

# Package ‘cophescan’

July 30, 2025

**Title** Adaptation of the Coloc Method for PheWAS

**Version** 1.4.2

**Maintainer** Ichcha Manipur <im504@cam.ac.uk>

**Description** A Bayesian method for Phenome-wide association studies (PheWAS) that identifies causal associations between genetic variants and traits, while simultaneously addressing confounding due to linkage disequilibrium. For details see Manipur et al (2024, Nature Communications) <doi:10.1038/s41467-024-49990-8>.

**License** GPL-3

**Encoding** UTF-8

**LazyData** true

**VignetteBuilder** knitr

**RoxygenNote** 7.2.3

**Depends** R (>= 3.5.0)

**URL** <https://github.com/ichcha-m/cophescan>,  
<https://ichcha-m.github.io/cophescan/>

**BugReports** <https://github.com/ichcha-m/cophescan/issues>

**Imports** Rcpp (>= 1.0.7), coloc, data.table, ggplot2, ggrepel,  
pheatmap, methods, viridis, stats, grDevices, magrittr, utils,  
matrixStats, dplyr

**Suggests** knitr, testthat (>= 3.0.0), rmarkdown, RColorBrewer, ggpubr

**Collate** 'cophescan-package.R' 'singlevar.R' 'multivarsusie.R'  
'multitrait.R' 'cophe\_hyp\_predict.R' 'copheplots.R'  
'testdata.R' 'RcppExports.R' 'metrop\_hier\_priors.R' 'zzz.R'

**LinkingTo** Rcpp, RcppArmadillo

**Config/testthat/edition** 3

**ByteCompile** true

**NeedsCompilation** yes

**Author** Ichcha Manipur [aut, cre],  
Chris Wallace [aut]

**Repository** CRAN

**Date/Publication** 2025-07-30 20:00:02 UTC

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|                   |                                 |
|-------------------|---------------------------------|
| cophescan-package | <i>The 'cophescan' package.</i> |
|-------------------|---------------------------------|

---

**Description**

Coloc adapted Phenome-wide Scans

---

|               |                      |
|---------------|----------------------|
| adjust_priors | <i>adjust_priors</i> |
|---------------|----------------------|

---

**Description**

adjust fixed priors when nsnp in region is high

**Usage**

```
adjust_priors(
  nsnp,
  pa = 3.82e-05,
  pc = 0.00182,
  p1 = NULL,
  p2 = NULL,
  p12 = NULL
)
```

**Arguments**

|      |                                                                                                                                                                                  |
|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| nsnp | number of SNPs                                                                                                                                                                   |
| pa   | prior probability that a non-query variant is causally associated with the query trait (cophescan prior), default 3.82e-5                                                        |
| pc   | prior probability that the query variant is causally associated with the query trait (cophescan prior), default 1.82e-3 (cophescan prior)                                        |
| p1   | prior probability a SNP is associated with trait 1, (coloc prior), pc derived by using $pc = p12/p1 + p12$ ; use p1, p2, p12 only when pa and pc are unavailable (See vignettes) |
| p2   | prior probability a SNP is associated with trait 2, (coloc prior), pa derived by using $pa = p2$                                                                                 |
| p12  | prior probability a SNP is associated with both traits, (coloc prior), pc derived by using $pc = p12/p1 + p12$                                                                   |

**Value**

vector of pn, pa and pc adjusted prior probabilities

---

|              |                                            |
|--------------|--------------------------------------------|
| average_piks | <i>Average of priors: pnk, pak and pck</i> |
|--------------|--------------------------------------------|

---

**Description**

Average of priors: pnk, pak and pck

**Usage**

```
average_piks(params, nsnp, covar_vec, nits, thin, covar = FALSE)
```

**Arguments**

|           |                                                             |
|-----------|-------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$       |
| nsnp      | number of snps                                              |
| covar_vec | Vector of the covariate                                     |
| nits      | Number of iterations run in mcmc                            |
| thin      | thinning                                                    |
| covar     | logical: was the covariate information used? default: False |

**Value**

average pik matrix of priors: pnk, pak and pck

---

|                   |                                                                         |
|-------------------|-------------------------------------------------------------------------|
| average_piks_list | <i>Average of priors: pnk, pak and pck from list (memory intensive)</i> |
|-------------------|-------------------------------------------------------------------------|

---

**Description**

Average of priors: pnk, pak and pck from list (memory intensive)

**Usage**

```
average_piks_list(params, nsnp, covar_vec, nits, thin, covar = FALSE)
```

**Arguments**

|           |                                                             |
|-----------|-------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$       |
| nsnp      | number of snps                                              |
| covar_vec | Vector of the covariate                                     |
| nits      | Number of iterations run in mcmc                            |
| thin      | thinning                                                    |
| covar     | logical: was the covariate information used? default: False |

**Value**

average pik matrix of priors: pnk, pak and pck

---

average\_posterior\_prob

*Average of posterior probabilities: Hn, Ha and Hc*

---

**Description**

Average of posterior probabilities: Hn, Ha and Hc

**Usage**

```
average_posterior_prob(
  params,
  lbf_mat,
  nsnp,
  covar_vec,
  nits,
  thin,
  covar = FALSE
)
```

**Arguments**

|           |                                                             |
|-----------|-------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$       |
| lbf_mat   | matrix of log bayes factors: LBF.Ha and LBF.Hc              |
| nsnp      | number of snps                                              |
| covar_vec | Vector of the covariate                                     |
| nits      | Number of iterations run in mcmc                            |
| thin      | thinning                                                    |
| covar     | logical: was the covariate information used? default: False |

**Value**

matrix with average of all the posterior probabilities: Hn, Ha and Hc

---

average\_posterior\_prob\_list

*Average of posterior probabilities: Hn, Ha and Hc from list (memory intensive)*

---

### Description

Average of posterior probabilities: Hn, Ha and Hc from list (memory intensive)

