.1198 0.9046 -0.4377 0.3873
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_OF_Sex_lnCVR <- rma.mv(yi, vi, mods = ~Sex-1, random = list(~1|Paper_ID,~1|Cohort_ID,~1|ES_ID
Adiposity_OF_Sex_lnCVR
##
## Multivariate Meta-Analysis Model (k = 46; method: REML)
## Variance Components:
##
                                              factor
##
              estim
                       sqrt nlvls fixed
## sigma^2.1 0.0592 0.2433
                                 6
                                       no
                                            Paper_ID
## sigma^2.2
             0.0329 0.1813
                                16
                                           Cohort_ID
                                       no
## sigma^2.3 0.0000 0.0000
                                46
                                               ES_ID
                                       no
##
## Test for Residual Heterogeneity:
## QE(df = 44) = 49.6421, p-val = 0.2585
```

```
##
## Test of Moderators (coefficient(s) 1:2):
## QM(df = 2) = 1.8913, p-val = 0.3884
##
## Model Results:
##
                                     estimate
                                                                                             zval
                                                                                                                  pval
                                                                                                                                   ci.lb
                                                                                                                                                             ci.ub
                                                                           se
                                        0.0154 0.1640
                                                                                    0.0940 0.9251 -0.3060 0.3368
## SexFemale
## SexMale
                                       -0.1752 0.1605 -1.0916 0.2750 -0.4899 0.1394
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
Adiposity_Exp_mods_lnCVR <- tibble(
     Exposure_Type = c("Multigenerational", "Multigenerational", "One off", "One off", "Multigenerational",
     mod = c("F0 Female", "F0 Male", "F0 Female", "F0 Male", "Female Offspring", "Male Offspring", "Female Offspring", "Female Offspring", "Fomale Offspring", "Fomale
    lnCVR = c(0.0503, -0.0858, -0.1104, 0.0600, 0.0267, -0.0252, 0.0154, -0.1752),
     ci.lb = c(-0.3711, -0.9736, -0.4445, -0.6657, -0.4326, -0.4377, -0.3060, -0.4899),
     ci.ub = c(0.4717, 0.8020, 0.2237, 0.7857, 0.4860, 0.3873, 0.3368, 0.1394)
plot_Adiposity_lnCVR_mods_exp <- ggplot(Adiposity_Exp_mods_lnCVR, aes(x=mod, y=lnCVR, colour=Exposure_T
     geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
     geom_point(aes(x = mod, y = lnCVR), position = position_dodge(0.3))+
     geom_hline(yintercept = 0, lty = "dotted") +
     labs(x = "mod", y = "lnCVR") +
     coord_flip()
plot_Adiposity_lnCVR_mods_exp
```



#Triglycerides lnRR

##

```
#Subsetting Data for lnRR
Triglycerides_lnRR <- subset(Effect_Size_All_Traits, Effect_Size_All_Traits$Trait == "Triglycerides")
Triglycerides_lnRR_MG <- subset(Triglycerides_lnRR, Triglycerides_lnRR$Exposure_Type == "Multigeneration Triglycerides_lnRR_OF <- subset(Triglycerides_lnRR, Triglycerides_lnRR$Exposure_Type == "One off")
#Subsetting Data for lmCVR
Triglycerides_lnCVR <- subset(Effect_Size_All_Traits_lnCVR, Effect_Size_All_Traits_lnCVR$Trait == "Triglycerides_lnCVR_MG <- subset(Triglycerides_lnCVR, Triglycerides_lnCVR$Exposure_Type == "Multigeneration")
Triglycerides_lnCVR_MG <- subset(Triglycerides_lnCVR, Triglycerides_lnCVR$Exposure_Type == "Multigeneration")</pre>
```

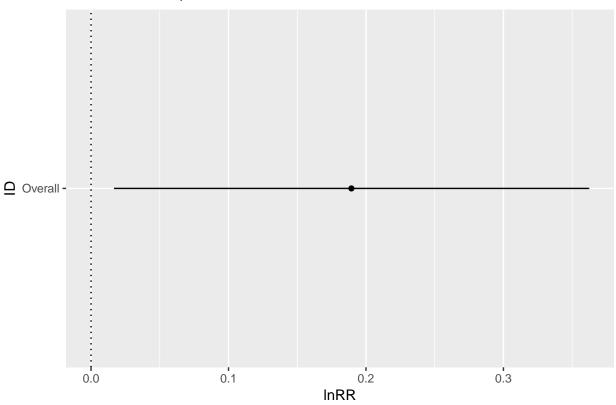
Overall Triglycerides analysis

