# Example R and Stata code for a Mendelian randomization analysis

Sanderson et al.

20 March 2023

#### R code

• Read in the data

```
library(tidyverse)
library(haven)
library(remotes)

dat <- read_dta("https://raw.github.com/remlapmot/mrrobust/master/dodata.dta")
dat <- dat %>% filter(ldlcp2 < 1e-8)</pre>
```

## Example code using the MendelianRandomization package

• Install the package and load it into memory

```
library(MendelianRandomization)
```

• Convert our data frame to the required class

```
datfmt <- mr_mvinput(
  bx = as.matrix(cbind(dat$ldlcbeta, dat$hdlcbeta, dat$tgbeta)),
  bxse = as.matrix(cbind(dat$ldlcse, dat$hdlcse, dat$tgse)),
  by = dat$chdbeta,
  byse = dat$chdse,
  exposure = "exposure",
  outcome = "outcome",
  snps = "snp",
  effect_allele = dat$a1,
  other_allele = dat$a2,
  eaf = NA
)</pre>
```

• Fit an MVMR/MVIVW model

```
mvivwfit <- mr_mvivw(datfmt)
mvivwfit</pre>
```

```
##
## Multivariable inverse-variance weighted method
## (variants uncorrelated, random-effect model)
```

• Fit an MVMR-Egger model

```
mvmreggerfit <- mr_mvegger(datfmt)</pre>
mvmreggerfit
##
## Multivariable MR-Egger method
## (variants uncorrelated, random-effect model)
##
## Orientated to exposure : 1
## Number of Variants : 73
## ---
      Exposure Estimate Std Error 95% CI
##
                                              p-value
##
    exposure 1 0.567
                        0.100 0.371, 0.764 0.000
##
    exposure_2 -0.136
                        0.133 -0.398, 0.125 0.306
                        0.125 0.030, 0.518 0.028
##
    exposure_3 0.274
## (intercept) -0.009
                        0.005 -0.020, 0.001
                                                0.084
## Residual standard error = 1.469
## Heterogeneity test statistic = 148.9290 on 69 degrees of freedom, (p-value = 0.0000)
```

#### Example code using the MVMR and RMVMR packages

• Install the package and load it into memory

```
if (!requireNamespace("MVMR", quietly = TRUE)) {
  remotes::install_github("WSpiller/MVMR")
}
library(MVMR)
```

• Create a data object of the required structure

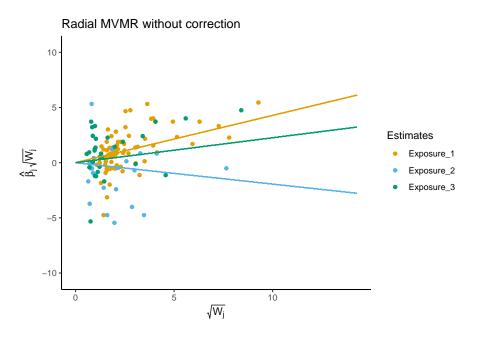
```
r_input <- format_mvmr(
    BXGs = dat[,c("ldlcbeta","hdlcbeta","tgbeta")],
    BYG = dat$chdbeta,
    seBXGs = dat[,c("ldlcse","hdlcse","tgse")],
    seBYG = dat$chdse,
    RSID = dat$rsid
)</pre>
```

