Package 'spaSim'

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Title Spatial point data simulator for tissue images

```
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Description A suite of functions for simulating spatial patterns of cells in
      tissue images. Output images are multitype point data in SingleCellExperiment
      format. Each point represents a cell, with its 2D locations and cell type.
      Potential cell patterns include background cells, tumour/immune cell clusters,
      immune rings, and blood/lymphatic vessels.
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```

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Description

A data frame containing 4915 background cells on 2000 by 2000 size window

Usage

bg1

Format

A data frame with 4915 rows (cells) and 3 columns:

Cell.X.Position X coordinate **Cell.Y.Position** Y coordinate

Cell.Type cell type

Description

Format a data frame object into a SpatialExperiment class object where the count assay is empty.

Usage

 $format_spe(df)$

Arguments

df

A data frame where each row contains information about a cell. The columns of the data frame will become the colData of the SpatialExperiment object.

Value

An SPE object

```
multiple_background_images
```

Simulate multiple background images (mixed cell types)

Description

Generate a set of background images with different proportions of mixed cell types all at once. The default values for the arguments give an example of multiple image simulation which enable an automatic multiple image simulation without the specification of any argument.

Usage

```
multiple_background_images(
  bg_sample = bg1,
  idents = c("Tumour", "Immune", "Others"),
  props = list(rep(0.1, 9), seq(0, 0.4, 0.05), seq(0.9, 0.5, -0.05)),
  plot_image = TRUE,
  plot_colours = NULL
)
```

Arguments

bg_sample A data frame or SpatialExperiment class object with locations of points rep-

resenting background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By

default use the internal bg1 background image.

idents String Vector. Names of the cell types to generate.

props List. Each element is a vector of proportions of the corresponding cell type. The

length of the vector is how many images to generate. All vectors should be of

the same length, also equal to the number of images.

plot_image Boolean. Whether plot the simulated images or not. Default is TRUE.

plot_colours String Vector. If plot_image is TRUE, this param is the corresponding colours

for the idents arg. Default is NULL, the predefined colour vector would be

used for plotting.

Value

A list of SPE objects

See Also

multiple_images_with_clusters for simulating multiple images with clusters, and multiple_images_with_immune for simulating multiple images with immune rings.

Other simulate multiple images functions: multiple_images_with_clusters(), multiple_images_with_immune_ri

Examples

```
idents <- c("Tumour","Immune","Others")
prop1 <- rep(0.1,9)
prop2 <- seq(0, 0.4, 0.05)
prop3 <- seq(0.9,0.5,-0.05)
set.seed(610)
bg_image_list <- multiple_background_images(bg_sample = bg1,
idents = idents, props = list(prop1, prop2, prop3), plot_image = FALSE)</pre>
```

multiple_images_with_clusters

Simulate multiple images with clusters

Description

Generate a set of images with different cluster properties. Trying out the default parameters is recommended for understanding this function. The default values for the arguments give an example of multiple image simulation which enable an automatic multiple image simulation without the specification of any argument.

Usage