```
TG_overall_lnRR_Omods <- rma.mv(yi, vi,random = list(~1|Cohort_ID,~1|ES_ID), data=Triglycerides_lnRR, m
summary(TG_overall_lnRR_Omods)

##
## Multivariate Meta-Analysis Model (k = 33; method: REML)</pre>
```

```
logLik Deviance
                           AIC
                                     BIC
                                               AICc
## -13.9786
              27.9572 33.9572
                                 38.3544
                                           34.8143
##
## Variance Components:
##
##
                        sqrt nlvls fixed
                                              factor
              estim
## sigma^2.1 0.1439 0.3793
                                22
                                           Cohort ID
                                       no
## sigma^2.2 0.0092 0.0961
                                               ES_ID
                                33
                                       no
##
## Test for Heterogeneity:
## Q(df = 32) = 262.1206, p-val < .0001
## Model Results:
##
## estimate
               se
                      zval
                              pval
                                     ci.lb ci.ub
   0.1894 0.0881 2.1502 0.0315 0.0168 0.3621 *
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
TG_overall_lnRR <- tibble(</pre>
 ID = "Overall",
 lnRR = 0.1894,
  ci.lb = 0.0168,
  ci.ub = 0.3621
plot_tg_overall <- ggplot(TG_overall_lnRR, aes(x=ID, y=lnRR)) +</pre>
  geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
  geom_point(aes(x = ID, y = lnRR), position = position_dodge(0.3))+
  geom_hline(yintercept = 0, lty = "dotted") +
  labs(title ="TG differences, overall", x = "ID", y = "lnRR") +
  coord_flip()
plot_tg_overall
```

TG differences, overall



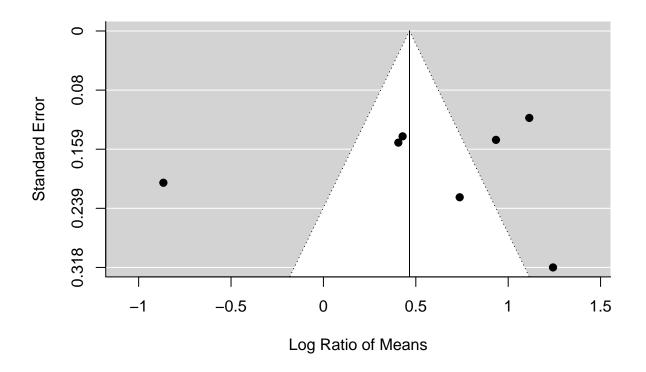
#Triglycerides analysis (split by exposure type)

```
Triglycerides_overall_lnRR_MG <- rma.mv(yi, vi,random = list(~1|Cohort_ID,~1|ES_ID), data=Triglycerides
summary(Triglycerides_overall_lnRR_MG)</pre>
```

```
##
## Multivariate Meta-Analysis Model (k = 7; method: REML)
##
##
    logLik Deviance
                         AIC
                                          AICc
                                  BIC
##
   -4.7574
             9.5149
                     15.5149
                              14.8902
                                       27.5149
##
## Variance Components:
##
                     sqrt nlvls fixed
##
             estim
                                          factor
## sigma^2.1 0.6116 0.7821
                              5
                                       Cohort_ID
## sigma^2.2 0.0038 0.0618
                                           ES_ID
                              7
                                    no
##
## Test for Heterogeneity:
## Q(df = 6) = 83.2645, p-val < .0001
##
## Model Results:
##
## estimate
               se
                    zval
                            pval
                                   ci.lb
##
    ##
```