• Fit an MVMR model

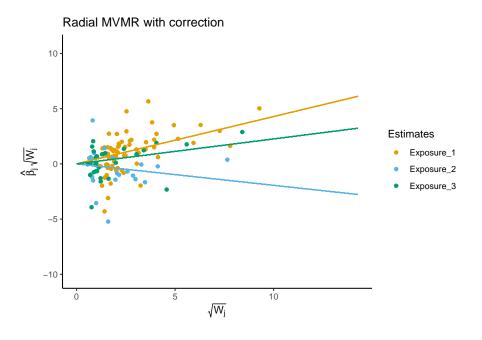
```
mvmrfit <- ivw_mvmr(r_input)</pre>
##
## Multivariable MR
##
##
               Estimate Std. Error t value
## exposure1 0.4286200 0.0609661 7.030464 1.099077e-09
## exposure2 -0.1941989 0.1308289 -1.484372 1.421994e-01
## exposure3 0.2260456 0.1232828 1.833554 7.097168e-02
##
## Residual standard error: 1.49 on 70 degrees of freedom
   • Conditional F-statistics for instrument strength (Sanderson, Spiller, and Bowden 2021)
strength_mvmr(r_input)
## Warning in strength_mvmr(r_input): Covariance between effect of genetic variants on each exposure
## not specified. Fixing covariance at 0.
##
## Conditional F-statistics for instrument strength
##
##
               exposure1 exposure2 exposure3
## F-statistic 126.7447 35.29937 39.32731
               exposure1 exposure2 exposure3
## F-statistic 126.7447 35.29937 39.32731
   • Fit a radial MVMR model
if (!requireNamespace("RMVMR", quietly = TRUE)) {
  remotes::install github("WSpiller/RMVMR")
library(RMVMR)
rmvmr_input <- mrmvinput_to_rmvmr_format(datfmt)</pre>
rmvmr_fit <- ivw_rmvmr(rmvmr_input, summary = TRUE)</pre>
##
## Radial Multivariable MR
##
##
              Estimate Std. Error t value
                                                 Pr(>|t|)
## exposure1 0.4286200 0.0609661 7.030464 1.099077e-09
## exposure2 -0.1941989 0.1308289 -1.484372 1.421994e-01
## exposure3 0.2260456 0.1232828 1.833554 7.097168e-02
## Residual standard error: 1.49 on 70 degrees of freedom
rmvmr_fit$coef
##
               Estimate Std. Error t value
                                                 Pr(>|t|)
## exposure1 0.4286200 0.0609661 7.030464 1.099077e-09
## exposure2 -0.1941989 0.1308289 -1.484372 1.421994e-01
## exposure3 0.2260456 0.1232828 1.833554 7.097168e-02
```

#### • Plot the radial MVMR models

```
plt_rmvmr <- plot_rmvmr(rmvmr_input, rmvmr = rmvmr_fit)
plt_rmvmr$p1</pre>
```



#### plt\_rmvmr\$p2



ullet Heterogeneity statistics

```
pleio_rmvmr <- pleiotropy_rmvmr(rmvmr_input, rmvmr = rmvmr_fit)
pleio_rmvmr$gq</pre>
```

#### head(pleio\_rmvmr\$qdat)

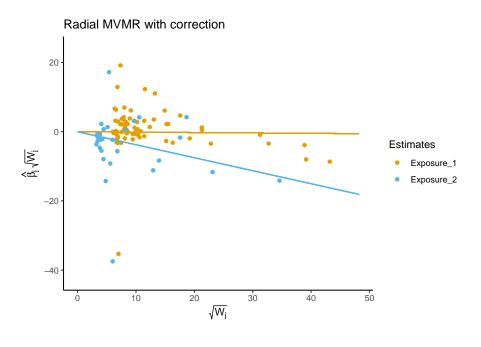
```
qj_p ref_exposure
##
                 wj corrected_beta
       snp
## 1 snp_1 2.414215
                         0.3141338 0.031643343 0.85881269
                                                             Exposure_1
## 2 snp_2 3.938760
                         0.3821443 0.008507667 0.92650973
                                                             Exposure_1
## 3 snp_3 3.246657
                        -0.6057993 3.473998077 0.06234046
                                                             Exposure_1
## 4 snp_4 2.001191
                         0.3273484 0.020524059 0.88608310
                                                             Exposure_1
## 5 snp_5 9.278826
                         0.5414293 0.118081748 0.73112437
                                                             Exposure_1
## 6 snp_6 1.317671
                         1.0713541 0.544339092 0.46064001
                                                             Exposure_1
```

• Conditional F-statistics for instrument strength (Sanderson, Spiller, and Bowden 2021)

```
str_rmvmr <- strength_rmvmr(rmvmr_input)</pre>
```

## Warning in MVMR::strength\_mvmr(r\_input, gencov): Covariance between effect of genetic variants on ## each exposure not specified. Fixing covariance at 0.

```
str_rmvmr$plot[[2]]
```



#### str\_rmvmr\$qstat[[2]]

#### Stata code

• Load the Statamarkdown package to enable Stata code chunks in an R Markdown file