### Usage

```
average_posterior_prob_list(
  params,
  lbf_mat,
  nsnp,
  covar_vec,
  nits,
  thin,
  covar = FALSE
)
```

### Arguments

|           |                                                             |
|-----------|-------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$       |
| lbf_mat   | matrix of log bayes factors: lbf.Ha and lbf.Hc              |
| nsnp      | number of snps                                              |
| covar_vec | Vector of the covariate                                     |
| nits      | Number of iterations run in mcmc                            |
| thin      | thinning                                                    |
| covar     | logical: was the covariate information used? default: False |

### Value

matrix with average of all the posterior probabilities: Hn, Ha and Hc

---

combine.bf

*combine.bf*

---

### Description

Calculate posterior probabilities for all the configurations

## Usage

```
combine.bf(lBF_df, pn, pa, pc)
```

## Arguments

|        |                                                                                                    |
|--------|----------------------------------------------------------------------------------------------------|
| lBF_df | dataframe with log bayes factors of hypothesis Ha and Hn: column names should be lBF.Ha and lBF.Hc |
| pn     | prior probability that none of the SNPs/variants in the region are associated with the query trait |
| pa     | prior probability that a non-query variant is causally associated with the query trait             |
| pc     | prior probability that the query variant is causally associated with the query trait               |

## Value

named numeric vector of posterior probabilities and bayes factors

## Author(s)

Ichcha Manipur

---

|                   |                                                             |
|-------------------|-------------------------------------------------------------|
| cophe.hyp.predict | <i>Predict cophescan hypothesis for tested associations</i> |
|-------------------|-------------------------------------------------------------|

---

## Description

Predict cophescan hypothesis for tested associations

## Usage

```
cophe.hyp.predict(
  cophe.res,
  grouping.vars = c("querysnp", "querytrait"),
  Hc.cutoff = 0.6,
  Hn.cutoff = 0.2
)
```

## Arguments

|               |                                                                                                                                                                                                                           |
|---------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| cophe.res     | results obtained from cophe.single, cophe.susie or cophe.multitrait or data.frame with the following columns: PP.Hn, PP.Hc, PP.Ha, querysnp, querytrait                                                                   |
| grouping.vars | This is important for results from cophe.susie where there are multiple signals. These will be collapsed into one call. If you want to return all signals set this to a single variable eg: grouping.vars = c('querysnp') |
| Hc.cutoff     | threshold for PP.Hc above which the associations are called Hc                                                                                                                                                            |
| Hn.cutoff     | threshold for PP.Hn above which the associations are called Hn                                                                                                                                                            |

**Value**

returns dataframe with posterior probabilities of Hn, Hc and Ha with the predicted hypothesis based on the provided cut.off.s.

**See Also**

[cophe.single](#), [cophe.susie](#), [cophe.multitrait](#), [multitrait.simplify](#)

---

|                               |                                                 |
|-------------------------------|-------------------------------------------------|
| <code>cophe.multitrait</code> | <i>Run cophescan on multiple traits at once</i> |
|-------------------------------|-------------------------------------------------|

---

**Description**

Run cophescan on multiple traits at once

**Usage**

```
cophe.multitrait(
  trait.dat,
  querysnpid,
  querytrait.names,
  LDmat = NULL,
  method = "single",
  simplify = FALSE,
  predict.hyp = TRUE,
  Hn.cutoff = 0.2,
  Hc.cutoff = 0.6,
  est.fdr.based.cutoff = FALSE,
  fdr = 0.05,
  ...
)
```

**Arguments**

|                               |                                                                                                                                                |
|-------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>trait.dat</code>        | Named(traits) list of coloc structured data for k traits (Total number of traits)                                                              |
| <code>querysnpid</code>       | vector of query variant ids = length(trait.dat), if the same variant                                                                           |
| <code>querytrait.names</code> | vector of names for the query traits, if the names of the multi.dat list contain the trait names please pass querytrait.names=names(multi.dat) |
| <code>LDmat</code>            | LD matrix                                                                                                                                      |
| <code>method</code>           | either 'single' for <code>cophe.single</code> or 'susie' for <code>cophe.susie</code>                                                          |
| <code>simplify</code>         | if TRUE removes intermediate results from output using 'multitrait.simplify'                                                                   |
| <code>predict.hyp</code>      | if TRUE predicts the hypothesis based on the provided thresholds for pp.Hc and pp.Hn (overrides simplify) using <code>cophe.hyp.predict</code> |
| <code>Hn.cutoff</code>        | threshold for PP.Hc above which the associations are called Hc                                                                                 |

Hc.cutoff            threshold for PP.Hc above which the associations are called Hn  
 est.fdr.based.cutoff            if True calculates the Hc.cutoff using  $1 - \text{mean}(\text{PP.Hc}) | \text{PP.Hc} > \text{cutoff}$   
 fdr            fdr threshold to estimate Hc.cutoff  
 ...            additional arguments of priors for cophe.susie or cophe.single

## Value

if simplify is False returns multi-trait list of lists, each with:

- a summary data.frame of the cophescan results
- priors used
- querysnp
- querytrait

if simplify is TRUE only returns dataframe with posterior probabilities of Hn, Hc and Ha with no intermediate results if predict.hyp is TRUE returns a dataframe with output of simplify and the predicted hypotheses for all associations

## Author(s)

Ichcha Manipur

---

|              |                                                                    |
|--------------|--------------------------------------------------------------------|
| cophe.single | <i>Bayesian cophescan analysis using Approximate Bayes Factors</i> |
|--------------|--------------------------------------------------------------------|

