

RStudio Markdown File: Working memory and reaction time variability mediate the relationship between polygenic risk and ADHD traits in a general population sample

Load packages

```
library(data.table)
library(ggplot2)
library(ggpubr)
library(visreg)
library(mice)
```

```
##  
## Attaching package: 'mice'
```

```
## The following object is masked from 'package:stats':  
##  
##     filter
```

```
## The following objects are masked from 'package:base':  
##  
##     cbind, rbind
```

```
library(lme4)
```

```
## Loading required package: Matrix
```

```
## Registered S3 methods overwritten by 'lme4':  
##   method           from  
##   cooks.distance.influence.merMod car  
##   influence.merMod      car  
##   dfbeta.influence.merMod    car  
##   dfbetas.influence.merMod   car
```

```
library(lmerTest)
```

```
##  
## Attaching package: 'lmerTest'
```

```
## The following object is masked from 'package:lme4':  
##  
##     lmer
```

```
## The following object is masked from 'package:stats':  
##  
##      step
```

```
library(JWileymisc)  
library(multilevelTools)  
library(emmeans)  
library(MeMoBootR)
```

```
## Loading required package: boot
```

```
## Loading required package: diagram
```

```
## Loading required package: shape
```

```
library(FSA)
```

```
## Registered S3 methods overwritten by 'FSA':  
##   method      from  
##   confint.boot car  
##   hist.boot    car
```

```
## ## FSA v0.9.1. See citation('FSA') if used in publication.  
## ## Run fishR() for related website and fishR('IFAR') for related book.
```

```
library(lm.beta)  
library(sjmisc)  
library(pscl)
```

```
## Classes and Methods for R developed in the  
## Political Science Computational Laboratory  
## Department of Political Science  
## Stanford University  
## Simon Jackman  
## hurdle and zeroinfl functions by Achim Zeileis
```

```
library(maczac)
```

```
## Loading required package: MASS
```

```
library(mediation)
```

```
## Loading required package: mvtnorm
```

```
## Loading required package: sandwich
```

```
## mediation: Causal Mediation Analysis  
## Version: 4.5.0
```

```
library(foreign)  
library(MASS)  
library(AER)
```

```
## Loading required package: car
```

```
## Loading required package: carData
```

```
##  
## Attaching package: 'car'
```

```
## The following object is masked from 'package:FSA':  
##  
##     bootCase
```

```
## The following object is masked from 'package:boot':  
##  
##     logit
```

```
## Loading required package: lmtest
```

```
## Loading required package: zoo
```

```
##  
## Attaching package: 'zoo'
```

```
## The following objects are masked from 'package:base':  
##  
##     as.Date, as.Date.numeric
```

```
## Loading required package: survival
```

```
##  
## Attaching package: 'survival'
```

```
## The following object is masked from 'package:boot':  
##  
##     aml
```

```
library(boot)  
library(performance)  
library(MBESS)
```

```
##  
## Attaching package: 'MBESS'
```

```
## The following objects are masked from 'package:JWileymisc':  
##  
##     cor2cov, smd
```

```
library(EnvStats)
```

```
##  
## Attaching package: 'EnvStats'
```

```
## The following object is masked from 'package:MBESS':  
##  
##     CV
```

```
## The following object is masked from 'package:car':  
##  
##     qqPlot
```

```
## The following object is masked from 'package:MASS':  
##  
##     boxcox
```

```
## The following object is masked from 'package:Matrix':  
##  
##     print
```

```
## The following objects are masked from 'package:stats':  
##  
##     predict, predict.lm
```

```
## The following object is masked from 'package:base':  
##  
##     print.default
```

```
library(EFA.dimensions)  
library(psych)
```

```
##  
## Attaching package: 'psych'
```

```
## The following object is masked from 'package:MBESS':  
##  
##     cor2cov
```

```
## The following object is masked from 'package:car':  
##  
##      logit
```

```
## The following object is masked from 'package:mediation':  
##  
##      mediate
```

```
## The following object is masked from 'package:FSA':  
##  
##      headtail
```

```
## The following object is masked from 'package:boot':  
##  
##      logit
```

```
## The following object is masked from 'package:JWileymisc':  
##  
##      cor2cov
```

```
## The following objects are masked from 'package:ggplot2':  
##  
##      %+%, alpha
```

```
library(plyr)
```

```
##  
## Attaching package: 'plyr'
```

```
## The following object is masked from 'package:FSA':  
##  
##      mapvalues
```

```
## The following object is masked from 'package:ggpubr':  
##  
##      mutate
```

Load data

```

# Polygenic risk scores (PRS) for ADHD
data_PRS_ADHD <- read.table("ADHD_PRSice_SCORES_AT_ALL_THRESHOLDS.txt",
                             header = TRUE, sep = " ", stringsAsFactors = FALSE)
names(data_PRS_ADHD)[1] <- 'src_subject_id'

# PRS for Autism Spectrum Disorder
data_PRS_ASD <- read.table("AUT_PRSice_SCORES_AT_ALL_THRESHOLDS.txt",
                           header = TRUE, sep = " ", stringsAsFactors = FALSE)
names(data_PRS_ASD)[1] <- 'src_subject_id'

# PRS for Bipolar Disorder
data_PRS_BIP <- read.table("BIP_PRSice_SCORES_AT_ALL_THRESHOLDS.txt",
                           header = TRUE, sep = " ", stringsAsFactors = FALSE)
names(data_PRS_BIP)[1] <- 'src_subject_id'

# PRS for Major Depressive Disorder
data_PRS_MDD <- read.table("MDD_PRSice_SCORES_AT_ALL_THRESHOLDS.txt",
                           header = TRUE, sep = " ", stringsAsFactors = FALSE)
names(data_PRS_MDD)[1] <- 'src_subject_id'

# PRS for Schizophrenia
data_PRS_SCZ <- read.table("SCZ_PRSice_SCORES_AT_ALL_THRESHOLDS.txt",
                           header = TRUE, sep = " ", stringsAsFactors = FALSE)
names(data_PRS_SCZ)[1] <- 'src_subject_id'

# Genetic PCs covariates (PC1, PC2, PC3)
data_PCs <- read.table("ABCD_eur_covariatesPC3_PLINK.txt",
                       header = TRUE, sep = "\t", stringsAsFactors = FALSE)
names(data_PCs)[1] <- 'src_subject_id'
data_PCs <- dplyr::select(data_PCs, c('src_subject_id', 'PC1', 'PC2', 'PC3'))

# Child Behaviour Checklist (CBCL)
data_CBCL <- read.table("abcd_cbcls01.txt", header = TRUE, sep = "\t",
                        stringsAsFactors = FALSE)

# Early Adolescent Temperament Questionnaire Revised (EATQ-R) and CBCL items
data_EATQR_CBCL <- read.table("CBCL_EATQ_BISBAS_2year.txt", header = TRUE,
                               sep = "\t", stringsAsFactors = FALSE)

# Working Memory (WM)
data_WM <- read.table("abcd_mrinback02.txt",
                      header = TRUE, sep = "\t", stringsAsFactors = FALSE)

# Stop Signal Reaction Time (SSRT)
data_SSRT <- read.table("abcd_sst02.txt",
                        header = TRUE, sep = "\t", stringsAsFactors = FALSE)

# Reaction Time Variability (RTV)
data_RTV <- read.table("abcd_sst02.txt",
                       header = TRUE, sep = "\t", stringsAsFactors = FALSE)

```

Select 2-year follow-up and change character to numeric

```

# CBCL at 2-year follow-up
data_CBCL_2y <- filter(data_CBCL, eventname == "2_year_follow_up_y_arm_1")
data_CBCL_2y[, 10:79] <- lapply(data_CBCL_2y[, 10:79], as.numeric)

# WM at 2-year follow-up
data_WM_2y <- filter(data_WM, eventname == "2_year_follow_up_y_arm_1")
data_WM_2y[, 13:590] <- lapply(data_WM_2y[, 13:590], as.numeric)

# SSRT at 2-year follow-up
data_SSRT_2y <- filter(data_SSRT, eventname == "2_year_follow_up_y_arm_1")
data_SSRT_2y[, 13:104] <- lapply(data_SSRT_2y[, 13:104], as.numeric)

# RTV at 2-year follow-up
data_RTV_2y <- filter(data_RTV, eventname == "2_year_follow_up_y_arm_1")
data_RTV_2y[, 13:104] <- lapply(data_RTV_2y[, 13:104], as.numeric)

```

Quality control measures

```

# Quality control for WM data
data_WM_2y_QC <- filter(data_WM_2y,
                         tfMRI_nback_beh_performflag == 1, # performance good
                         !is.na(tfMRI_nb_all_beh_c2b_rate)) # remove missing

# Quality control for SSRT data
data_SSRT_2y_QC <- filter(data_SSRT_2y,
                            tfMRI_sst_beh_performflag == 1, # performance good
                            tfMRI_sst_beh_violatorflag == 0, # racing assum met
                            tfMRI_sst_beh_glitchflag == 0, # no issues run task
                            between(tfMRI_sst_all_beh_incrs_rt, .25, .75), # prop incor
                            tfMRI_sst_all_beh_nrgo_rt < .3, # go omission < 30%
                            tfMRI_sst_all_beh_total_issrt >= 120, # reliable scores
                            !is.na(tfMRI_sst_all_beh_total_issrt)) # remove missing

# Quality control for RTV data
data_RTV_2y_QC <- filter(data_RTV_2y,
                           tfMRI_sst_beh_performflag == 1, # performance good
                           tfMRI_sst_beh_glitchflag == 0, # no issues run task
                           tfMRI_sst_all_beh_nrgo_rt < .3, # go omission < 30%
                           !is.na(tfMRI_sst_all_beh_crgo_stdrt)) # remove missing

```

Run exploratory factor analysis to create an outcome variable

```

# Data selection - Effortful control (EC) items from the EATQ-R and AP items from CBCL
# note - this data is already at the 2 year follow-up
data_EFA <- dplyr::select(data_EATQR_CBCL, c('src_subject_id', 'interview_age', 'sex', 'cbcl_q01_p',
, 'cbcl_q04_p', 'cbcl_q08_p', 'cbcl_q10_p', 'cbcl_q13_p', 'cbcl_q17_p', 'cbcl_q41_p', 'cbcl_q61_p',
, 'cbcl_q78_p', 'cbcl_q80_p', 'eatq_finish_p', 'eatq_deal_p', 'eatq_turn_taking_p', 'eatq_open_present_p',
'eatq_before_hw_p', 'eatq_concentrate_p', 'eatq_right_away_p', 'eatq_distracted_p', 'eatq_impulse_p',
'eatq_try_focus_p', 'eatq_finish_hw_p', 'eatq_early_start_p', 'eatq_peripheral_p', 'eatq_puts_off_p',
'eatq_laugh_control_p', 'eatq_sidetracked_p', 'eatq_stick_to_plan_p', 'eatq_close_attention_p'))

# remove missing data
comp_data_EFA <- data_EFA[complete.cases(data_EFA), ]
# take out subject ID, age and sex
data_for_EFA <- comp_data_EFA[, c(4:31)]

# Kaiser-Meyer-Olkin test
KMO(data_for_EFA) # we have enough participants

```

```

## Kaiser-Meyer-Olkin factor adequacy
## Call: KMO(r = data_for_EFA)
## Overall MSA = 0.95
## MSA for each item =
##          cbcl_q01_p      cbcl_q04_p      cbcl_q08_p
##            0.98          0.97          0.94
##          cbcl_q10_p      cbcl_q13_p      cbcl_q17_p
##            0.94          0.91          0.94
##          cbcl_q41_p      cbcl_q61_p      cbcl_q78_p
##            0.95          0.97          0.95
##          cbcl_q80_p      eatq_finish_p   eatq_deal_p
##            0.91          0.97          0.94
##          eatq_turn_taking_p  eatq_open_present_p  eatq_before_hw_p
##            0.94          0.91          0.96
##          eatq_concentrate_p  eatq_right_away_p  eatq_distracted_p
##            0.98          0.97          0.95
##          eatq_impulse_p      eatq_try_focus_p  eatq_finish_hw_p
##            0.94          0.97          0.96
##          eatq_early_start_p  eatq_peripheral_p  eatq_puts_off_p
##            0.93          0.96          0.95
##          eatq_laugh_control_p  eatq_sidetracked_p  eatq_stick_to_plan_p
##            0.92          0.97          0.96
##          eatq_close_attention_p
##            0.96

```

```

# Bartlett's test for sphericity
cortest.bartlett(data_for_EFA) # highly significant

```

```

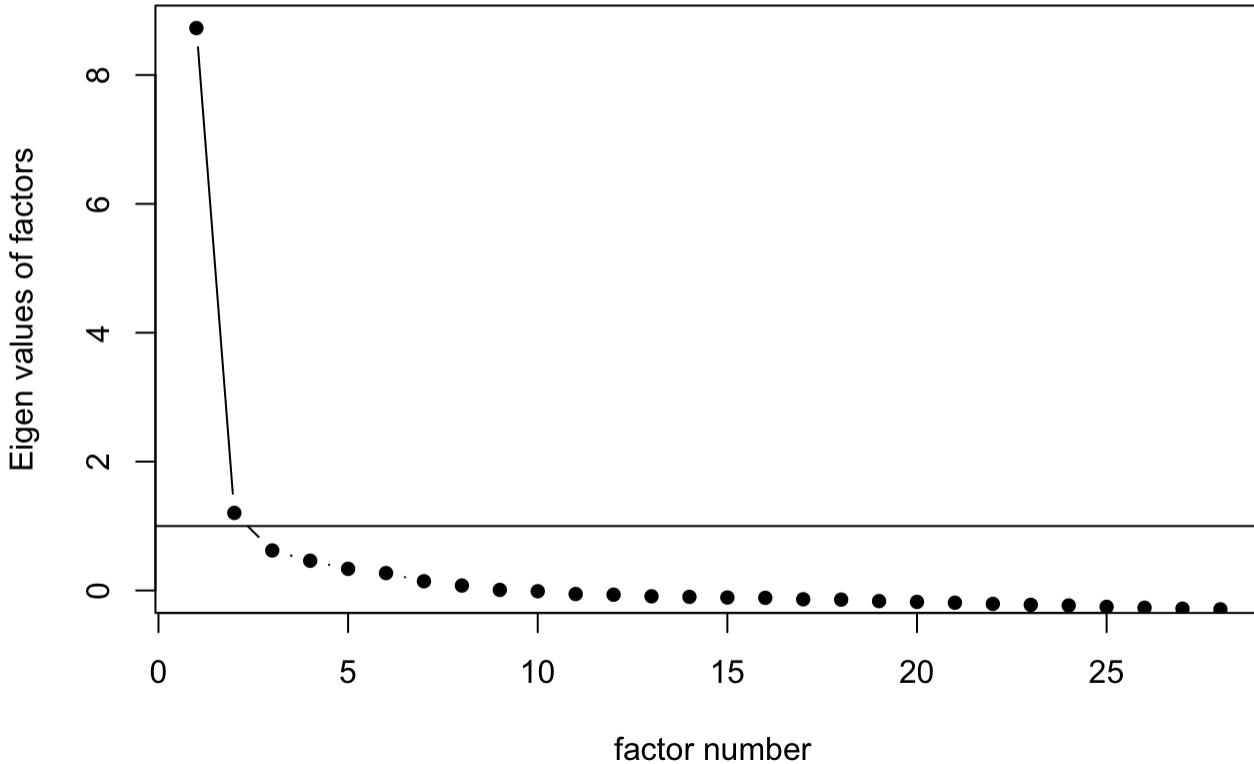
## R was not square, finding R from data

```

```
## $chisq  
## [1] 63437.46  
##  
## $p.value  
## [1] 0  
##  
## $df  
## [1] 378
```

```
# Scree plot to determine the number of factors to extract  
scree(data_for_EFA, pc = FALSE)
```

Scree plot



```
# Extract one factor and assess factor loadings  
oneF_fit <- factanal(data_for_EFA, 1, rotation = "promax")  
oneF_loads <- oneF_fit$loadings  
oneF_loads
```

```

## 
## Loadings:
##                               Factor1
## cbcl_q01_p                  -0.420
## cbcl_q04_p                  -0.709
## cbcl_q08_p                  -0.768
## cbcl_q10_p                  -0.550
## cbcl_q13_p                  -0.321
## cbcl_q17_p                  -0.436
## cbcl_q41_p                  -0.542
## cbcl_q61_p                  -0.547
## cbcl_q78_p                  -0.755
## cbcl_q80_p                  -0.358
## eatq_finish_p                0.777
## eatq_deal_p                 0.300
## eatq_turn_taking_p          0.418
## eatq_open_present_p          0.291
## eatq_before_hw_p             0.507
## eatq_concentrate_p          0.487
## eatq_right_away_p            0.529
## eatq_distracted_p           0.509
## eatq_impulse_p               0.413
## eatq_try_focus_p              0.639
## eatq_finish_hw_p              0.652
## eatq_early_start_p           0.697
## eatq_peripheral_p             0.666
## eatq_puts_off_p               0.671
## eatq_laugh_control_p          0.197
## eatq_sidetracked_p            0.659
## eatq_stick_to_plan_p          0.587
## eatq_close_attention_p         0.617
##
##                               Factor1
## SS loadings      8.721
## Proportion Var   0.311

```

```

# Create 'ADHD Traits' Bartlett Factor Scores
ADHD_Traits_scores_create <- factanal(data_for_EFA, 1, scores = "Bartlett",
                                         rotation = "promax")
ADHD_Traits_scores <- ADHD_Traits_scores_create$scores

# Add factor scores back into table
comp_data_EFA[, "ADHD_Traits"] <- ADHD_Traits_scores