#### library(Statamarkdown)

• Read in the data and create an indicator variable to select observations with p-value between the genotype and LDL-C < 10-8

```
use https://raw.github.com/remlapmot/mrrobust/master/dodata, clear
gen byte sel1 = (ldlcp2 < 1e-8)</pre>
```

#### Example code using the mrrobust package

• Install the mrrobust package using the github package

```
// Note: output suppressed
net install mrrobust, from("https://raw.github.com/remlapmot/mrrobust/master/")
mrdeps
```

• Fit IVW (with fixed effect standard errors)

```
mregger chdbeta ldlcbeta [aw=1/(chdse^2)] if sel1==1, ivw fe
```

```
| Number of genotypes = 73
| Residual standard error constrained at 1
| Coefficient Std. err. z P>|z| [95% conf. interval]
| Coefficient Std. err. z P>|z| [95% conf. interval]
| Coefficient Std. err. z P>|z| [95% conf. interval]
```

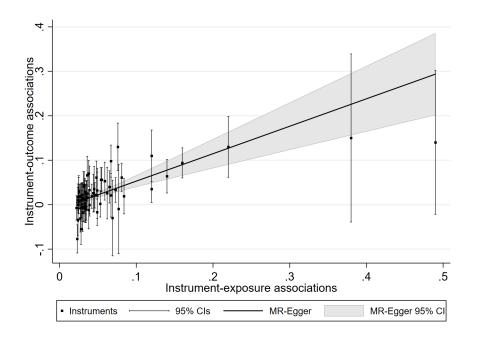
• Fit MR-Egger reporting I2GX statistic and heterogeneity Q-test

```
mregger chdbeta ldlcbeta [aw=1/(chdse^2)] if sel1==1, gxse(ldlcse) heterogi
```

```
Q_GX statistic (weighted) = 3454.26
                                     I^2_GX statistic (weighted) = 97.92%
                                                 Number of genotypes = 73
                                         Residual standard error = 1.548
              Ruecker's Q for heterogeneity; chi2(71) = 170.11 (p = 0.0000)
                          I-squared statistic = 58.3% (95% CI 45.8%, 67.8%)
           | Coefficient Std. err. z P>|z| [95% conf. interval]
       - 1
chdbeta
      slope | .6173131 .1034573 5.97 0.000
                                                              .8200858
                                                    .4145405
      _cons | -.0087706 .0054812
                                    -1.60 0.110
                                                    -.0195136
                                                               .0019723
```

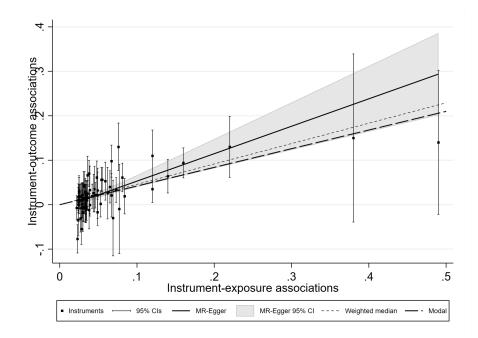
• Simple plot of IVW and MR-Egger univariate estimates

```
mreggerplot chdbeta chdse ldlcbeta ldlcse if sel1==1
qui gr export mreggerplot-1.png, replace
```



• Adding the modal and median estimates onto the plot

```
Number of genotypes = 73
                                             Replications = 1000
    | Coefficient Std. err. z P>|z|
                                            [95% conf. interval]
       .4582573
                 .0633137
                             7.24 0.000
                                            .3341648
                                                      .5823499
beta |
                                        Number of genotypes = 73
                                             Replications = 1000
                                                      Phi = .25
    | Coefficient Std. err. z P>|z| [95% conf. interval]
                                                    .8629471
       .4198713 .2260632 1.86 0.063
beta |
                                           -.0232045
```



• Fit and MVMR model with phenotypes LDL-c and HDL-c (Burgess, Dudbridge, and Thompson 2015).