```
multiple_images_with_clusters(
   bg_sample = bg1,
   cluster_shape = 2,
   prop_infiltration = 0.1,
   cluster_size = seq(200, 1000, 100),
   cluster_loc_x = 0,
   cluster_loc_y = 0,
   plot_image = TRUE,
   plot_categories = NULL,
   plot_colours = NULL
)
```

Arguments

bg_sample

A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By default use the internal bg1 background image.

cluster_shape

Number. Choose from one of the following pre-designed shapes (1 or 2 for tumour cluster or 3 for immune cluster). The pre-designed shape contains information of the cell names of the cluster, the infiltration cell types, the proportions of infiltration, the cluster size, and the cluster centre locations. In order

to simulate a set of images, use the arguments below to specify the ranges of the properties. The predefined cell types can not be changed, while users can change them manually after the simulation.

prop_infiltration

Numeric Vector. The degree of infiltration. If numeric, all simulated images have the same infiltration degree. If vector, images with a range of different infiltration proportions will be simulated.

cluster_size Numeric Vector. The size of the cluster. If numeric, all simulated images have the same cluster size. If vector, images with a range of different cluster sizes

will be simulated. The size should not exceed the limit of the image sides.

cluster_loc_x Numeric or Vector. The X location of the cluster center offset. If numeric, all simulated images have the same center X location. If vector, images with a

range of different center locations will be simulated.

cluster_loc_y Numeric or Vector of the same length of cluster_loc_x. The Y location of the

cluster center offset.

plot_image Boolean Whether plot the simulated images or not.Default is TRUE.

plot_categories

String Vector specifying the order of the cell categories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for

plotting.

 $\verb|plot_colours| String Vector specifying the order of the colours that correspond to the \verb|plot_categories| \\$

arg. Default is NULL - the predefined colour vector would be used for plotting.

Value

A list of SPE objects

See Also

multiple_background_images for simulating multiple mixed background images, and multiple_images_with_immur for simulating multiple images with immune rings.

Other simulate multiple images functions: multiple_background_images(), multiple_images_with_immune_rings

Examples

```
set.seed(610)
cluster_image_list <- multiple_images_with_clusters(bg_sample=bg1,
cluster_shape=2, prop_infiltration=0.1, cluster_size = seq(200,1000,100),
cluster_loc_x = 0, cluster_loc_y = 0, plot_image = TRUE)</pre>
```

```
multiple_images_with_immune_rings
```

Simulate multiple images with immune rings

Description

Generate a set of images with different immune ring properties. The default values for the arguments give an example of multiple image simulation which enable an automatic multiple image simulation without the specification of any argument.

Usage

```
multiple_images_with_immune_rings(
  bg_sample = bg1,
  cluster_size = 200,
  ring_shape = 1,
  prop_infiltration = 0,
  ring_width = seq(50, 100, 10),
  cluster_loc_x = 0,
  cluster_loc_y = 0,
  prop_ring_infiltration = seq(0, 0.2, 0.05),
  plot_image = TRUE,
  plot_categories = NULL,
  plot_colours = NULL
)
```

Arguments

bg_sample

A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By default use the internal bg1 background image.

cluster_size

Numeric Vector. The size of the cluster. If numeric, all simulated images have the same cluster size. If vector, images with a range of different cluster sizes will be simulated. The size should not exceed the limit of the image sides.

ring_shape

Number. Choose from one of the following pre-designed shapes (1,2 or 3). The pre-designed shape contains information of the cell names of the cluster, the infiltration cell types, the proportions of infiltration, the cluster size, the ring width, the proportions of infiltrated cells into immune rings and the cluster centre locations. In order to simulate a set of images, use the arguments below to specify the ranges of the properties. The predefined cell types can not be changed, while users can change them manually after the simulation.

prop_infiltration

Numeric Vector. The degree of infiltration. If numeric, all simulated images have the same infiltration degree. If vector, images with a range of different infiltration proportions will be simulated.

ring_width

Numeric Vector. The width of the immune ring. If numeric, all simulated images have the same ring width. If vector, images with a range of different ring widths will be simulated.

cluster_loc_x

Numeric or Vector. The X location of the cluster center offset. If numeric, all simulated images have the same center X location. If vector, images with a range of different center locations will be simulated.

cluster_loc_y

Numeric or Vector of the same length of cluster_loc_x. The Y location of the cluster center offset.

prop_ring_infiltration

Numeric or Vector. The degree of tumour infiltration in the region of immune rings.

plot_image

Boolean Whether plot the simulated images or not. Default is TRUE.

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```
plot_categories
```

String Vector specifying the order of the cell cateories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for plotting.

plot_colours

String Vector specifying the order of the colours that correspond to the plot_categories arg. Default is NULL - the predefined colour vector would be used for plotting.

Value

A list of spe objects

See Also

multiple_background_images for simulating multiple mixed background images, and multiple_images_with_clust for simulating multiple images with clusters.

