# Package 'MGDrivE2'

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Type Package **Title** Mosquito Gene Drive Explorer 2 Version 2.1.0 Maintainer Agastya Mondal <amondal13@gmail.com> URL https://marshalllab.github.io/MGDrivE/docs\_v2/index.html, https://www.marshalllab.com/ BugReports https://github.com/MarshallLab/MGDrivE/issues

**Description** A simulation modeling framework which significantly extends capabilities from the 'MGDrivE' simulation package via a new mathematical and computational frame-

work based on stochastic Petri nets.

For more information about 'MGDrivE', see our publication: Sánchez et al. (2019) <doi:10.1111/2041-210X.13318>

Some of the notable capabilities of 'MGDrivE2' include: incorporation of human populations, epidemiological dynamics, time-varying parameters, and a continuous-time simulation framework with various sampling algorithms for both deterministic and stochastic interpretations. 'MGDrivE2' relies on the genetic inheritance structures provided in package 'MGDrivE', so we suggest installing that package initially.

License GPL-3 **Encoding UTF-8** ByteCompile true LazyData true **Depends** R (>= 3.1.0) Imports Matrix, deSolve, statmod Suggests MGDrivE, knitr, rmarkdown, markdown, ggplot2 VignetteBuilder knitr RoxygenNote 7.2.3 NeedsCompilation no Author Sean L. Wu [aut], Jared B. Bennett [aut],

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add\_interventions

This set of functions modifies mosquito life history parameters in the presence of adult interventions - indoor residual spraying (IRS) and insecticide treated nets (ITN) This is based on the work of Le Menach et al (2007) and Griffin et al (2010). We vary three parameters in the presence of interventions: Egg laying rate (beta), Adult mortality (muF), Mosquito biting rate (av0)

## Description

This set of functions modifies mosquito life history parameters in the presence of adult interventions - indoor residual spraying (IRS) and insecticide treated nets (ITN) This is based on the work of Le Menach et al (2007) and Griffin et al (2010). We vary three parameters in the presence of interventions: Egg laying rate (beta), Adult mortality (muF), Mosquito biting rate (av0)

#### Usage

```
add_interventions(params, IRS_cov, LLIN_cov)
```

#### **Arguments**

params a named list of parameters

IRS\_cov proportion of humans in the node receiving IRS

LLIN\_cov proportion of humans in the node receiving LLIN

#### Value

a vector of the equilibrium number of females in each SEI stage

base\_aquatic\_geno

Base Aquatic Function for Genotype Summary

### **Description**

This function takes a given aquatic (egg, larval, pupal) stage and sums over the Erlang-distributed stages, returning summary trajectories by genotype.

## Usage

```
base_aquatic_geno(out, spn_P, elp)
```

#### **Arguments**

out	the output of sim_trajectory_R
spn_P	the places of the SPN, see details

elp stage to summarize, one of: "egg", "larvae", "pupae"

base\_aquatic\_stage 5

#### **Details**

This function is the base function for summarize\_eggs\_geno, summarize\_larvae\_geno, and summarize\_pupae\_geno.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, genotype, and value.

#### Value

a 3 to 5 column dataframe for plotting with ggplot2

base\_aquatic\_stage Base Aquatic Function for Erlang-Stage Summary

## **Description**

This function takes a given aquatic (egg, larval, pupal) stage and sums over the genotypes, returning summary trajectories by Erlang-distributed stage.

#### Usage

```
base_aquatic_stage(out, spn_P, elp)
```

## Arguments

out	the output of sim_trajectory_R
spn_P	the places of the SPN, see details
elp	stage to summarize, one of: "egg", "larvae", "pupae"

### **Details**

This function is the base function for summarize\_eggs\_stage, summarize\_larvae\_stage, and summarize\_pupae\_stage.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, Erlang-stage, and value.

## Value

a 3 to 5 column dataframe for plotting with ggplot2

base\_erlang\_F

base_erlang	Base Summary of Erlang Stages for Aquatic Life Stages	
base_erlang	Base Summary of Erlang Stages for Aquatic Life Stages	

## **Description**

This function takes the given aquatic stage and summarizes them by Erlang-distributed dwell times, writing output to provided folders.