---

## Description

Bayesian cophescan analysis under single causal variant assumption

## Usage

```
cophe.single(
  dataset,
  querysnpid,
  querytrait,
  MAF = NULL,
  pa = 3.82e-05,
  pc = 0.00182,
  p1 = NULL,
  p2 = NULL,
  p12 = NULL
)
```

**Arguments**

|            |                                                                                                                                                                                  |
|------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| dataset    | a list with specifically named elements defining the query trait dataset to be analysed.                                                                                         |
| quersnpid  | Id of the query variant, (id in dataset\$snp)                                                                                                                                    |
| querytrait | Query trait name                                                                                                                                                                 |
| MAF        | Minor allele frequency vector                                                                                                                                                    |
| pa         | prior probability that a non-query variant is causally associated with the query trait (cophescan prior), default 3.82e-5                                                        |
| pc         | prior probability that the query variant is causally associated with the query trait (cophescan prior), default 1.82e-3 (cophescan prior)                                        |
| p1         | prior probability a SNP is associated with trait 1, (coloc prior), pc derived by using $pc = p12/p1 + p12$ ; use p1, p2, p12 only when pa and pc are unavailable (See vignettes) |
| p2         | prior probability a SNP is associated with trait 2, (coloc prior), pa derived by using $pa = p2$                                                                                 |
| p12        | prior probability a SNP is associated with both traits, (coloc prior), pc derived by using $pc = p12/p1 + p12$                                                                   |

**Details**

This function calculates posterior probabilities of different causal variant configurations under the assumption of a single causal variant for each trait.

If regression coefficients and variances are available, it calculates Bayes factors for association at each SNP. If only p values are available, it uses an approximation that depends on the SNP's MAF and ignores any uncertainty in imputation. Regression coefficients should be used if available. Find more input data structure details in the coloc package

**Value**

a list of two data.frames:

- summary is a vector giving the number of SNPs analysed, and the posterior probabilities of Hn (no shared causal variant), Ha (two distinct causal variants) and Hc (one common causal variant)
- results is an annotated version of the input data containing log Approximate Bayes Factors and intermediate calculations, and the posterior probability SNP.PP.Hc of the SNP being causal for the shared signal *if* Hc is true. This is only relevant if the posterior support for Hc in summary is convincing.

**Author(s)**

Ichcha Manipur

## Examples

```
library(cophescan)
data(cophe_multi_trait_data)
query_trait_1 <- cophe_multi_trait_data$summ_stat[['Trait_1']]
querysnpid <- cophe_multi_trait_data$querysnpid
res.single <- cophe.single(query_trait_1, querysnpid = querysnpid, querytrait='Trait_1')
summary(res.single)
```

---

|                  |                         |
|------------------|-------------------------|
| cophe.single.lbf | <i>cophe.single.lbf</i> |
|------------------|-------------------------|

---

## Description

Calculate log bayes factors for each hypothesis (Single causal variant assumption)

## Usage

```
cophe.single.lbf(dataset, querysnpid, querytrait, MAF = NULL)
```

## Arguments

|            |                                                                                          |
|------------|------------------------------------------------------------------------------------------|
| dataset    | a list with specifically named elements defining the query trait dataset to be analysed. |
| querysnpid | Id of the query variant, (id in dataset\$snp)                                            |
| querytrait | Query trait name                                                                         |
| MAF        | Minor allele frequency vector                                                            |

## Value

data frame with log bayes factors for Hn and Ha hypotheses

## Author(s)

Ichcha Manipur

## See Also

[cophe.single](#)

## Examples

```
library(cophescan)
data(cophe_multi_trait_data)
query_trait_1 <- cophe_multi_trait_data$summ_stat[['Trait_1']]
querysnpid <- cophe_multi_trait_data$querysnpid
res.single.lbf <- cophe.single.lbf(query_trait_1, querysnpid = querysnpid, querytrait='Trait_1')
res.single.lbf
```

---

|             |                                                               |
|-------------|---------------------------------------------------------------|
| cophe.susie | <i>run cophe.susie using susie to detect separate signals</i> |
|-------------|---------------------------------------------------------------|

---

## Description

Check if a variant causally associated in one trait might be causal in another trait

## Usage

```
cophe.susie(
  dataset,
  querysnpid,
  querytrait,
  pa = 3.82e-05,
  pc = 0.00182,
  p1 = NULL,
  p2 = NULL,
  p12 = NULL,
  susie.args = list()
)
```

## Arguments

|            |                                                                                                                                                                                  |
|------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| dataset    | <i>either</i> a list with specifically named elements defining the dataset to be analysed.<br>(see <a href="#">check_dataset</a> )                                               |
| querysnpid | Id of the query variant                                                                                                                                                          |
| querytrait | Query trait name                                                                                                                                                                 |
| pa         | prior probability that a non-query variant is causally associated with the query trait (cophescan prior), default 3.82e-5                                                        |
| pc         | prior probability that the query variant is causally associated with the query trait (cophescan prior), default 1.82e-3                                                          |
| p1         | prior probability a SNP is associated with trait 1, (coloc prior), pc derived by using $pc = p12/p1 + p12$ ; use p1, p2, p12 only when pa and pc are unavailable (See vignettes) |
| p2         | prior probability a SNP is associated with trait 2, (coloc prior), pa derived by using $pa = p2$                                                                                 |
| p12        | prior probability a SNP is associated with both traits, (coloc prior), pc derived by using $pc = p12/p1 + p12$                                                                   |
| susie.args | a named list of additional arguments to be passed to <a href="#">runsusie</a>                                                                                                    |

## Value

a list, containing elements

- summary a data.table of posterior probabilities of each global hypothesis, one row per pairwise comparison of signals from the two traits

