Package 'CRTspat'

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Title Workflow for Cluster Randomised Trials with Spillover
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Description

aggregateCRT aggregates data from a "CRTsp" object or trial data frame containing multiple records with the same location, and outputs a list of class "CRTsp" containing single values for each location, for both the coordinates and the auxiliary variables.

Usage

```
aggregateCRT(trial, auxiliaries = NULL)
```

Arguments

An object of class "CRTsp" containing locations (x,y) and variables to be summed auxiliaries vector of names of auxiliary variables to be summed across each location

Details

Variables that in the trial dataframe that are not included in auxiliaries are retained in the output algorithm "CRTsp" object, with the value corresponding to that of the first record for the location in the input data frame

Value

A list of class "CRTsp"

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Examples

```
{
trial <- readdata('example_site.csv')
trial$base_denom <- 1
aggregated <- aggregateCRT(trial, auxiliaries = c("RDT_test_result","base_denom"))
}</pre>
```

anonymize_site

Anonymize locations of a trial site

Description

anonymize_site transforms coordinates to remove potential identification information.

Usage

```
anonymize_site(trial, ID = NULL, latvar = "lat", longvar = "long")
```

Arguments

trial "CRTsp" object or trial data frame with co-ordinates of households

ID name of column used as an identifier for the points
 latvar name of column containing latitudes in decimal degrees
 longvar name of column containing longitudes in decimal degrees

Details

The coordinates are transformed to support confidentiality of information linked to households by replacing precise geo-locations with transformed co-ordinates which preserve distances but not positions. The input may have either lat long or x,y coordinates. The function first searches for any lat long co-ordinates and converts these to x,y Cartesian coordinates. These are then are rotated by a random angle about a random origin. The returned object has transformed co-ordinates re-centred at the origin. Centroids stored in the "CRTsp" object are removed. Other data are unchanged.

Value

```
A list of class "CRTsp".
```

```
#Rotate and reflect test site locations
transformedTestlocations <- anonymize_site(trial = readdata("exampleCRT.txt"))</pre>
```

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coef.CRTanalysis

Extract model coefficients

Description

coef.CRTanalysis method for extracting model fitted values

Usage

```
## S3 method for class 'CRTanalysis'
coef(object, ...)
```

Arguments

```
object CRTanalysis object
... other arguments
```

Value

the model coefficients returned by the statistical model run within the CRTanalysis function

Examples

```
{example <- readdata('exampleCRT.txt')
exampleGEE <- CRTanalysis(example, method = "GEE")
coef(exampleGEE)
}</pre>
```

compute_distance

Compute distance or surround values for a cluster randomized trial

Description

compute_distance computes distance or surround values for a cluster randomized trial (CRT)

Usage

```
compute_distance(
   trial,
   distance = "nearestDiscord",
   scale_par = NULL,
   auxiliary = NULL
)
```

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Arguments

trial an object of class "CRTsp" or a data frame containing locations in (x,y) coordi-

nates, cluster assignments (factor cluster), and arm assignments (factor arm).

distance the quantity(s) to be computed. Options are:

"nearestDiscord" distance to nearest discordant location (km)

"distanceAssigned" distance to the nearest pixel in the assigned cluster (km)

"disc" disc

"kern" kernel-based measure
"hdep" Tukey half space depth
"sdep" simplicial depth

scale_par scale parameter equal to the disc radius in km if distance = "disc" or to the

standard deviance of the kernels if distance = "kern"

auxiliary "CRTsp" object containing external cluster and or arm assignments.

Details

For each selected distance measure, the function first checks whether the variable is already present, and carries out the calculations only if the corresponding field is absent from the trial data frame.

If distance = "nearestDiscord" is selected the computed values are Euclidean distances assigned a positive sign for the intervention arm of the trial, and a negative sign for the control arm.

If distance = "distanceAssigned" is selected the computed values are Euclidean distances to the nearest pixel in the auxiliary "CRTsp" object.

If distance = "disc" is specified, the disc statistic is computed for each location as the number of locations within the specified radius that are in the intervention arm (Anaya-Izquierdo & Alexander(2020)). The input value of scale_par is stored in the design list of the output "CRTsp" object. Recalculation is carried out if the input value of scale_par differs from the one in the input design list. The value of the the surround calculated based on intervened locations is divided by the value of the surround calculated on the basis of all locations, so the value returned is a proportion.

If distance = "kern" is specified, the Normal curve with standard deviation scale_par is used to simulate diffusion of the intervention effect by Euclidean distance. For each location in the trial, the contributions of all intervened locations are summed. As with distance = "disc", when distance = "kern" the surround calculated based on intervened locations is divided by the value of the surround calculated on the basis of all locations, so the value returned is a proportion.

If either distance = "hdep" or distance = "sdep" is specified then both the simplicial depth and Tukey half space depth are calculated using the algorithm of Rousseeuw & Ruts(1996). The half-depth probability within the intervention cloud (di) is computed with respect to other locations in the intervention arm (Anaya-Izquierdo & Alexander(2020)). The half-depth within the half-depth within the control cloud (dc) is also computed. CRTspat returns the proportion di/(dc + di).