```

Create smaller tables with relevant variables

```

# Note - excluding PRS for SCZ and BIP as they showed no association to the ADHD Traits factor scores in PRSice

# ADHD PRS table with ID & pT = .132
data_PRS_ADHD_small <- dplyr::select(data_PRS_ADHD, c('src_subject_id', 'pT_0.132'))
names(data_PRS_ADHD_small)[2] <- 'ADHD_PRS'

# MDD PRS table with ID & pT = .0945
data_PRS_MDD_small <- dplyr::select(data_PRS_MDD, c('src_subject_id', 'pT_0.0945'))
names(data_PRS_MDD_small)[2] <- 'MDD_PRS'

# ASD PRS table with ID & pT = .003
data_PRS_ASD_small <- dplyr::select(data_PRS_ASD, c('src_subject_id', 'pT_0.003'))
names(data_PRS_ASD_small)[2] <- 'ASD_PRS'

# merge all PRS tables
data_PRS <- merge(data_PRS_ADHD_small, data_PRS_MDD_small)
data_PRS <- merge(data_PRS, data_PRS_ASD_small)

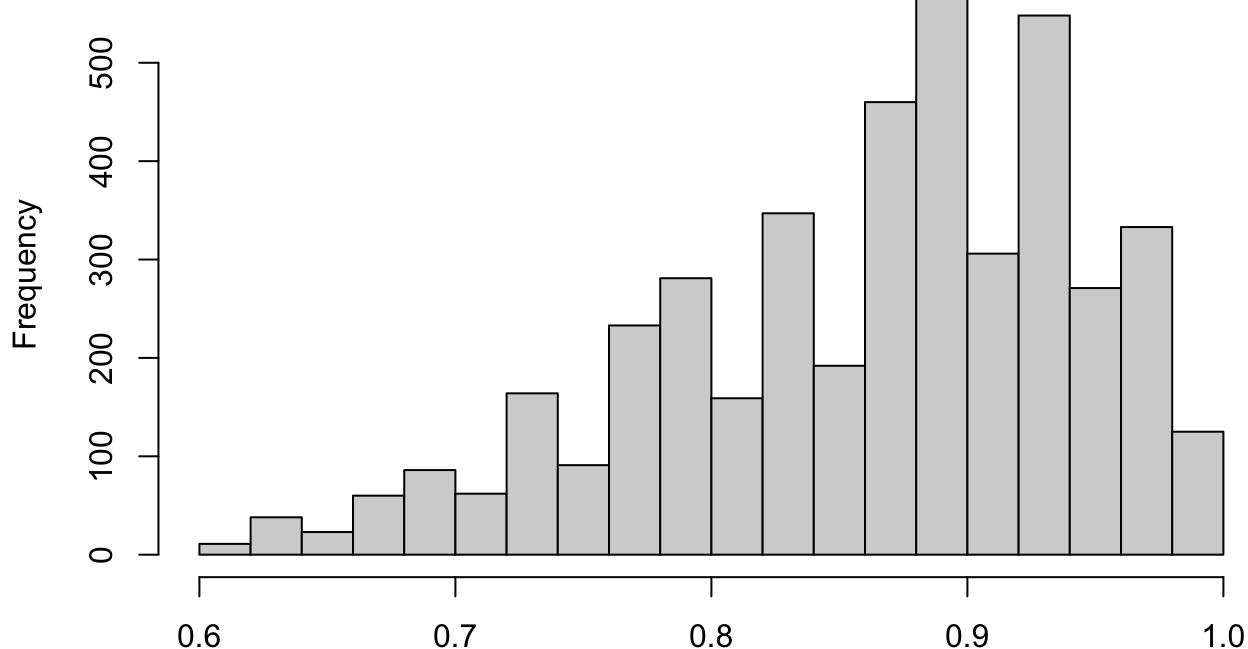
# add PCs to the PRS table
data_PRS <- merge(data_PRS, data_PCs)

# ADHD Traits table with ID, ADHD traits, sex, age (change to years), then add age2 and sex*age
data_ADHD_Traits_small <- dplyr::select(comp_data_EFA, c('src_subject_id', 'interview_age', 'sex', 'ADHD_Traits'))
Age <- (data_ADHD_Traits_small$interview_age)/12
data_ADHD_Traits_small[, "Age"] <- Age
Age2 <- (data_ADHD_Traits_small$Age)^2
data_ADHD_Traits_small[, "Age2"] <- Age2
data_ADHD_Traits_small$Sex <- ifelse(data_ADHD_Traits_small$sex=="M", 1, 2)
AgeSex <- (data_ADHD_Traits_small$Age)*(data_ADHD_Traits_small$Sex)
data_ADHD_Traits_small[, "AgeSex"] <- AgeSex
data_ADHD_Traits_small <- data_ADHD_Traits_small[, c('src_subject_id', 'ADHD_Traits', 'Age', 'Age2', 'Sex', 'AgeSex')]

# WM table with ID & WM
data_WM_2y_small <- dplyr::select(data_WM_2y_QC, c('src_subject_id', 'tfMRI_nb_all_beh_c2b_rate'))
# Assess distribution
hist(data_WM_2y_small$tfMRI_nb_all_beh_c2b_rate)

```

Histogram of data_WM_2y_small\$tfMRI_nb_all_beh_c2b_rate



```
skewness(data_WM_2y_small$tfMRI_nb_all_beh_c2b_rate)
```

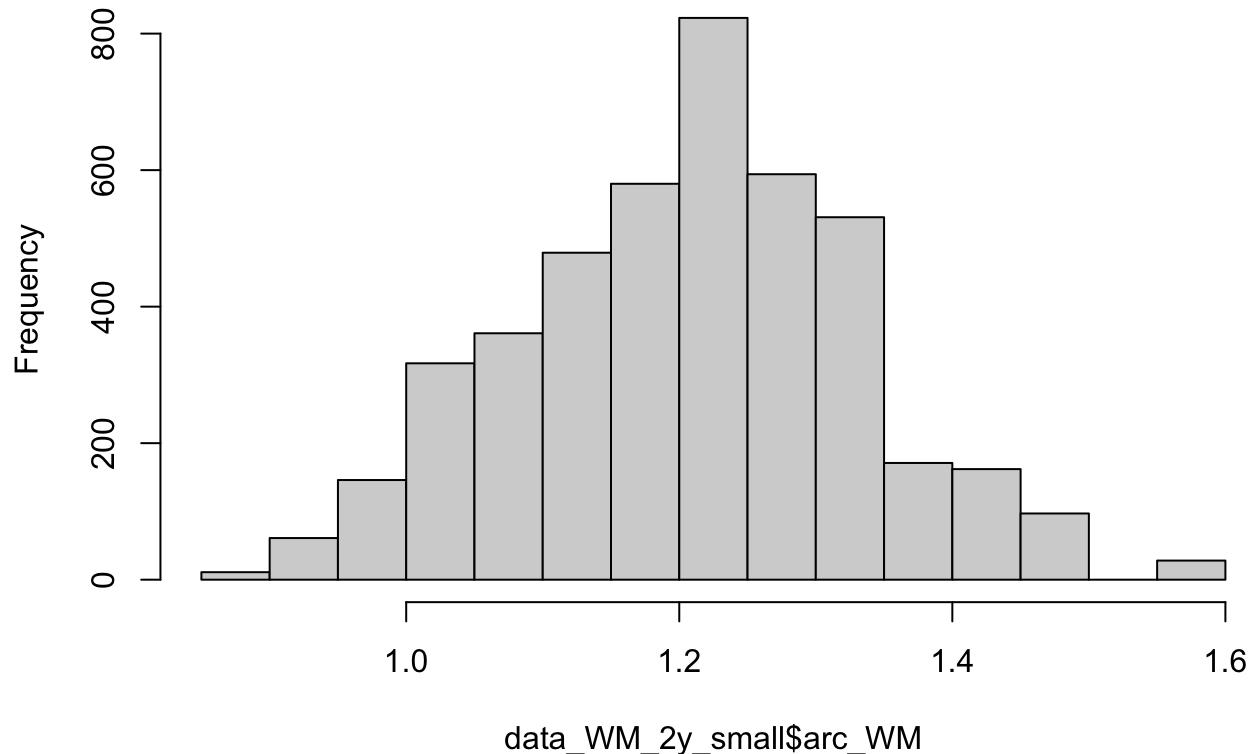
```
## [1] -0.6963714
```

```
kurtosis(data_WM_2y_small$tfMRI_nb_all_beh_c2b_rate)
```

```
## [1] -0.1255527
```

```
# Apply an arcsine transformation
arc_WM <- asin(sqrt(data_WM_2y_small$tfMRI_nb_all_beh_c2b_rate))
data_WM_2y_small[, "arc_WM"] <- arc_WM
hist(data_WM_2y_small$arc_WM)
#Assess the distribution
hist(data_WM_2y_small$arc_WM)
```

Histogram of data_WM_2y_small\$arc_WM



```
skewness(data_WM_2y_small$arc_WM)
```

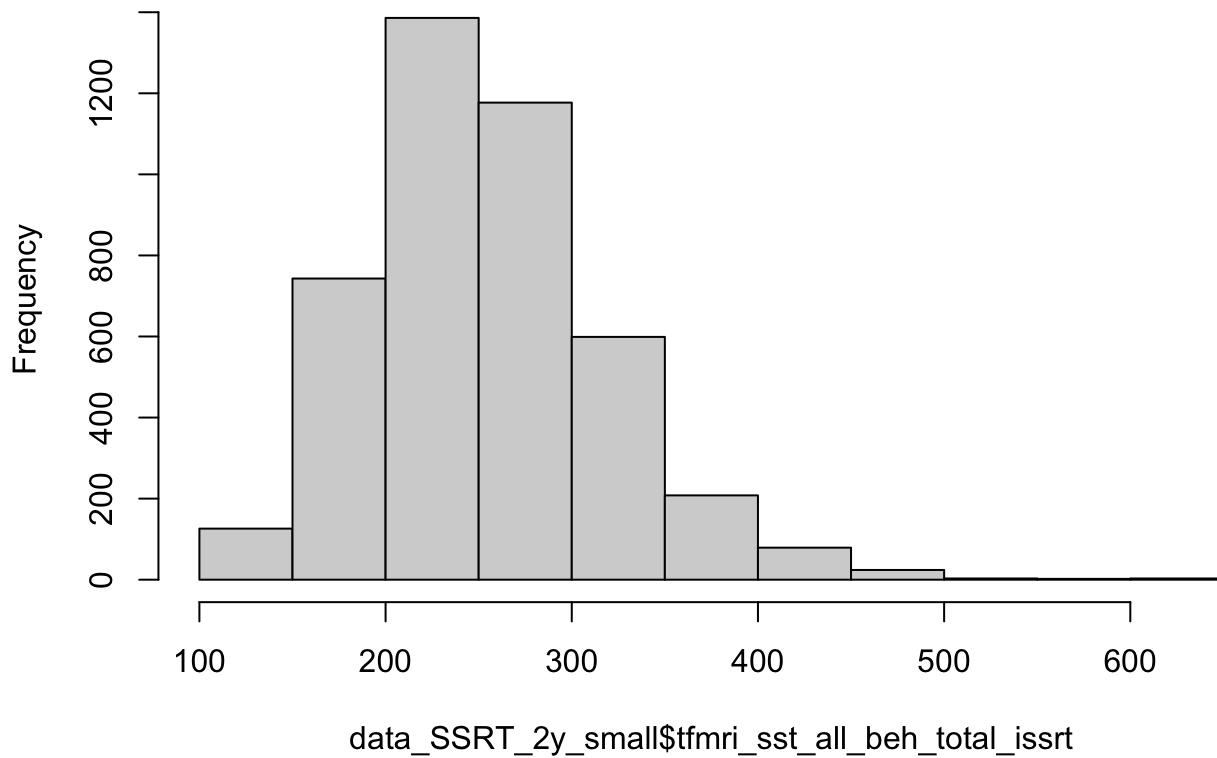
```
## [1] -0.05909523
```

```
kurtosis(data_WM_2y_small$arc_WM)
```

```
## [1] -0.3189293
```

```
# SSRT table with ID & SSRT (and then add a log(SSRT))
data_SSRT_2y_small <- dplyr::select(data_SSRT_2y_QC, c('src_subject_id', 'tfMRI_sst_all_beh_total_issrt'))
# Assess distribution
hist(data_SSRT_2y_small$tfMRI_sst_all_beh_total_issrt)
```

Histogram of data_SSRT_2y_small\$tfMRI_sst_all_beh_total_issrt



```
skewness(data_SSRT_2y_small$tfMRI_sst_all_beh_total_issrt)
```

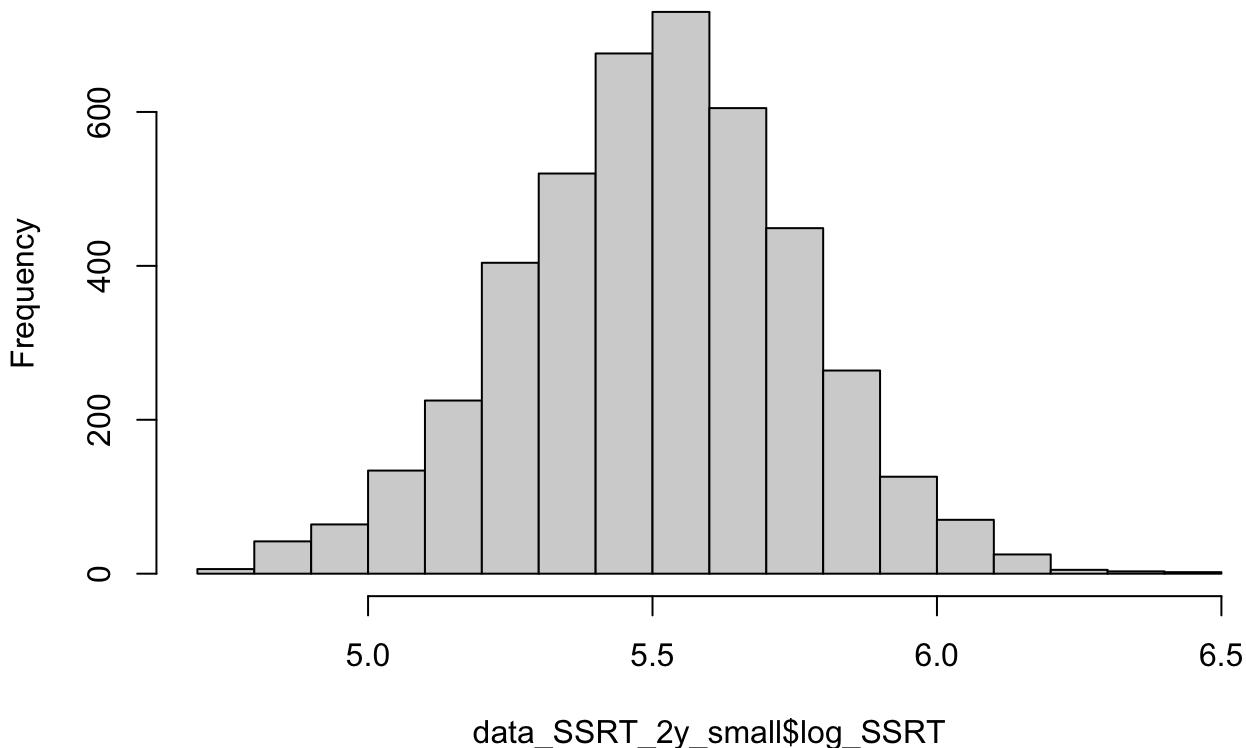
```
## [1] 0.7534291
```

```
kurtosis(data_SSRT_2y_small$tfMRI_sst_all_beh_total_issrt)
```

```
## [1] 1.34823
```

```
#Apply a natural log transformation
log_SSRT <- log(data_SSRT_2y_small$tfMRI_sst_all_beh_total_issrt)
data_SSRT_2y_small[, "log_SSRT"] <- log_SSRT
#Assess distribution
hist(data_SSRT_2y_small$log_SSRT)
```

Histogram of data_SSRT_2y_small\$log_SSRT



```
skewness(data_SSRT_2y_small$log_SSRT)
```

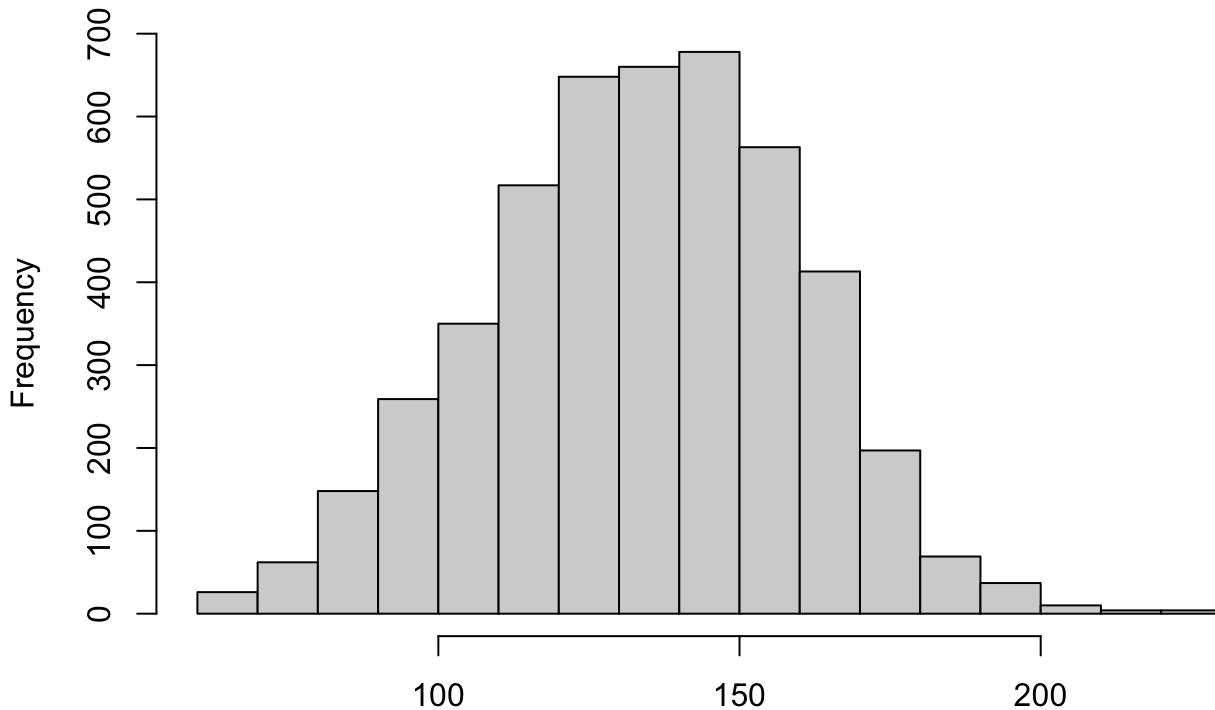
```
## [1] -0.07521215
```

```
kurtosis(data_SSRT_2y_small$log_SSRT)
```

```
## [1] 0.08714305
```

```
# RTV table with ID & RTV (and then add a log(RTV))
data_RTV_2y_small <- dplyr::select(data_RTV_2y_QC, c('src_subject_id', 'tfMRI_sst_all_beh_crgo_stdrt'))
# Assess distribution
hist(data_RTV_2y_small$tfMRI_sst_all_beh_crgo_stdrt)
```

Histogram of data_RTV_2y_small\$tfMRI_SST_all_beh_crgo_stdrt



data_RTV_2y_small\$tfMRI_SST_all_beh_crgo_stdrt

```
skewness(data_RTV_2y_small$tfMRI_SST_all_beh_crgo_stdrt)
```

```
## [1] -0.09476349
```

```
kurtosis(data_RTV_2y_small$tfMRI_SST_all_beh_crgo_stdrt)
```

```
## [1] -0.1794368
```

```
names(data_RTV_2y_small)[2] <- 'RTV'
```

Create final tables

```
# Join PRS and CBCL
data_final_genbehav <- merge(data_PRS, data_ADHD_Traits_small)

# Final WM table
data_finalWM_group <- merge(data_final_genbehav, data_WM_2y_small)

# Final SSRT table
data_finalSSRT_group <- merge(data_final_genbehav, data_SSRT_2y_small)

# Final RTV table
data_finalRTV_group <- merge(data_final_genbehav, data_RTV_2y_small)
```

Participant demographics

```
# participants at 2-year follow-up with attention problems according to CBCL  
CBCL_clinADHD <- filter(data_CBCL_2y, cbcl_scr_syn_attention_t >= 65)  
# 308/5823 = 5%  
  
# WM group  
mean(data_finalWM_group$Age)
```

```
## [1] 11.93426
```

```
sd(data_finalWM_group$Age)
```

```
## [1] 0.642018
```

```
table(data_finalWM_group$Sex)
```

```
##  
##      1      2  
## 1232   989
```

```
# SSRT group  
mean(data_finalSSRT_group$Age)
```

```
## [1] 11.92548
```

```
sd(data_finalSSRT_group$Age)
```

```
## [1] 0.6422761
```

```
table(data_finalSSRT_group$Sex)
```

```
##  
##      1      2  
## 1095   909
```

```
# RTV group  
mean(data_finalRTV_group$Age)
```

```
## [1] 11.93312
```

```
sd(data_finalRTV_group$Age)
```

```
## [1] 0.6425747
```

```
table(data_finalRTV_group$Sex)
```

```
##  
##      1     2  
## 1165  957
```