## Usage

```
base_erlang(fileVec, outList, genos, nGenos, nErlang, times, nTimes, nNodes)
```

## Arguments

fileVec	Vector of files for analysis
outList	List of files, organized by repetition, to write output
genos	Genotypes to summarize by
nGenos	Number of genotypes
nErlang	Number of Erlang stages
times	Vector of sampling times
nTimes	Number of sampled times
nNodes	Number of nodes in the network

## **Details**

This function is a base function used in split\_aggregate\_CSV.

## Value

None

## Description

This function takes ALL of the adult female stages and summarized them by Erlang-distributed latent infection, writing output to provided folders.

## Usage

```
base_erlang_F(fileList, outList, nGenos, nErlang, times, nTimes, nNodes)
```

base\_gen 7

## Arguments

fileList	Length 3 list holding 'FS', 'FE', and 'FI' files for analysis
outList	List of files, organized by repetition, to write output
nGenos	Number of genotypes
nErlang	Number of Erlang stages
times	Vector of sampling times
nTimes	Number of sampled times
nNodes	Number of nodes in the network

## **Details**

This function is a base function used in split\_aggregate\_CSV.

#### Value

None

Base Summary for Eggs, Larvae, Pupae, Susceptible Females, and
Infectious Females

## Description

This function takes a given stage and summarizes them by genotype, writing output to provided folders.

## Usage

```
base_gen(fileVec, outList, genos, nGenos, nIDX1, times, nTimes, nNodes)
```

## **Arguments**

fileVec	Vector of files for analysis
outList	List of files, organized by repetition, to write output
genos	Genotypes to summarize by
nGenos	Number of genotypes
nIDX1	First index to expand over, nE/nL/nP for aquatic stages, 1 for the rest
times	Vector of sampling times
nTimes	Number of sampled times
nNodes	Number of nodes in the network

## **Details**

This function is a base function used in split\_aggregate\_CSV.

8 base\_gen\_FE

## Value

None

base\_gen\_FE Base Summary for Latent Females

## Description

This function takes 'E' stage females and summarizes them by genotype, writing output to provided folders.

## Usage

```
base_gen_FE(fileVec, outList, genos, nGenos, nIDX1, times, nTimes, nNodes)
```

## Arguments

fileVec	Vector of files for analysis
outList	List of files, organized by repetition, to write output
genos	Genotypes to summarize by
nGenos	Number of genotypes
nIDX1	First index to expand over, nE/nL/nP for aquatic stages, 1 for the rest
times	Vector of sampling times
nTimes	Number of sampled times
nNodes	Number of nodes in the network

## **Details**

This function is a base function used in split\_aggregate\_CSV.

## Value

None

base\_MQ

base\_MQ Base Summary Function

## Description

This function does the actual calculations for summarize\_stats\_CSV. It calculates mean and quantiles, writing output to the appropriate folder.

## Usage

```
base_MQ(
fList,
oDir,
sName,
nodeNames,
nNodes,
genos,
nGenos,
times,
nTimes,
num_repss,
mean,
quantiles,
oDepth
)
```

## Arguments

fList	File list, all files for this stage, organized by repetition
oDir	Output directory
sName	Stage signifier
nodeNames	Properly formatted vector of node names for printing
nNodes	Number of nodes in the simulation
genos	Vector of genotypes for the header
nGenos	Number of genotypes
times	Vector of sampling times
nTimes	Number of sampled times
num_repss	Number of repetitions from the simulation
mean	Boolean, calculate mean or not
quantiles	Vector of quantiles to calculate, or NULL

Max(1, number of quantiles)

### Value

None

oDepth

base\_MUH Base Summary for Males, Unmated Females, and Humans

## **Description**

This function takes a given stage (males, unmated females, or humans) and summarizes them by genotype (infection status for humans), writing output to provided folders.

## Usage

```
base_MUH(fileVec, outList, genos, nGenos, nTimes, nNodes)
```

## **Arguments**

fileVec Vector of files for analysis

outList List of files, organized by repetition, to write output

genos Genotypes to summarize by

nGenos Number of genotypes nTimes Number of sampled times

nNodes Number of nodes in the network

#### **Details**

This function is a base function used in split\_aggregate\_CSV.