If an auxiliary auxiliary "CRTsp" object is specified then either distanceAssigned or nearestDiscord

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(the default) is computed with respect to the assignments in the auxiliary. If the auxiliary is a grid with design\$geometry set to 'triangle', 'square' or 'hexagon' then the distance is computed to the edge of the nearest grid pixel in the discordant arm (using a circular approximation for the perimeter) rather than to the point location itself.

Value

The input "CRTsp" object with additional column(s) added to the trial data frame with variable name corresponding to the input value of distance.

Examples

```
{
# Calculate the disc with a radius of 0.5 km
exampletrial <- compute_distance(trial = readdata('exampleCRT.txt'),
distance = 'disc', scale_par = 0.5)
}</pre>
```

compute_mesh

Create INLA mesh for spatial analysis

Description

compute_mesh create objects required for INLA analysis of an object of class "CRTsp".

Usage

```
compute_mesh(
  trial = trial,
  offset = -0.1,
  max.edge = 0.25,
  inla.alpha = 2,
  maskbuffer = 0.5,
  pixel = 0.5
)
```

Arguments

```
an object of class "CRTsp" or a data frame containing locations in (x,y) coordinates, cluster assignments (factor cluster), and arm assignments (factor arm) and outcome.

offset see inla.mesh.2d documentation

max.edge see inla.mesh.2d documentation

inla.alpha parameter related to the smoothness (see inla documentation)

maskbuffer numeric: width of buffer around points (km)

pixel numeric: size of pixel (km)
```

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Details

compute_mesh carries out the computationally intensive steps required for setting-up an INLA analysis of an object of class "CRTsp", creating the prediction mesh and the projection matrices. The mesh can be reused for different models fitted to the same geography. The computational resources required depend largely on the resolution of the prediction mesh. The prediction mesh is thinned to include only pixels centred at a distance less than maskbuffer from the nearest point.

A warning may be generated if the Matrix library is not loaded.

Value

list

- prediction Data frame containing the prediction points and covariate values
- A projection matrix from the observations to the mesh nodes.
- Ap projection matrix from the prediction points to the mesh nodes.
- indexs index set for the SPDE model
- spde SPDE model
- pixel pixel size (km)

Examples

```
{
# low resolution mesh for test dataset
library(Matrix)
example <- readdata('exampleCRT.txt')
exampleMesh=compute_mesh(example, pixel = 0.5)
}</pre>
```

CRTanalysis

Analysis of cluster randomized trial with spillover

Description

CRTanalysis carries out a statistical analysis of a cluster randomized trial (CRT).

Usage

```
CRTanalysis(
   trial,
   method = "GEE",
   distance = "nearestDiscord",
   scale_par = NULL,
   cfunc = "L",
   link = "logit",
   numerator = "num",
   denominator = "denom",
```

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```
excludeBuffer = FALSE,
      alpha = 0.05,
      baselineOnly = FALSE,
      baselineNumerator = "base_num",
      baselineDenominator = "base_denom",
      personalProtection = FALSE,
      clusterEffects = TRUE,
      spatialEffects = FALSE,
      requireMesh = FALSE,
      inla_mesh = NULL
    )
Arguments
    trial
                      an object of class "CRTsp" or a data frame containing locations in (x,y) coordi-
                      nates, cluster assignments (factor cluster), and arm assignments (factor arm)
                      and outcome data (see details).
    method
                      statistical method with options:
                   "EMP"
                              simple averages of the data
                    "T"
                             comparison of cluster means by t-test
                    "GEE"
                              Generalised Estimating Equations
                    "LME4"
                             Generalized Linear Mixed-Effects Models
                    "INLA"
                             Integrated Nested Laplace Approximation (INLA)
                    "MCMC"
                             Markov chain Monte Carlo using "JAGS"
                    "WCA"
                              Within cluster analysis
    distance
                      Measure of distance or surround with options:
                 "nearestDiscord"
                                      distance to nearest discordant location (km)
                 "disc"
                 "kern"
                                      surround based on sum of normal kernels
                 "hdep"
                                      Tukey half space depth
                 "sdep"
                                      simplicial depth
    scale_par
                      numeric: pre-specified value of the spillover parameter or disc radius for models
                      where this is fixed (cfunc = "R").
    cfunc
                      transformation defining the spillover function with options:
```

"Z" arm effects not considered reference model

"X" spillover not modelled the only valid value of cfunc for methods "EMP", "T" and "GEE"

"L" inverse logistic (sigmoid) the default for "INLA" and "MCMC" methods

"P" inverse probit (error function) available with "INLA" and "MCMC" methods

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"S"	piecewise linear	only available with the "MCMC" method
"E"	estimation of scale factor	only available with distance = "disc" or distance = "kern"
"R"	rescaled linear	

link link function with options:

"logit" (the default). numerator has a binomial distribution with denominator denominator.

"log" numerator is Poisson distributed with an offset of log(denominator).

"cloglog" numerator is Bernoulli distributed with an offset of log(denominator).

The outcome is numerator/denominator with a normally distributed error function.

numerator string: name of numerator variable for outcome

denominator string: name of denominator variable for outcome data (if present)

excludeBuffer logical: indicator of whether any buffer zone (records with buffer=TRUE) should

be excluded from analysis

alpha numeric: confidence level for confidence intervals and credible intervals

baselineOnly logical: indicator of whether required analysis is of effect size or of baseline

only

baselineNumerator

string: name of numerator variable for baseline data (if present)

baselineDenominator

string: name of denominator variable for baseline data (if present)

personalProtection

logical: indicator of whether the model includes local effects with no spillover

clusterEffects logical: indicator of whether the model includes cluster random effects

spatialEffects logical: indicator of whether the model includes spatial random effects (avail-

able only for method = "INLA")

requireMesh logical: indicator of whether spatial predictions are required (available only for

method = "INLA")

inla_mesh string: name of pre-existing INLA input object created by compute_mesh()

Details

CRTanalysis is a wrapper for the statistical analysis packages: geepack, INLA, jagsUI, and the t.test function of package stats.

The wrapper does not provide an interface to the full functionality of these packages. It is specific for typical analyses of cluster randomized trials with geographical clustering. Further details are provided in the vignette.

The key results of the analyses can be extracted using a summary() of the output list. The model_object in the output list is the usual output from the statistical analysis routine, and can be also be inspected with summary(), or analysed using stats::fitted() for purposes of evaluation of model fit etc..

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For models with a complementary log-log link function specified with link = "cloglog". the numerator must be coded as 0 or 1. Technically the binomial denominator is then 1. The value of denominator is used as a rate multiplier.

With the "INLA" and "MCMC" methods 'iid' random effects are used to model extra-Poisson variation.

Interval estimates for the coefficient of variation of the cluster level outcome are calculated using the method of Vangel (1996).

Value

list of class CRTanalysis containing the following results of the analysis:

• description : description of the dataset

• method: statistical method

• pt_ests: point estimates

• int_ests: interval estimates

• model_object : object returned by the fitting routine

• spillover: function values and statistics describing the estimated spillover

Examples