#### mvmr chdbeta ldlcbeta hdlcbeta [aw=1/(chdse^2)] if sel1==1

Number of genotypes = 73 Number of phenotypes = 2 Standard errors: Random effect Residual standard error = 1.514

	Coefficient				2 - 7 - 7	interval]
chdbeta ldlcbeta		.0581901	8.03 -2.42	0.000	.3530214	.5811224 0554921

• Additionally include a third phenotype – triglycerides.

#### mvmr chdbeta ldlcbeta hdlcbeta tgbeta [aw=1/(chdse^2)] if sel1==1

Number of genotypes = 73 Number of phenotypes = 3 Standard errors: Random effect Residual standard error = 1.490

	Coefficient		z 	P> z	[95% conf.	interval]
chdbeta						
ldlcbeta	.42862	.0609661	7.03	0.000	.3091286	.5481113
hdlcbeta	1941989	.1308289	-1.48	0.138	4506189	.0622211
tgbeta	.2260456	.1232828	1.83	0.067	0155842	.4676755

• Report the QA statistic for instrument validity and the conditional F-statistics for instrument strength for each phenotype (Sanderson et al. 2019; Sanderson, Spiller, and Bowden 2021).

# mvmr chdbeta ldlcbeta hdlcbeta tgbeta [aw=1/(chdse^2)] if sel1==1, gxse(ldlcse hdlcse tgse)

> e hdlcse tgse)

Number of genotypes = 73 Number of phenotypes = 3Standard errors: Random effect Residual standard error = 1.490

	Coefficient		z			interval]
chdbeta	I					
ldlcbeta	.42862	.0609661	7.03	0.000	.3091286	.5481113
hdlcbeta	1941989	.1308289	-1.48	0.138	4506189	.0622211
tgbeta	.2260456	.1232828	1.83	0.067	0155842	.4676755

Q A statistic for instrument validity; chi2(70) = 152.88 (p = 0.0000) Conditional F-statistics for instrument strength:

 $F_x1 = 130.31$  (ldlcbeta) F\_x2 = 36.29 (hdlcbeta) F\_x3 = 40.44 (tgbeta)

• Fit an MVMR-Egger regression (Rees, Wood, and Burgess 2017), orienting the model with respect to the first phenotype in the main varlist.

#### mrmvegger chdbeta ldlcbeta hdlcbeta tgbeta [aw=1/(chdse^2)] if sel1==1

MVMR-Egger model oriented wrt: ldlcbeta Number of genotypes = 73 Number of phenotypes = 3Residual standard error = 1.469 | Coefficient Std. err. z P>|z| [95% conf. interval] \_\_\_\_\_\_ chdbeta | ldlcbeta | .5672993 .1002611 5.66 0.000 .370791 .7638075

hdlcbeta | -.1364113 .1332727 -1.02 0.306 -.3976209 .1247983 .5183735 cgbeta | .2739803 .1246927 2.20 0.028 \_cons | -.0093655 .0054187 -1.73 0.084 .0295871 tgbeta | -.019986 .001255

We can also orient the model with respect to HDL-C.

```
mrmvegger chdbeta ldlcbeta hdlcbeta tgbeta [aw=1/(chdse^2)] if sel1==1, orient(2)
```

> t(2)

MVMR-Egger model oriented wrt: hdlcbeta Number of genotypes = 73 Number of phenotypes = 3Residual standard error = 1.501 | Coefficient Std. err. z P>|z| [95% conf. interval] chdbeta |

```
    ldlcbeta | .4286398
    .0614056
    6.98
    0.000
    .308287
    .5489926

    hdlcbeta | -.1989637
    .1541909
    -1.29
    0.197
    -.5011723
    .1032449

    tgbeta | .2256794
    .1243221
    1.82
    0.069
    -.0179875
    .4693463

    _cons | .0002155
    .0036218
    0.06
    0.953
    -.006883
    .0073141
```

Or we can orient the model with respect to triglycerides.