### Value

None

base\_summarize\_humans Base Function for Human Summary

## Description

This function takes a given infection ('S','E','I','R') status and returns a summary trajectory

## Usage

```
base_summarize_humans(out, infState)
```

## Arguments

out the output of sim\_trajectory\_R

infState type of humans to summarize: 'S','E','I','R'

base\_sum\_F

## **Details**

This function is the base function for summarize\_humans\_epiSIS, summarize\_humans\_epiSEIR.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, inf, genotype, and value.

#### Value

a 4 to 6 column dataframe for plotting with ggplot2

base\_sum\_F

Base Summary of Infection (SEI) Stages for Adult Females

## Description

This function takes ALL of the adult female stages and summarized them by Erlang-distributed latent infection, writing output to provided folders.

#### Usage

```
base_sum_F(fileList, outList, genos, nGenos, nErlang, times, nTimes, nNodes)
```

Length 3 list holding 'FS', 'FE', and 'FI' files for analysis

## Arguments

fileList

11102100	Zengur 5 not nothing 15,12, and 11 mes for unaryons
outList	List of files, organized by repetition, to write output
genos	Genotypes to summarize by
nGenos	Number of genotypes
nErlang	Number of Erlang stages
times	Vector of sampling times
nTimes	Number of sampled times
nNodes	Number of nodes in the network

#### **Details**

This function is a base function used in split\_aggregate\_CSV.

### Value

None

batch\_migration

batch\_migration

Sample Batch Migration Events

#### **Description**

Sample batch migration events for simulation given rates of occurance and probability of destination for each patch. Batch migration can be simulated for the aquatic life stages (eggs, larvae, pupae), adult females, and/or adult males. To simulate batch migration, each life stage needs all 3 of its arguments specified. If any arguments are left unspecified (NULL), batch migration for that life stage will not be sampled. The output of this function should be passed to sim\_trajectory\_R or sim\_trajectory\_CSV as the argument batch. Calls the internal function batch\_migration\_stage.

## Usage

```
batch_migration(
   SPN_P,
   tmax,
   ELPrates = NULL,
   ELPmove = NULL,
   Frates = NULL,
   Frates = NULL,
   Fmove = NULL,
   Fmove = NULL,
   Mrates = NULL,
   Mmove = NULL,
   Mprob = NULL,
   stage = NULL
)
```

#### **Arguments**

SPN_P	places of the SPN
tmax	maximum time of the simulation
ELPrates	rate at which aquatic stage batch migration occurs for each node (nodes without mosquitoes should be set to NaN or NA)
ELPmove	movement matrix for destinations of aquatic stage batch migration events (diagonal will be set to zero and off-diagonal elements normalized)
ELPprob	probability for each individual to be chosen for aquatic stage batch migration events (must be same length as ELPrates)
Frates	rate at which adult female batch migration occurs for each node (nodes without mosquitoes should be set to NaN or NA)
Fmove	movement matrix for destinations of adult female batch migration events (diagonal will be set to zero and off-diagonal elements normalized)
Fprob	probability for each individual to be chosen for adult female batch migration events (must be same length as Frates)

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Mrates	rate at which adult male batch migration occurs for each node (nodes without mosquitoes should be set to NaN or NA)
Mmove	movement matrix for destinations of adult male batch migration events (diagonal will be set to zero and off-diagonal elements normalized)
Mprob	probability for each individual to be chosen for adult male batch migration events (must be same length as Mrates)
stage	either NULL or "E", "L", or "P". If not NULL and migration for aquatic stages is specified by ELPrates, only the aquatic stage specified here will move

## Value

vector of lists describing all batch migration events, segmented by life stage.

 ${\tt batch\_migration\_stage} \quad \textit{Internal function to sample and set up data structure for batch migration} \\$ 

## Description

Internal function to sample and set up data structure for batch migration

## Usage

```
batch_migration_stage(SPN_P, rates, move, prob, stage, tmax)
```

## Arguments

SPN_P	a set of Petri net places
rates	a vector of rates for each node
move	a movement matrix (where do the batches go?)
prob	the probability vector for each individual moving in each batch
stage	the life stage (one of 'ELP', 'F', 'M')
tmax	maximum simulation time

calc\_move\_rate

Calculate Outbound Movement Rate

#### **Description**

Given P, the cumulative probability of moving before dying, and mu, the daily mortality rate, calculate the movement rate gamma to get P. The equation comes from integrating the competing risks and solving for gamma.