Regressions - cognition regressed onto behaviour

```
# WM - ADHD  
regression_WM_ADHD <- lm(ADHD_Traits ~ arc_WM +  
                           Sex + Age + Age2 + AgeSex, data = data_finalWM_group)  
summary(regression_WM_ADHD)
```

```
##  
## Call:  
## lm(formula = ADHD_Traits ~ arc_WM + Sex + Age + Age2 + AgeSex,  
##      data = data_finalWM_group)  
##  
## Residuals:  
##       Min     1Q   Median     3Q    Max  
## -3.6838 -0.6051  0.1535  0.7385  2.0102  
##  
## Coefficients:  
##             Estimate Std. Error t value Pr(>|t|)  
## (Intercept) 1.484191  8.127849  0.183  0.8551  
## arc_WM      1.545891  0.182609  8.466 <2e-16 ***  
## Sex        -1.070207  0.801952 -1.335  0.1822  
## Age        -0.418563  1.349199 -0.310  0.7564  
## Age2        0.007199  0.056187  0.128  0.8981  
## AgeSex      0.124949  0.067149  1.861  0.0629 .  
## ---  
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1  
##  
## Residual standard error: 1.005 on 2215 degrees of freedom  
## Multiple R-squared:  0.06736,    Adjusted R-squared:  0.06525  
## F-statistic: 31.99 on 5 and 2215 DF,  p-value: < 2.2e-16
```

```
effectsize::standardize_parameters(regression_WM_ADHD)
```

```

## Registered S3 methods overwritten by 'parameters':
##   method                  from
##   as.double.parameters_kurtosis    datawizard
##   as.double.parameters_skewness     datawizard
##   as.double.parameters_smoothness  datawizard
##   as.numeric.parameters_kurtosis   datawizard
##   as.numeric.parameters_skewness   datawizard
##   as.numeric.parameters_smoothness datawizard
##   print.parameters_distribution   datawizard
##   print.parameters_kurtosis       datawizard
##   print.parameters_skewness       datawizard
##   summary.parameters_kurtosis    datawizard
##   summary.parameters_skewness    datawizard

```

Parameter <chr>	Std_Coefficient <dbl>	CI <dbl>	CI_low <dbl>	CI_high <dbl>
1 (Intercept)	-2.730355e-16	0.95	-0.04023082	0.04023082
2 arc_WM	1.781271e-01	0.95	0.13686417	0.21938993
3 Sex	-5.119269e-01	0.95	-1.26419815	0.24034426
4 Age	-2.585805e-01	0.95	-1.89312260	1.37596165
5 Age2	1.062943e-01	0.95	-1.52069242	1.73328103
6 AgeSex	7.163444e-01	0.95	-0.03860148	1.47129022

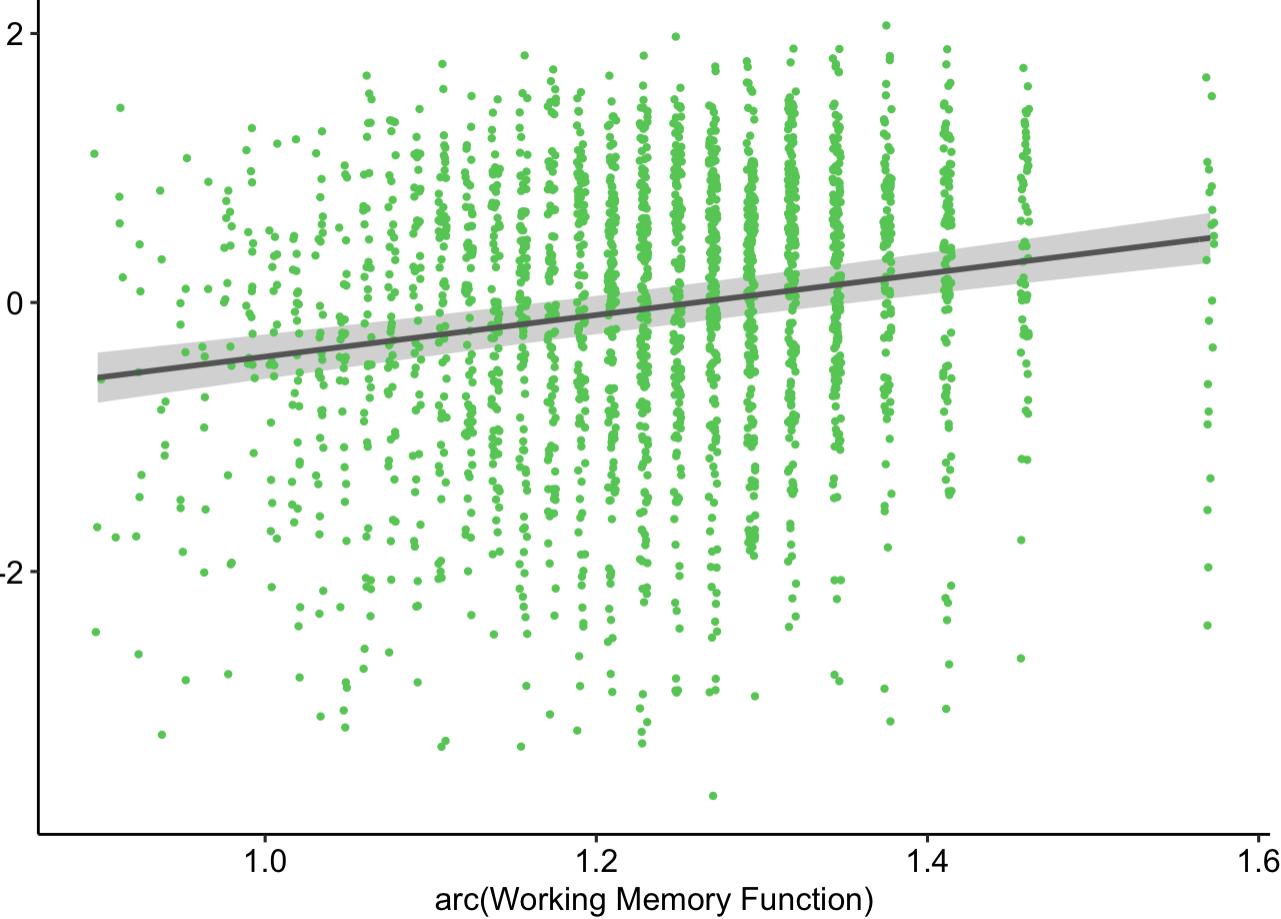
6 rows

```

# graph
visreg(regression_WM_ADHD, xvar = "arc_WM", rug = FALSE, jitter = TRUE, overlay = TRUE, gg = TRUE,
line=list(col="#666666"), points=list(col="#66CC66")) + theme_pubr() + labs(x = "arc(Working Memory Function)", y = "ADHD-Like Traits")

```

ADHD-Like Traits



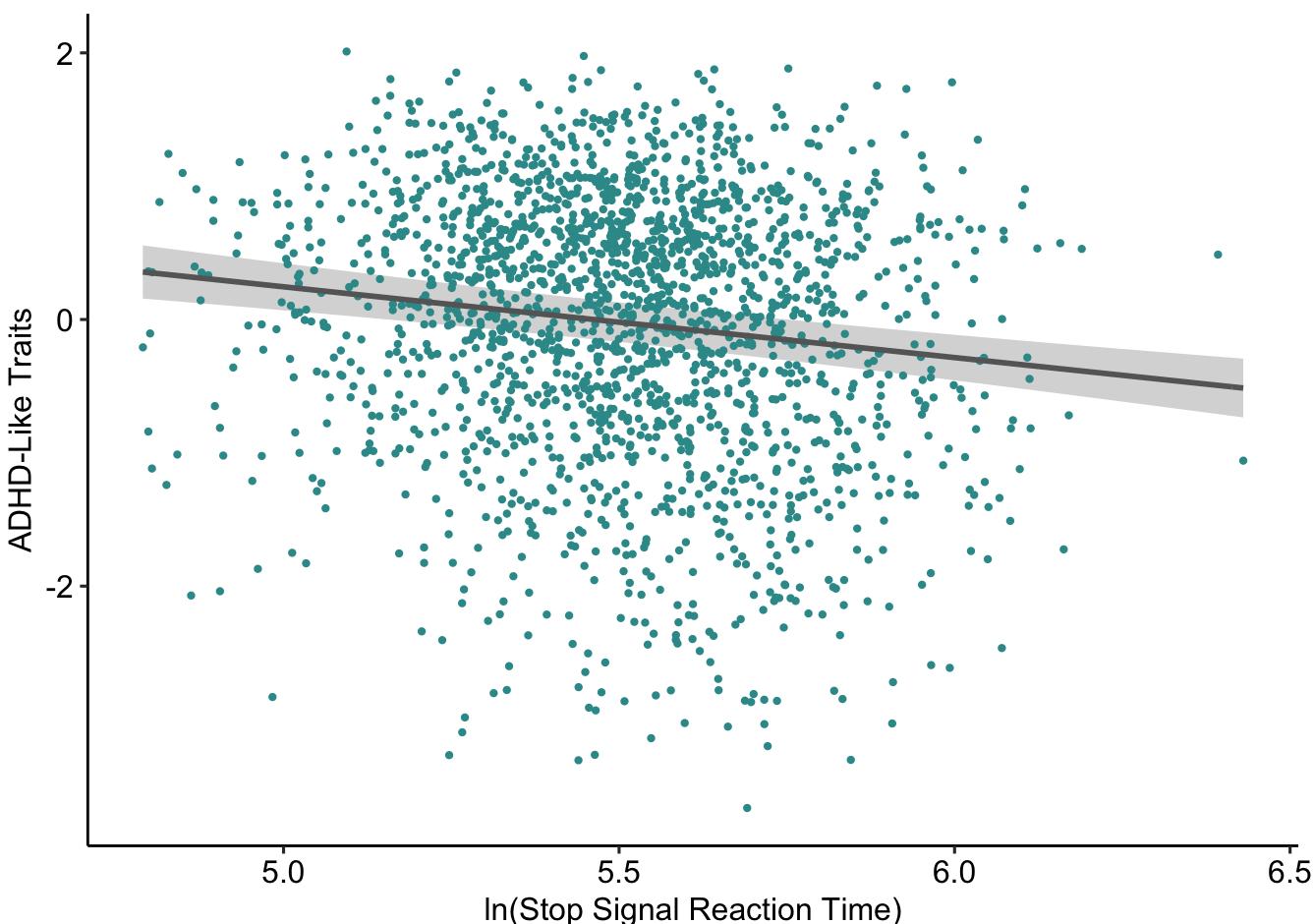
```
# SSRT - ADHD
regression_SSRT_ADHD <- lm(ADHD_Traits ~ log_SSRT +
                           Sex + Age + Age2 + AgeSex, data = data_finalSSRT_group)
summary(regression_SSRT_ADHD)
```

```
##
## Call:
## lm(formula = ADHD_Traits ~ log_SSRT + Sex + Age + Age2 + AgeSex,
##      data = data_finalSSRT_group)
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -3.5427 -0.6001  0.1504  0.7377  2.0627
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)
## (Intercept) 7.51452   8.51984   0.882   0.378
## log_SSRT    -0.53091   0.09242  -5.745 1.06e-08 ***
## Sex         -1.12975   0.83840  -1.348   0.178
## Age        -0.63837   1.41070  -0.453   0.651
## Age2        0.01752   0.05865   0.299   0.765
## AgeSex      0.12739   0.07026   1.813   0.070 .
## ---
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9992 on 1998 degrees of freedom
## Multiple R-squared:  0.05485,    Adjusted R-squared:  0.05249
## F-statistic: 23.19 on 5 and 1998 DF,  p-value: < 2.2e-16
```

```
effectsize::standardize_parameters(regression_SSRT_ADHD)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	4.321936e-16	0.95	-0.04264374	0.04264374
2 log_SSRT	-1.254864e-01	0.95	-0.16832636	-0.08264635
3 Sex	-5.480611e-01	0.95	-1.34570224	0.24958003
4 Age	-3.994264e-01	0.95	-2.13048849	1.33163563
5 Age2	2.619632e-01	0.95	-1.45825824	1.98218474
6 AgeSex	7.385383e-01	0.95	-0.06029435	1.53737103
6 rows				

```
# graph
visreg(regression_SSRT_ADHD, xvar = "log_SSRT", rug = FALSE, jitter = TRUE, overlay = TRUE, gg = TRUE,
line=list(col="#666666"), points=list(col="#339999")) + theme_pubr() + labs(x = "ln(Stop Signal Reaction Time)", y = "ADHD-Like Traits")
```



```
# RTV - ADHD
regression_RTV_ADHD <- lm(ADHD_Traits ~ RTV +
                           Sex + Age + Age2 + AgeSex, data = data_finalRTV_group)
summary(regression_RTV_ADHD)
```

```

## 
## Call:
## lm(formula = ADHD_Traits ~ RTV + Sex + Age + Age2 + AgeSex, data = data_finalRTV_group)
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -3.6283 -0.6137  0.1836  0.7307  2.0571
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 2.8715056  8.3030316  0.346   0.7295    
## RTV        -0.0065023  0.0008544 -7.611  4.08e-14 ***  
## Sex        -1.2330599  0.8156540 -1.512   0.1307    
## Age        -0.1843683  1.3775785 -0.134   0.8935    
## Age2       -0.0026904  0.0572837 -0.047   0.9625    
## AgeSex      0.1382039  0.0683134  2.023   0.0432 *   
## ---      
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.9996 on 2116 degrees of freedom
## Multiple R-squared:  0.0658, Adjusted R-squared:  0.0636 
## F-statistic: 29.81 on 5 and 2116 DF,  p-value: < 2.2e-16

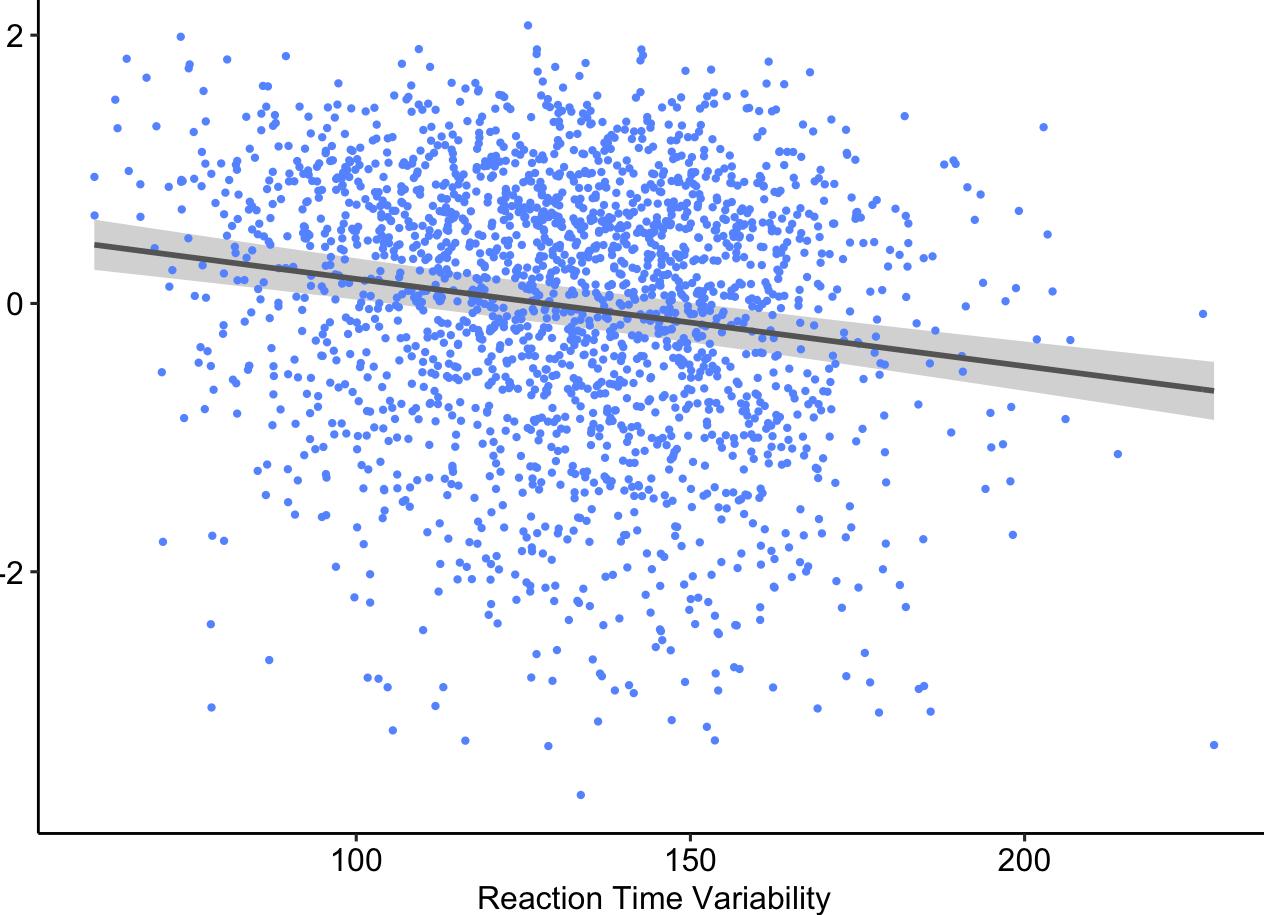
```

```
effectsize::standardize_parameters(regression	RTV_AdHD)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-2.209132e-16	0.95	-0.04119602	0.04119602
2 RTV	-1.622620e-01	0.95	-0.20407356	-0.12045045
3 Sex	-5.940983e-01	0.95	-1.36478291	0.17658628
4 Age	-1.146854e-01	0.95	-1.79517105	1.56580023
5 Age2	-4.000604e-02	0.95	-1.71045596	1.63044388
6 AgeSex	7.964207e-01	0.95	0.02440798	1.56843335
6 rows				

```
# graph
visreg(regression_RTV_AdHD, xvar = "RTV", rug = FALSE, jitter = TRUE, overlay = TRUE, gg = TRUE, line=list(col="#666666"), points=list(col="#6699FF")) + theme_pubr() + labs(x = "Reaction Time Variability", y = "ADHD-Like Traits")
```

ADHD-Like Traits



```
# Correction for multiple testing
# Extract p-values from each table
# WM_ADHD
s_regression_WM_ADHD <- summary(regression_WM_ADHD)
p_WM_ADHD <- s_regression_WM_ADHD$coefficients["arc_WM", "Pr(>|t|)"]
# SSRT_ADHD
s_regression_SSRT_ADHD <- summary(regression_SSRT_ADHD)
p_SSRT_ADHD <- s_regression_SSRT_ADHD$coefficients["log_SSRT", "Pr(>|t|)"]
# RTV_ADHD
s_regression_RTV_ADHD <- summary(regression_RTV_ADHD)
p_RTV_ADHD <- s_regression_RTV_ADHD$coefficients["RTV", "Pr(>|t|)"]
# Create a vector of p-values
corr_cog_beh <- c(p_WM_ADHD, p_SSRT_ADHD, p_RTV_ADHD)
# Apply bonferroni correction
p.adjust(corr_cog_beh, method = "bonferroni") # all significant
```

```
## [1] 1.370402e-16 3.189053e-08 1.224591e-13
```