### Session information for reproducibility

#### R session information

cli

cluster

```
library(sessioninfo)
session_info()
- Session info ------
setting value
version R version 4.2.3 (2023-03-15 ucrt)
os Windows 10 x64 (build 22621)
system x86 64, mingw32
ui
      RTerm
language (EN)
collate English_United Kingdom.utf8
ctype English_United Kingdom.utf8
tz
date
       Europe/London
       2023-03-20
pandoc 2.19.2 @ C:/Program Files/RStudio/resources/app/bin/quarto/bin/tools/ (via rmarkdown)
- Packages ------
                  * version date (UTC) lib source
package
arrangements
                    1.1.9 2020-09-13 [1] CRAN (R 4.2.2)
                    1.4.1 2021-12-13 [1] CRAN (R 4.2.0)
backports
base64enc
                   0.1-3 2015-07-28 [1] CRAN (R 4.2.0)
                    2.1.0 2022-04-21 [1] CRAN (R 4.2.1)
checkmate
```

2.1.4

3.6.0 2023-01-09 [1] CRAN (R 4.2.2)

2022-08-22 [2] CRAN (R 4.2.3)

```
0.2-19
                                   2023-02-01 [2] CRAN (R 4.2.3)
codetools
                                   2023-01-23 [1] CRAN (R 4.2.2)
colorspace
                          2.1-0
curl
                          5.0.0
                                   2023-01-12 [1] CRAN (R 4.2.2)
data.table
                          1.14.8
                                   2023-02-17 [1] CRAN (R 4.2.2)
                                   2022-04-03 [1] CRAN (R 4.2.0)
DEoptimR
                          1.0-11
digest
                          0.6.31
                                   2022-12-11 [1] CRAN (R 4.2.2)
                                   2023-01-29 [1] CRAN (R 4.2.2)
                        * 1.1.0
dplyr
                          0.3.2
                                   2021-04-29 [1] CRAN (R 4.2.0)
ellipsis
                                   2023-01-17 [1] CRAN (R 4.2.2)
evaluate
                          0.20
                                   2023-01-22 [1] CRAN (R 4.2.2)
fansi
                          1.0.4
                                   2022-07-06 [1] CRAN (R 4.2.1)
farver
                          2.1.1
                          1.1.1
                                   2023-02-24 [1] CRAN (R 4.2.2)
fastmap
forcats
                        * 1.0.0
                                   2023-01-29 [1] CRAN (R 4.2.2)
                                   2022-02-02 [1] CRAN (R 4.2.2)
                          1.5.2
foreach
                                   2022-12-06 [2] CRAN (R 4.2.3)
                          0.8-84
foreign
Formula
                          1.2-5
                                   2023-02-24 [1] CRAN (R 4.2.2)
                          0.1.3
                                   2022-07-05 [1] CRAN (R 4.2.1)
generics
                        * 3.4.1
                                   2023-02-10 [1] CRAN (R 4.2.2)
ggplot2
                                   2022-11-27 [1] CRAN (R 4.2.2)
glmnet
                          4.1 - 6
                                   2022-02-24 [1] CRAN (R 4.2.0)
glue
                          1.6.2
                          0.7 - 1
                                   2023-02-07 [1] CRAN (R 4.2.2)
gmp
                          2.3
                                   2017-09-09 [1] CRAN (R 4.2.1)
gridExtra
                          0.3.2
                                   2023-03-17 [1] CRAN (R 4.2.3)
gtable
haven
                        * 2.5.2
                                   2023-02-28 [1] CRAN (R 4.2.2)
Hmisc
                        * 5.0-1
                                   2023-03-08 [1] CRAN (R 4.2.2)
                                   2022-08-19 [1] CRAN (R 4.2.1)
hms
                          1.1.2
htmlTable
                          2.4.1
                                   2022-07-07 [1] CRAN (R 4.2.1)
                                   2022-12-07 [1] CRAN (R 4.2.2)
htmltools
                          0.5.4
                          1.6.2
                                   2023-03-17 [1] CRAN (R 4.2.3)
htmlwidgets
                                   2023-02-24 [1] CRAN (R 4.2.2)
                          1.4.5
httr
                                   2022-02-05 [1] CRAN (R 4.2.2)
iterators
                          1.0.14
iterpc
                          0.4.2
                                   2020-01-10 [1] CRAN (R 4.2.2)
jsonlite
                          1.8.4
                                   2022-12-06 [1] CRAN (R 4.2.2)
                          1.42.5
                                   2023-03-14 [1] Github (yihui/knitr@c50d307)
knitr
                          0.4.2
                                   2020-10-20 [1] CRAN (R 4.2.0)
labeling
                                   2021-09-22 [2] CRAN (R 4.2.3)