#### Usage

```
calc_move_rate(mu, P)
```

#### **Arguments**

mu daily mortality rate

P cumulative probability to move before dying

## Value

numeric probability of movement

## **Examples**

```
convert_prevalence_to_eir
```

Generally, pathogen prevalence is a more accessible metric for users, but the Imperial equilibrium function requires an annual EIR. This function converts a given pathogen prevalence to an EIR

## Description

Generally, pathogen prevalence is a more accesible metric for users, but the Imperial equilibrium function requires an annual EIR. This function converts a given pathogen prevalence to an EIR

### Usage

```
convert_prevalence_to_eir(prevalence, age_vector, ft, params)
```

## **Arguments**

prevalence desired prevalence value

age\_vector age distribution of the population

ft proportion treated

params entomological and epidemiological parameters

#### Value

a vector of the equilibrium number of humans in each SIS stage

equilibrium\_Imperial\_decoupled

This function calculates the human and mosquito equilibrium values for the decoupled Imperial model. Currently this only works in one node.

## Description

This function calculates the human and mosquito equilibrium values for the decoupled Imperial model. Currently this only works in one node.

## Usage

```
equilibrium_Imperial_decoupled(age_vector, ft, eir, theta, cube, spn_P)
```

#### **Arguments**

age\_vector age structure of population (see vignette for example)

ft proportion of population seeking treatment

eir desired annual EIR

theta parameters

cube inheritance cube

spn\_P places of the stochastic petri net

#### Value

a matrix of the equilibrium number of humans in each Imperial stage by age, and immunity. mosquito equilibrium values, and full theta vector

16 equilibrium\_lifeycle

```
equilibrium_Imperial_decoupled_human
```

This function calculates the human equilibrium values for the decoupled Imperial model. Requires the age structure of the population Currently this only works in one node.

## Description

This function calculates the human equilibrium values for the decoupled Imperial model. Requires the age structure of the population Currently this only works in one node.

#### Usage

```
equilibrium_Imperial_decoupled_human(age_vector, ft, EIR, model_param_list)
```

### **Arguments**

```
age_vector age structure of population (see vignette for example)

ft proportion of population seeking treatment

EIR desired annual EIR

model_param_list

parameters for the
```

### Value

a matrix of the equilibrium number of humans in each Imperial stage by age, and immunity

```
equilibrium_lifeycle Calculate Equilibrium for Lifecycle Model (Logistic or Lotka-Volterra)
```

## Description

This function calculates deterministic equilibria for the mosquito lifecycle model.

## Usage

```
equilibrium_lifeycle(
  params,
  NF,
  phi = 0.5,
  log_dd = TRUE,
  spn_P,
  pop_ratio_Aq = NULL,
  pop_ratio_F = NULL,
```

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```
pop_ratio_M = NULL,
  cube
)
```

### Arguments

params a named list of parameters (see details)

NF vector of female mosquitoes at equilibrium, for every population in the environ-

ment

phi sex ratio of mosquitoes at emergence

log\_dd Boolean: TRUE implies logistic density dependence, FALSE implies Lotka-

Volterra model

spn\_P the set of places (P) (see details)

pop\_ratio\_Aq May be empty; if not, a named vector or matrix. (see details)
pop\_ratio\_F May be empty; if not, a named vector or matrix. (see details)
pop\_ratio\_M May be empty; if not, a named vector or matrix. (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

Equilibrium can be calculated using one of two models: classic logistic dynamics or following the Lotka-Volterra competition model. This is determined by the parameter log\_dd, and it changes elements of the return list: K is returned for logistic dynamics, or gamma is returned for Lotka-Volterra dynamics.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The initial population genotype ratios are set by supplying the pop\_ratio\_Aq, pop\_ratio\_F, and pop\_ratio\_M values. The default value is NULL, and the function will use the wild-type alleles provided in the cube object. However, one can supply several different objects to set the initial genotype ratios. All genotypes provided must exist in the cube (this is checked by the function). If a single, named vector is provided, then all patches will be initialized with the same ratios. If a matrix is provided, with the number of columns (and column names) giving the initial genotypes, and a row for each patch, each patch can be set to a different initial ratio. The three parameters do not need to match each other.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This is used to set the initial population distribution and during the simulation to maintain equilibrium. params must include the following named parameters:

- qE: inverse of mean duration of egg stage
- nE: shape parameter of Erlang-distributed egg stage
- qL: inverse of mean duration of larval stage
- nL: shape parameter of Erlang-distributed larval stage
- qP: inverse of mean duration of pupal stage
- nP: shape parameter of Erlang-distributed pupal stage

- muE: egg mortality
- muL: density-independent larvae mortality
- muP: pupae mortality
- · muF: adult female mortality
- muM: adult male mortality
- beta: egg-laying rate, daily
- nu: mating rate of unmated females

The return list contains all of the params parameters, along with the density-dependent parameter, either K or gamma. These are the parameters necessary later in the simulations. This was done for compatibility with equilibrium\_SEI\_SIS, which requires several extra parameters not required further in the simulations.

For equilibrium with epidemiological parameters, see equilibrium\_SEI\_SIS. For equilibrium with latent humans (SEIR dynamics), see equilibrium\_SEI\_SEIR.

#### Value

a list with 3 elements: init a matrix of equilibrium values for every life-cycle stage, params a list of parameters for the simulation, M0 a vector of initial conditions