```
example <- readdata('exampleCRT.txt')
# Analysis of test dataset by t-test
exampleT <- CRTanalysis(example, method = "T")
summary(exampleT)
# Standard GEE analysis of test dataset ignoring spillover
exampleGEE <- CRTanalysis(example, method = "GEE")
summary(exampleGEE)
# LME4 analysis with error function spillover function
exampleLME4 <- CRTanalysis(example, method = "LME4", cfunc = "P")
summary(exampleLME4)</pre>
```

CRTpower

Power and sample size calculations for a cluster randomized trial

Description

CRTpower carries out power and sample size calculations for cluster randomized trials.

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Usage

```
CRTpower(
  trial = NULL,
  locations = NULL,
  alpha = 0.05,
  desiredPower = 0.8,
  effect = NULL,
 yC = NULL,
  outcome_type = "d",
  sigma2 = NULL,
  denominator = 1,
 N = 1,
  ICC = NULL,
  cv_percent = NULL,
  c = NULL,
  sd_h = 0,
  spillover_interval = 0,
  contaminate_pop_pr = 0,
  distance_distribution = "normal"
)
```

Arguments

dataframe or 'CRTsp' object: optional list of locations trial numeric: total number of units available for randomization (required if trial is locations not specified) alpha numeric: confidence level numeric: desired power desiredPower effect numeric: required effect size yC numeric: baseline (control) value of outcome character: with options - 'y': continuous; 'n': count; 'e': event rate; 'p': outcome_type proportion; 'd': dichotomous. numeric: variance of the outcome (required for outcome_type = 'y') sigma2 numeric: rate multiplier (for outcome_type = 'n' or outcome_type = 'e') denominator numeric: mean of the denominator for proportions (for outcome_type = 'p') Ν ICC numeric: Intra-cluster correlation numeric: Coefficient of variation of the outcome (expressed as a percentage) cv_percent integer: number of clusters in each arm (required if trial is not specified) С numeric: standard deviation of number of units per cluster (required if trial is sd_h not specified) spillover_interval numeric: 95% spillover interval (km) contaminate_pop_pr

numeric: Proportion of the locations within the 95% spillover interval.

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distance_distribution

numeric: algorithm for computing distribution of spillover, with options - 'empirical': empirical distribution; 'normal': normal distribution.

Details

Power and sample size calculations are for an unmatched two-arm trial. For counts or event rate data the formula of Hayes & Bennett, 1999 is used. This requires as an input the between cluster coefficient of variation (cv_percent). For continuous outcomes and proportions the formulae of Hemming et al, 2011 are used. These make use of the intra-cluster correlation in the outcome (ICC) as an input. If the coefficient of variation and not the ICC is supplied then the intra-cluster correlation is computed from the coefficient of variation using the formulae from Hayes & Moulton. If incompatible values for ICC and cv_percent are supplied then the value of the ICC is used.

The calculations do not consider any loss in power due to loss to follow-up and by default there is no adjustment for effects of spillover.

Spillover bias can be allowed for using a diffusion model of mosquito movement. If no location or arm assignment information is available then contaminate_pop_pr is used to parameterize the model using a normal approximation for the distribution of distance to discordant locations.

If a trial data frame or 'CRTsp' object is input then this is used to determine the number of locations. If this input object contains cluster assignments then the numbers and sizes of clusters in the input data are used to estimate the power. If spillover_interval > 0 and distance_distribution = 'empirical' then effects of spillover are incorporated into the power calculations based on the empirical distribution of distances to the nearest discordant location. (If distance_distribution is not equal to 'empirical' then the distribution of distances is assumed to be normal.

If geolocations are not input then power and sample size calculations are based on the scalar input parameters.

If buffer zones have been specified in the 'CRTsp' object then separate calculations are made for the core area and for the full site.

The output is an object of class 'CRTsp' containing any input trial data frame and values for:

- The required numbers of clusters to achieve the specified power.
- The design effect based on the input ICC.
- Calculations of the power ignoring any bias caused by loss to follow-up etc.
- Calculations of delta, the expected spillover bias.

Value

A list of class 'CRTsp' object comprising the input data, cluster and arm assignments, trial description and results of power calculations

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Examples

