# Package 'UKB.COVID19'

August 17, 2024
Type Package
<b>Title</b> UK Biobank COVID-19 Data Processing and Risk Factor Association Tests
Version 0.1.6
<b>Date</b> 2024-08-17
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<b>Description</b> Process UK Biobank COVID-19 test result data for susceptibility, severity and mortality analyses, perform potential nongenetic COVID-19 risk factor and comorbidity association tests. Wang et al. (2021) <doi:10.5281 zenodo.5174381="">.</doi:10.5281>
Imports questionr, data.table, tidyverse, magrittr, here, dplyr
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<pre>URL https://github.com/bahlolab/UKB.COVID19</pre>
Encoding UTF-8
RoxygenNote 7.3.1
Suggests knitr, rmarkdown, testthat (>= 3.0.0)
Config/testthat/edition 3
NeedsCompilation no
Repository CRAN
<b>Date/Publication</b> 2024-08-17 10:20:59 UTC
VignetteBuilder knitr
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comorbidity\_asso

Generate comorbidity association result file

#### Description

Association tests between each co-morbidity and given phenotype (susceptibility, mortality or severity) with the adjustment of covariates.

#### Usage

```
comorbidity_asso(
  pheno,
  covariates,
  cormorbidity,
  population = "all",
  cov.name = c("sex", "age", "bmi"),
  phe.name,
  ICD10.file
)
```

#### Arguments

pheno	phenotype dataframe - output from makePheno function
covariates	covariate dataframe - output from risk.factor function. Optional.
cormorbidity	Comorbidity summary generated from comorbidity.summary.
population	Choose self-report population/ethnic background group from "all", white", "black", "asian", "mixed", or "other". By default, population="all", include all ethnic groups.
cov.name	Selected covariates names. By default, cov.name=c("sex","age","bmi"), covariates are sex age and BMI.
phe.name	Phenotype name.
ICD10.file	The ICD10 code file, which is included in the package.

#### Value

Outputs a comorbidity association test result with OR, 95% CI and p-value.

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#### **Examples**

```
## Not run:
comorb.asso <- comorbidity_asso(pheno=phe,
covariates=covar,
cormorbidity=comorb,
population="white",
cov.name=c("sex","age","bmi","SES","smoke","inAgedCare"),
phe.name="hospitalisation",
ICD10.file=covid_example("ICD10.coding19.txt.gz"))
## End(Not run)</pre>
```

comorbidity\_summary

Create comorbidity summary file

#### **Description**

summarise disease history records of each individual from the hospital inpatient diagnosis data.

#### Usage

```
comorbidity_summary(
  ukb.data,
  hesin.file,
  hesin_diag.file,
  primary = FALSE,
  ICD10.file,
  Date.start = NULL,
  Date.end = NULL
)
```

#### **Arguments**

ukb.data tab delimited UK Biobank phenotype file, containing sample qc fields (with

default UKBiobank codes as column names)

hesin.file Latest hospital inpatient master file.

hesin\_diag.file

Latest hospital inpatient diagnosis file.

primary TRUE: include primary diagnosis only; FALSE: include all diagnoses.

ICD10. file The ICD10 code file, which is included in the package.

Date.start Date, dd/mm/yyyy, select the start date of hospital inpatient record period.

Date . end Date, dd/mm/yyyy, select the end date of hospital inpatient record period.

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#### Value

Outputs comorbidity summary table, named comorbidity\_<Date.start>\_<Date.end>.RData, including phenotype, non-genetic risk factors and all comorbidities, which will be used in the comorbidity association tests.

#### **Examples**

```
## Not run:
comorb <- comorbidity_summary(ukb.data=covid_example("sim_ukb.tab.gz"),
hesin.file=covid_example("sim_hesin.txt.gz"),
hesin_diag.file=covid_example("sim_hesin_diag.txt.gz"),
ICD10.file=covid_example("ICD10.coding19.txt.gz"),
primary = FALSE,
Date.start = "16/03/2020")
## End(Not run)</pre>
```

covid\_example

Provide working directory for UKB.COVID19 example files

#### Description

Provide working directory for UKB.COVID19 example files

#### Usage

```
covid_example(path)
```

#### **Arguments**

path path to file

#### Value

Outputs the working directory for UKB.COVID19 example files.

```
covid_example('results/covariate.txt')
```

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Reform variables		
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#### Description

Reform variables

#### Usage

```
data_reform(res, type)
```

#### **Arguments**

res Merged data of phenotype from makePhenotypes or comorbidity\_summary and

covariates from risk\_factor.

type Data type: susceptibility, severity, mortality or comorbidity.