                          0.20-45
lattice
                                   2019-03-15 [1] CRAN (R 4.2.1)
lazyeval
                          0.2.2
                          1.0.3
                                   2022-10-07 [1] CRAN (R 4.2.1)
lifecycle
                        * 1.9.2
                                   2023-02-10 [1] CRAN (R 4.2.2)
lubridate
                          2.0.3
                                   2022-03-30 [1] CRAN (R 4.2.0)
magrittr
                          7.3-58.3 2023-03-07 [1] CRAN (R 4.2.2)
MASS
Matrix
                          1.5 - 3
                                   2022-11-11 [1] CRAN (R 4.2.2)
                                   2022-09-11 [1] CRAN (R 4.2.1)
MatrixModels
                          0.5 - 1
MendelianRandomization * 0.7.0
                                   2023-01-09 [1] CRAN (R 4.2.2)
                          0.5.0
                                   2018-06-12 [1] CRAN (R 4.2.0)
munsell
                                   2023-03-20 [1] Github (WSpiller/MVMR@3a0b54b)
MVMR
                        * 0.3
                          7.3-18
                                   2022-09-28 [2] CRAN (R 4.2.3)
nnet
pillar
                          1.8.1
                                   2022-08-19 [1] CRAN (R 4.2.1)
                                   2019-09-22 [1] CRAN (R 4.2.0)
                          2.0.3
pkgconfig
                                   2022-11-07 [1] CRAN (R 4.2.2)
                          4.10.1
plotly
                        * 1.0.1
                                   2023-01-10 [1] CRAN (R 4.2.2)
purrr
                          5.94
                                   2022-07-20 [1] CRAN (R 4.2.1)
quantreg
                          2.5.1
                                   2021-08-19 [1] CRAN (R 4.2.0)
R6
RadialMR
                          1.0
                                   2023-02-17 [1] Github (WSpiller/RadialMR@d63d3fc)
                                   2023-01-22 [1] CRAN (R 4.2.2)
                          1.0.10
Rcpp
                                   2023-02-10 [1] CRAN (R 4.2.2)
readr
                        * 2.1.4
                        * 2.4.2
                                   2021-11-30 [1] CRAN (R 4.2.0)
remotes
rjson
                          0.2.21
                                   2022-01-09 [1] CRAN (R 4.2.0)
```

```
2023-03-14 [1] CRAN (R 4.2.2)
rlang
                      1.1.0
                      2.20
                               2023-01-19 [1] CRAN (R 4.2.2)
rmarkdown
RMVMR
                    * 0.2
                               2023-03-20 [1] Github (WSpiller/RMVMR@a162f9e)
                     0.95-0 2022-04-02 [1] CRAN (R 4.2.2)
robustbase
                     4.1.19 2022-10-21 [2] CRAN (R 4.2.3)
rpart
rstudioapi
                     0.14
                               2022-08-22 [1] CRAN (R 4.2.1)
                      1.2.1 2022-08-20 [1] CRAN (R 4.2.1)
scales
                * 1.2.2
sessioninfo
                               2021-12-06 [1] CRAN (R 4.2.1)
                     1.4.6
                               2021-05-19 [1] CRAN (R 4.2.0)
shape
                     1.81
                               2021-02-18 [1] CRAN (R 4.2.0)
SparseM
Statamarkdown
                   * 0.7.2
                               2023-02-15 [1] CRAN (R 4.2.2)
                      1.7.12 2023-01-11 [1] CRAN (R 4.2.2)
stringi
                               2022-12-02 [1] CRAN (R 4.2.2)
stringr
                    * 1.5.0
survival
                      3.5-5
                               2023-03-12 [2] CRAN (R 4.2.2)
                              2023-03-20 [1] CRAN (R 4.2.3)
                    * 3.2.1
tibble
                    * 1.3.0
tidyr
                               2023-01-24 [1] CRAN (R 4.2.2)
tidyselect
                      1.2.0
                             2022-10-10 [1] CRAN (R 4.2.1)
                   * 2.0.0 2023-02-22 [1] CRAN (R 4.2.2)
tidyverse
                              2023-01-11 [1] CRAN (R 4.2.2)
timechange
                     0.2.0
                     0.3.0
                               2022-03-28 [1] CRAN (R 4.2.0)
tzdb
utf8
                      1.2.3
                               2023-01-31 [1] CRAN (R 4.2.2)
                      0.6.0
                               2023-03-16 [1] CRAN (R 4.2.2)
vctrs
                      0.4.1
                               2022-08-22 [1] CRAN (R 4.2.1)
viridisLite
withr
                      2.5.0 2022-03-03 [1] CRAN (R 4.2.0)
xfun
                      0.37
                             2023-01-31 [1] CRAN (R 4.2.2)
                      2.3.7 2023-01-23 [1] CRAN (R 4.2.2)
yaml
```