Other simulate multiple images functions: multiple_background_images(), multiple_images_with_clusters()

Examples

```
set.seed(610)
ring_image_list <- multiple_images_with_immune_rings(bg_sample = bg1,
ring_shape = 1, prop_infiltration = 0, ring_width = seq(50,100,10),
cluster_size = 200, cluster_loc_x = 0, cluster_loc_y = 0,
prop_ring_infiltration = seq(0, 0.2,0.05), plot_image = TRUE)</pre>
```

plot_cells

plot_cells

Description

Produces a scatter plot of the cells in the tissue. Cells are coloured categorically by Cell.Type column. Cell categories not specified will be coloured "lightgrey" and labled "Unspecified".

Usage

```
plot_cells(
   spe_object,
   categories_of_interest = NULL,
   colour_vector = NULL,
   feature_colname = "Cell.Type"
)
```

Arguments

feature_colname

String specifying the column the cell categories belong to.

Value

A plot is returned

Description

Simulate cell locations. The 2D locations of the cells are simulated and plotted in a rectangular window. Users can specify the window size, cell number and the minimum distance between two cells. All cells have the same cell type, specified by the "Cell.Type" param.

Usage

```
simulate_background_cells(
    n_cells,
    width,
    height,
    method = "Hardcore",
    min_d,
    oversampling_rate = 1.2,
    jitter = 0.3,
    Cell.Type = "Others",
    plot_image = TRUE
)
```

Arguments

n_cerrs Numeric. Number of cens to simulate in the backgroun	n_cells	Numeric. Number of cells to simulate i	n the backgroun
--	---------	--	-----------------

width, height Numeric. The width and height of the image.

method String. The distribution model for the background cells. Options are "Hardcore"

for tumour tissues and "Even" for normal tissues. Default is "Hardcore".

min_d (OPTIONAL) Numeric. Use when method is "Hardcore". The minimum dis-

tance between two cells.

oversampling_rate

(OPTIONAL) Numeric. Use when method is "Hardcore". The multiplier for oversampling. Without oversampling, the simulation deletes cells that are within min_d from each other, resulting in a less total number of cells than n_cells. Default is 1.2 (this should be set based on n_cells and min_d; should always

be larger than 1).

jitter (OPTIONAL) Numeric. Use when method is "Even". The uniform distribution

parameter to generate the jitter distance for each cell from the vertices of the

hexagon.

Cell. Type (OPTIONAL) String. The name of the background cell type. Default is "Others"

since there shouldn't be any identity of the background cells.

plot_image (OPTIONAL) Boolean. Default is TRUE.

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Details

There are two options for the background cell distribution model. 1) Hardcore model for tumour tissues. This model uses rHardcore spatstat.random. Since rHardcore deletes cells that are within the distance of min_d to another cell, resulting in fewer cell specified by users, we here introduce parameter oversampling_rate to generate more cells than specified. 2) Normal tissues use an evenly-spaced model where the cells are distributed approximately according to the vertices of a hexagon. The function accomplishes this by generating cells on a hexagonal grid and individually applying a bounded uniform random jitter. In our algorithm, jitter is the parameter to define the uniform distribution of the jitter of the cells from the hexagon vertices. If there is a reference image, jitter can be estimated by comparing the average minimum distance between cells of the simulated image and the reference image. If without a reference image, We suggest 0.3 as the default value of jitter as this gives a sensible outcome.

Value

A data frame of the simulated background image

See Also

```
simulate_mixing for mixed background simulation, simulate_clusters for cluster simulation, simulate_immune_rings/simulate_double_rings for immune ring simulation, and simulate_stripes for vessel simulation.
```

```
Other simulate pattern functions: simulate_clusters(), simulate_double_rings(), simulate_immune_rings(), simulate_mixing(), simulate_stripes()
```

Examples

simulate_clusters

Simulate clusters

Description

Based on an existing background image, simulate clusters of cells where the same type of cells aggregate. The default values for the arguments give an example of cluster simulation which enable an automatic simulation of clusters without the specification of any argument.