```
{# Power calculations for a binary outcome without input geolocations
examplePower1 <- CRTpower(locations = 3000, ICC = 0.10, effect = 0.4, alpha = 0.05,
    outcome_type = 'd', desiredPower = 0.8, yC=0.35, c = 20, sd_h = 5)
summarv(examplePower1)
# Power calculations for a rate outcome without input geolocations
examplePower2 <- CRTpower(locations = 2000, cv_percent = 40, effect = 0.4, denominator = 2.5,
    alpha = 0.05, outcome_type = 'e', desiredPower = 0.8, yC = 0.35, c = 20, sd_h=5)
summary(examplePower2)
# Example with input geolocations
examplePower3 <- CRTpower(trial = readdata('example_site.csv'), desiredPower = 0.8,
    effect=0.4, yC=0.35, outcome_type = 'd', ICC = 0.05, c = 20)
summary(examplePower3)
# Example with input geolocations, randomisation, and spillover
example4 <- randomizeCRT(specify_clusters(trial = readdata('example_site.csv'), c = 20))</pre>
examplePower4 <- CRTpower(trial = example4, desiredPower = 0.8,
    effect=0.4, yC=0.35, outcome_type = 'd', ICC = 0.05, contaminate_pop_pr = 0.3)
summary(examplePower4)
```

CRTsp

Create or update a "CRTsp" object

Description

CRTsp coerces data frames containing co-ordinates and location attributes into objects of class "CRTsp" or creates a new "CRTsp" object by simulating a set of Cartesian co-ordinates for use as the locations in a simulated trial site

Usage

```
CRTsp(
  x = NULL,
  design = NULL,
  geoscale = NULL,
  locations = NULL,
  kappa = NULL,
  mu = NULL,
  geometry = "point"
)
```

Arguments

Χ

an object of class "CRTsp" or a data frame containing locations in (x,y) coordinates, cluster assignments (factor cluster), and arm assignments (factor arm). Optionally specification of a buffer zone (logical buffer); any other variables required for subsequent analysis.

design

list: an optional list containing the requirements for the power of the trial

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geoscale numeric: standard deviation of random displacement from each settlement clus-

ter center (for new objects)

locations integer: number of locations in population (for new objects)

kappa numeric: intensity of Poisson process of settlement cluster centers (for new ob-

jects)

mu numeric: mean number of points per settlement cluster (for new objects)

geometry with valid values 'point' (the default, corresponding to point locations), 'triangle',

'square' and 'hexagon' corresponding to grids constructed from pixels of reg-

ular polygons.

Details

If a data frame or "CRTsp" object is input then the output "CRTsp" object is validated, a description of the geography is computed and power calculations are carried out.

If geoscale, locations, kappa and mu are specified then a new trial dataframe is constructed corresponding to a novel simulated human settlement pattern. This is generated using the Thomas algorithm (rThomas) in spatstat.random allowing the user to defined the density of locations and degree of spatial clustering. The resulting trial data frame comprises a set of Cartesian coordinates centred at the origin.

Value

A list of class "CRTsp" containing the following components:

design list: parameters required for power calculations geom_full list: summary statistics describing the site

geom_core list: summary statistics describing the core area (when a buffer is specified)

trial data frame: rows correspond to geolocated points, as follows:

x numeric vector: x-coordinates of locations y numeric vector: y-coordinates of locations cluster factor: assignments to cluster of each location

arm factor: assignments to "control" or "intervention" for each location nearestDiscord numeric vector: Euclidean distance to nearest discordant location (km)

buffer logical: indicator of whether the point is within the buffer

... other objects included in the input "CRTsp" object or data frame

```
{# Generate a simulated area with 10,000 locations
example_area = CRTsp(geoscale = 1, locations=10000, kappa=3, mu=40)
summary(example_area)
}
```

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CRTwrite

Export of GIS layer from 'CRTsp'

Description

CRTwrite exports a simple features object in a GIS format

Usage