Regressions - ADHD PRS regressed onto cognition

```
# ADHD PRS - WM
regression_ADHD_PRS_WM <- lm(formula = arc_WM ~ ADHD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalWM_group)
summary(regression_ADHD_PRS_WM)
```

```

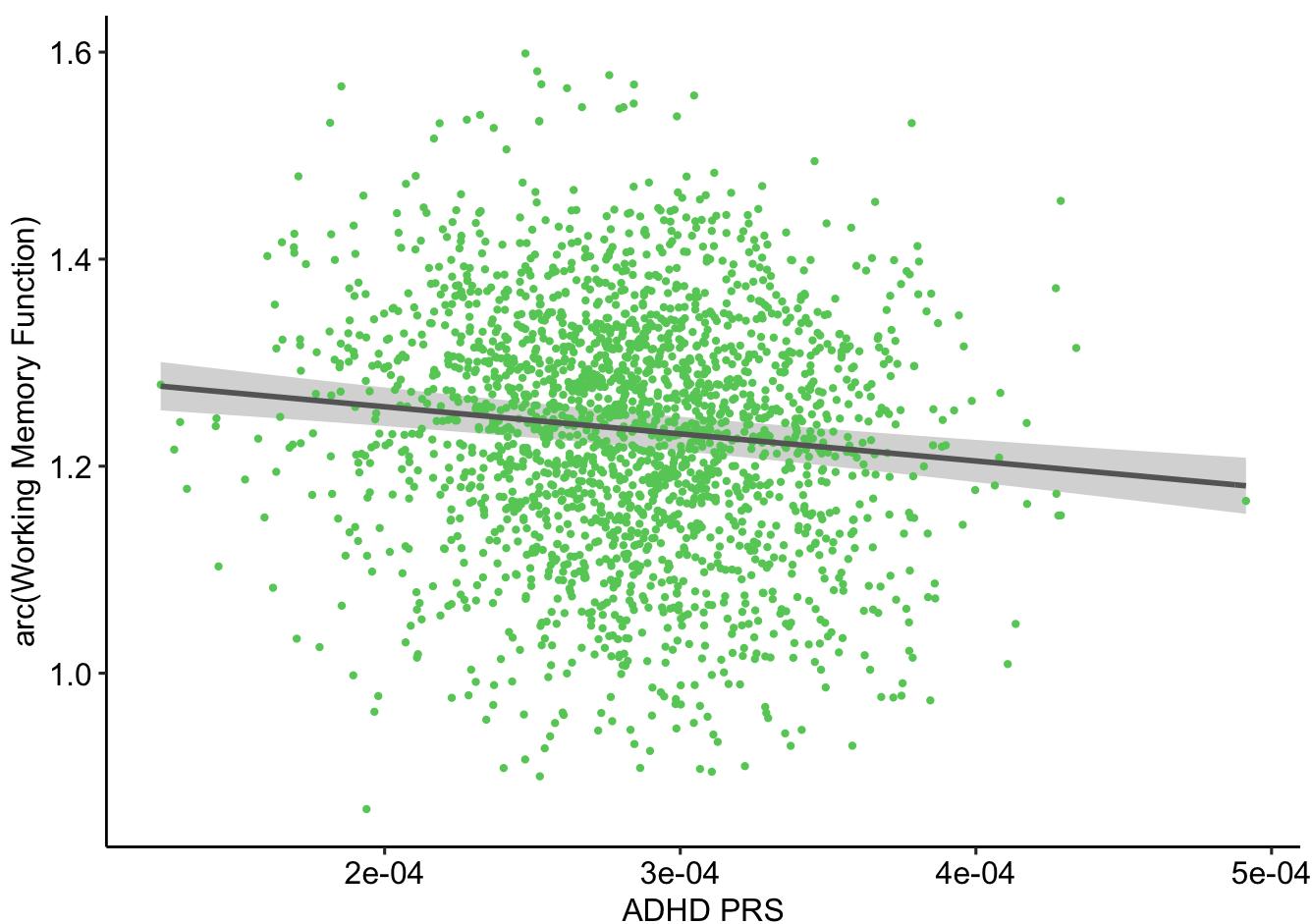
## 
## Call:
## lm(formula = arc_WM ~ ADHD_PRS + Sex + Age + Age2 + AgeSex +
##     PC1 + PC2 + PC3, data = data_finalWM_group)
##
## Residuals:
##      Min        1Q    Median        3Q       Max
## -0.39029 -0.07435  0.00737  0.07943  0.35379
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 1.893e+00  9.413e-01  2.011   0.0445 *  
## ADHD_PRS    -2.620e+02  5.288e+01 -4.955 7.78e-07 *** 
## Sex         -4.651e-02  9.288e-02 -0.501   0.6166    
## Age        -1.317e-01  1.563e-01 -0.843   0.3994    
## Age2        6.972e-03  6.509e-03  1.071   0.2843    
## AgeSex      2.573e-03  7.778e-03  0.331   0.7408    
## PC1         -9.702e-01  1.394e+00 -0.696   0.4867    
## PC2         6.810e-02  1.678e+00  0.041   0.9676    
## PC3         -1.160e+00  1.091e+00 -1.063   0.2878    
## ---        
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.1163 on 2212 degrees of freedom
## Multiple R-squared:  0.05983,    Adjusted R-squared:  0.05643 
## F-statistic: 17.6 on 8 and 2212 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(regression_ADHD_PRS_WM)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-3.098663e-16	0.95	-0.04042018	0.04042018
2 ADHD_PRS	-1.030782e-01	0.95	-0.14387298	-0.06228332
3 Sex	-1.930972e-01	0.95	-0.94921710	0.56302275
4 Age	-7.063704e-01	0.95	-2.34992793	0.93718719
5 Age2	8.934378e-01	0.95	-0.74237765	2.52925330
6 AgeSex	1.280405e-01	0.95	-0.63082734	0.88690831
7 PC1	-1.688960e-02	0.95	-0.06449512	0.03071592
8 PC2	8.472297e-04	0.95	-0.04008809	0.04178255
9 PC3	-2.600880e-02	0.95	-0.07397629	0.02195870
9 rows				

```
# graph
visreg(regression_ADHD_PRS_WM, xvar = "ADHD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg =
TRUE, line=list(col="#666666"), points=list(col="#66CC66")) + theme_pubr() + labs(x = "ADHD PRS",
y = "arc(Working Memory Function)")
```



```
# ADHD PRS - SSRT
regression_ADHD_PRS_SSRT <- lm(formula = log_SSRT ~ ADHD_PRS +
                                    Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalSSRT_group)
summary(regression_ADHD_PRS_SSRT)
```

```

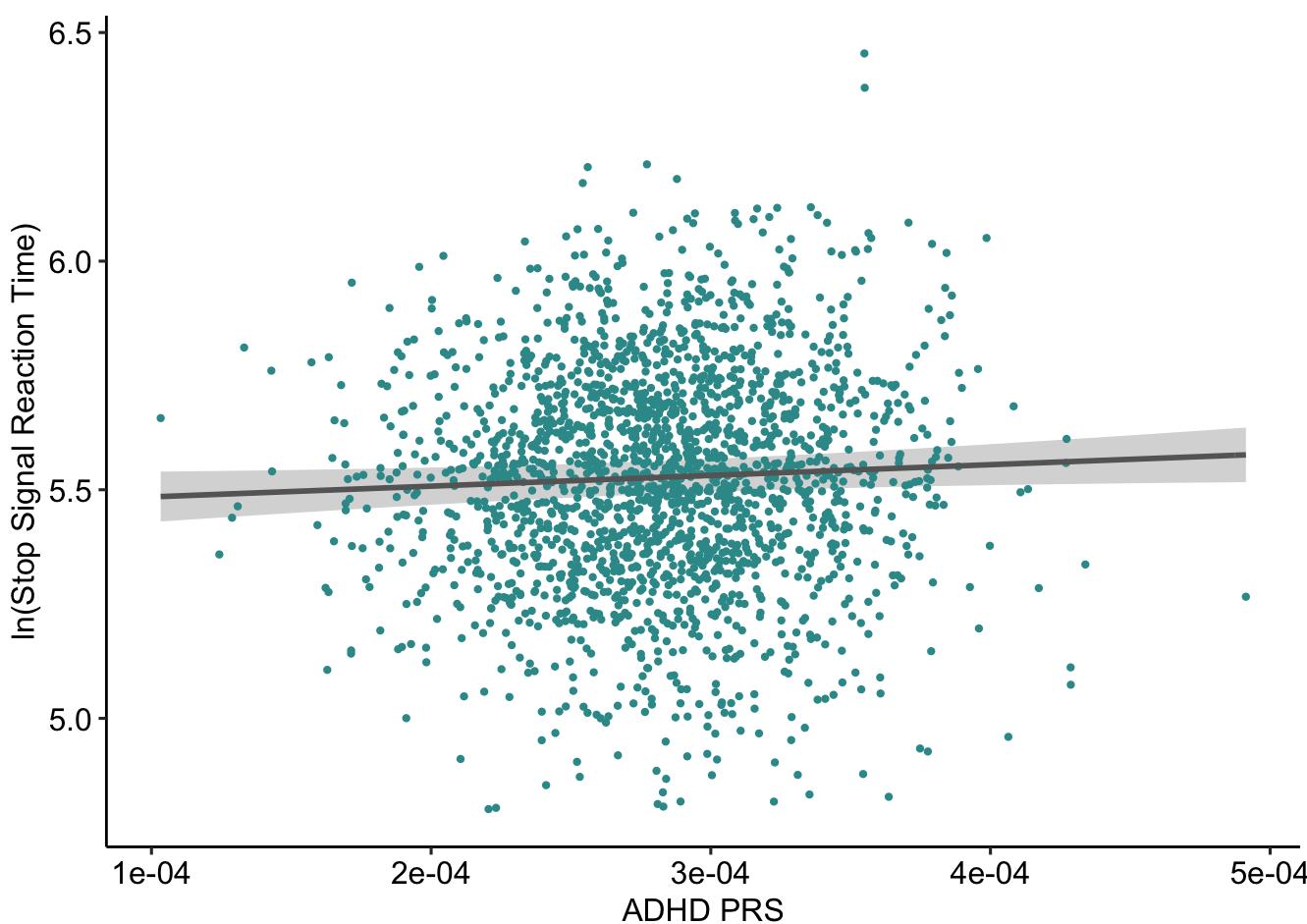
## 
## Call:
## lm(formula = log_SSRT ~ ADHD_PRS + Sex + Age + Age2 + AgeSex +
##     PC1 + PC2 + PC3, data = data_finalSSRT_group)
##
## Residuals:
##      Min        1Q    Median        3Q       Max
## -0.72037 -0.15908  0.00456  0.16191  0.91007
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 5.654676  2.061173  2.743   0.00613 **  
## ADHD_PRS    234.929573 116.522361  2.016   0.04392 *   
## Sex         -0.123574  0.202966 -0.609   0.54270    
## Age          0.019286  0.341780  0.056   0.95501    
## Age2         -0.002696  0.014211 -0.190   0.84954    
## AgeSex       0.008674  0.017011  0.510   0.61016    
## PC1          2.276855  3.037419  0.750   0.45358    
## PC2          -2.687410  3.705798 -0.725   0.46842    
## PC3          -0.005208  2.382833 -0.002   0.99826    
## --- 
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.2417 on 1995 degrees of freedom
## Multiple R-squared:  0.01139,   Adjusted R-squared:  0.00743 
## F-statistic: 2.874 on 8 and 1995 DF,  p-value: 0.0035

```

```
effectsize::standardize_parameters(regression_ADHD_PRS_SSRT)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-8.611489e-16	0.95	-0.043645891	0.04364589
2 ADHD_PRS	4.536510e-02	0.95	0.001238028	0.08949217
3 Sex	-2.536292e-01	0.95	-1.070600724	0.56334227
4 Age	5.105415e-02	0.95	-1.723350048	1.82545834
5 Age2	-1.705967e-01	0.95	-1.933976416	1.59278298
6 AgeSex	2.127674e-01	0.95	-0.605529436	1.03106424
7 PC1	1.976192e-02	0.95	-0.031940402	0.07146424
8 PC2	-1.637340e-02	0.95	-0.060652471	0.02790566
9 PC3	-5.804428e-05	0.95	-0.052142275	0.05202619
9 rows				

```
# graph
visreg(regression_ADHD_PRS_SSRT, xvar = "ADHD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg =
= TRUE, line=list(col="#666666"), points=list(col="#339999")) +
  theme_pubr() +
  labs(x = "ADHD PRS", y = "ln(Stop Signal Reaction Time)")
```



```
# ADHD PRS - RTV
regression_ADHD_PRS_RTV <- lm(formula = RTV ~ ADHD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalRTV_group)
summary(regression_ADHD_PRS_RTV)
```

```

## 
## Call:
## lm(formula = RTV ~ ADHD_PRS + Sex + Age + Age2 + AgeSex + PC1 +
##     PC2 + PC3, data = data_finalRTV_group)
##
## Residuals:
##      Min        1Q    Median        3Q       Max
## -70.187 -17.204   0.667  16.563  94.053
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 36.423    209.463   0.174  0.861971    
## ADHD_PRS    67838.223  11847.216   5.726 1.17e-08 *** 
## Sex         -5.023    20.549  -0.244  0.806914    
## Age          20.047    34.739   0.577  0.563944    
## Age2         -1.151    1.444  -0.797  0.425710    
## AgeSex       0.488     1.721   0.284  0.776779    
## PC1          49.077    308.655   0.159  0.873681    
## PC2          -480.926   374.343  -1.285  0.199031    
## PC3          841.029    241.480   3.483  0.000506 *** 
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 25.17 on 2113 degrees of freedom
## Multiple R-squared:  0.05005,    Adjusted R-squared:  0.04645 
## F-statistic: 13.92 on 8 and 2113 DF,  p-value: < 2.2e-16

```

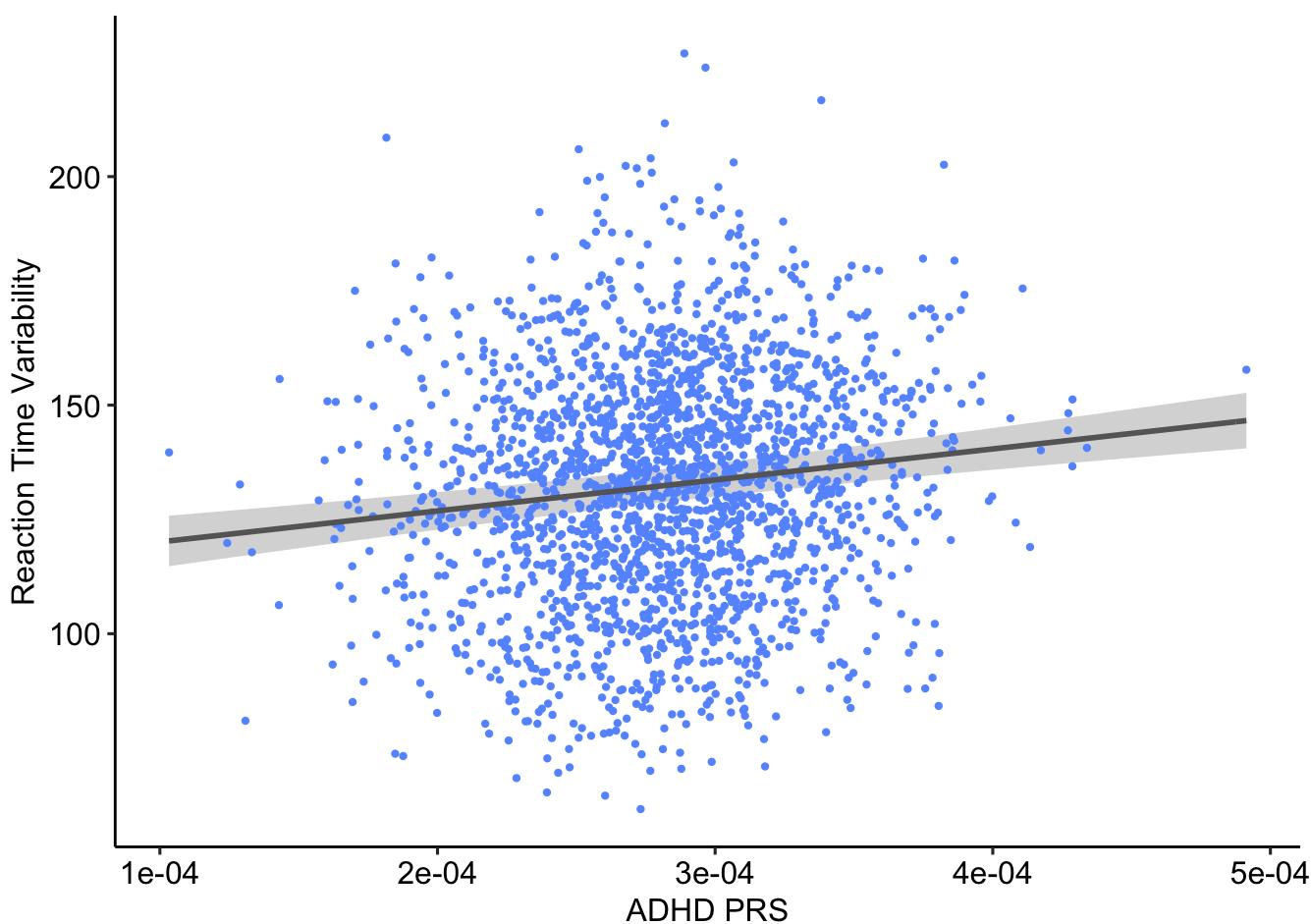
```
effectsize::standardize_parameters(regression_ADHD_PRS_RTV)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-1.127441e-15	0.95	-0.04157144	0.04157144
2 ADHD_PRS	1.226384e-01	0.95	0.08063687	0.16463996
3 Sex	-9.697950e-02	0.95	-0.87502512	0.68106612
4 Age	4.997249e-01	0.95	-1.19847346	2.19792329
5 Age2	-6.857673e-01	0.95	-2.37377363	1.00223905
6 AgeSex	1.126960e-01	0.95	-0.66671672	0.89210876
7 PC1	3.986752e-03	0.95	-0.04518416	0.05315766
8 PC2	-2.761511e-02	0.95	-0.06976879	0.01453857
9 PC3	8.791372e-02	0.95	0.03841171	0.13741574
9 rows				

```

# graph
visreg(regression_ADHD_PRS_RTV, xvar = "ADHD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg
= TRUE, line=list(col="#666666"), points=list(col="#6699FF")) +
theme_pubr() +
labs(x = "ADHD PRS", y = "Reaction Time Variability")

```



```

# Correction for multiple testing
# Extract p-values from each table
# ADHD_PRS_WM
s_regression_ADHD_PRS_WM <- summary(regression_ADHD_PRS_WM)
p_ADHD_PRS_WM <- s_regression_ADHD_PRS_WM$coefficients["ADHD_PRS", "Pr(>|t|)"]
# ADHD_PRS_SSRT
s_regression_ADHD_PRS_SSRT <- summary(regression_ADHD_PRS_SSRT)
p_ADHD_PRS_SSRT <- s_regression_ADHD_PRS_SSRT$coefficients["ADHD_PRS", "Pr(>|t|)"]
# ADHD_PRS_RTV
s_regression_ADHD_PRS_RTV <- summary(regression_ADHD_PRS_RTV)
p_ADHD_PRS_RTV <- s_regression_ADHD_PRS_RTV$coefficients["ADHD_PRS", "Pr(>|t|)"]
# Create a vector of p-values
corr_gen_cog <- c(p_ADHD_PRS_WM, p_ADHD_PRS_SSRT, p_ADHD_PRS_RTV)
# Apply bonferroni correction
p.adjust(corr_gen_cog, method = "bonferroni") # WM and RTV significant

```