- results a data.table of detailed results giving the posterior probability for each snp to be jointly causal for both traits *assuming Hc is true*. Please ignore this column if the corresponding posterior support for H4 is not high.
- priors a vector of the priors used for the analysis

### Author(s)

Ichcha Manipur

### Examples

```
library(cophescan)
data(cophe_multi_trait_data)
query_trait_1 <- cophe_multi_trait_data$summ_stat[['Trait_1']]
querysnpid <- cophe_multi_trait_data$querysnpid
query_trait_1$LD <- cophe_multi_trait_data$LD
res.susie <- cophe.susie(query_trait_1, querysnpid = querysnpid, querytrait='Trait_1')
summary(res.susie)
```

---

cophe.susie.lbf

*cophe.susie.lbf*

---

### Description

Calculate log bayes factors for each hypothesis (SuSIE - multiple causal variant assumption)

### Usage

```
cophe.susie.lbf(
  dataset,
  querysnpid,
  querytrait,
  switch = TRUE,
  susie.args = list(),
  MAF = NULL
)
```

### Arguments

|            |                                                                                          |
|------------|------------------------------------------------------------------------------------------|
| dataset    | a list with specifically named elements defining the query trait dataset to be analysed. |
| querysnpid | Id of the query variant, (id in dataset\$snp)                                            |
| querytrait | Query trait name                                                                         |
| switch     | Set switch=TRUE to obtain single BF when credible sets not found with SuSIE              |
| susie.args | a named list of additional arguments to be passed to <a href="#">runsusie</a>            |
| MAF        | Minor allele frequency vector                                                            |

**Value**

data frame with log bayes factors for Hn and Ha hypotheses

**Author(s)**

Ichcha Manipur

**See Also**

[cophe.susie](#)

**Examples**

```
library(cophescan)
data(cophe_multi_trait_data)
query_trait_1 <- cophe_multi_trait_data$summ_stat[['Trait_1']]
query_trait_1$LD <- cophe_multi_trait_data$LD
querysnpid <- cophe_multi_trait_data$querysnpid
res.susie.lbf <- cophe.susie.lbf(query_trait_1, querysnpid = querysnpid,
                                querytrait='Trait_1', switch=T)

res.susie.lbf
```

---

cophe\_heatmap

*Heatmap of multi-trait cophescan results*

---

**Description**

Heatmap of multi-trait cophescan results

**Usage**

```
cophe_heatmap(
  multi.dat,
  querysnpid,
  query_trait_names,
  thresh_Hc = 0.5,
  thresh_Ha = 0.5,
  ...
)
```

**Arguments**

|                   |                                                                                                                                              |
|-------------------|----------------------------------------------------------------------------------------------------------------------------------------------|
| multi.dat         | multi trait cophescan results returned from <code>cophe.multitrait</code> or formatted in the same way with <code>multitrait.simplify</code> |
| querysnpid        | query variant                                                                                                                                |
| query_trait_names | names of phenotypes corresponding to the multi.dat results                                                                                   |

|           |                                                               |
|-----------|---------------------------------------------------------------|
| thresh_Hc | Hc threshold to be displayed                                  |
| thresh_Ha | Ha threshold to be displayed                                  |
| ...       | additional arguments to be passed to <a href="#">pheatmap</a> |

**Value**

heatmap of posterior probabilities of the phentypes above the set threshold

---

|                                   |
|-----------------------------------|
| cophe_multi_trait_data            |
| <i>Simulated multi-trait data</i> |

---

**Description**

Simulated multi-trait data

**Usage**

```
data(cophe_multi_trait_data)
```

**Format**

list of coloc structred datasets for 24 traits (cophe\_multi\_trait\_data\$summ\_stat), LD matrix (cophe\_multi\_trait\_data\$LD) and the id of the query snp (cophe\_multi\_trait\_data\$querysnpid). #' The trait dataset are simulated summary statistics (1000 SNPs) for 10 Hn, 10 Ha and 10 Hc.

---

|            |                                                                                                   |
|------------|---------------------------------------------------------------------------------------------------|
| cophe_plot | <i>cophe_plots showing the Ha and Hc of all traits and labelled above the specified threshold</i> |
|------------|---------------------------------------------------------------------------------------------------|

---

**Description**

cophe\_plots showing the Ha and Hc of all traits and labelled above the specified threshold

**Usage**

```
cophe_plot(  
  multi.dat,  
  querysnpid,  
  query_trait_names,  
  thresh_Hc = 0.5,  
  thresh_Ha = 0.5,  
  beta_p = NULL,  
  traits.dat = NULL,  
  group_pheno = NULL  
)
```

**Arguments**

|                   |                                                                                                                                                                                                                |
|-------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| multi.dat         | multi trait cophescan results returned from <code>cophe.multitrait</code> or <code>multitrait.simplify</code>                                                                                                  |
| querysnpid        | query variant (only a single variant for PheWAS plots)                                                                                                                                                         |
| query_trait_names | list of phenotype names                                                                                                                                                                                        |
| thresh_Hc         | Hc threshold to be displayed                                                                                                                                                                                   |
| thresh_Ha         | Ha threshold to be displayed                                                                                                                                                                                   |
| beta_p            | data.frame (from the <code>get.beta</code> function) with four columns : 1. "beta_plot": indicating beta direction (p or n) 2. "beta_plot": -log10(pval) of the queried variant 3. "querysnp" 4. "querytrait". |
| traits.dat        | list of multi-trait coloc structured datasets                                                                                                                                                                  |
| group_pheno       | Vector with additional grouping of phenotypes                                                                                                                                                                  |

**Value**

cophescan plots of Ha and Hc

**See Also**

[cophe.single](#), [cophe.susie](#), [cophe.multitrait](#), [multitrait.simplify](#)