[1] C:/Users/tom/AppData/Local/R/win-library/4.2

[2] C:/Program Files/R/R-4.2.3/library

-----

#### Stata session information

# about ado describe mrrobust

Stata/MP 17.0 for Windows (64-bit x86-64) Revision 08 Mar 2023

Copyright 1985-2021 StataCorp LLC

Total physical memory: 32.00 GB Available physical memory: 15.27 GB

Stata license: Unlimited-user 2-core network, expiring 21 Jan 2024

Serial number: 501709378202 Licensed to: Tom Palmer

University of Bristol

\_\_\_\_\_

[47] package mrrobust from https://raw.githubusercontent.com/remlapmot/mrrobust

> /master

-----

TITLE

```
DESCRIPTION/AUTHOR(S)
      Author: Tom Palmer
      Distribution-Date: 20210917
INSTALLATION FILES
      m\mrmedian.ado
      m\mrmedian.sthlp
      m\mrmedianobs.ado
      m\mrmedianobs_work.ado
      m\mrmedianobs.sthlp
      m\mregger.ado
      m\mregger.sthlp
      m\mrrobust.sthlp
      m\mreggerplot.ado
      m\mreggerplot.sthlp
      m\mrmodal.ado
      m\mrmodal.sthlp
      m\mrratio.ado
      m\mrratio.sthlp
      m\mrivests.ado
      m\mrivests.sthlp
      m\mrforest.ado
      m\mrforest.sthlp
      m\mreggersimex.ado
      m\mreggersimex.sthlp
      m\mreggersimexonce.ado
      m\mrmodalplot.ado
      m\mrmodalplot.sthlp
      m\mrfunnel.ado
      m\mrfunnel.sthlp
      m\mrdeps.ado
      m\mrdeps.sthlp
      m\mr.ado
      m\mr.sthlp
      m\mrmvivw.ado
      m\mrmvivw.sthlp
      m\mvivw.ado
      m\mvivw.sthlp
      m\mvmr.ado
      m\mvmr.sthlp
      m\mrmvegger.ado
      m\mrmvegger.sthlp
      m\mrleaveoneout.ado
      m\mrleaveoneout.sthlp
      m\mrrobust-author.ihlp
INSTALLED ON
     17 Feb 2023
```

'mrrobust': Stata package for two-sample Mendelian randomization analyses

#### References

Burgess, S, F Dudbridge, and SG Thompson. 2015. "Multivariable Mendelian randomization: the use of pleiotropic genetic variants to estimate causal effects." *American Journal of Epidemiology* 181: 251--260. https://doi.org/10.1093/aje/kwu283.

- Rees, J, A Wood, and S Burgess. 2017. "Extending the MR-Egger method for multivariable Mendelian randomization to correct for both measured and unmeasured pleiotropy." *Statistics in Medicine* 36: 4705–18. https://doi.org/10.1002/sim.7492.
- Sanderson, E, G Davey Smith, F Windmeijer, and J Bowden. 2019. "An examination of multivariable Mendelian randomization in the single-sample and two-sample summary data settings." *International Journal of Epidemiology* 48: 713–27. https://doi.org/10.1093/ije/dyy262.
- Sanderson, E, W Spiller, and J Bowden. 2021. "Testing and Correcting for Weak and Pleiotropic Instruments in Two-Sample Multivariable Mendelian Randomization." Statistics in Medicine. https://doi.org/10.1002/sim.9133.