```
## [1] 2.333895e-06 1.317462e-01 3.523493e-08
```

Regressions - ASD PRS & MDD PRS regressed onto cognition

```

# ASD PRS - WM
regression_ASDD_PRS_WM <- lm(formula = arc_WM ~ ASD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalWM_group)
summary(regression_ASDD_PRS_WM)

```

```

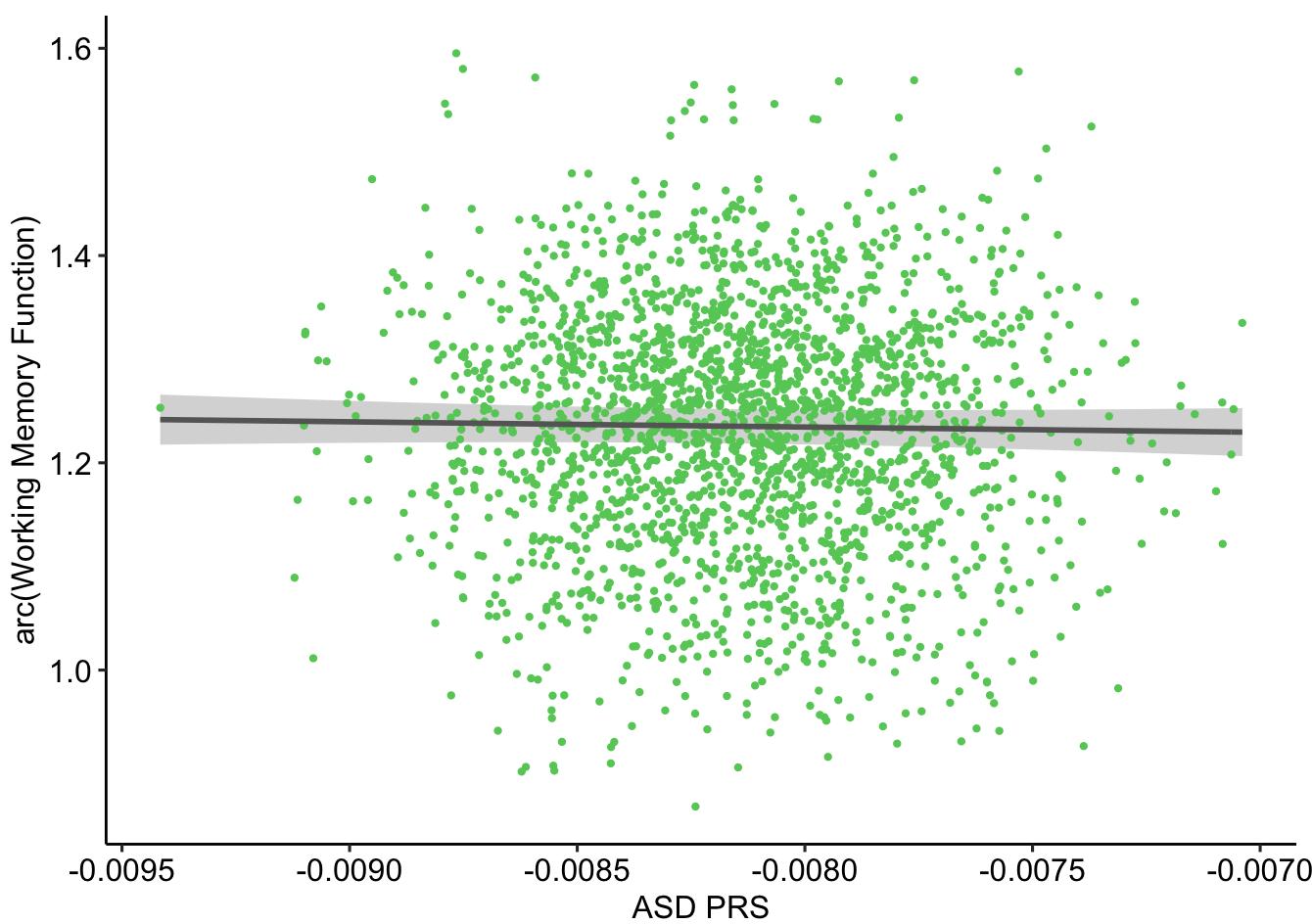
## 
## Call:
## lm(formula = arc_WM ~ ASD_PRS + Sex + Age + Age2 + AgeSex + PC1 +
##     PC2 + PC3, data = data_finalWM_group)
##
## Residuals:
##      Min      1Q Median      3Q      Max
## -0.36739 -0.07429  0.00743  0.07916  0.35680
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 1.913059  0.947218  2.020   0.0435 *  
## ASD_PRS     -5.059243  7.226279 -0.700   0.4839    
## Sex        -0.039459  0.093381 -0.423   0.6727    
## Age       -0.155669  0.157145 -0.991   0.3220    
## Age2       0.008003  0.006544  1.223   0.2215    
## AgeSex     0.001905  0.007819  0.244   0.8075    
## PC1        -1.319244  1.401063 -0.942   0.3465    
## PC2        -0.360364  1.685299 -0.214   0.8307    
## PC3        -0.893772  1.096983 -0.815   0.4153    
## ---      
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.1169 on 2212 degrees of freedom
## Multiple R-squared:  0.04961,    Adjusted R-squared:  0.04617 
## F-statistic: 14.43 on 8 and 2212 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(regression_ASD_PRS_WM)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-2.288175e-16	0.95	-0.04063938	0.04063938
2 ASD_PRS	-1.455697e-02	0.95	-0.05533120	0.02621727
3 Sex	-1.638063e-01	0.95	-0.92401588	0.59640338
4 Age	-8.346126e-01	0.95	-2.48684180	0.81761670
5 Age2	1.025543e+00	0.95	-0.61890496	2.66999147
6 AgeSex	9.480127e-02	0.95	-0.66812320	0.85772574
7 PC1	-2.296564e-02	0.95	-0.07079525	0.02486397
8 PC2	-4.483170e-03	0.95	-0.04559873	0.03663239
9 PC3	-2.003526e-02	0.95	-0.06825825	0.02818774
9 rows				

```
# graph
visreg(regression_ASD_PRS_WM, xvar = "ASD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg = TRUE,
line=list(col="#666666"), points=list(col="#66CC66")) +
  theme_pubr() +
  labs(x = "ASD PRS", y = "arc(Working Memory Function)")
```



```
# MDD PRS - WM
regression_MDD_PRS_WM <- lm(formula = arc_WM ~ MDD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalWM_group)
summary(regression_MDD_PRS_WM)
```

```

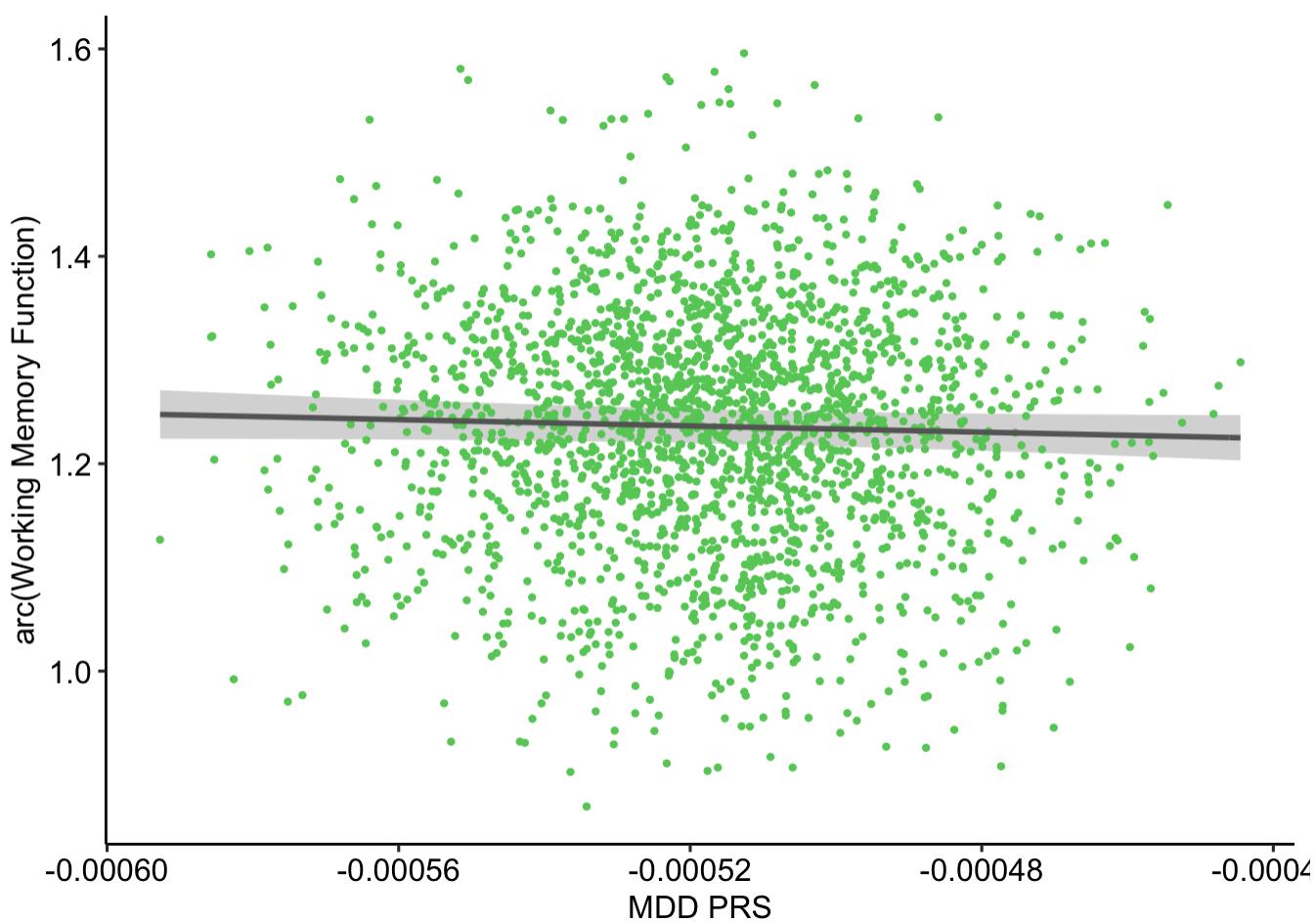
## 
## Call:
## lm(formula = arc_WM ~ MDD_PRS + Sex + Age + Age2 + AgeSex + PC1 +
##     PC2 + PC3, data = data_finalWM_group)
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -0.36915 -0.07423  0.00803  0.07906  0.36048
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 1.845e+00  9.484e-01  1.946   0.0518 .  
## MDD_PRS    -1.514e+02  1.058e+02 -1.432   0.1524    
## Sex        -4.672e-02  9.345e-02 -0.500   0.6171    
## Age        -1.494e-01  1.571e-01 -0.951   0.3416    
## Age2        7.705e-03  6.541e-03  1.178   0.2389    
## AgeSex      2.477e-03  7.824e-03  0.317   0.7516    
## PC1        -1.234e+00  1.401e+00 -0.881   0.3785    
## PC2        -3.680e-01  1.685e+00 -0.218   0.8271    
## PC3        -8.351e-01  1.095e+00 -0.763   0.4458    
## ---      
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.1169 on 2212 degrees of freedom
## Multiple R-squared:  0.05028,   Adjusted R-squared:  0.04684 
## F-statistic: 14.64 on 8 and 2212 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(regression_MDD_PRS_WM)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-2.407405e-16	0.95	-0.04062507	0.04062507
2 MDD_PRS	-2.974362e-02	0.95	-0.07048567	0.01099843
3 Sex	-1.939574e-01	0.95	-0.95472428	0.56680948
4 Age	-8.010959e-01	0.95	-2.45258865	0.85039691
5 Age2	9.873923e-01	0.95	-0.65638403	2.63116863
6 AgeSex	1.232531e-01	0.95	-0.64017921	0.88668535
7 PC1	-2.147540e-02	0.95	-0.06929070	0.02633991
8 PC2	-4.578670e-03	0.95	-0.04567556	0.03651822
9 PC3	-1.871925e-02	0.95	-0.06685360	0.02941510
9 rows				

```
# graph
visreg(regression_MDD_PRS_WM, xvar = "MDD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg = TRUE,
line=list(col="#666666"), points=list(col="#66CC66")) +
  theme_pubr() +
  labs(x = "MDD PRS", y = "arc(Working Memory Function)")
```



```
# ASD PRS - SSRT
regression_ASD_PRS_SSRT <- lm(formula = log_SSRT ~ ASD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalSSRT_group)
summary(regression_ASD_PRS_SSRT)
```

```

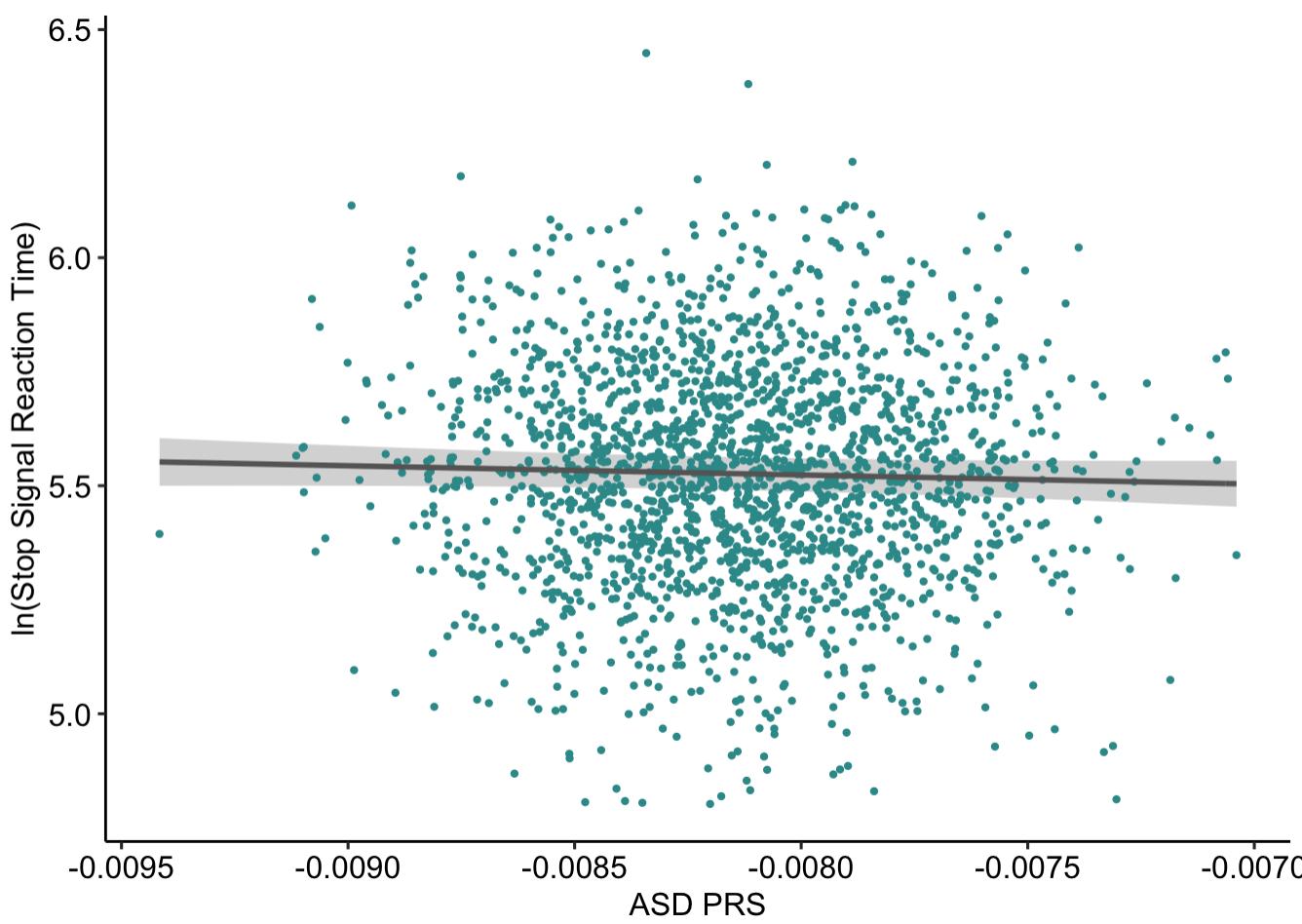
## 
## Call:
## lm(formula = log_SSRT ~ ASD_PRS + Sex + Age + Age2 + AgeSex +
##     PC1 + PC2 + PC3, data = data_finalSSRT_group)
##
## Residuals:
##      Min      1Q Median      3Q      Max 
## -0.7270 -0.1587  0.0035  0.1626  0.9180 
## 
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 5.481895  2.063995  2.656  0.00797 **  
## ASD_PRS     -20.080397 15.627508 -1.285  0.19896    
## Sex        -0.124676  0.203101 -0.614  0.53938    
## Age         0.032086  0.341887  0.094  0.92524    
## Age2       -0.003237  0.014215 -0.228  0.81989    
## AgeSex      0.008940  0.017020  0.525  0.59948    
## PC1        2.503211  3.035808  0.825  0.40972    
## PC2        -2.425443  3.704511 -0.655  0.51272    
## PC3        -0.469279  2.386267 -0.197  0.84412    
## --- 
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1 
## 
## Residual standard error: 0.2419 on 1995 degrees of freedom
## Multiple R-squared:  0.0102, Adjusted R-squared:  0.00623 
## F-statistic:  2.57 on 8 and 1995 DF,  p-value: 0.008657

```

```
effectsize::standardize_parameters(regression_ASD_PRS_SSRT)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-8.445662e-16	0.95	-0.04367227	0.04367227
2 ASD_PRS	-2.874324e-02	0.95	-0.07261294	0.01512647
3 Sex	-2.558913e-01	0.95	-1.07340529	0.56162265
4 Age	8.493968e-02	0.95	-1.69002295	1.85990231
5 Age2	-2.048098e-01	0.95	-1.96873066	1.55911113
6 AgeSex	2.192806e-01	0.95	-0.59947956	1.03804076
7 PC1	2.172657e-02	0.95	-0.02994833	0.07340147
8 PC2	-1.477734e-02	0.95	-0.05904103	0.02948636
9 PC3	-5.230364e-03	0.95	-0.05738966	0.04692894
9 rows				

```
# graph
visreg(regression_ASD_PRS_SSRT, xvar = "ASD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg =
TRUE, line=list(col="#666666"), points=list(col="#339999")) +
  theme_pubr() +
  labs(x = "ASD PRS", y = "ln(Stop Signal Reaction Time)")
```



```
# MDD PRS - SSRT
regression_MDD_PRS_SSRT <- lm(formula = log_SSRT ~ MDD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalSSRT_group)
summary(regression_MDD_PRS_SSRT)
```

```

## 
## Call:
## lm(formula = log_SSRT ~ MDD_PRS + Sex + Age + Age2 + AgeSex +
##     PC1 + PC2 + PC3, data = data_finalSSRT_group)
##
## Residuals:
##      Min        1Q    Median        3Q       Max
## -0.72595 -0.15729  0.00578  0.16289  0.92791
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 5.771845  2.068916  2.790  0.00532 **  
## MDD_PRS     250.008592 228.647957  1.093  0.27434    
## Sex        -0.119059  0.203308 -0.586  0.55821    
## Age         0.031969  0.341961  0.093  0.92553    
## Age2        -0.003202  0.014219 -0.225  0.82185    
## AgeSex      0.008448  0.017037  0.496  0.62003    
## PC1         2.551037  3.035417  0.840  0.40077    
## PC2        -2.309102  3.704174 -0.623  0.53311    
## PC3        -0.272652  2.381135 -0.115  0.90885    
## ---        
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.2419 on 1995 degrees of freedom
## Multiple R-squared:  0.009973, Adjusted R-squared:  0.006003 
## F-statistic: 2.512 on 8 and 1995 DF,  p-value: 0.01024