---

get\_beta

---

*Extract beta and p-values of queried variant*


---

**Description**

Extract beta and p-values of queried variant

**Usage**

```
get_beta(traits.dat, querysnpid, querytrait)
```

**Arguments**

|            |                                  |
|------------|----------------------------------|
| traits.dat | list of coloc structured dataset |
| querysnpid | vector of querysnpid             |
| querytrait | vector of querytrait names       |

**Value**

data.frame with one column named `beta_plot`: indicating beta direction (n/p) and another column named `pval_plot` with -log10(pval) of the queried variant

---

|                    |                                                           |
|--------------------|-----------------------------------------------------------|
| get_posterior_prob | <i>Calculation of the posterior prob of Hn, Ha and Hc</i> |
|--------------------|-----------------------------------------------------------|

---

**Description**

Calculation of the posterior prob of Hn, Ha and Hc

**Usage**

```
get_posterior_prob(params, lbf_mat, nsnp, covar_vec, covar = FALSE)
```

**Arguments**

|           |                                                                   |
|-----------|-------------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$             |
| lbf_mat   | matrix of log bayes factors: IBF.Ha and IBF.Hc                    |
| nsnp      | number of snps                                                    |
| covar_vec | Vector of the covariate                                           |
| covar     | logical: should the covariate information be used? default: False |

**Value**

posterior prob of Hn, Ha and Hc

---

|               |                                                    |
|---------------|----------------------------------------------------|
| Hc.cutoff.fdr | <i>Estimate the Hc.cutoff for the required FDR</i> |
|---------------|----------------------------------------------------|

---

**Description**

Estimate the Hc.cutoff for the required FDR

**Usage**

```
Hc.cutoff.fdr(ppHc, fdr = 0.05, return_plot = TRUE)
```

**Arguments**

|             |                                                                                              |
|-------------|----------------------------------------------------------------------------------------------|
| ppHc        | a vector containing the PP.Hc (the posterior probability of causal association) of all tests |
| fdr         | FDR default: 0.05                                                                            |
| return_plot | default: TRUE, plot the fdr estimated at the different Hc.cutoff                             |

**Value**

the Hc.cutoff value for the specified FDR, if return\_plot is True returns a plot showing the FDR calculated at different Hc thresholds

---

|                   |                          |
|-------------------|--------------------------|
| hypothesis.priors | <i>hypothesis.priors</i> |
|-------------------|--------------------------|

---

**Description**

Estimate priors for each hypothesis

**Usage**

```
hypothesis.priors(nsnps, pn, pa, pc)
```

**Arguments**

|       |                                                                                                    |
|-------|----------------------------------------------------------------------------------------------------|
| nsnps | number of SNPs                                                                                     |
| pn    | prior probability that none of the SNPs/variants in the region are associated with the query trait |
| pa    | prior probability that a non-query variant is causally associated with the query trait             |
| pc    | prior probability that the query variant is causally associated with the query trait               |

**Value**

hypotheses priors

**Author(s)**

Ichcha Manipur

---

|            |                        |
|------------|------------------------|
| logd_alpha | <i>dnorm for alpha</i> |
|------------|------------------------|

---

**Description**

dnorm for alpha

**Usage**

```
logd_alpha(a, alpha_mean = -10, alpha_sd = 0.5)
```

**Arguments**

|            |                                           |
|------------|-------------------------------------------|
| a          | current alpha                             |
| alpha_mean | prior for the mean of alpha               |
| alpha_sd   | prior for the standard deviation of alpha |

**Value**

log dnorm

---

|           |                        |
|-----------|------------------------|
| logd_beta | <i>dgamma for beta</i> |
|-----------|------------------------|

---

**Description**

dgamma for beta

**Usage**

logd\_beta(b, beta\_shape = 2, beta\_scale = 2)

**Arguments**

- b                      current beta
- beta\_shape           prior for the shape (gamma distibution) of beta
- beta\_scale           prior for the scale of beta

**Value**

log dgamma

---

|            |                         |
|------------|-------------------------|
| logd_gamma | <i>dgamma for gamma</i> |
|------------|-------------------------|

---

**Description**

dgamma for gamma

**Usage**

logd\_gamma(g, gamma\_shape = 2, gamma\_scale = 2)

**Arguments**

- g                      current gamma
- gamma\_shape          prior for the shape (gamma distibution) of gamma
- gamma\_scale          prior for the scale of gamma

**Value**

log dgamma

---

|        |                                   |
|--------|-----------------------------------|
| loglik | <i>Log likelihood calculation</i> |
|--------|-----------------------------------|

---

**Description**

Log likelihood calculation

**Usage**

```
loglik(params, lbf_mat, nsnp, covar_vec, covar = FALSE)
```

**Arguments**

|           |                                                                   |
|-----------|-------------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$             |
| lbf_mat   | matrix of log bayes factors: IBF.Ha and IBF.Hc                    |
| nsnp      | number of snps                                                    |
| covar_vec | Vector of the covariate                                           |
| covar     | logical: should the covariate information be used? default: False |

**Value**

logpost flog of the posteriors

---

|         |                                  |
|---------|----------------------------------|
| logpost | <i>Log posterior calculation</i> |
|---------|----------------------------------|

---

**Description**

Log posterior calculation

**Usage**

```
logpost(params, lbf_mat, nsnp, covar_vec, covar = FALSE)
```

**Arguments**

|           |                                                                   |
|-----------|-------------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$             |
| lbf_mat   | matrix of log bayes factors: IBF.Ha and IBF.Hc                    |
| nsnp      | number of snps                                                    |
| covar_vec | Vector of the covariate                                           |
| covar     | logical: should the covariate information be used? default: False |