Usage

```
simulate_clusters(
  bg_sample = bg1,
  n_clusters = 2,
  bg_type = "Others",
  cluster_properties = list(C1 = list(name_of_cluster_cell = "Tumour", size = 300, shape
```

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```
= "Oval", centre_loc = data.frame(x = 500, y = 500), infiltration_types =
    c("Immune1", "Others"), infiltration_proportions = c(0.1, 0.05)), C2 =
    list(name_of_cluster_cell = "Immune1", size = 500, shape = "Irregular", centre_loc =
    data.frame(x = 1500, y = 500), infiltration_types = c("Immune2", "Others"),
    infiltration_proportions = c(0.1, 0.05))),
    plot_image = TRUE,
    plot_categories = NULL,
    plot_colours = NULL
)
```

Arguments

bg_sample (OPTIONAL) A data frame or SpatialExperiment class object with locations

of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions".

By default use the internal bg1 background image.

n_clusters Numeric. Number of clusters. This must match the length(cluster_properties).

bg_type (OPTIONAL) String. The name of the background cell type if the background

sample does not have a "Cell.Type" column. By default is "Others".

cluster_properties

List of properties of the clusters. See examples for the format of this arg.

plot_image Boolean. Whether the simulated image is plotted.

plot_categories

String Vector specifying the order of the cell categories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for

plotting.

plot_colours String Vector specifying the order of the colours that correspond to the plot_categories

arg. Default is NULL - the predefined colour vector would be used for plotting.

Value

A data.frame of the simulated image

See Also

simulate_background_cells for all cell simulation, simulate_mixing for mixed background simulation, simulate_immune_rings/simulate_double_rings for immune ring simulation, and simulate_stripes for vessel simulation.

Other simulate pattern functions: simulate_background_cells(), simulate_double_rings(), simulate_immune_rings(), simulate_mixing(), simulate_stripes()

Examples

```
set.seed(610)
cluster_image <- simulate_clusters(bg_sample = bg1,
n_clusters=2, cluster_properties=list(C1=list(name_of_cluster_cell="Tumour",
size=300, shape="0val", centre_loc=data.frame("x"=500, "y"=500),
infiltration_types=c("Immune1", "Others"), infiltration_proportions=c(0.1, 0.05)),
C2=list(name_of_cluster_cell="Immune1", size=500, shape="Irregular",
centre_loc=data.frame("x"=1500,"y"=500), infiltration_types=c("Immune2", "Others"),
infiltration_proportions=c(0.1, 0.05))))</pre>
```

simulate_double_rings Simulate double immune rings

Description

Based on an existing background image, simulate double rings of immune cells that surround tumour clusters. The inner ring is the internal margin of a tumour cluster. The outer ring is the external tumour margin. The tumour clusters and the double immune rings are simulated at the same time. The default values for the arguments give an example of double ring simulation which enable an automatic simulation of double rings without the specification of any argument.

Usage

```
simulate_double_rings(
  bg_sample = bg1,
 bg_type = "Others",
 n_dr = 2,
 dr_properties = list(D1 = list(name_of_cluster_cell = "Tumour", size = 300, shape =
    "Circle", centre_loc = data.frame(x = 1000, y = 1000), infiltration_types =
  c("Immune1", "Immune2", "Others"), infiltration_proportions = c(0.15, 0.05, 0.05),
    name_of_ring_cell = "Immune1", immune_ring_width = 150,
  immune_ring_infiltration_types = c("Others"), immune_ring_infiltration_proportions =
    c(0.15), name_of_double_ring_cell = "Immune2", double_ring_width = 100,
  double_ring_infiltration_types = c("Others"), double_ring_infiltration_proportions =
    c(0.15)),
    D2 = list(name_of_cluster_cell = "Tumour", size = 300, shape =
    "Oval", centre_loc = data.frame(x = 1200, y = 1200), infiltration_types =
  c("Immune1", "Immune2", "Others"), infiltration_proportions = c(0.15, 0.05, 0.05),
    name_of_ring_cell = "Immune1", immune_ring_width = 150,
  immune_ring_infiltration_types = c("Others"), immune_ring_infiltration_proportions =
    c(0.15), name_of_double_ring_cell = "Immune2", double_ring_width = 100,
  double_ring_infiltration_types = c("Others"), double_ring_infiltration_proportions =
    c(0.15)),
  plot_image = TRUE,
 plot_categories = NULL,
 plot_colours = NULL
```

Arguments

bg_sample	(OPTIONAL) A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By default use the internal bg1 background image.
bg_type	(OPTIONAL) String The name of the background cell type. By default is "Others".
n_dr	Number of double immune rings. This must match the length(dr_properties).
dr_properties	List of properties of the double immune rings. Please refer to the examples for the structure of dr_properties.