#### Value

Reformed data for association tests using logistic regression models.

log_cov	Perform association tests between phenotype and covariates

#### Description

Perform association tests between phenotype and covariates

#### Usage

```
log_cov(pheno, covariates, phe.name, cov.name = c("sex", "age", "bmi"))
```

#### **Arguments**

pheno phenotype dataframe - output from makePhenotypes function

covariates covariate dataframe - output from risk\_factor function.

phe. name Phenotype name in the data.

cov.name Selected covariate names in the data. By default, cov.name=c("sex", "age", "bmi"),

covariates include sex, age and BMI.

#### Value

Outputs association test results with OR, 95% CI, and p-value.

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#### **Examples**

#### **Description**

Generate files for GWAS Software. SAIGE and Plink currently supported.

#### Usage

```
makeGWASFiles(
  ukb.data,
  pheno,
  covariates,
  phe.name,
  cov.name = NULL,
  includeSampsFile = NULL,
  software = "SAIGE",
  outDir = "",
  prefix
)
```

#### Arguments

default UKBiobank codes as column names)

pheno phenotype dataframe - output from makePhenotype function covariates covariate dataframe - output from risk.factor function. Optional.

phe.name phenotypes to be included in outputted data. multiple phenotypes can be speci-

fied as a vector. if null, all phenotypes will be outputted.

cov. name covariates to be included in outputted data. Optional. multiple covariates can be

specified as a vector. if null, all covariates in file will be outputted

includeSampsFile

list of samples to be included GWAS. File with the first column containing sample IDs to be kept. Can contain other columns. output from sampleQC function

may be used. Optional - if null, all samples will be outputted.

software specify "SAIGE" or "plink" - defaults to "SAIGE"

outDir specify directory to output file

prefix prefix for file - optional

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#### Value

outputs file, suitable for reading by chosen GWAS software

#### **Examples**

```
## Not run:
makeGWASFiles(ukb.data=covid_example("sim_ukb.tab.gz"),
pheno=phe,
covariates=covar,
phe.name="hospitalisation",
cov.name=NULL,
includeSampsFile=NULL,
software="SAIGE",
outDir=covid_example("results"),
prefix="hospitalisation")
## End(Not run)
```

makePhenotypes

Generate COVID-19 phenotypes

#### **Description**

Generate COVID-19 phenotypes

#### Usage

```
makePhenotypes(
  ukb.data,
  res.eng,
  res.wal = NULL,
  res.sco = NULL,
  death.file,
  death.cause.file,
  hesin_diag.file,
  hesin_oper.file,
  hesin_critical.file,
  code.file,
  pheno.type = "severity",
  Date = NULL
)
```

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#### **Arguments**

ukb.data	tab delimited UK Biobank phenotype file.
res.eng	Latest covid result file/files for England.
res.wal	Latest covid result file/files for Wales. Only available for downloads after April 2021.
res.sco	Latest covid result file/files for Scotland. Only available for downloads after April 2021.
death.file	Latest death register file.
death.cause.fil	e
	Latest death cause file.
hesin.file	Latest hospital inpatient master file.
hesin_diag.file	
	Latest hospital inpatient diagnosis file.
hesin_oper.file	
	Latest hospital inpatient operation file.
hesin_critical.	file
	Latest hospital inpatient critical care file.
code.file	The operation code file, which is included in the package.
pheno.type	The phenotype options, which include "susceptibility", "severity", and "mortality".
Date	Date, ddmmyyyy, select the results until a certain date. By default, Date = NULL, the latest hospitalization date.

#### Value

Returns a data.frame with phenotypes for COVID-19 susceptibility, severity and mortality.

```
## Not run:
pheno <- makePhenotypes(ukb.data=covid_example("sim_ukb.tab.gz"),
res.eng=covid_example("sim_result_england.txt.gz"),
death.file=covid_example("sim_death.txt.gz"),
death.cause.file=covid_example("sim_death_cause.txt.gz"),
hesin.file=covid_example("sim_hesin.txt.gz"),
hesin_diag.file=covid_example("sim_hesin_diag.txt.gz"),
hesin_oper.file=covid_example("sim_hesin_oper.txt.gz"),
hesin_critical.file=covid_example("sim_hesin_critical.txt.gz"),
code.file=covid_example("coding240.txt.gz"),
pheno.type = "severity")

## End(Not run)</pre>
```

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risk_factor Generate covariate file
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#### Description

This function formats and outputs a covariate table, used for input for other functions.

