Package 'CRTspat'

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Title Workflow for Cluster Randomised Trials with Spillover

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Description Design, workflow and statistical analysis of Cluster Randomised Trials of (health) interventions where there may be spillover between the arms (see <<u>https://thomasasmith.github.io/index.html</u>>).

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aggregateCRT

Aggregate data across records with duplicated locations

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

trial	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"

anonymize_site

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".

Examples

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

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
)
```

Arguments

trial	5	f class "CRTsp" or a data frame containing locations in (x,y) coordier assignments (factor cluster), and arm assignments (factor arm).
distance	the quantity	v(s) to be computed. Options are:
"nearestD "distance "disc" "kern" "hdep" "sdep"	iscord" Assigned"	distance to nearest discordant location (km) distance to the nearest pixel in the assigned cluster (km) disc kernel-based measure Tukey half space depth simplicial depth
scale_par	-	neter equal to the disc radius in km if distance = "disc" or to the viance of the kernels if distance = "kern"

auxiliarv	"CRTsp" o	biect containing external	l 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 halfdepth 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

(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

trial	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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CRTanalysis

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)
library(sp)
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",
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	nates	ect of class "CRTsp" or a data frame containing locations in (x,y) coordi- cluster assignments (factor cluster), and arm assignments (factor arm) tcome data (see details).	
metho	d statis	cal method with options:	
	"EMP" "T" "GEE" "LME4" "INLA" "MCMC" "WCA"	simple averages of the data comparison of cluster means by t-test Generalised Estimating Equations Generalized Linear Mixed-Effects Models Integrated Nested Laplace Approximation (INLA) Markov chain Monte Carlo using "JAGS" Within cluster analysis	
dista	nce Meas	re of distance or surround with options:	
	"nearest["disc" "kern" "hdep" "sdep"	scord" distance to nearest discordant location (km) disc surround based on sum of normal kernels Tukey half space depth simplicial depth	
scale	•	c: pre-specified value of the spillover parameter or disc radius for models this is fixed (cfunc = "R").	
cfunc	trans	rmation defining the spillover function with options:	
"Z" "X" "L"	arm effects not co spillover not mod inverse logistic (s	lled the only valid value of cfunc for methods "EMP", "T" and "GEE	"

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CRTanalysis

"P" "S" "E" "R"	piecewise	of scale factor only available with distance = "disc" or distance = "kern"						
link		link function with options:						
"logit" "log" "cloglo "identi	nume og" nume	default). numerator has a binomial distribution with denominator denominator. erator is Poisson distributed with an offset of log(denominator). erator is Bernoulli distributed with an offset of log(denominator). outcome is numerator/denominator with a normally distributed error function.						
numer	ator	string: name of numerator variable for outcome						
denom	inator	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						
basel	ineOnly	logical: indicator of whether required analysis is of effect size or of baseline only						
basel	ineNumera	tor						
		string: name of numerator variable for baseline data (if present)						
basel	ineDenomi							
	10 (string: name of denominator variable for baseline data (if present)						
perso	nalProtect	tion logical: indicator of whether the model includes local effects with no spillover						
aluat	arEffacta	logical: indicator of whether the model includes cluster random effects						
		-						
spati	aleffects	logical: indicator of whether the model includes spatial random effects (avail- able only for method = "INLA")						
requi	reMesh	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: gee, 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..

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.

CRTpower

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

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

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), Int. J. Epi., 28(2) pp319–326 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

CRTsp

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,</pre>
    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,</pre>
    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

х	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.
design	list: an optional list containing the requirements for the power of the trial

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 objects)
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:

ecified)
n location
on (km)
e
ł

Examples

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

CRTwrite

Description

CRTwrite exports a simple features object in a GIS format

Usage

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

Arguments

object	object of	class 'CRTsp'
dsn	dataset na	me (relative path) for output objects
feature	feature to be exported, options are:	
	'cluster' 'arms' 'buffer' 'mask'	cluster assignments arm assignments buffer zone or spillover zone 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

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

Examples

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

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latlong_as_xy

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 class "CRTsp" containing latitudes and longitudes in decimal degrees
latvar	name of column containing latitudes in decimal degrees
longvar	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:

geom_full	list:	summary statistics describing the site
trial	data frame:	rows correspond to geolocated points, as follows:
	х	numeric vector: x-coordinates of locations
	У	numeric vector: y-coordinates of locations
		other objects included in the input "CRTsp" object or data frame

Examples

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

plotCRT

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

object	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

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

'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	
showClusterLab	els	
	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 legend.positior	size of cluster number labels	
	(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:

- 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)
}
```

predict.CRTanalysis Model predictions

Description

predict.CRTanalysis method for extracting model predictions

Usage

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

randomizeCRT

Arguments

object	CRTanalysis object
	other arguments

Value

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

Examples

```
{example <- readdata('exampleCRT.txt')
exampleGEE <- CRTanalysis(example, method = "GEE")
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.	
matchedPair	logical: indicator of whether pair-matching on the baseline data should be used in randomization	
baselineNumerator		
	name of numerator variable for baseline data (required for matched-pair ran- domization)	
baselineDenominator		
	name of denominator variable for baseline data (required for matched-pair ran- domization)	

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:
	х	numeric vector: x-coordinates of locations
	У	numeric vector: y-coordinates of locations
	cluster	factor: assignments to cluster of each location
	pair	factor: assigned matched pair of each location (for matchedPair randomisations)
	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

Examples

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

residuals.CRTanalysis Extract model residuals

Description

residuals.CRTanalysis 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",
```

```
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	
generateBaselin	ne	
	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'.	
baselineNumerat	or	
	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

trial	data frame:	rows correspond to geolocated points, as follows:
	х	numeric vector: x-coordinates of locations
	У	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)
	propensity	numeric vector: propensity for each location
	base_denom	numeric vector: denominator for baseline
	base_num	numeric vector: numerator for baseline
	denom	numeric vector: denominator for the outcome
	num	numeric vector: numerator for the outcome
		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) coordinates, 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)

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specify_clusters

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:
	Х	numeric vector: x-coordinates of locations
	У	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)
	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)</pre>
```

specify_clusters Assign locations to clusters in a CRT

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
С	integer: number of clusters in each arm
h	integer: number of locations per cluster
algorithm	algorithm for cluster boundaries, with options:

specify_clusters

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:
	х	numeric vector: x-coordinates of locations
	У	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

Examples

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) coordinates, 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	

Value

No return value, write text to the console.

Examples

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

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