```
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

funnel (Triglycerides_overall_lnRR_MG)
```

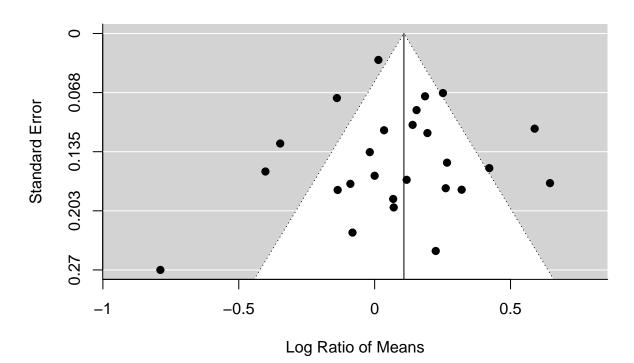


Triglycerides_overall_lnRR_OF <- rma.mv(yi, vi,random = list(~1|Cohort_ID,~1|ES_ID), data=Triglycerides
summary(Triglycerides_overall_lnRR_OF)</pre>

```
##
## Multivariate Meta-Analysis Model (k = 26; method: REML)
##
##
     logLik Deviance
                            AIC
                                       BIC
                                                AICc
    -2.6854
               5.3709
                        11.3709
                                   15.0275
                                             12.5137
##
##
## Variance Components:
##
               estim
                        sqrt
                              nlvls
                                     fixed
                                                factor
## sigma^2.1 0.0390 0.1974
                                  17
                                             Cohort_ID
                                         no
## sigma^2.2 0.0114 0.1067
                                  26
                                                 ES_ID
                                         no
## Test for Heterogeneity:
## Q(df = 25) = 101.6759, p-val < .0001
## Model Results:
```

```
##
## estimate se zval pval ci.lb ci.ub
## 0.1077 0.0605 1.7797 0.0751 -0.0109 0.2262 .
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

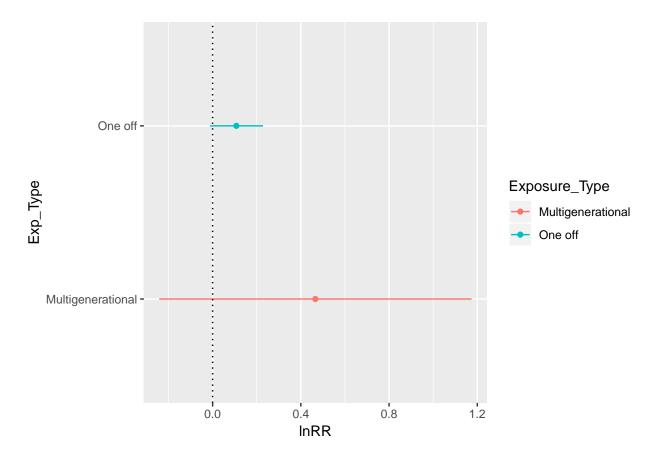
funnel (Triglycerides_overall_lnRR_OF)



```
Triglycerides_Exp_lnRR <- tibble(</pre>
  Exposure_Type = c("Multigenerational", "One off"),
  lnRR = c(0.4658, 0.1077),
  ci.lb = c(-0.2411, -0.0109),
  ci.ub = c(1.1727, 0.2262)
)
Triglycerides_Exp_lnRR
## # A tibble: 2 x 4
                                ci.lb ci.ub
##
     Exposure_Type
                        lnRR
                        <dbl>
                                <dbl> <dbl>
## 1 Multigenerational 0.466 -0.241 1.17
                       0.108 -0.0109 0.226
## 2 One off
```

```
#Plotting when split by exp type

plot_Triglycerides_exp_type <- ggplot(Triglycerides_Exp_lnRR, aes(x=Exposure_Type, y=lnRR, colour = Exp
    geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
    geom_point(aes(x = Exposure_Type, y = lnRR), position = position_dodge(0.3))+
    geom_hline(yintercept = 0, lty = "dotted") +
    labs(x = "Exp_Type", y = "lnRR") +
    coord_flip()
plot_Triglycerides_exp_type</pre>
```