```
CRTwrite(
  object,
  dsn,
  feature = "clusters",
  buffer_width,
  maskbuffer = 0.2,
  ...
)
```

Arguments

```
object of class 'CRTsp'
object
dsn
                  dataset name (relative path) for output objects
feature
                  feature to be exported, options are:
                'cluster'
                             cluster assignments
                'arms'
                              arm assignments
                'buffer'
                             buffer zone or spillover zone
                 'mask'
                              mask for areas that are distant from habitations
buffer_width
                  width of buffer between discordant locations (km)
maskbuffer
                  radius of buffer drawn around inhabited areas (km)
                  other arguments passed to 'sf::write_sf'
. . .
```

Details

'sf::write_sf' is used to format the output. The function returns TRUE on success, FALSE on failure, invisibly.

If the input object contains a 'centroid' then this is used to compute lat long coordinates, which are assigned the "WGS84" coordinate reference system. Otherwise the objects have equirectangular co-ordinates with centroid (0,0).

If feature = 'buffer' then buffer width determination is as described under plotCRT().

The output vector objects are constructed by forming a Voronoi tessellation of polygons around

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each of the locations and combining these polygons. The polygons on the outside of the study area extend outwards to an external rectangle. The 'mask' is used to mask out the areas of these polygons that are at a distance > maskbuffer from the nearest location.

Value

```
obj, invisibly
```

Examples

```
tmpdir = tempdir()
dsn <- paste0(tmpdir,'/arms')
CRTwrite(readdata('exampleCRT.txt'), dsn = dsn, feature = 'arms',
driver = 'ESRI Shapefile', maskbuffer = 0.2)</pre>
```

fitted.CRTanalysis

Extract model fitted values

Description

fitted.CRTanalysis method for extracting model fitted values

Usage

```
## S3 method for class 'CRTanalysis'
fitted(object, ...)
```

Arguments

```
object CRTanalysis object
... other arguments
```

Value

the fitted values returned by the statistical model run within the CRTanalysis function

```
{example <- readdata('exampleCRT.txt')
exampleGEE <- CRTanalysis(example, method = "GEE")
fitted_values <- fitted(exampleGEE)
}</pre>
```

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Convert lat long co-ordinates to x,y	latlong_as_xy
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Description

latlong_as_xy converts co-ordinates expressed as decimal degrees into x,y

Usage

```
latlong_as_xy(trial, latvar = "lat", longvar = "long")
```

Arguments

trial	A trial dataframe or list of o	class "CRTsp" o	containing latitudes	and longitudes in
-------	--------------------------------	-----------------	----------------------	-------------------

decimal degrees

latvar name of column containing latitudes in decimal degreeslongvar name of column containing longitudes in decimal degrees

Details

The output object contains the input locations replaced with Cartesian coordinates in units of km, centred on (0,0), corresponding to using the equirectangular projection (valid for small areas). Other data are unchanged.

Value

A list of class "CRTsp" containing the following components:

```
examplexy <- latlong_as_xy(readdata("example_latlong.csv"))</pre>
```

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plotCRT

Graphical displays of the geography of a CRT

Description

plotCRT returns graphical displays of the geography of a CRT or of the results of statistical analyses of a CRT

Usage

```
plotCRT(
  object,
  map = FALSE,
  distance = "nearestDiscord",
  fill = "arms",
  showLocations = FALSE,
  showClusterBoundaries = TRUE,
  showClusterLabels = FALSE,
  showBuffer = FALSE,
  cpalette = NULL,
  buffer_width = NULL,
  maskbuffer = 0.2,
  labelsize = 4,
  legend.position = NULL
)
```

Arguments

fill

```
object of class 'CRTanalysis' produced by CRTanalysis()
map logical: indicator of whether a map is required
```

distance measure of distance or surround with options:

```
"nearestDiscord" distance to nearest discordant location (km)
"disc" disc
"hdep" Tukey's half space depth
"sdep" simplicial depth
```

fill layer of map with options:

```
'cluster' cluster assignment
'arms' arm assignment
'nearestDiscord' distance to the nearest discordant location
'disc' disc measure of surround
'hdep' Tukey's half space depth
```

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'sdep' simplicial depth

'prediction' model prediction of the outcome

'none' No fill

showLocations logical: determining whether locations are shown

showClusterBoundaries

logical: determining whether cluster boundaries are shown

showClusterLabels

logical: determining whether the cluster numbers are shown

showBuffer logical: whether a buffer zone should be overlayed

cpalette colour palette (to use different colours for clusters this must be at least as long

as the number of clusters.

buffer_width width of buffer zone to be overlayed (km)
maskbuffer radius of buffer around inhabited areas (km)

labelsize size of cluster number labels

legend.position

(using ggplot2::themes syntax)

Details

If map = FALSE and the input is a trial data frame or a CRTsp object, containing a randomisation to arms, a stacked bar chart of the outcome grouped by the specified distance is produced. If the specified distance has not yet been calculated an error is returned.

If map = FALSE and the input is a CRTanalysis object a plot of the estimated spillover function is generated. The fitted spillover function is plotted as a continuous blue line against the measure the surround or of the distance to the nearest discordant location. Using the same axes, data summaries are plotted for ten categories of distance from the boundary. Both the average of the outcome and confidence intervals are plotted.

- For analyses with logit link function the outcome is plotted as a proportion.
- For analyses with log or cloglog link function the data are plotted on a scale of the Williams mean (mean of exp(log(x + 1))) 1) rescaled so that the median matches the fitted curve at the midpoint.

If map = TRUE a thematic map corresponding to the value of fill is generated.

- fill = 'clusters' or leads to thematic map showing the locations of the clusters
- fill = 'arms' leads to a thematic map showing the geography of the randomization
- fill = 'distance' leads to a raster plot of the distance to the nearest discordant location.
- fill = 'prediction' leads to a raster plot of predictions from an 'INLA' model.

If showBuffer = TRUE the map is overlaid with a grey transparent layer showing which areas are within a defined distance of the boundary between the arms. Possibilities are:

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- If the trial has not been randomised or if showBuffer = FALSE no buffer is displayed
- If buffer_width takes a positive value then buffers of this width are displayed irrespective of any pre-specified or spillover limits.
- If the input is a 'CRTanalysis' and spillover limits have been estimated by an 'LME4' or 'INLA' model then these limits are used to define the displayed buffer.
- If buffer_width is not specified and no spillover limits are available, then any pre-specified buffer (e.g. one generated by specify_buffer()) is displayed.

A message is output indicating which of these possibilities applies.

Value

graphics object produced by the ggplot2 package

Examples

```
{example <- readdata('exampleCRT.txt')</pre>
#Plot of data by distance
plotCRT(example)
#Map of locations only
plotCRT(example, map = TRUE, fill = 'none', showLocations = TRUE,
           showClusterBoundaries=FALSE, maskbuffer=0.2)
#show cluster boundaries and number clusters
plotCRT(example, map = TRUE, fill ='none', showClusterBoundaries=TRUE,
           showClusterLabels=TRUE, maskbuffer=0.2, labelsize = 2)
#show clusters in colour
plotCRT(example, map = TRUE, fill = 'clusters', showClusterLabels = TRUE,
          labelsize=2, maskbuffer=0.2)
#show arms
plotCRT(example, map = TRUE,
fill = 'arms', maskbuffer=0.2, legend.position=c(0.8,0.8))
#spillover plot
analysis <- CRTanalysis(example)</pre>
plotCRT(analysis, map = FALSE)
}
```

Description

predict. CRT analysis method for extracting model predictions

Usage

```
## S3 method for class 'CRTanalysis'
predict(object, ...)
```

randomizeCRT 21

Arguments

```
CRTanalysis object
object
                  other arguments
. . .
```

Value

the model predictions returned by the statistical model run within the CRTanalysis function

Examples

```
{example <- readdata('exampleCRT.txt')</pre>
exampleGEE <- CRTanalysis(example, method = "GEE")</pre>
predictions <- predict(exampleGEE)</pre>
}#'
```

randomizeCRT

Randomize a two-armed cluster trial

Description

randomizeCRT carries out randomization of clusters and augments the trial data frame with assignments to arms

Usage

```
randomizeCRT(
  trial,
 matchedPair = FALSE,
 baselineNumerator = "base_num",
 baselineDenominator = "base_denom"
)
```

Arguments

trial an object of class "CRTsp" or a data frame containing locations in (x,y) coordi-

> nates, cluster assignments (factor cluster), and arm assignments (factor arm). Optionally: specification of a buffer zone (logical buffer); any other variables

required for subsequent analysis.

logical: indicator of whether pair-matching on the baseline data should be used matchedPair

in randomization

baselineNumerator

name of numerator variable for baseline data (required for matched-pair randomization)

baselineDenominator

name of denominator variable for baseline data (required for matched-pair randomization)

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Value

A list of class "CRTsp" containing the following components:

parameters required for power calculations design list: geom_full list: summary statistics describing the site geom_core list: summary statistics describing the core area (when a buffer is specified) rows correspond to geolocated points, as follows: trial data frame: numeric vector: x-coordinates of locations numeric vector: y-coordinates of locations cluster factor: assignments to cluster of each location factor: assigned matched pair of each location (for matchedPair randomisations) pair arm factor: assignments to "control" or "intervention" for each location other objects included in the input "CRTsp" object or data frame . . .

Examples

```
# Randomize the clusters in an example trial
exampleCRT <- randomizeCRT(trial = readdata('exampleCRT.txt'), matchedPair = TRUE)</pre>
```

readdata

Read example dataset

Description

readdata reads a file from the package library of example datasets

Usage

```
readdata(filename)
```

Arguments

filename

name of text file stored within the package

Details

The input file name should include the extension (either .csv or .txt). The resulting object is a data frame if the extension is .csv.

Value

R object corresponding to the text file

```
exampleCRT <- readdata('exampleCRT.txt')</pre>
```

residuals.CRTanalysis

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residuals.CRTanalysis Extract model residuals

Description

residuals. CRT analysis method for extracting model residuals

Usage