```
plot_image Boolean. Whether the simulated image is plotted.

plot_categories

String Vector specifying the order of the cell categories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for plotting.

plot_colours

String Vector specifying the order of the colours that correspond to the plot_categories arg. Default is NULL - the predefined colour vector would be used for plotting.
```

Value

A data.frame of the simulated image

See Also

simulate_background_cells for all cell simulation, simulate_mixing for mixed background simulation, simulate_clusters for cluster simulation, simulate_immune_rings for single immune ring simulation, and simulate_stripes for vessel simulation.

Other simulate pattern functions: simulate_background_cells(), simulate_clusters(), simulate_immune_rings simulate_mixing(), simulate_stripes()

Examples

```
# manually define the properties of the immune ring
dr_properties <- list(D1 = list(name_of_cluster_cell = "Tumour", size = 300,</pre>
shape = "Circle",centre_loc = data.frame("x" = 1000, "y" = 1000),infiltration_types
= c("Immune1", "Immune2", "Others"), infiltration_proportions = <math>c(0.15, 0.05, 0.05),
name_of_ring_cell = "Immune1",immune_ring_width = 150,immune_ring_infiltration_types
= c("Others"),immune_ring_infiltration_proportions = c(0.15),name_of_double_ring_cell
= "Immune2", double_ring_width = 100, double_ring_infiltration_types = c("Others"),
double_ring_infiltration_proportions = c(0.15)),
D2 = list(name_of_cluster_cell = "Tumour", size = 300, shape = "Oval", centre_loc
= data.frame("x" = 1200, "y" = 1200), infiltration_types = c("Immune1", "Immune2", "Others"),
infiltration\_proportions = c(0.15, 0.05, 0.05), name\_of\_ring\_cell = "Immune1", other infiltration\_proportions = c(0.15, 0.05, 0.05), name\_of\_ring\_cell = "Immune1", other infiltration\_proportions = c(0.15, 0.05, 0.05), name\_of\_ring\_cell = "Immune1", other infiltration\_proportions = c(0.15, 0.05, 0.05), name\_of\_ring\_cell = "Immune1", other infiltration\_proportions = c(0.15, 0.05), other infiltration\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_proportion\_propo
immune_ring_width = 150,immune_ring_infiltration_types = c("Others"),
immune\_ring\_infiltration\_proportions = c(0.15), name\_of\_double\_ring\_cell = "Immune2",
double_ring_width = 100,double_ring_infiltration_types = c("Others"),
double_ring_infiltration_proportions = c(0.15))
double_ring_image <- simulate_double_rings(bg_sample = bg1,</pre>
n_dr = 2, dr_properties = dr_properties)
```

```
simulate_immune_rings simulate_immune_rings
```

Description

Based on an existing background image, simulate rings of immune cells that surround tumour clusters. The tumour clusters and immune rings are simulated at the same time. The default values for the arguments give an example of immune ring simulation which enable an automatic simulation of immune rings without the specification of any argument.

Usage