${\bf Trigly cerides}\ {\bf lnCVR}$

##

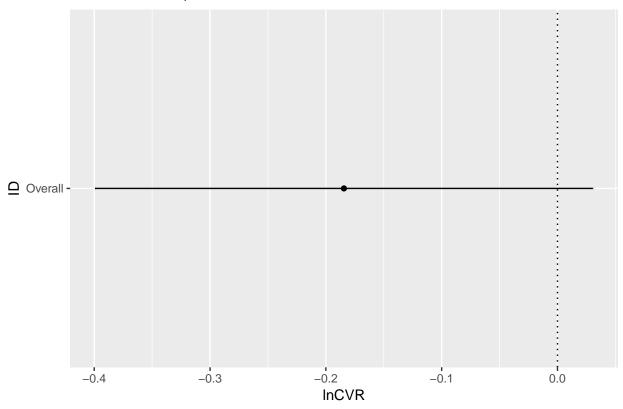
Overall Triglycerides analysis

Multivariate Meta-Analysis Model (k = 33; method: REML)

```
TG_overall_lnCVR_Omods <- rma.mv(yi, vi,random = list(~1|Cohort_ID,~1|ES_ID), data=Triglycerides_lnCVR,
summary(TG_overall_lnCVR_Omods)</pre>
```

```
logLik Deviance
                           AIC
                                      BIC
                                               AICc
## -26.2586
             52.5171
                       58.5171
                                  62.9143
                                            59.3743
##
## Variance Components:
##
##
                        sqrt nlvls fixed
                                               factor
               estim
## sigma^2.1 0.1394 0.3734
                                 22
                                           Cohort ID
                                        no
## sigma^2.2 0.0000 0.0001
                                                ES_ID
                                 33
                                        no
##
## Test for Heterogeneity:
## Q(df = 32) = 56.1655, p-val = 0.0052
## Model Results:
##
## estimate
                se
                        zval
                                pval
                                        ci.lb
                                              ci.ub
## -0.1844 0.1096 -1.6820 0.0926 -0.3993 0.0305
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
TG_overall_lnCVR <- tibble(</pre>
 ID = "Overall",
 lnCVR = -0.1844,
  ci.lb = -0.3993,
  ci.ub = 0.0305
plot_tg_overall_lnCVR <- ggplot(TG_overall_lnCVR, aes(x=ID, y=lnCVR)) +</pre>
  geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
  geom_point(aes(x = ID, y = lnCVR), position = position_dodge(0.3))+
  geom_hline(yintercept = 0, lty = "dotted") +
  labs(title ="TG differences, overall", x = "ID", y = "lnCVR") +
  coord_flip()
plot_tg_overall_lnCVR
```