```

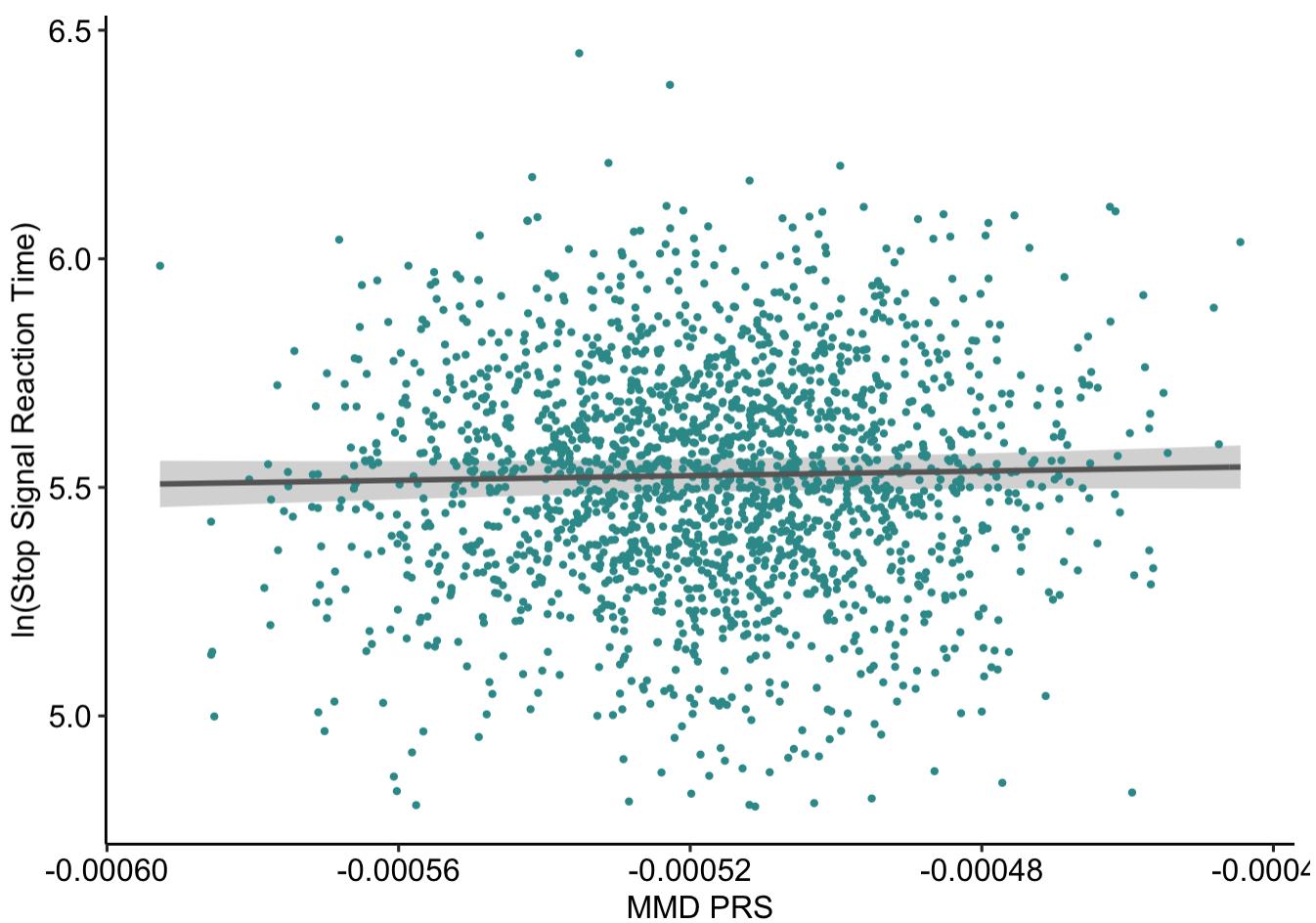
```
effectsize::standardize_parameters(regression_MDD_PRS_SSRT)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-8.634033e-16	0.95	-0.04367725	0.04367725
2 MDD_PRS	2.442870e-02	0.95	-0.01938646	0.06824386
3 Sex	-2.443615e-01	0.95	-1.06271133	0.57398825
4 Age	8.463068e-02	0.95	-1.69071469	1.85997606
5 Age2	-2.026007e-01	0.95	-1.96702117	1.56181973
6 AgeSex	2.072244e-01	0.95	-0.61232447	1.02677325
7 PC1	2.214167e-02	0.95	-0.02952657	0.07380991
8 PC2	-1.406852e-02	0.95	-0.05832818	0.03019114
9 PC3	-3.038852e-03	0.95	-0.05508596	0.04900826
9 rows				

```

# graph
visreg(regression_MDD_PRS_SSRT, xvar = "MDD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg =
TRUE, line=list(col="#666666"), points=list(col="#339999")) +
  theme_pubr() +
  labs(x = "MMD PRS", y = "ln(Stop Signal Reaction Time)")

```



```
# ASD PRS - RTV
regression_ASD_PRS_RTV <- lm(formula = RTV ~ ASD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalRTV_group)
summary(regression_ASD_PRS_RTV)
```

```

## 
## Call:
## lm(formula = RTV ~ ASD_PRS + Sex + Age + Age2 + AgeSex + PC1 +
##     PC2 + PC3, data = data_finalRTV_group)
##
## Residuals:
##      Min      1Q Median      3Q      Max
## -70.356 -17.430   0.761  17.322  95.485
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 18.2498   211.1790   0.086  0.93114    
## ASD_PRS     647.0931  1605.0385   0.403  0.68687    
## Sex        -5.9429   20.7083  -0.287  0.77415    
## Age         27.4213   34.9982   0.784  0.43342    
## Age2        -1.4643    1.4553  -1.006  0.31443    
## AgeSex       0.5936    1.7343   0.342  0.73220    
## PC1         147.5590  310.6978   0.475  0.63489    
## PC2        -370.9201  376.8337  -0.984  0.32508    
## PC3         776.1168  243.5353   3.187  0.00146 **  
## ---      
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 25.37 on 2113 degrees of freedom
## Multiple R-squared:  0.03538,   Adjusted R-squared:  0.03173 
## F-statistic: 9.689 on 8 and 2113 DF,  p-value: 2.798e-13

```

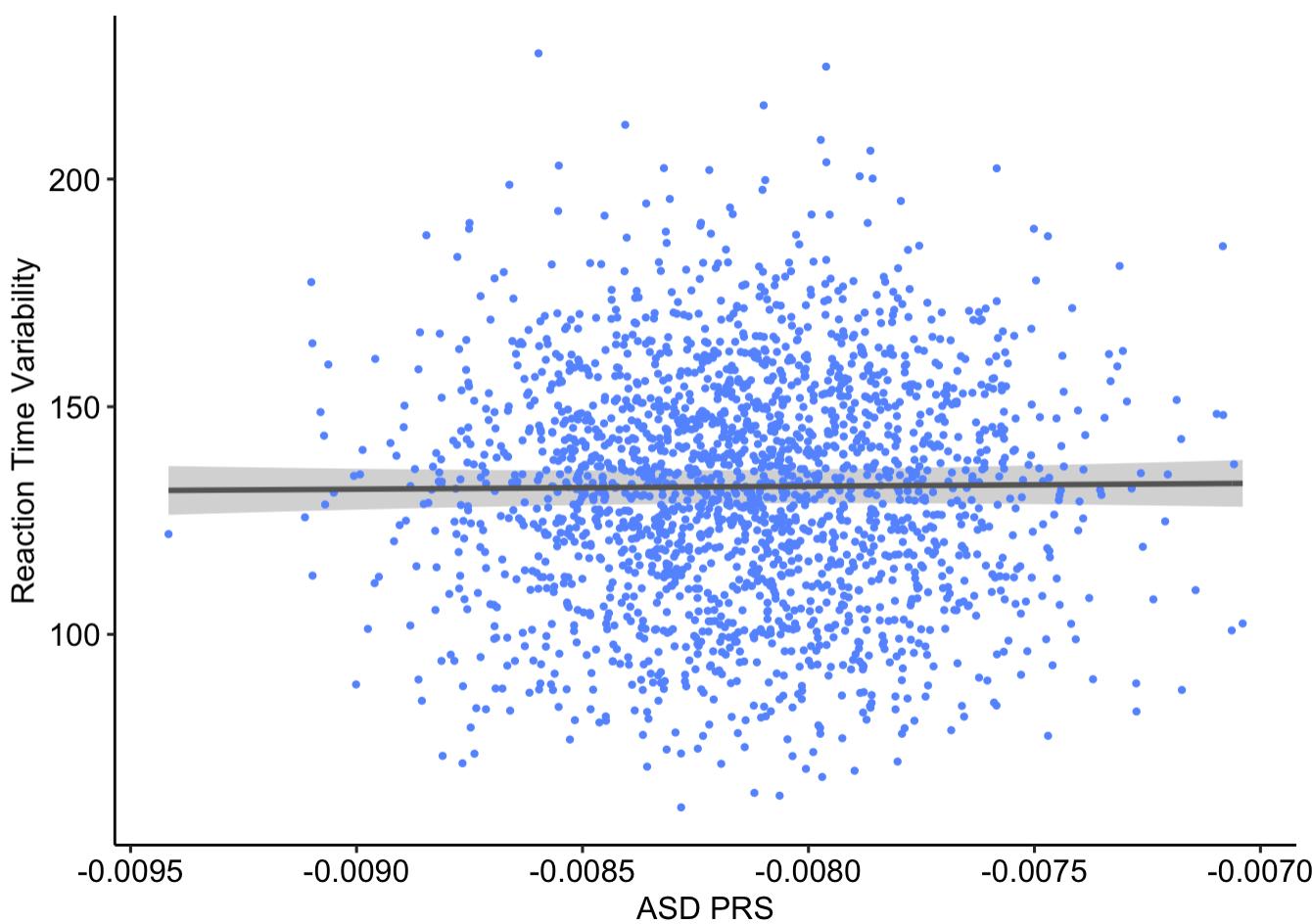
```
effectsize::standardize_parameters(regression_ASD_PRS_RTV)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-1.302076e-15	0.95	-0.04189113	0.04189113
2 ASD_PRS	8.645613e-03	0.95	-0.03340879	0.05070001
3 Sex	-1.147416e-01	0.95	-0.89882948	0.66934619
4 Age	6.835315e-01	0.95	-1.02732343	2.39438635
5 Age2	-8.725537e-01	0.95	-2.57314733	0.82803999
6 AgeSex	1.370699e-01	0.95	-0.64835205	0.92249177
7 PC1	1.198681e-02	0.95	-0.03750954	0.06148316
8 PC2	-2.129849e-02	0.95	-0.06373259	0.02113561
9 PC3	8.112833e-02	0.95	0.03120491	0.13105175
9 rows				

```

# graph
visreg(regression_ASD_PRS_RTV, xvar = "ASD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg =
TRUE, line=list(col="#666666"), points=list(col="#6699FF")) +
  theme_pubr() +
  labs(x = "ASD PRS", y = "Reaction Time Variability")

```



```
# MDD PRS - RTV
regression_MDD_PRS_RTV <- lm(formula = RTV ~ MDD_PRS +
                                Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3, data = data_finalRTV_group)
summary(regression_MDD_PRS_RTV)
```

```

## 
## Call:
## lm(formula = RTV ~ MDD_PRS + Sex + Age + Age2 + AgeSex + PC1 +
##     PC2 + PC3, data = data_finalRTV_group)
##
## Residuals:
##      Min      1Q Median      3Q      Max
## -69.973 -17.191   0.841  17.092  95.119
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 55.0660   211.5139   0.260  0.79463    
## MDD_PRS    53051.3779  23261.3437   2.281  0.02267 *  
## Sex        -3.5755    20.7051  -0.173  0.86291    
## Age         24.6214    34.9592   0.704  0.48133    
## Age2        -1.3358    1.4538  -0.919  0.35828    
## AgeSex       0.4046    1.7340   0.233  0.81553    
## PC1         123.5961   310.3393   0.398  0.69048    
## PC2        -366.0114   376.3212  -0.973  0.33086    
## PC3         765.9162   242.7289   3.155  0.00163 ** 
## ---
## Signif. codes:  0 '****' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 25.34 on 2113 degrees of freedom
## Multiple R-squared:  0.03768,    Adjusted R-squared:  0.03404 
## F-statistic: 10.34 on 8 and 2113 DF,  p-value: 2.715e-14

```

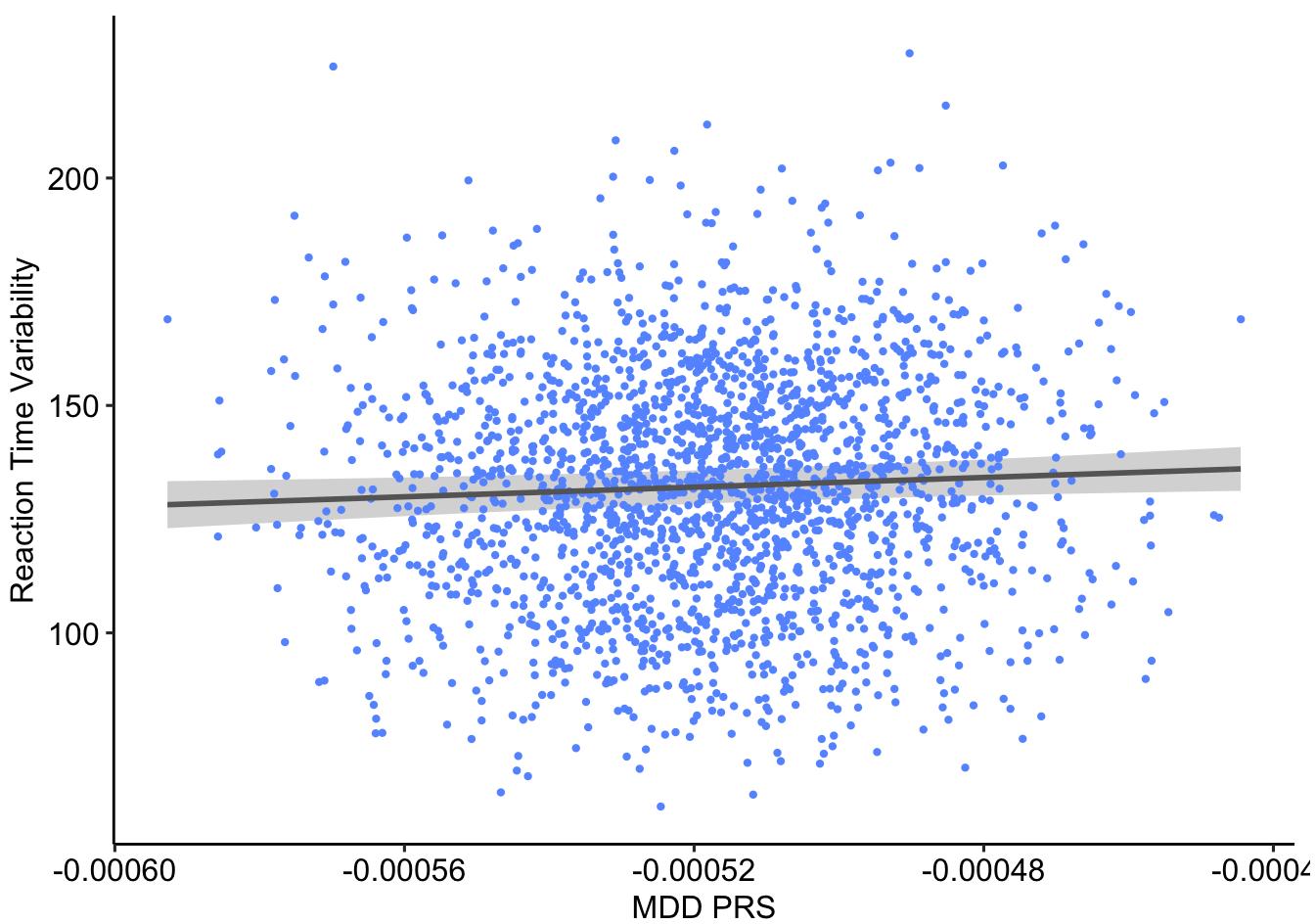
```
effectsize::standardize_parameters(regression_MDD_PRS_RTV)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-1.149956e-15	0.95	-0.04184127	0.04184127
2 MDD_PRS	4.881602e-02	0.95	0.00684037	0.09079167
3 Sex	-6.903361e-02	0.95	-0.85300168	0.71493446
4 Age	6.137388e-01	0.95	-1.09520936	2.32268696
5 Age2	-7.959694e-01	0.95	-2.49480343	0.90286457
6 AgeSex	9.342630e-02	0.95	-0.69182832	0.87868093
7 PC1	1.004021e-02	0.95	-0.03939902	0.05947944
8 PC2	-2.101662e-02	0.95	-0.06339302	0.02135977
9 PC3	8.006205e-02	0.95	0.03030393	0.12982016
9 rows				

```

# graph
visreg(regression_MDD_PRS_RTV, xvar = "MDD_PRS", rug = FALSE, jitter = TRUE, overlay = TRUE, gg =
TRUE, line=list(col="#666666"), points=list(col="#6699FF")) +
  theme_pubr() +
  labs(x = "MDD PRS", y = "Reaction Time Variability")

```



```

# Correction for multiple testing
# Extract p-values from each table
# ASD_PRS_WM
s_regression_ASD_PRS_WM <- summary(regression_ASD_PRS_WM)
p_ASD_PRS_WM <- s_regression_ASD_PRS_WM$coefficients["ASD_PRS", "Pr(>|t|)"]
# MDD_PRS_WM
s_regression_MDD_PRS_WM <- summary(regression_MDD_PRS_WM)
p_MDD_PRS_WM <- s_regression_MDD_PRS_WM$coefficients["MDD_PRS", "Pr(>|t|)"]
# ASD_PRS_SSRT
s_regression_ASD_PRS_SSRT <- summary(regression_ASD_PRS_SSRT)
p_ASD_PRS_SSRT <- s_regression_ASD_PRS_SSRT$coefficients["ASD_PRS", "Pr(>|t|)"]
# MDD_PRS_SSRT
s_regression_MDD_PRS_SSRT <- summary(regression_MDD_PRS_SSRT)
p_MDD_PRS_SSRT <- s_regression_MDD_PRS_SSRT$coefficients["MDD_PRS", "Pr(>|t|)"]
# ASD_PRS_RTV
s_regression_ASD_PRS_RTV <- summary(regression_ASD_PRS_RTV)
p_ASD_PRS_RTV <- s_regression_ASD_PRS_RTV$coefficients["ASD_PRS", "Pr(>|t|)"]
# MDD_PRS_RTV
s_regression_MDD_PRS_RTV <- summary(regression_MDD_PRS_RTV)
p_MDD_PRS_RTV <- s_regression_MDD_PRS_RTV$coefficients["MDD_PRS", "Pr(>|t|)"]
# Create a vector of p-values
corr_spec <- c(p_ASD_PRS_WM, p_MDD_PRS_WM, p_ASD_PRS_SSRT, p_MDD_PRS_SSRT,
                p_ASD_PRS_RTV, p_MDD_PRS_RTV)
# Apply bonferroni correction
p.adjust(corr_spec, method = "bonferroni") # none are significant

```