```
simulate_immune_rings(
 bg_sample = bg1,
 bg_type = "Others",
 n_{ir} = 2,
 ir_properties = list(I1 = list(name_of_cluster_cell = "Tumour", size = 600, shape =
    "Circle", centre_loc = data.frame(x = 930, y = 1000), infiltration_types =
  c("Immune1", "Immune2", "Others"), infiltration_proportions = <math>c(0.15, 0.05, 0.05),
    name_of_ring_cell = "Immune1", immune_ring_width = 150,
  immune_ring_infiltration_types = c("Others"), immune_ring_infiltration_proportions =
  c(0.15)), I2 = list(name_of_cluster_cell = "Tumour", size = 500, shape = "Oval",
   centre_loc = data.frame(x = 1330, y = 1100), infiltration_types = c("Immune1",
    "Immune2", "Others"), infiltration_proportions = c(0.15, 0.05, 0.05),
    name_of_ring_cell = "Immune1", immune_ring_width = 150,
  immune_ring_infiltration_types = c("Others"), immune_ring_infiltration_proportions =
    c(0.15)),
  plot_image = TRUE,
 plot_categories = NULL,
 plot_colours = NULL
)
```

Arguments

bg_sample	(OPTIONAL) A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By default use the internal bg1 background image.			
bg_type	(OPTIONAL) String The name of the background cell type. By default is "Others".			
n_ir	Number of immune rings. This must match the arg length(ir_properties).			
ir_properties	List of properties of the immune rings. Please refer to the examples for the structure of ir_properties.			
plot_image	Boolean. Whether the simulated image is plotted.			
plot_categories				
	String Vector specifying the order of the cell categories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for plotting.			
plot_colours	String Vector specifying the order of the colours that correspond to the plot_categories arg. Default is NULL - the predefined colour vector would be used for plotting.			

Value

A data.frame of the simulated image

See Also

simulate_background_cells for all cell simulation, simulate_mixing for mixed background simulation, simulate_clusters for cluster simulation, simulate_double_rings for double immune ring simulation, and simulate_stripes for vessel simulation.

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Other simulate pattern functions: simulate_background_cells(), simulate_clusters(), simulate_double_rings simulate_mixing(), simulate_stripes()

Examples

```
set.seed(610)
# manually define the properties of the immune ring
ir_properties <- list(I1=list(name_of_cluster_cell="Tumour", size=600,
shape="Circle",centre_loc=data.frame("x"=930, "y"=1000),
infiltration_types=c("Immune1", "Immune2", "Others"), infiltration_proportions
=c(0.15, 0.05, 0.05), name_of_ring_cell="Immune1", immune_ring_width=150,
immune_ring_infiltration_types=c("Others"), immune_ring_infiltration_proportions=c(0.15)),
I2=list(name_of_cluster_cell="Tumour", size=500, shape="Oval",
centre_loc=data.frame("x"=1330, "y"=1100), infiltration_types=c("Immune1", "Immune2", "Others"),
infiltration_proportions=c(0.15, 0.05, 0.05), name_of_ring_cell="Immune1",
immune_ring_width=150, immune_ring_infiltration_types=c("Others"),
immune_ring_infiltration_proportions=c(0.15)))
# simulate immune rings (`n_ir` should match the length of `ir_properties`)
immune_ring_image <- simulate_immune_rings(bg_sample=bg1,
n_ir=2, ir_properties=ir_properties)</pre>
```

simulate_mixing

Simulate mixed background image

Description

Based on an existing background image, simulate mixed cell types with specified cell types and proportions. The default values for the arguments give an example of mixed cell type simulation which enable an automatic simulation of mixed cell types without the specification of any argument.

Usage