TG differences, overall



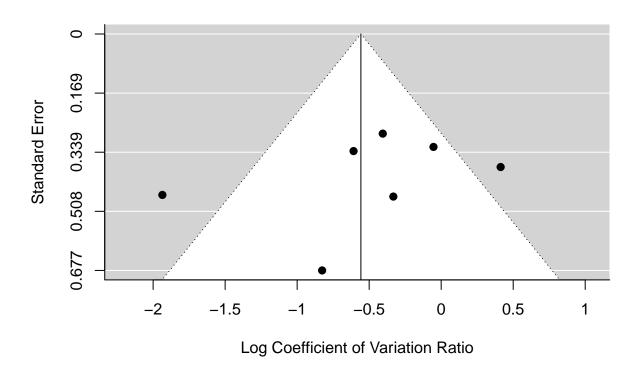
#Triglycerides analysis (split by exposure type)

```
Triglycerides_overall_lnCVR_MG <- rma.mv(yi, vi,random = list(~1|Cohort_ID,~1|ES_ID), data=Triglyceride
summary(Triglycerides_overall_lnCVR_MG)</pre>
```

```
##
## Multivariate Meta-Analysis Model (k = 7; method: REML)
##
##
     logLik Deviance
                            AIC
                                       BIC
                                                AICc
##
   -6.0827
              12.1653
                        18.1653
                                  17.5406
                                             30.1653
##
## Variance Components:
##
##
               {\tt estim}
                        sqrt nlvls fixed
                                                factor
## sigma^2.1 0.5408 0.7354
                                  5
                                             Cohort_ID
## sigma^2.2 0.0000 0.0000
                                                 ES_ID
                                  7
                                        no
##
## Test for Heterogeneity:
## Q(df = 6) = 17.5981, p-val = 0.0073
##
## Model Results:
##
## estimate
                 se
                        zval
                                pval
                                         ci.lb
  -0.5575  0.3744  -1.4891  0.1365  -1.2913  0.1763
##
```

```
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

funnel (Triglycerides_overall_lnCVR_MG)
```

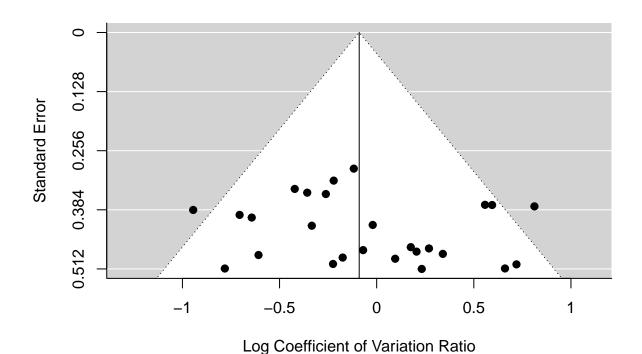


Triglycerides_overall_lnCVR_OF <- rma.mv(yi, vi,random = list(~1|Cohort_ID,~1|ES_ID), data=Triglyceride
summary(Triglycerides_overall_lnCVR_OF)</pre>

```
##
## Multivariate Meta-Analysis Model (k = 26; method: REML)
##
     logLik Deviance
                            AIC
                                       BIC
                                                AICc
## -17.4330
              34.8660
                        40.8660
                                   44.5226
                                             42.0089
##
## Variance Components:
##
               estim
                        sqrt
                              nlvls
                                     fixed
                                                factor
## sigma^2.1 0.0712 0.2668
                                             Cohort_ID
                                  17
                                         no
## sigma^2.2 0.0000
                      0.0000
                                  26
                                                 ES_ID
                                         no
## Test for Heterogeneity:
## Q(df = 25) = 34.2371, p-val = 0.1029
## Model Results:
```

```
##
## estimate se zval pval ci.lb ci.ub
## -0.0900 0.1057 -0.8516 0.3944 -0.2972 0.1171
##
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1

funnel (Triglycerides_overall_lnCVR_OF)
```



Triglycerides_Exp_lnCVR <- tibble(
 Exposure_Type = c("Multigenerational", "One off"),
 lnCVR = c(-0.5575,-0.0900),
 ci.lb = c(-1.2913,-0.2972),
 ci.ub = c(0.1763,0.1171)
)</pre>
Triglycerides_Exp_lnCVR

```
#Plotting when split by exp type

plot_Triglycerides_exp_type <- ggplot(Triglycerides_Exp_lnCVR, aes(x=Exposure_Type, y=lnCVR, colour = E
    geom_errorbar(aes(ymin=ci.lb, ymax=ci.ub), width = 0, position = position_dodge(0.3)) +
    geom_point(aes(x = Exposure_Type, y = lnCVR), position = position_dodge(0.3))+
    geom_hline(yintercept = 0, lty = "dotted") +
    labs(x = "Exp_Type", y = "lnCVR") +
    coord_flip()
plot_Triglycerides_exp_type</pre>
```

