

# Package: MVMRcML (via r-universe)

July 4, 2024

**Type** Package

**Title** Multivariable Mendelian randomization based on constrained maximum likelihood

**Version** 0.1.0

**Author** Zhaotong Lin

**Maintainer** Zhaotong Lin <z123k@fsu.edu>

**Description** Robust multivariable Mendelian randomization based on constrained maximum likelihood.

**License** MIT + file LICENSE

**Encoding** UTF-8

**LazyData** true

**Imports** numDeriv, Rcpp, RcppArmadillo

**LinkingTo** Rcpp, RcppArmadillo

**RoxygenNote** 7.3.1

**Repository** <https://mrcieu.r-universe.dev>

**RemoteUrl** <https://github.com/ZhaotongL/MVMR-cML>

**RemoteRef** HEAD

**RemoteSha** 20d09a54637e70a35f31a43d56c1e6276c99bb17

## Contents

invcov_mvnr . . . . .	2
MVcML_SdTheta . . . . .	2
MVmr_cML . . . . .	3
MVmr_cML_DP . . . . .	4

<b>Index</b>	<b>5</b>
--------------	----------

---

invcov_mvmmr	<i>Generate the list of inverse of covariance matrices used in MVMR-cML-DP</i>
--------------	--

---

**Description**

Generate the list of inverse of covariance matrices used in MVMR-cML-DP

**Usage**

```
invcov_mvmmr(se_bx, se_by, rho_mat)
```

**Arguments**

se_bx	A m*L matrix of standard errors of SNP-exposure association
se_by	A vector of standard errors of SNP-outcome association
rho_mat	A (L+1)×(L+1) correlation matrix among the L exposures and the outcome, note that the upper-left L×L submatrix corresponding to the L exposures.

**Value**

A list of inverse of covariance matrices with respect to each genetic variant, retaining the ordering in se\_bx

---

MVcML_SdTheta	<i>Standard error estimate for MVMR-cML-BIC</i>
---------------	---

---

**Description**

This is based on the profile likelihood of the set of valid IVs, which is not robust to uncertainty in model selection.

**Usage**

```
MVcML_SdTheta(b_exp, b_out, Sig_inv_l, theta, zero_ind, r_vec = NULL)
```

**Arguments**

b_exp	A matrix of SNP effects on the exposure variable.
b_out	A vector of SNP effects on the outcome variable.
Sig_inv_l	A list of inverse of covariance matrix.
theta	A vector of final estimates of causal effect of each exposure by MVMR-cML-BIC obtained from MVmmr_cML_DP.
zero_ind	A vector of the index of valid IVs.
r_vec	A vector of estimated horizontal pleiotropic effects.

**Value**

A vector

---

MVmr\_cML

*MVMRcML method with BIC selection*

---

**Description**

This is the main function of MVMR-cML.

**Usage**

```
MVmr_cML(
  b_exp,
  b_out,
  se_bx,
  Sig_inv_l,
  n,
  K_vec = as.numeric(c()),
  random_start = 1L,
  maxit = 100L,
  thres = 1e-04
)
```

**Arguments**

b_exp	A m*L matrix of SNP effects on the exposure variable.
b_out	A m*1 matrix of SNP effects on the outcome variable.
se_bx	A m*L matrix of standard errors of b_exp.
Sig_inv_l	A list of the inverse of m covariance matrices, generated by <code>invcov_mvmmr()</code> .
n	The smallest sample size of the L+1 GWAS dataset.
K_vec	Sets of candidate K's, the constraint parameter representing number of invalid IVs.
random_start	Number of random start points, default is 1.
maxit	Maximum number of iterations for each optimization, default is 100.
thres	Threshold for convergence criterion.

**Value**

A list

**BIC\_theta** Estimated causal effect from MVMR-cML-BIC

**BIC\_invalid** Invalid IVs selected by MVMR-cML-BIC

**K\_vec** K\_vec to be searched from

**Converge** 0 if converge; 1 otherwise

**BIC\_vec** BIC vector corresponding to K\_vec

MVmr\_cML\_DP

*MVMRcML method with Data Perturbation***Description**

This is the main function of MVMR-cML.

**Usage**

```
MVmr_cML_DP(
  b_exp,
  b_out,
  se_bx,
  Sig_inv_l,
  n,
  K_vec = as.numeric(c()),
  random_start = 1L,
  num_pert = 100L,
  maxit = 100L,
  thres = 1e-04
)
```

**Arguments**

b_exp	A m*L matrix of SNP effects on the exposure variable.
b_out	A m*1 matrix of SNP effects on the outcome variable.
se_bx	A m*L matrix of standard errors of b_exp.
Sig_inv_l	A list of the inverse of m covariance matrices, generated by <code>invcov_mvmmr()</code> .
n	The smallest sample size of the L+1 GWAS dataset.
K_vec	Sets of candidate K's, the constraint parameter representing number of invalid IVs.
random_start	Number of random start points, default is 1.
num_pert	Number of perturbation, default is 100.
maxit	Maximum number of iterations for each optimization, default is 100.
thres	Threshold for convergence criterion.

**Value**

A list

**BIC\_theta** Estimated causal effect from MVMR-cML-BIC  
**BIC\_invalid** Invalid IVs selected by MVMR-cML-BIC  
**BIC\_DP\_theta** Estimated causal effect from MVMR-cML-DP  
**BIC\_DP\_se** Estimate standard error for BIC\_DP\_theta  
**eff\_DP\_B** Data perturbation with successful convergence

# Index

[invcov\\_mvnr](#), 2

[MVcML\\_SdTheta](#), 2

[MVnr\\_cML](#), 3

[MVnr\\_cML\\_DP](#), 4