```
simulate_mixing(
  bg_sample = bg1,
  idents = c("Tumour", "Immune", "Others"),
  props = c(0.2, 0.4, 0.4),
  plot_image = TRUE,
  plot_colours = NULL
)
```

Arguments

bg_sample	(OPTIONAL) A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By default use the internal bg1 background image.
idents	String Vector of the mixed cell types.
props	Numeric Vector of the proportions of the mixed cell types.
plot_image	Boolean. Whether the simulated image is plotted.
plot colours	String Vector specifying the order of the colours that correspond to the idents

arg. Default is NULL - the predefined colour vector would be used for plotting..

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Value

A data.frame of the simulated image

See Also

simulate_background_cells for all cell simulation, simulate_clusters for cluster simulation, simulate_immune_rings/simulate_double_rings for immune ring simulation, and simulate_stripes for vessel simulation.

Other simulate pattern functions: simulate_background_cells(), simulate_clusters(), simulate_double_rings simulate_immune_rings(), simulate_stripes()

Examples

```
set.seed(610)
mix_background <- simulate_mixing(bg_sample=bg1,
idents=c("Tumour","Immune", "Others"), props=c(0.2, 0.4, 0.4),
plot_image=TRUE)</pre>
```

simulate_stripes

simulate_stripes

Description

Based on an existing background image, simulate stripes of cells representing vessels. The cell types and widths of the stripes can be specified. The locations of the stripes are randomly simulated. Please refer to the examples to check what properties of the stripes can be specified. The default values for the arguments give an example of vessel simulation which enable an automatic simulation of vessels without the specification of any argument.

Usage

```
simulate_stripes(
  bg_sample = bg1,
  n_stripe_type = 2,
  stripe_properties = list(S1 = list(number_of_stripes = 1, name_of_stripe_cell =
    "Others", width_of_stripe = 80, infiltration_types = c("Immune"),
    infiltration_proportions = c(0.08)), S2 = list(number_of_stripes = 1,
    name_of_stripe_cell = "Others", width_of_stripe = 80, infiltration_types =
    c("Immune"), infiltration_proportions = c(0.08))),
    plot_image = TRUE,
    plot_categories = NULL,
    plot_colours = NULL
)
```

Arguments

bg_sample

(OPTIONAL) A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords() of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions". By default use the internal bg1 background image.

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n_stripe_type Number of stripe types. Should be the same as length(stripe_properties.

stripe_properties

List of the properties of the stripes. See examples for the format of the properties. Please refer to the examples for the structure of stripe_properties.

plot_image Boolean. Whether the simulated image is plotted. Default is TRUE.

plot_categories

String Vector specifying the order of the cell categories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for plotting.

plot_colours String Vector specifying the order of the colours that correspond to the plot_categories arg. Default is NULL - the predefined colour vector would be used for plotting.

Value

A data.frame of the simulated image

See Also

simulate_background_cells for all cell simulation, simulate_mixing for mixed background simulation, simulate_clusters for cluster simulation, and simulate_immune_rings/simulate_double_rings for immune ring simulation

Other simulate pattern functions: simulate_background_cells(), simulate_clusters(), simulate_double_rings simulate_immune_rings(), simulate_mixing()

Examples

```
stripe_properties <- list(</pre>
S1 = list(
  number_of_stripes = 1,
  name_of_stripe_cell = "Others",
 width_of_stripe = 80,
  infiltration_types = c("Immune"),
  infiltration\_proportions = c(0.08)
), S2 = list(
  number_of_stripes = 1,
  name_of_stripe_cell = "Others",
  width_of_stripe = 80,
  infiltration_types = c("Immune"),
  infiltration\_proportions = c(0.08)))
set.seed(610)
stripe_image <- simulate_stripes(bg_sample = bg1, n_stripe_type=2,</pre>
stripe_properties = stripe_properties, plot_image = TRUE)
```

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Description

Tissue Image Simulator (TIS) integrates the basic simulation functions in spaSim, including simulating (mixed) background image, clusters, immune rings, double immune rings and stripes. The patterns are simulated on separate layers sequentially (e.g. immune rings are simulated after/on top of background cells). And each layer is also plot sequentially.

Pattern properties (e.g. properties_of_clusters) contain the properties of a pattern in the format of list where each element is one pattern. These properties need to be manually defined. Details about the format of the properties see the examples in simulate_clusters simulate_immune_rings simulate_double_rings simulate_stripes

Usage