```
## [1] 1.0000000 0.9143127 1.0000000 1.0000000 1.0000000 0.1360039
```

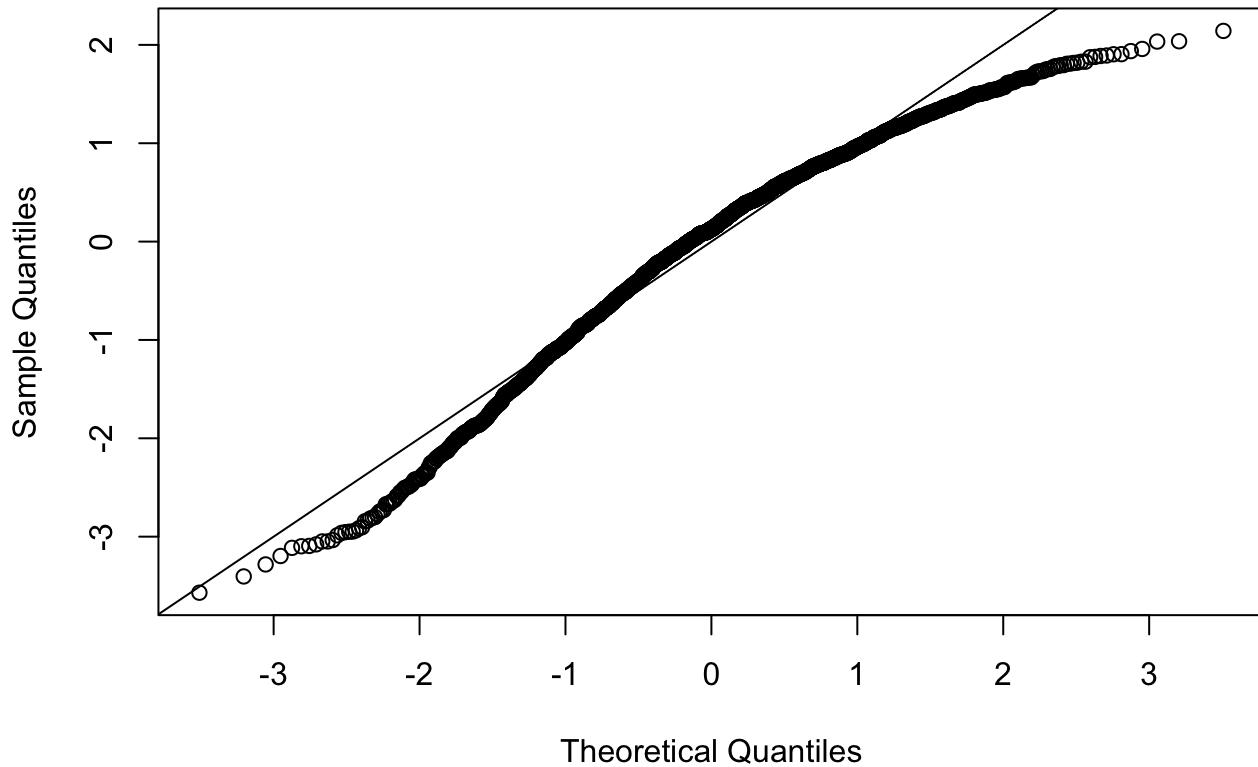
Mediation - ADHD PRS | working memory | ADHD traits

```
#Set a seed for the sake of replicability when bootstrapping  
set.seed(5)
```

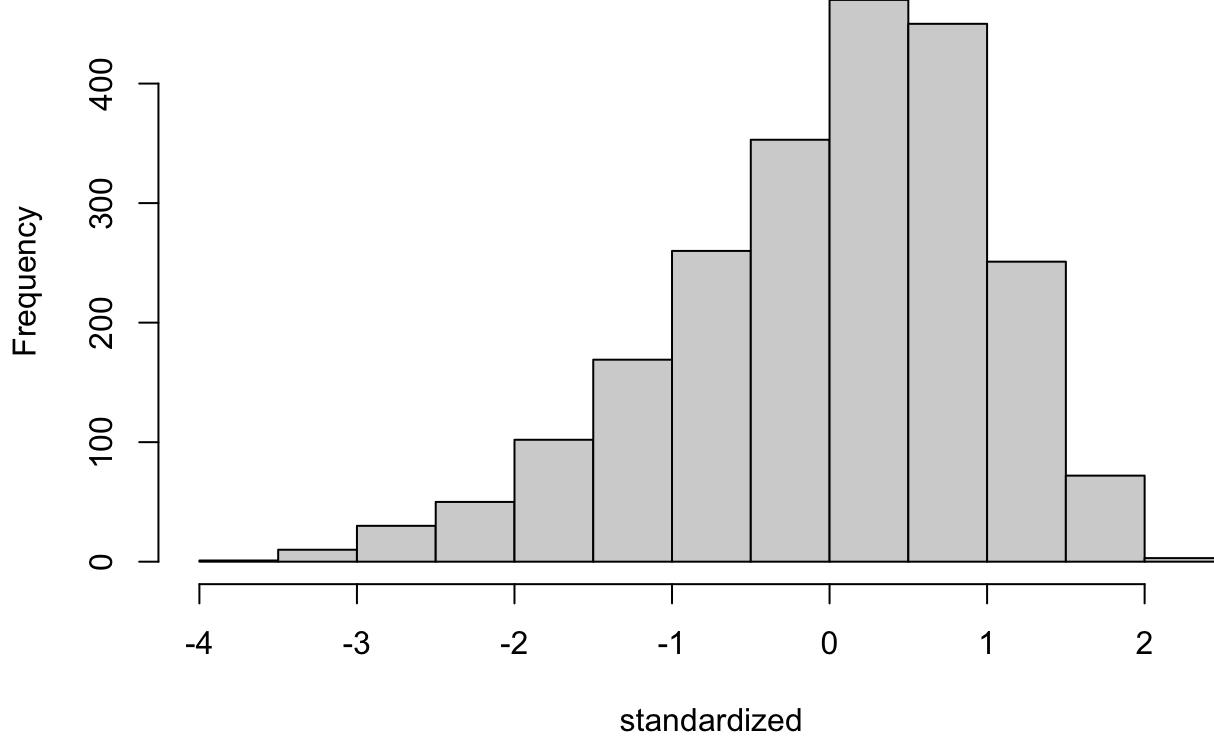
```
# Create the model
```

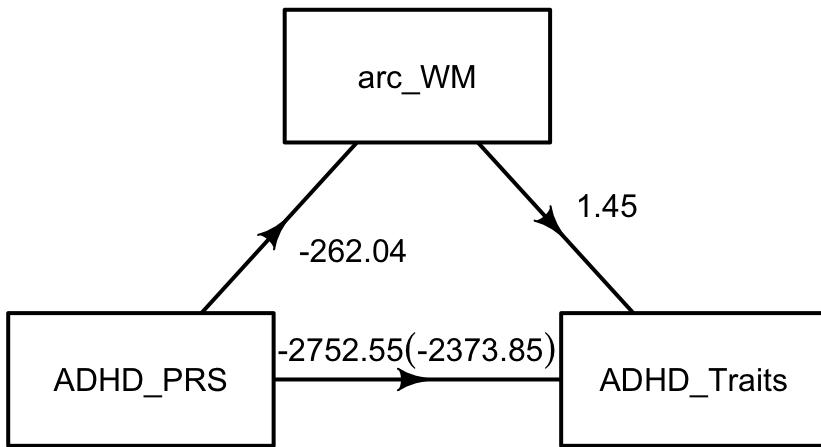
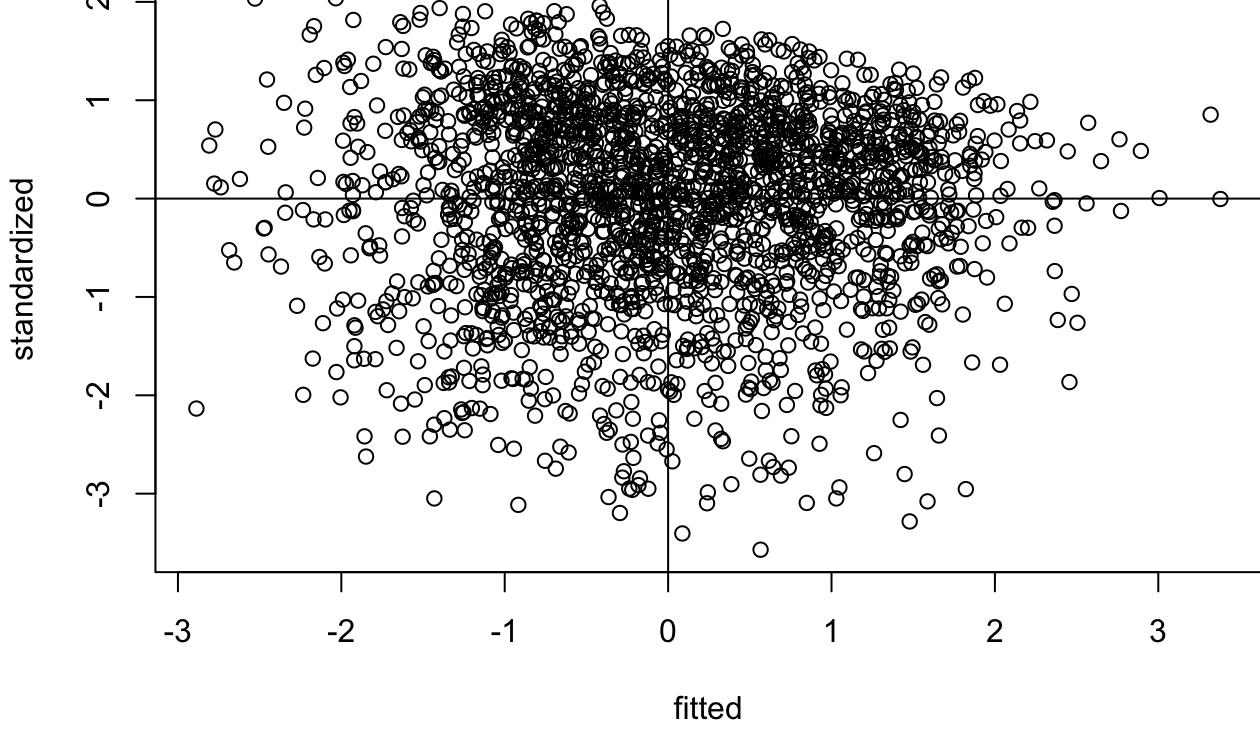
```
WM_meditation_model <- mediation1(y = "ADHD_Traits", #DV  
                                     x = "ADHD_PRS", #IV  
                                     m = "arc_WM", #Mediator  
                                     cvs = c("Age", "Sex", "Age2", "AgeSex", "PC1", "PC2", "PC3"), #Covariates  
                                     df = data_finalWM_group, #Datafram  
                                     with_out = T, #Including outliers  
                                     nboot = 5000, #Number of bootstraps  
                                     conf_level = .95 #CI width  
                                     )
```

Normal Q-Q Plot



Histogram of standardized





```
#bootstrapped indirect effect  
WM_mediation_model$boot.results
```

```
##  
## ORDINARY NONPARAMETRIC BOOTSTRAP  
##  
##  
## Call:  
## boot(data = finaldata, statistic = indirectmed, R = nboot, formula2 = allformulas$eq2,  
##       formula3 = allformulas$eq3, x = x, med.var = m)  
##  
##  
## Bootstrap Statistics :  
##      original     bias    std. error  
## t1* -378.701 -1.689526   92.15222
```

```
#bootstrapped CI (test of significance)  
WM_meditation_model$boot.ci # doesn't include 0 = significant
```

```
## BOOTSTRAP CONFIDENCE INTERVAL CALCULATIONS  
## Based on 5000 bootstrap replicates  
##  
## CALL :  
## boot.ci(boot.out = bootresults, conf = conf_level, type = "norm")  
##  
## Intervals :  
## Level      Normal  
## 95%  (-557.6, -196.4 )  
## Calculations and Intervals on Original Scale
```

```
# STANDARDIZE results  
# fit the models  
# a path  
mediation_1 <- lm(arc_WM ~ ADHD_PRS + Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3,  
                     data = data_finalWM_group)  
summary(mediation_1)
```

```

## 
## Call:
## lm(formula = arc_WM ~ ADHD_PRS + Sex + Age + Age2 + AgeSex +
##     PC1 + PC2 + PC3, data = data_finalWM_group)
##
## Residuals:
##      Min        1Q    Median        3Q       Max
## -0.39029 -0.07435  0.00737  0.07943  0.35379
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 1.893e+00  9.413e-01  2.011   0.0445 *  
## ADHD_PRS   -2.620e+02  5.288e+01 -4.955 7.78e-07 *** 
## Sex        -4.651e-02  9.288e-02 -0.501   0.6166    
## Age        -1.317e-01  1.563e-01 -0.843   0.3994    
## Age2        6.972e-03  6.509e-03  1.071   0.2843    
## AgeSex      2.573e-03  7.778e-03  0.331   0.7408    
## PC1        -9.702e-01  1.394e+00 -0.696   0.4867    
## PC2         6.810e-02  1.678e+00  0.041   0.9676    
## PC3        -1.160e+00  1.091e+00 -1.063   0.2878    
## ---        
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.1163 on 2212 degrees of freedom
## Multiple R-squared:  0.05983,    Adjusted R-squared:  0.05643 
## F-statistic: 17.6 on 8 and 2212 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(mediation_1)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-3.098663e-16	0.95	-0.04042018	0.04042018
2 ADHD_PRS	-1.030782e-01	0.95	-0.14387298	-0.06228332
3 Sex	-1.930972e-01	0.95	-0.94921710	0.56302275
4 Age	-7.063704e-01	0.95	-2.34992793	0.93718719
5 Age2	8.934378e-01	0.95	-0.74237765	2.52925330
6 AgeSex	1.280405e-01	0.95	-0.63082734	0.88690831
7 PC1	-1.688960e-02	0.95	-0.06449512	0.03071592
8 PC2	8.472297e-04	0.95	-0.04008809	0.04178255
9 PC3	-2.600880e-02	0.95	-0.07397629	0.02195870
9 rows				

```

# b and c' paths
mediation_2 <- lm(ADHD_Traits ~ ADHD_PRS + arc_WM + Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3,
                    data = data_finalWM_group)
summary(mediation_2)

```

```

## 
## Call:
## lm(formula = ADHD_Traits ~ ADHD_PRS + arc_WM + Sex + Age + Age2 +
##      AgeSex + PC1 + PC2 + PC3, data = data_finalWM_group)
##
## Residuals:
##    Min      1Q  Median      3Q     Max 
## -3.5472 -0.6125  0.1272  0.7384  2.1350 
##
## Coefficients:
##              Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 1.180e+00  8.096e+00  0.146   0.8841    
## ADHD_PRS    -2.374e+03  4.569e+02 -5.195 2.23e-07 *** 
## arc_WM       1.445e+00  1.827e-01  7.910 4.03e-15 *** 
## Sex          -1.129e+00  7.982e-01 -1.415   0.1573    
## Age          -2.354e-01  1.343e+00 -0.175   0.8609    
## Age2         -5.029e-04  5.595e-02 -0.009   0.9928    
## AgeSex       1.306e-01  6.683e-02  1.953   0.0509 .  
## PC1          -3.515e-01  1.198e+01 -0.029   0.9766    
## PC2          6.369e+00  1.442e+01  0.442   0.6587    
## PC3          -6.776e+00  9.379e+00 -0.723   0.4701    
## --- 
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1 
##
## Residual standard error: 0.9995 on 2211 degrees of freedom
## Multiple R-squared:  0.0787, Adjusted R-squared:  0.07495 
## F-statistic: 20.99 on 9 and 2211 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(mediation_2)
```

	Parameter	Std_Coefficient	CI	CI_low	CI_high
	<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1	(Intercept)	-3.351996e-16	0.95	-0.040021609	0.04002161
2	ADHD_PRS	-1.075976e-01	0.95	-0.148213692	-0.06698145
3	arc_WM	1.665248e-01	0.95	0.125240062	0.20780954
4	Sex	-5.401231e-01	0.95	-1.288829558	0.20858344
5	Age	-1.454197e-01	0.95	-1.773031933	1.48219249
6	Age2	-7.426210e-03	0.95	-1.627531349	1.61267893
7	AgeSex	7.484834e-01	0.95	-0.002920066	1.49988684
8	PC1	-7.049989e-04	0.95	-0.047846249	0.04643625
9	PC2	9.129313e-03	0.95	-0.031402373	0.04966100
10	PC3	-1.750314e-02	0.95	-0.065009782	0.03000349

1-10 of 10 rows

```
# c path
mediation_3 <- lm(ADHD_Traits ~ ADHD_PRS + Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3,
                   data = data_finalWM_group)
summary(mediation_3)
```

```
##
## Call:
## lm(formula = ADHD_Traits ~ ADHD_PRS + Sex + Age + Age2 + AgeSex +
##      PC1 + PC2 + PC3, data = data_finalWM_group)
##
## Residuals:
##    Min      1Q  Median      3Q     Max 
## -3.5023 -0.6172  0.1371  0.7520  2.0681 
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 3.916e+00 8.201e+00  0.478   0.6330    
## ADHD_PRS   -2.753e+03 4.607e+02 -5.975 2.68e-09 *** 
## Sex        -1.196e+00 8.091e-01 -1.479   0.1394    
## Age        -4.258e-01 1.362e+00 -0.313   0.7546    
## Age2        9.573e-03 5.671e-02  0.169   0.8660    
## AgeSex      1.343e-01 6.776e-02  1.982   0.0476 *   
## PC1        -1.754e+00 1.215e+01 -0.144   0.8852    
## PC2         6.467e+00 1.462e+01  0.442   0.6582    
## PC3        -8.453e+00 9.506e+00 -0.889   0.3740    
## ---      
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 1.013 on 2212 degrees of freedom
## Multiple R-squared:  0.05263,    Adjusted R-squared:  0.0492 
## F-statistic: 15.36 on 8 and 2212 DF,  p-value: < 2.2e-16
```

```
effectsize::standardize_parameters(mediation_3)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-3.868000e-16	0.95	-0.040574748	0.04057475
2 ADHD_PRS	-1.247626e-01	0.95	-0.165713473	-0.08381181
3 Sex	-5.722785e-01	0.95	-1.331289868	0.18673281
4 Age	-2.630479e-01	0.95	-1.912890456	1.38679464
5 Age2	1.413533e-01	0.95	-1.500717516	1.78342421
6 AgeSex	7.698053e-01	0.95	0.008035556	1.53157505
7 PC1	-3.517536e-03	0.95	-0.051305097	0.04427002
8 PC2	9.270398e-03	0.95	-0.031821462	0.05036226
9 PC3	-2.183425e-02	0.95	-0.069985177	0.02631667
9 rows				

```
# standardized indirect effect
WM_M1 <- lm.beta(mediation_1) # standardized a path
WM_M2 <- lm.beta(mediation_2) # standardized b path
WM_ie <- (WM_M1$standardized.coefficients["ADHD_PRS"])*(WM_M2$standardized.coefficients["arc_WM"])
# times the standardized a path by the standardized b path
WM_ie
```