```
## S3 method for class 'CRTanalysis'
residuals(object, ...)
```

Arguments

```
object CRTanalysis object
... other arguments
```

Value

the residuals from the statistical model run within the CRTanalysis function

Examples

```
{example <- readdata('exampleCRT.txt')
exampleGEE <- CRTanalysis(example, method = "GEE")
residuals <- residuals(exampleGEE)
}</pre>
```

simulateCRT

Simulation of cluster randomized trial with spillover

Description

simulateCRT generates simulated data for a cluster randomized trial (CRT) with geographic spillover between arms.

Usage

```
simulateCRT(
   trial = NULL,
   effect = 0,
   outcome0 = NULL,
   generateBaseline = TRUE,
   matchedPair = TRUE,
   scale = "proportion",
   baselineNumerator = "base_num",
```

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```
baselineDenominator = "base_denom",
denominator = NULL,
ICC_inp = NULL,
kernels = 200,
sigma_m = NULL,
spillover_interval = NULL,
tol = 0.005
)
```

Arguments

trial an object of class "CRTsp" or a data frame containing locations in (x,y) coordi-

nates, cluster assignments (factor cluster), and arm assignments (factor arm).

Each location may also be assigned a propensity (see details).

effect numeric. The simulated effect size (defaults to 0)

outcome0 numeric. The anticipated value of the outcome in the absence of intervention

generateBaseline

logical. If TRUE then baseline data and the propensity will be simulated

matchedPair logical. If TRUE then the function tries to carry out randomization using pair-

matching on the baseline data (see details)

scale measurement scale of the outcome. Options are: 'proportion' (the default);

'count'; 'continuous'.

baselineNumerator

optional name of numerator variable for pre-existing baseline data

baselineDenominator

optional name of denominator variable for pre-existing baseline data

denominator optional name of denominator variable for the outcome

ICC_inp numeric. Target intra cluster correlation, provided as input when baseline data

are to be simulated

kernels number of kernels used to generate a de novo propensity

sigma_m numeric. standard deviation of the normal kernel measuring spatial smoothing

leading to spillover

spillover_interval

numeric. input spillover interval

tol numeric. tolerance of output ICC

Details

Synthetic data are generated by sampling around the values of variable propensity, which is a numerical vector (taking positive values) of length equal to the number of locations. There are three ways in which propensity can arise:

- 1. propensity can be provided as part of the input trial object.
- 2. Baseline numerators and denominators (values of baselineNumerator and baselineDenominator may be provided. propensity is then generated as the numerator:denominator ratio for each location in the input object

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3. Otherwise propensity is generated using a 2D Normal kernel density. The OOR::StoSOO is used to achieve an intra-cluster correlation coefficient (ICC) that approximates the value of 'ICC_inp' by searching for an appropriate value of the kernel bandwidth.

num[i], the synthetic outcome for location i is simulated with expectation:

```
E(num[i]) = outcome0[i]*propensity[i]*denom[i]*(1-effect*I[i])/mean(outcome0[]*propensity[])
```

The sampling distribution of num[i] depends on the value of scale as follows:

- scale='continuous': Values of num are sampled from a Normal distributions with means E(num[i]) and variance determined by the fitting to ICC_inp.
- scale='count': Simulated events are allocated to locations via multivariate hypergeometric distributions parameterised with E(num[i]).
- scale='proportion': Simulated events are allocated to locations via multinomial distributions parameterised with E(num[i]).

denominator may specify a vector of numeric (non-zero) values in the input "CRTsp" or data. frame which is returned as variable denom. It acts as a scale-factor for continuous outcomes, rate-multiplier for counts, or denominator for proportions. For discrete data all values of denom must be > 0.5 and are rounded to the nearest integer in calculations of num.

By default, denom is generated as a vector of ones, leading to simulation of dichotomous outcomes if scale='proportion'.

If baseline numerators and denominators are provided then the output vectors base_denom and base_num are set to the input values. If baseline numerators and denominators are not provided then the synthetic baseline data are generated by sampling around propensity in the same way as the outcome data, but with the effect size set to zero.

If matchedPair is TRUE then pair-matching on the baseline data will be used in randomization providing there are an even number of clusters. If there are an odd number of clusters then matched pairs are not generated and an unmatched randomization is output.

Either sigma_m or spillover_interval must be provided. If both are provided then the value of sigma_m is overwritten by the standard deviation implicit in the value of spillover_interval. Spillover is simulated as arising from a diffusion-like process.