#### Usage

```
risk_factor(
  ukb.data,
  ABO.data = NULL,
  hesin.file,
  res.eng,
  res.wal = NULL,
  res.sco = NULL,
  fields = NULL,
  field.names = NULL)
```

#### **Arguments**

ukb.data	tab delimited UK Biobank phenotype file. The file should include fields of gender, year of birth, BMI, ethnic background, SES, and smoking.
ABO.data	Latest yyyymmdd_covid19_misc.txt file.
hesin.file	Latest yyyymmdd_hesin.txt file.
res.eng	Latest covid result file/files for England.
res.wal	Latest covid result file/files for Wales. Only available for downloads after April 2021.
res.sco	Latest covid result file/files for Scotland. Only available for downloads after April 2021.
fields	User specified field codes from ukb.data file.
field.names	User specified field names.

#### Value

Outputs a covariate table, used for input for other functions. Automatically returns sex, age at birthday in 2020, SES, self-reported ethnicity, most recently reported BMI, most recently reported pack-years, whether they reside in aged care (based on hospital admissions data, and covid test data) and blood type. Function also allows user to specify fields of interest (field codes, provided by UK Biobank), and allows the users to specify more intuitive names, for selected fields.

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#### **Examples**

```
## Not run:
covars <- risk_factor(ukb.data=covid_example("sim_ukb.tab.gz"),
ABO.data=covid_example("sim_covid19_misc.txt.gz"),
hesin.file=covid_example("sim_hesin.txt.gz"),
res.eng=covid_example("sim_result_england.txt.gz"))
## End(Not run)</pre>
```

sampleQC

Sample QC for genetic analyses

#### **Description**

Sample QC for genetic analyses

#### Usage

```
sampleQC(ukb.data, withdrawnFile, ancestry = "all", software = "SAIGE", outDir)
```

#### **Arguments**

ukb.data tab delimited UK Biobank phenotype file, containing sample qc fields (with

default UKBiobank codes as column names)

withdrawnFile csv file with withdrawn IDs from UK Biobank
ancestry specify "WhiteBritish" or "all" - defaults to "all"
software specify "SAIGE" or "plink" - defaults to "SAIGE"

outDir specify directory for sample QC file and inclusion/exclusion lists

#### Value

outputs sample QC file, and sample inclusion / exclusion lists for specified software

```
## Not run:
sampleQC(ukb.data=covid_example("sim_ukb.tab.gz"),
withdrawnFile=covid_example("sim_withdrawn.csv.gz"),
ancestry="all",
software="SAIGE",
outDir=covid_example("results"))
## End(Not run)
```

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variantQC	Variant QC for Genetic Analyses	

#### Description

Variant QC for Genetic Analyses

#### Usage

```
variantQC(snpQcFile, mfiDir, mafFilt = 0.001, infoFilt = 0.5, outDir)
```

#### Arguments

mfiDir directory where the per chromosome UKBiobank MAF/INFO files (ukb_mfi_chr*_v3.txt) are located  mafFilt minor allele frequency filter - default 0.001  infoFilt imputation quality (INFO) score filter - default 0.5  outDir output directory	snpQcFile	file containing SNP QC info (ukb_snp_qc.txt)
infoFilt imputation quality (INFO) score filter - default 0.5	mfiDir	,
	mafFilt	minor allele frequency filter - default 0.001
outDir output directory	infoFilt	imputation quality (INFO) score filter - default 0.5
	outDir	output directory

#### Value

outputs SNP inclusion lists (SNPID and rsID formats) for given MAF/INFO filters. Also outputs list of SNPs to be used for genetic Relatedness Matrix (GRM) calculations.

```
## Not run:
variantQC(snpQcFile=covid_example("sim_ukb_snp_qc.txt.gz"),
mfiDir=covid_example("alleleFreqs"),
mafFilt=0.001,
infoFilt=0.5,
outDir=covid_example("results"))
## End(Not run)
```

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