```
## ADHD_PRS
## -0.01716507
```

```
# proportion mediated
WM_M3 <- lm.beta(mediation_3) # standardized c (total) path
WM_ie/(WM_M3$standardized.coefficients["ADHD_PRS"])# divide the standardized indirect effect by the standardized total effect
```

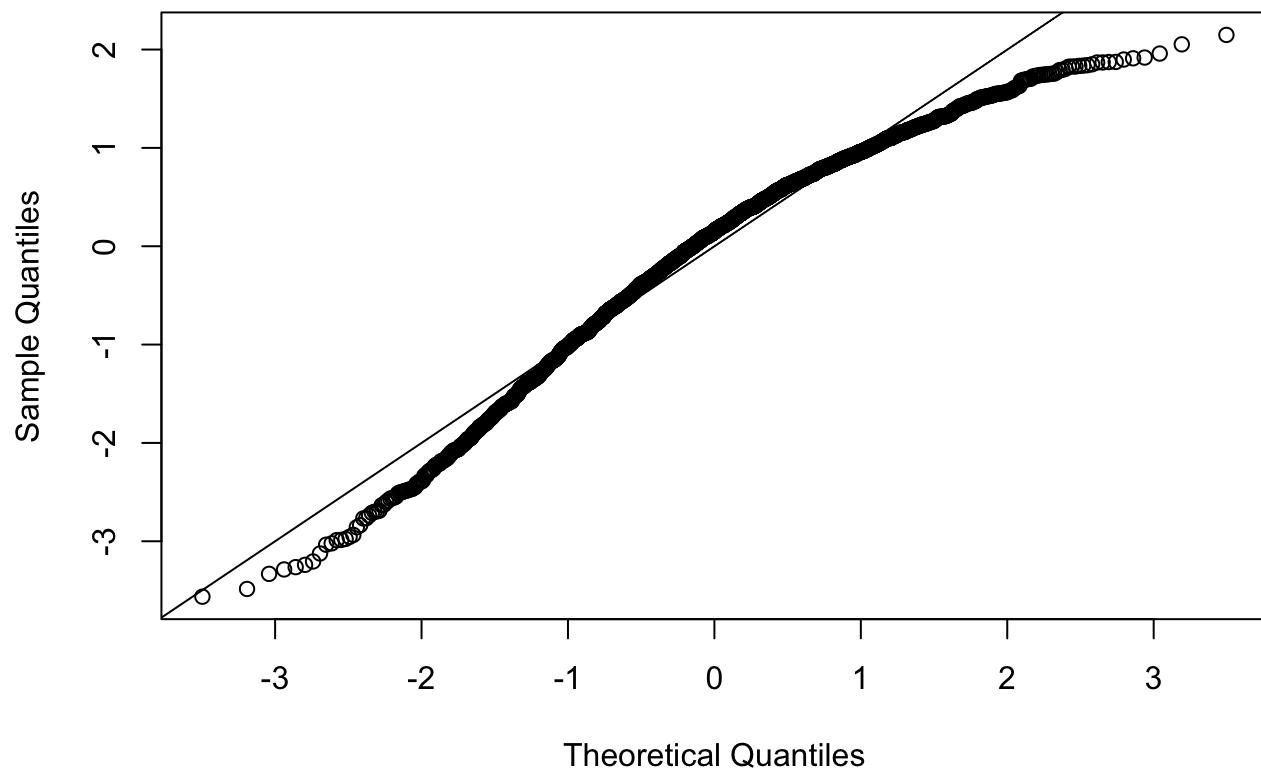
```
## ADHD_PRS
## 0.1375818
```

Mediation - ADHD PRS | reaction time variability | ADHD traits

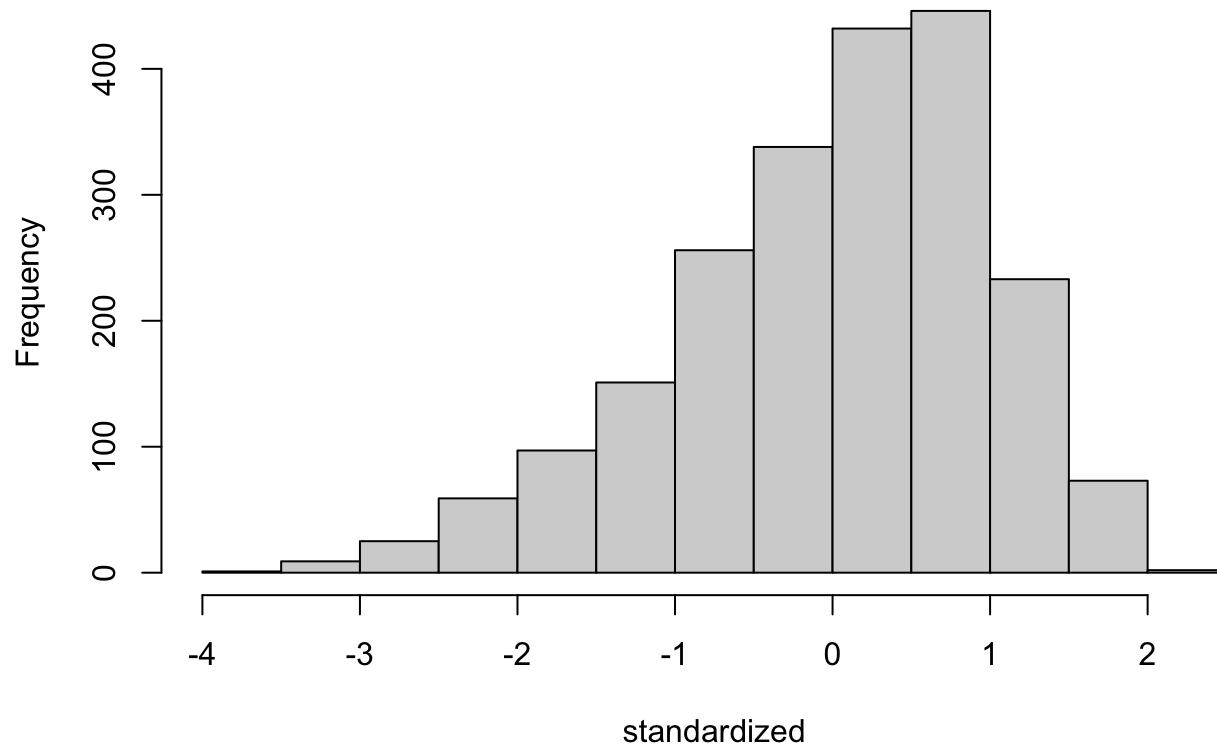
```
#Set a seed for the sake of replicability when bootstrapping
set.seed(5)
```

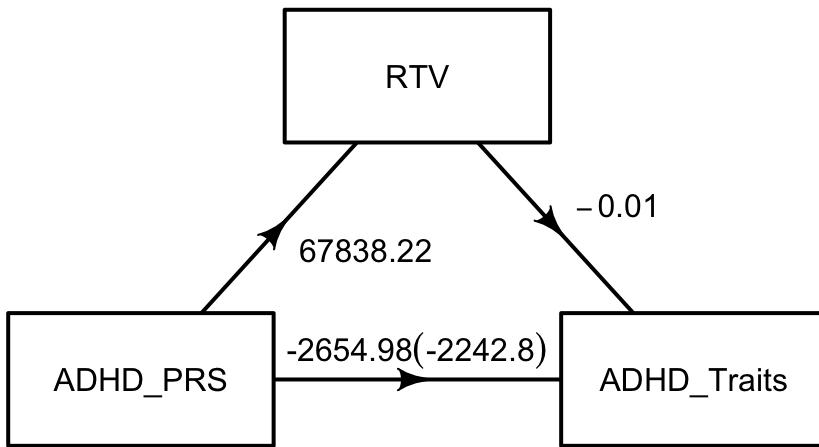
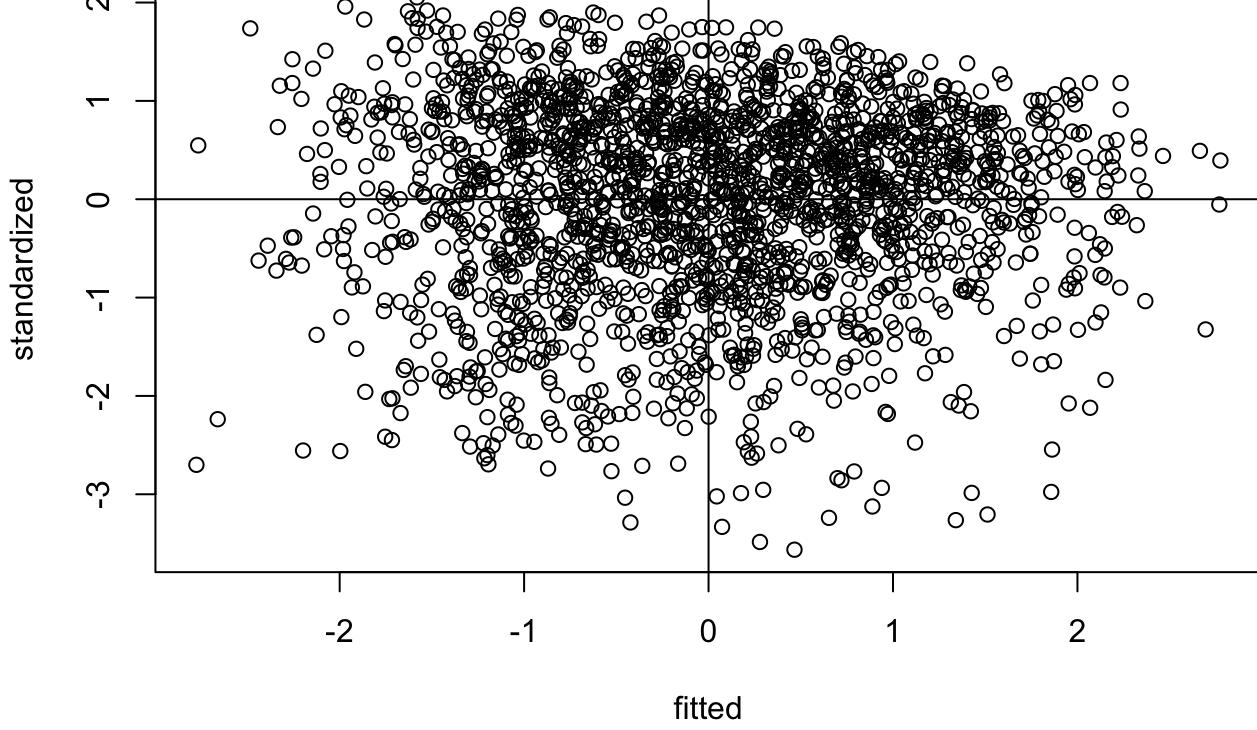
```
# Create the model
RTV_meditation_model <- mediation1(y = "ADHD_Traits", #DV
                                      x = "ADHD_PRS", #IV
                                      m = "RTV", #Mediator
                                      cvs = c("Age", "Sex", "Age2", "AgeSex", "PC1", "PC2", "PC3"), #Covariates
                                      df = data_finalRTV_group, #Datafram
                                      with_out = T, #Including outliers
                                      nboot = 5000, #Number of bootstraps
                                      conf_level = .95 #CI width
                                      )
```


Normal Q-Q Plot



Histogram of standardized





```
#bootstrapped indirect effect  
RTV_meditation_model$boot.results
```

```
##  
## ORDINARY NONPARAMETRIC BOOTSTRAP  
##  
##  
## Call:  
## boot(data = finaldata, statistic = indirectmed, R = nboot, formula2 = allformulas$eq2,  
##       formula3 = allformulas$eq3, x = x, med.var = m)  
##  
##  
## Bootstrap Statistics :  
##      original    bias   std. error  
## t1* -412.1706 2.930065   91.64532
```

```
#bootstrapped CI (test of significance)  
RTV_meditation_model$boot.ci # doesn't include 0 = significant
```

```
## BOOTSTRAP CONFIDENCE INTERVAL CALCULATIONS  
## Based on 5000 bootstrap replicates  
##  
## CALL :  
## boot.ci(boot.out = bootresults, conf = conf_level, type = "norm")  
##  
## Intervals :  
## Level      Normal  
## 95%  (-594.7, -235.5 )  
## Calculations and Intervals on Original Scale
```

```
# STANDARDIZE results  
# fit the modelss  
  
# a path  
mediation_4 <- lm(RTV ~ ADHD_PRS + Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3,  
                   data = data_finalRTV_group)  
summary(mediation_4)
```

```

## 
## Call:
## lm(formula = RTV ~ ADHD_PRS + Sex + Age + Age2 + AgeSex + PC1 +
##     PC2 + PC3, data = data_finalRTV_group)
##
## Residuals:
##      Min      1Q Median      3Q      Max
## -70.187 -17.204   0.667  16.563  94.053
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 36.423    209.463   0.174  0.861971    
## ADHD_PRS    67838.223  11847.216   5.726 1.17e-08 *** 
## Sex         -5.023    20.549  -0.244  0.806914    
## Age          20.047    34.739   0.577  0.563944    
## Age2         -1.151    1.444  -0.797  0.425710    
## AgeSex       0.488     1.721   0.284  0.776779    
## PC1          49.077    308.655   0.159  0.873681    
## PC2          -480.926   374.343  -1.285  0.199031    
## PC3          841.029    241.480   3.483  0.000506 *** 
## --- 
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 25.17 on 2113 degrees of freedom
## Multiple R-squared:  0.05005,    Adjusted R-squared:  0.04645 
## F-statistic: 13.92 on 8 and 2113 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(mediation_4)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-1.127441e-15	0.95	-0.04157144	0.04157144
2 ADHD_PRS	1.226384e-01	0.95	0.08063687	0.16463996
3 Sex	-9.697950e-02	0.95	-0.87502512	0.68106612
4 Age	4.997249e-01	0.95	-1.19847346	2.19792329
5 Age2	-6.857673e-01	0.95	-2.37377363	1.00223905
6 AgeSex	1.126960e-01	0.95	-0.66671672	0.89210876
7 PC1	3.986752e-03	0.95	-0.04518416	0.05315766
8 PC2	-2.761511e-02	0.95	-0.06976879	0.01453857
9 PC3	8.791372e-02	0.95	0.03841171	0.13741574
9 rows				

```
# b and c' paths
mediation_5 <- lm(ADHD_Traits ~ ADHD_PRS + RTV + Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3,
                    data = data_finalRTV_group)
summary(mediation_5)
```

```

## 
## Call:
## lm(formula = ADHD_Traits ~ ADHD_PRS + RTV + Sex + Age + Age2 +
##      AgeSex + PC1 + PC2 + PC3, data = data_finalRTV_group)
##
## Residuals:
##    Min      1Q  Median      3Q     Max 
## -3.5238 -0.6041  0.1460  0.7296  2.1301 
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 2.219e+00  8.279e+00  0.268   0.7887    
## ADHD_PRS    -2.243e+03  4.719e+02 -4.753  2.14e-06 *** 
## RTV        -6.076e-03  8.598e-04 -7.066  2.16e-12 *** 
## Sex        -1.261e+00  8.122e-01 -1.552   0.1208    
## Age         3.590e-02  1.373e+00  0.026   0.9791    
## Age2       -1.199e-02  5.710e-02 -0.210   0.8337    
## AgeSex     1.414e-01  6.803e-02  2.079   0.0378 *  
## PC1        7.712e+00  1.220e+01  0.632   0.5273    
## PC2       -3.207e+00  1.480e+01 -0.217   0.8285    
## PC3        6.621e+00  9.572e+00  0.692   0.4892    
## ---      
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1 
##
## Residual standard error: 0.9949 on 2112 degrees of freedom
## Multiple R-squared:  0.07628,    Adjusted R-squared:  0.07234 
## F-statistic: 19.38 on 9 and 2112 DF,  p-value: < 2.2e-16

```

```
effectsize::standardize_parameters(mediation_5)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-3.547633e-16	0.95	-0.04100324	0.04100324
2 ADHD_PRS	-1.011799e-01	0.95	-0.14292760	-0.05943229
3 RTV	-1.516189e-01	0.95	-0.19369840	-0.10953945
4 Sex	-6.073684e-01	0.95	-1.37479051	0.16005375
5 Age	2.233428e-02	0.95	-1.65278509	1.69745365
6 Age2	-1.783398e-01	0.95	-1.84352450	1.48684492
7 AgeSex	8.149329e-01	0.95	0.04615860	1.58370728
8 PC1	1.563400e-02	0.95	-0.03286513	0.06413313
9 PC2	-4.595043e-03	0.95	-0.04618880	0.03699872
10 PC3	1.727212e-02	0.95	-0.03169325	0.06623749

1-10 of 10 rows

```
# c path
mediation_6 <- lm(ADHD_Traits ~ ADHD_PRS + Sex + Age + Age2 + AgeSex + PC1 + PC2 + PC3,
                     data = data_finalRTV_group)
summary(mediation_6)
```

```
##
## Call:
## lm(formula = ADHD_Traits ~ ADHD_PRS + Sex + Age + Age2 + AgeSex +
##      PC1 + PC2 + PC3, data = data_finalRTV_group)
##
## Residuals:
##    Min      1Q  Median      3Q     Max 
## -3.5304 -0.6174  0.1403  0.7455  2.0590 
##
## Coefficients:
##             Estimate Std. Error t value Pr(>|t|)    
## (Intercept) 1.997e+00  8.374e+00  0.239   0.8115    
## ADHD_PRS   -2.655e+03  4.737e+02 -5.605 2.35e-08 *** 
## Sex        -1.230e+00  8.215e-01 -1.497   0.1345    
## Age        -8.590e-02  1.389e+00 -0.062   0.9507    
## Age2       -5.001e-03  5.775e-02 -0.087   0.9310    
## AgeSex     1.385e-01  6.881e-02  2.012   0.0443 *   
## PC1        7.414e+00  1.234e+01  0.601   0.5480    
## PC2       -2.848e-01  1.497e+01 -0.019   0.9848    
## PC3        1.511e+00  9.654e+00  0.157   0.8756    
## ---      
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 1.006 on 2113 degrees of freedom
## Multiple R-squared:  0.05444,    Adjusted R-squared:  0.05086 
## F-statistic: 15.21 on 8 and 2113 DF,  p-value: < 2.2e-16
```

```
effectsize::standardize_parameters(mediation_6)
```

Parameter	Std_Coefficient	CI	CI_low	CI_high
<chr>	<dbl>	<dbl>	<dbl>	<dbl>
1 (Intercept)	-1.838220e-16	0.95	-0.04147526	0.04147526
2 ADHD_PRS	-1.197742e-01	0.95	-0.16167862	-0.07786987
3 Sex	-5.926645e-01	0.95	-1.36890999	0.18358108
4 Age	-5.343348e-02	0.95	-1.74770290	1.64083594
5 Age2	-7.436449e-02	0.95	-1.75846546	1.60973648
6 AgeSex	7.978461e-01	0.95	0.02023660	1.57545558
7 PC1	1.502953e-02	0.95	-0.03402762	0.06408668
8 PC2	-4.080701e-04	0.95	-0.04246422	0.04164808
9 PC3	3.942737e-03	0.95	-0.04544475	0.05333022
9 rows				

```
# standardized indirect effect
RTV_M4 <- lm.beta(mediation_4)
RTV_M5 <- lm.beta(mediation_5)
RTV_ie <- (RTV_M4$standardized.coefficients["ADHD_PRS"])*(RTV_M5$standardized.coefficients["RTV"])
# times the standardized a path by the standardized b path
RTV_ie
```

```
## ADHD_PRS
## -0.0185943
```

```
# proportion mediated
RTV_M6 <- lm.beta(mediation_6) # standardized c (total) path
RTV_ie/(RTV_M6$standardized.coefficients["ADHD_PRS"])# divide the standardized indirect effect by
# the standardized total effect
```

```
## ADHD_PRS
## 0.1552446
```