**Value**

logpost flog of the posteriors

---

`logpriors`*Calculate log priors*

---

**Description**

Calculate log priors

**Usage**

```
logpriors(  
  params,  
  covar = FALSE,  
  alpha_mean = -10,  
  alpha_sd = 0.5,  
  beta_shape = 2,  
  beta_scale = 2,  
  gamma_shape = 2,  
  gamma_scale = 2  
)
```

**Arguments**

|                          |                                                                   |
|--------------------------|-------------------------------------------------------------------|
| <code>params</code>      | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$             |
| <code>covar</code>       | logical: Should the covariate information be used? default: False |
| <code>alpha_mean</code>  | prior for the mean of alpha                                       |
| <code>alpha_sd</code>    | prior for the standard deviation of alpha                         |
| <code>beta_shape</code>  | prior for the shape (gamma distribution) of beta                  |
| <code>beta_scale</code>  | prior for the scale of beta                                       |
| <code>gamma_shape</code> | prior for the shape (gamma distribution) of gamma                 |
| <code>gamma_scale</code> | prior for the scale of gamma                                      |

**Value**

log priors

---

|        |               |
|--------|---------------|
| logsum | <i>logsum</i> |
|--------|---------------|

---

**Description**

Internal function, logsum Function directly taken from coloc This function calculates the log of the sum of the exponentiated logs taking out the max, i.e. insuring that the sum is not Inf

**Usage**

logsum(x)

**Arguments**

x                      numeric vector

**Value**

$\max(x) + \log(\text{sum}(\exp(x - \max(x))))$

---

|           |                |
|-----------|----------------|
| logsumexp | <i>Log sum</i> |
|-----------|----------------|

---

**Description**

Log sum

**Usage**

logsumexp(x)

**Arguments**

x                      vector of log scale values to be added

**Value**

log sum of input

---

`metrop_run`*Run the hierarchical mcmc model to infer priors*

---

**Description**

Run the hierarchical mcmc model to infer priors

**Usage**

```
metrop_run(  
  lbf_mat,  
  nsnp,  
  covar_vec,  
  covar = FALSE,  
  nits = 10000L,  
  thin = 1L,  
  alpha_mean = -10,  
  alpha_sd = 0.5,  
  beta_shape = 2,  
  beta_scale = 2,  
  gamma_shape = 2,  
  gamma_scale = 2  
)
```

**Arguments**

|                          |                                                                   |
|--------------------------|-------------------------------------------------------------------|
| <code>lbf_mat</code>     | matrix of log bayes factors: IBF.Ha and IBF.Hc                    |
| <code>nsnp</code>        | number of snps                                                    |
| <code>covar_vec</code>   | Vector of the covariate                                           |
| <code>covar</code>       | logical: Should the covariate information be used? default: False |
| <code>nits</code>        | Number of iterations run in mcmc                                  |
| <code>thin</code>        | thinning                                                          |
| <code>alpha_mean</code>  | prior for the mean of alpha                                       |
| <code>alpha_sd</code>    | prior for the standard deviation of alpha                         |
| <code>beta_shape</code>  | prior for the shape (gamma distribution) of beta                  |
| <code>beta_scale</code>  | prior for the scale of beta                                       |
| <code>gamma_shape</code> | prior for the shape (gamma distribution) of gamma                 |
| <code>gamma_scale</code> | prior for the scale of gamma                                      |

**Value**

named list of log likelihood (ll) and parameters: alpha, beta and gamma

---

|                                  |                                                                                                                                  |
|----------------------------------|----------------------------------------------------------------------------------------------------------------------------------|
| <code>multitrait.simplify</code> | <i>Simplifying the output obtained from <code>cophe.multitrait</code>, <code>cophe.single</code> or <code>cophe.susie</code></i> |
|----------------------------------|----------------------------------------------------------------------------------------------------------------------------------|

---

**Description**

Simplifying the output obtained from `cophe.multitrait`, `cophe.single` or `cophe.susie`

**Usage**

```
multitrait.simplify(multi.dat, only_BF = FALSE)
```

**Arguments**

|                        |                                                                                                            |
|------------------------|------------------------------------------------------------------------------------------------------------|
| <code>multi.dat</code> | output obtained from <code>cophe.multitrait</code> , <code>cophe.single</code> or <code>cophe.susie</code> |
| <code>only_BF</code>   | return only bayes factors and not posterior probabilities (default=FALSE)                                  |

**Value**

dataframe with posterior probabilities of  $H_n$ ,  $H_c$  and  $H_a$

---

|                       |                                                                                                                                                            |
|-----------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <code>pars2pik</code> | <i>Conversion of parameters <math>\alpha</math>, <math>\beta</math> and <math>\gamma</math> to <math>pnk</math>, <math>pak</math> and <math>pck</math></i> |
|-----------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------|

---

**Description**

Conversion of parameters  $\alpha$ ,  $\beta$  and  $\gamma$  to  $pnk$ ,  $pak$  and  $pck$

**Usage**

```
pars2pik(params, nsnp, covar_vec, covar = FALSE)
```

**Arguments**

|                        |                                                                   |
|------------------------|-------------------------------------------------------------------|
| <code>params</code>    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$             |
| <code>nsnp</code>      | number of snps                                                    |
| <code>covar_vec</code> | Vector of the covariate                                           |
| <code>covar</code>     | logical: should the covariate information be used? default: False |

**Value**

pik matrix of priors:  $pnk$ ,  $pak$  and  $pck$

---

|           |                                           |
|-----------|-------------------------------------------|
| pars_init | Initiate parameters alpha, beta and gamma |
|-----------|-------------------------------------------|

---

**Description**

Initiate parameters alpha, beta and gamma

**Usage**

```
pars_init(  
  covar = FALSE,  
  alpha_mean = -10,  
  alpha_sd = 0.5,  
  beta_shape = 2,  
  beta_scale = 2,  
  gamma_shape = 2,  
  gamma_scale = 2  
)
```