For further details see Multerer (2021)

Value

A list of class "CRTsp" containing the following components:

geom_full list: summary statistics describing the site cluster assignments, and randomization design list: values of input parameters to the design

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trial data frame: rows correspond to geolocated points, as follows:

x numeric vector: x-coordinates of locations
y numeric vector: y-coordinates of locations
cluster factor: assignments to cluster of each location
arm factor: assignments to control or intervention for each location
nearestDiscord numeric vector: signed Euclidean distance to nearest discordant location (km)

... other objects included in the input "CRTsp" object or data. frame

Examples

```
{smalltrial <- readdata('smalltrial.csv')
simulation <- simulateCRT(smalltrial,
  effect = 0.25,
  ICC_inp = 0.05,
  outcome0 = 0.5,
  matchedPair = FALSE,
  scale = 'proportion',
  sigma_m = 0.6,
  tol = 0.05)
summary(simulation)
}</pre>
```

specify_buffer

Specification of buffer zone in a cluster randomized trial

Description

specify_buffer specifies a buffer zone in a cluster randomized trial (CRT) by flagging those locations that are within a defined distance of those in the opposite arm.

Usage

```
specify_buffer(trial, buffer_width = 0)
```

Arguments

trial an object of class "CRTsp" or a data frame containing locations in (x,y) coordi-

nates, cluster assignments (factor cluster), and arm assignments (factor arm).

buffer_width minimum distance between locations in opposing arms for them to qualify to be

included in the core area (km)

specify_clusters 27

Value

A list of class "CRTsp" containing the following components:

```
geom_full
              list:
                                   summary statistics describing the site, cluster assignments, and randomization.
geom_core
              list:
                                   summary statistics describing the core area
trial
              data frame:
                                   rows correspond to geolocated points, as follows:
                                   numeric vector: x-coordinates of locations
                                   numeric vector: y-coordinates of locations
              cluster
                                   factor: assignments to cluster of each location
                                   factor: assignments to "control" or "intervention" for each location
              arm
                                  numeric vector: signed Euclidean distance to nearest discordant location (km)
              nearestDiscord
              buffer
                                   logical: indicator of whether the point is within the buffer
                                   other objects included in the input "CRTsp" object or data frame
               . . .
```

Examples

```
#Specify a buffer of 200m
exampletrial <- specify_buffer(trial = readdata('exampleCRT.txt'), buffer_width = 0.2)

specify_clusters

Assign locations to clusters in a CRT</pre>
```

Description

specify_clusters algorithmically assigns locations to clusters by grouping them geographically

Usage

```
specify_clusters(
  trial = trial,
  c = NULL,
  h = NULL,
  algorithm = "NN",
  reuseTSP = FALSE,
  auxiliary = NULL
)
```

Arguments

trial A CRT object or data frame containing (x,y) coordinates of households
c integer: number of clusters in each arm
h integer: number of locations per cluster
algorithm algorithm for cluster boundaries, with options:

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NN	Nearest neighbour: assigns equal numbers of locations to each cluster
kmeans	kmeans clustering: aims to partition locations so that each belongs to the cluster with the nearest centroid.
TSP	travelling salesman problem heuristic: Assigns locations sequentially along a travelling salesman path.

reuseTSP logical: indicator of whether a pre-existing path should be used by the TSP

algorithm

auxiliary "CRTsp" object containing external cluster and or arm assignments.

Details

Either c or h must be specified. If both are specified the input value of c is ignored.

The reuseTSP parameter is used to allow the path to be reused for creating alternative allocations with different cluster sizes.

If an auxiliary auxiliary "CRTsp" object is specified then the other options are ignored and the cluster assignments (and arm assignments if available) are taken from the auxiliary object. The trial data frame is augmented with a column "nearestPixel" containing the distance to boundary of the nearest grid pixel in the auxiliary. If the auxiliary is a grid with design\$geometry set to 'triangle', 'square' or 'hexagon' then the distance is computed to the edge of the nearest grid pixel in the discordant arm (using a circular approximation for the perimeter) rather than to the point location itself. If the point is within the pixel then the distance is given a negative sign.

Value

A list of class "CRTsp" containing the following components:

```
geom_full list: summary statistics describing the site, and cluster assignments.

trial data frame: rows correspond to geolocated points, as follows:

x numeric vector: x-coordinates of locations
y numeric vector: y-coordinates of locations
cluster factor: assignments to cluster of each location
... other objects included in the input "CRTsp" object or data frame
```

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summary.CRTanalysis

Summary of the results of a statistical analysis of a CRT

Description

summary. CRTanalysis generates a summary of a CRTanalysis including the main results

Usage

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

Arguments

```
object an object of class "CRTanalysis"
... other arguments used by summary
```

Value

No return value, writes text to the console.

Examples

```
{example <- readdata('exampleCRT.txt')
exampleT <- CRTanalysis(example, method = "T")
summary(exampleT)
}</pre>
```

summary.CRTsp

Summary description of a "CRTsp" object

Description

```
summary. CRTsp provides a description of a "CRTsp" object
```

Usage

```
## S3 method for class 'CRTsp'
summary(object, maskbuffer = 0.2, ...)
```

Arguments

object an object of class "CRTsp" or a data frame containing locations in (x,y) coordi-

nates, cluster assignments (factor cluster), arm assignments (factor arm) and buffer zones (logical buffer), together with any other variables required for

subsequent analysis.

maskbuffer radius of area around a location to include in calculation of areas

. . . other arguments used by summary

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Value

No return value, write text to the console.

```
summary(CRTsp(readdata('exampleCRT.txt')))
```

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