```
TIS(
  bg_sample = NULL,
  n_{cells} = NULL,
  width = NULL,
  height = NULL,
  bg_method = NULL,
  min_d = NULL,
  oversampling_rate = 1.2,
  jitter = 0.3,
  names_of_bg_cells = NULL,
  proportions_of_bg_cells = NULL,
  n_clusters = NULL,
  properties_of_clusters = NULL,
  n_immune_rings = NULL,
  properties_of_immune_rings = NULL,
  n_double_rings = NULL,
  properties_of_double_rings = NULL,
  n_stripe_type = NULL,
  properties_of_stripes = NULL,
  image_name = NULL,
  plot_image = FALSE,
  plot_categories = NULL,
  plot_colours = NULL
```

Arguments

bg_sample	(OPTIONAL) A data frame or SpatialExperiment class object with locations of points representing background cells. Further cell types will be simulated based on this background sample. The data.frame or the spatialCoords of the SPE object should have colnames including "Cell.X.Positions" and "Cell.Y.Positions".
	By default use the internal bg1 background image.
n_cells	(OPTIONAL) Number of background cells to simulate. Only when bg_sample is NULL.
width	(OPTIONAL) Number The width of the image.
height	(OPTIONAL) Number The height of the image.
bg_method	(OPTIONAL) String specifying the distribution of the background cells. Choose from "Hardcore" and "Even".
min_d	(OPTIONAL) Number The minimum distance between two cells.

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oversampling_rate

(OPTIONAL) Numeric. The multiplier for oversampling. Without oversampling, the simulation deletes cells that are within min_d from each other, resulting in a less total number of cells than n_cells. Default is 1.2 (this should be set based on n_cells and min_d; should always be larger than 1).

jitter

(OPTIONAL) Numeric. Use when method is "Even". The uniform distribution parameter to generate the jitter distance for each cell from the vertices of the hexagon.

names_of_bg_cells

(OPTIONAL) Vector The cell types of the background cells. If NULL, the background cells are of one type.

proportions_of_bg_cells

(OPTIONAL) Vector The corresponding proportion of each cell type in the background cells.

 $\label{eq:nclusters} \mbox{n_clusters} \quad \mbox{(OPTIONAL) Number of cell clusters. If NULL, no clusters to simulate.} \\ \mbox{properties_of_clusters}$

(OPTIONAL) List of parameters to define the clusters.

 $\label{eq:n_immune_rings} \textbf{(OPTIONAL) Number of immune rings. If NULL, no immune rings to simulate.} \\ \textbf{properties_of_immune_rings}$

(OPTIONAL) List of parameters to define the immune rings.

n_double_rings (OPTIONAL) Number of double immune rings. If NULL, no double rings to simulate.

properties_of_double_rings

(OPTIONAL) List of parameters to define the double immune rings.

 $\label{eq:n_stripe_type} \textbf{(OPTIONAL) Number of stripe (vessel) types. If NULL, no stripes to simulate.} \\ \textbf{properties_of_stripes}$

(OPTIONAL) List of parameters to define the stripes.

image_name (OPTIONAL) String to name the output tissue image.

plot_image Boolean. Whether the simulated image is plotted.

plot_categories

String Vector specifying the order of the cell categories to be plotted. Default is NULL - the cell categories under the "Cell.Type" column would be used for plotting.

plot_colours

String Vector specifying the order of the colours that correspond to the plot_categories arg. Default is NULL - the predefined colour vector would be used for plotting.

Value

An spe object of the simulated image

Examples

```
set.seed(610)
double_ring_image <- TIS(bg_sample=bg1, n_clusters=1,
properties_of_clusters=list(C1=list( name_of_cluster_cell="Tumour",
size=300, shape="Oval", centre_loc=data.frame("x"=500, "y"=500),
infiltration_types=c("Immune1", "Others"), infiltration_proportions=c(0.1, 0.05))),
plot_image=TRUE)</pre>
```

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