**Arguments**

|             |                                                                   |
|-------------|-------------------------------------------------------------------|
| covar       | logical: Should the covariate information be used? default: False |
| alpha_mean  | prior for the mean of alpha                                       |
| alpha_sd    | prior for the standard deviation of alpha                         |
| beta_shape  | prior for the shape (gamma distribution) of beta                  |
| beta_scale  | prior for the scale of beta                                       |
| gamma_shape | prior for the shape (gamma distribution) of gamma                 |
| gamma_scale | prior for the scale of gamma                                      |

**Value**

params  $\alpha$ ,  $\beta$  and  $\gamma$

---

|                |                       |
|----------------|-----------------------|
| per.snp.priors | <i>per.snp.priors</i> |
|----------------|-----------------------|

---

**Description**

Estimate per snp priors

Usage

```
per.snp.priors(  
  nsnp,  
  pa = 3.82e-05,  
  pc = 0.00182,  
  p1 = NULL,  
  p2 = NULL,  
  p12 = NULL  
)
```

Arguments

|      |                                                                                                                                                                                  |
|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| nsnp | number of SNPs                                                                                                                                                                   |
| pa   | prior probability that a non-query variant is causally associated with the query trait (cophescan prior), default 3.82e-5                                                        |
| pc   | prior probability that the query variant is causally associated with the query trait (cophescan prior), default 1.82e-3 (cophescan prior)                                        |
| p1   | prior probability a SNP is associated with trait 1, (coloc prior), pc derived by using $pc = p12/p1 + p12$ ; use p1, p2, p12 only when pa and pc are unavailable (See vignettes) |
| p2   | prior probability a SNP is associated with trait 2, (coloc prior), pa derived by using $pa = p2$                                                                                 |
| p12  | prior probability a SNP is associated with both traits, (coloc prior), pc derived by using $pc = p12/p1 + p12$                                                                   |

Value

priors at the query variant

Author(s)

Ichcha Manipur

---

|      |                                                          |
|------|----------------------------------------------------------|
| piks | <i>List of priors: pn, pa and pc over all iterations</i> |
|------|----------------------------------------------------------|

---

Description

List of priors: pn, pa and pc over all iterations

Usage

```
piks(params, nsnp, covar_vec, covar = FALSE)
```

**Arguments**

|           |                                                             |
|-----------|-------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$       |
| nsnps     | number of snps                                              |
| covar_vec | Vector of the covariate                                     |
| covar     | logical: was the covariate information used? default: False |

**Value**

List of priors (len: iterations): pnk, pak and pck

---

|                   |                                                                           |
|-------------------|---------------------------------------------------------------------------|
| plot_trait_manhat | <i>Plot region Manhattan for a trait highlighting the queried variant</i> |
|-------------------|---------------------------------------------------------------------------|

---

**Description**

Plot region Manhattan for a trait highlighting the queried variant

**Usage**

```
plot_trait_manhat(trait.dat, querysnpid, alt.snpid = NULL)
```

**Arguments**

|            |                                                                           |
|------------|---------------------------------------------------------------------------|
| trait.dat  | dataset used as input for running cophescan                               |
| querysnpid | the id of the causal variant as present in trait.dat\$snp, plotted in red |
| alt.snpid  | the id of the other variants as a vector to be plotted, plotted in blue   |

**Value**

regional manhattan plot

---

|                |                                                                                                                     |
|----------------|---------------------------------------------------------------------------------------------------------------------|
| posterior_prob | <i>List of posterior probabilities: <math>H_n</math>, <math>H_a</math> and <math>H_c</math> over all iterations</i> |
|----------------|---------------------------------------------------------------------------------------------------------------------|

---

**Description**

List of posterior probabilities:  $H_n$ ,  $H_a$  and  $H_c$  over all iterations

**Usage**

```
posterior_prob(params, lbf_mat, nsnps, covar_vec, covar = FALSE)
```

**Arguments**

|           |                                                             |
|-----------|-------------------------------------------------------------|
| params    | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$       |
| lbf_mat   | matrix of log bayes factors: lbf.Ha and lbf.Hc              |
| nsnps     | number of snps                                              |
| covar_vec | Vector of the covariate                                     |
| covar     | logical: was the covariate information used? default: False |

**Value**

List of posterior probabilities (len: iterations): Hn, Ha and Hc

---

|                   |                                  |
|-------------------|----------------------------------|
| prepare_plot_data | <i>Prepare data for plotting</i> |
|-------------------|----------------------------------|

---

**Description**

Prepare data for plotting

**Usage**

```
prepare_plot_data(  
  multi.dat,  
  querysnpid,  
  query_trait_names,  
  thresh_Ha = 0.5,  
  thresh_Hc = 0.5,  
  hmp = FALSE,  
  cophe.plot = TRUE  
)
```

**Arguments**

|                   |                                                                                     |
|-------------------|-------------------------------------------------------------------------------------|
| multi.dat         | multi trait cophescan results returned from cophe.multitrait or multitrait.simplify |
| querysnpid        | query variant                                                                       |
| query_trait_names | vector of names of the query traits                                                 |
| thresh_Ha         | Ha threshold to be displayed                                                        |
| thresh_Hc         | Hc threshold to be displayed                                                        |
| hmp               | return for heatmap                                                                  |
| cophe.plot        | default: TRUE, return for cophe_plot                                                |

**Value**

plot list

**See Also**

[cophe\\_plot](#), [cophe.susie](#), [cophe.multitrait](#), [multitrait.simplify](#) default NULL

---

|         |                              |
|---------|------------------------------|
| propose | <i>Proposal distribution</i> |
|---------|------------------------------|

---

**Description**

Proposal distribution

**Usage**

```
propose(params, propsd = 0.5)
```

**Arguments**

|        |                                                       |
|--------|-------------------------------------------------------|
| params | Vector of parameters: $\alpha$ , $\beta$ and $\gamma$ |
| propsd | Standard deviation for the proposal                   |

**Value**

vector : proposal

---

|                   |                                                                       |
|-------------------|-----------------------------------------------------------------------|
| run_metrop_priors | <i>Run the hierarchical Metropolis Hastings model to infer priors</i> |
|-------------------|-----------------------------------------------------------------------|

---

**Description**

Run the hierarchical Metropolis Hastings model to infer priors

**Usage**

```
run_metrop_priors(
  multi.dat,
  covar = FALSE,
  covar_vec = NULL,
  is_covar_categorical = FALSE,
  nits = 10000,
  thin = 1,
  posterior = FALSE,
  avg_pik = TRUE,
  avg_posterior = TRUE,
  pik = FALSE,
  alpha_mean = -10,
  alpha_sd = 0.5,
```

```

    beta_shape = 2,
    beta_scale = 2,
    gamma_shape = 2,
    gamma_scale = 2
)

```

### Arguments

|                                   |                                                                                       |
|-----------------------------------|---------------------------------------------------------------------------------------|
| <code>multi.dat</code>            | matrix of bf values, rows=traits, named columns=("IBF.Ha", "IBF.Hc", "nsnps")         |
| <code>covar</code>                | whether to include covariates                                                         |
| <code>covar_vec</code>            | vector of covariates                                                                  |
| <code>is_covar_categorical</code> | only two categories supported (default=FALSE) - Experimental                          |
| <code>nits</code>                 | number of iterations                                                                  |
| <code>thin</code>                 | burnin                                                                                |
| <code>posterior</code>            | default: FALSE, estimate posterior probabilities of the hypotheses                    |
| <code>avg_pik</code>              | default: FALSE, estimate the average of the pik                                       |
| <code>avg_posterior</code>        | default: FALSE, estimate the average of the posterior probabilities of the hypotheses |
| <code>pik</code>                  | default: FALSE, inferred prior probabilities                                          |
| <code>alpha_mean</code>           | prior for the mean of alpha                                                           |
| <code>alpha_sd</code>             | prior for the standard deviation of alpha                                             |
| <code>beta_shape</code>           | prior for the shape (gamma distribution) of beta                                      |
| <code>beta_scale</code>           | prior for the scale of beta                                                           |
| <code>gamma_shape</code>          | prior for the shape (gamma distribution) of gamma                                     |
| <code>gamma_scale</code>          | prior for the scale of gamma                                                          |

### Value

List containing the posterior distribution of the parameters alpha, beta, gamma (if covariate included) and the loglikelihood

if `avg_posterior=TRUE` matrix with average of all the posterior probabilities of Hn, Ha and Hc

if `avg_pik=TRUE` matrix with average of all the priors: pn, pa and pc

`data`, `nits` and `thin` contain the input data, number of iterations and burnin respectively specified for the hierarchical model

---

|              |                     |
|--------------|---------------------|
| sample_alpha | <i>sample alpha</i> |
|--------------|---------------------|

---

**Description**

sample alpha

**Usage**

```
sample_alpha(alpha_mean = -10, alpha_sd = 0.5)
```

**Arguments**

|            |                                           |
|------------|-------------------------------------------|
| alpha_mean | prior for the mean of alpha               |
| alpha_sd   | prior for the standard deviation of alpha |

**Value**

sample from mnorm for  $\alpha$

---

|             |                    |
|-------------|--------------------|
| sample_beta | <i>sample beta</i> |
|-------------|--------------------|

---

**Description**

sample beta

**Usage**

```
sample_beta(beta_shape = 2, beta_scale = 2)
```

**Arguments**

|            |                                                  |
|------------|--------------------------------------------------|
| beta_shape | prior for the shape (gamma distribution) of beta |
| beta_scale | prior for the scale of beta                      |

**Value**

sample from rgamma for  $\beta$

---

|              |                     |
|--------------|---------------------|
| sample_gamma | <i>sample gamma</i> |
|--------------|---------------------|

---

**Description**

sample gamma

**Usage**

```
sample_gamma(gamma_shape = 2, gamma_scale = 2)
```

**Arguments**

|             |                                                   |
|-------------|---------------------------------------------------|
| gamma_shape | prior for the shape (gamma distribution) of gamma |
| gamma_scale | prior for the scale of gamma                      |

**Value**

sample from rgamma for  $\gamma$

---

|               |                                                                    |
|---------------|--------------------------------------------------------------------|
| summary.cophe | <i>print the summary of results from cophescan single or susie</i> |
|---------------|--------------------------------------------------------------------|

---

**Description**

print the summary of results from cophescan single or susie

**Usage**

```
## S3 method for class 'cophe'
summary(object, ...)
```

**Arguments**

|        |                                                      |
|--------|------------------------------------------------------|
| object | Result from either cophe.susie or cophe.single       |
| ...    | additional arguments affecting the summary produced. |

**Value**

log bayes and posterior probabilities

**See Also**

[cophe.single](#), [cophe.susie](#)

---

|        |                            |
|--------|----------------------------|
| target | <i>Target distribution</i> |
|--------|----------------------------|

---

**Description**

Target distribution

**Usage**

```
target(params, lbf_mat, nsnp, covar_vec, covar = FALSE)
```

**Arguments**

- params            Vector of parameters:  $\alpha$ ,  $\beta$  and  $\gamma$
- lbf\_mat          matrix of log bayes factors: lbf.Ha and lbf.Hc
- nsnp             number of snps
- covar\_vec        Vector of the covariate
- covar            logical: Should the covariate information be used? default: False

**Value**

target

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