```
equilibrium_SEI_decoupled_human
```

This function calculates the equilibrium values for the decoupled SIS human states. Currently this only works in one node.

### Description

This function calculates the equilibrium values for the decoupled SIS human states. Currently this only works in one node.

#### Usage

```
equilibrium_SEI_decoupled_human(params)
```

## Arguments

params a named list of parameters (see details)

#### Value

a vector of the equilibrium number of humans in each SIS stage

```
equilibrium_SEI_decoupled_mosy
```

Calculate Equilibrium for Decoupled Mosquito SEI model. Human states will be handled separately.

## **Description**

Given prevalence of disease in humans (modeled as an SIS: Susceptible-Infected-Susceptible process with birth and death) and entomological parameters of transmission, this function calculates the quasi-stationary distribution of adult female mosquitoes across SEI (Susceptible-Exposed-Infectious) stages, allowing for Erlang distributed E stage.

## Usage

```
equilibrium_SEI_decoupled_mosy(
  params,
  node_list = "b",
  NF = NULL,
  phi = 0.5,
  NH = NULL,
  log_dd = TRUE,
  spn_P,
  pop_ratio_Aq = NULL,
  pop_ratio_F = NULL,
  pop_ratio_M = NULL,
  pop_ratio_H = 1,
  cube
)
```

#### **Arguments**

params	a named list of parameters (see details)
node_list	a character vector specifying what type of nodes to create; $(m = a \text{ node with only mosquitoes}, h = a \text{ node with only humans}, b = a \text{ node with both humans and mosquitoes})$
NF	vector of female mosquitoes at equilibrium, for mosquito-only nodes
phi	sex ratio of mosquitoes at emergence
NH	vector of humans at equilibrium, for human-only nodes
log_dd	Boolean: TRUE implies logistic density dependence, FALSE implies Lotka-Volterra model
spn_P	the set of places (P) (see details)
pop_ratio_Aq	May be empty; if not, a named vector or matrix. (see details)
pop_ratio_F	May be empty; if not, a named vector or matrix. (see details)
pop_ratio_M	May be empty; if not, a named vector or matrix. (see details)
pop_ratio_H	Prevalence in human-only nodes
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

This function handles 3 types of nodes: Human only, mosquito only, and nodes with both. These nodes are set using the node\_list parameter. Mosquito-only node equilibrium calls equilibrium\_lifeycle, which follows one of two models: classic logistic dynamics or the Lotka-Volterra competition model. This is determined by the parameter log\_dd, and it changes elements of the return list: K is returned for logistic dynamics, or gamma is returned for Lotka-Volterra dynamics. This is parameterized with the NF parameter to define the adult female numbers. This parameter only needs to be supplied if there are mosquito-only nodes.

Human-only nodes don't require any equilibrium calculations. These nodes use the NH and pop\_ratio\_H to set adult human populations and infection rates in nodes. These two parameters only need to be supplied if there are human-only nodes.

For human and mosquito nodes, this function calls make\_Q\_SEI to construct the infinitesimal generator matrix which is used to solve for the quasi-stationary (stochastic) or equilibrium (deterministic) distribution of mosquitoes over stages. Parameters are provided by params.

For information on the method used to solve this distribution, see section "3.1.3 Nonsingularity of the Subintensity Matrix" of:

• Bladt, Mogens, and Bo Friis Nielsen. Matrix-exponential distributions in applied probability. Vol. 81. New York: Springer, 2017.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The initial population genotype ratios are set by supplying the pop\_ratio\_Aq, pop\_ratio\_F, and pop\_ratio\_M values. The default value is NULL, and the function will use the wild-type alleles provided in the cube object. However, one can supply several different objects to set the initial genotype ratios. All genotypes provided must exist in the cube (this is checked by the function). If a single, named vector is provided, then all patches will be initialized with the same ratios. If a matrix is provided, with the number of columns (and column names) giving the initial genotypes, and a row for each patch, each patch can be set to a different initial ratio. The three parameters do not need to match each other.

The params argument supplies all of the ecological and epidemiological parameters necessary to calculate equilibrium values. This is used to set the initial population distribution and during the simulation to maintain equilibrium. This params must include the following named parameters, noted as being the same as lifecycle parameters, or new for the epidemiological equilibrium

## • (Lifecycle parameters)

- qE: inverse of mean duration of egg stage
- nE: shape parameter of Erlang-distributed egg stage
- qL: inverse of mean duration of larval stage
- nL: shape parameter of Erlang-distributed larval stage
- qP: inverse of mean duration of pupal stage
- nP: shape parameter of Erlang-distributed pupal stage
- muE: egg mortality
- muL: density-independent larvae mortality
- muP: pupae mortality
- muF: adult female mortality

- muM: adult male mortality
- beta: egg-laying rate, daily
- nu: mating rate of unmated females

## • (Epidemiological parameters)

- NH: number of humans, can be a vector
- X: prevalence in humans, can be a vector
- NFX: number of female mosquitoes, only required if any prevalence (X) is zero
- b: mosquito to human transmission efficiency, can be a vector
- c: human to mosquito transmission efficiency, can be a vector
- r: rate of recovery in humans (1/duration of infectiousness)
- muH: death rate of humans (1/avg lifespan)
- f: rate of blood feeding
- 0: human blood index
- qEIP: related to scale parameter of Gamma distributed EIP (1/qEIP is mean length of EIP)
- nEIP: shape parameter of Gamma distributed EIP

The return list contains all of the parameters necessary later in the simulations.

For equilibrium without epidemiological parameters, see equilibrium\_lifeycle. For equilibrium with latent humans (SEIR dynamics), see equilibrium\_SEI\_SEIR.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

### Value

a vector of the equilibrium number of females in each SEI stage

```
equilibrium_SEI_Imperial
```

Calculate Equilibrium for Mosquito SEI - Human Imperial Model

## **Description**

In decoupled sampling, human states are handled separately from mosquito states. The function equilibrium\_Imperial\_decoupled\_human calculates the distribution of humans at equilibrium required for the Imperial model of malaria transmission. Here we use parameters from that model to calculate the equilibrium states of Susceptible-Exposed-Infectious (SEI) female mosquitoes

## Usage

```
equilibrium_SEI_Imperial(
  params,
  node_list = "b",
  NF = NULL,
  phi = 0.5,
```

```
NH = NULL,
log_dd = TRUE,
spn_P,
pop_ratio_Aq = NULL,
pop_ratio_F = NULL,
pop_ratio_M = NULL,
pop_ratio_H = 1,
cube
)
```

#### **Arguments**

params a named list of parameters (see details) node\_list list of geospatial nodes NF number of female mosquitoes phi sex ratio of mosquitoes at emergence NH vector of humans at equilibrium Boolean: TRUE implies logistic density dependence, FALSE implies Lotkalog\_dd Volterra model the set of places (P) (see details) spn\_P pop\_ratio\_Aq May be empty; if not, a named vector or matrix. (see details) pop\_ratio\_F May be empty; if not, a named vector or matrix. (see details) May be empty; if not, a named vector or matrix. (see details) pop\_ratio\_M Prevalence in human-only nodes pop\_ratio\_H an inheritance cube from the MGDrivE package (e.g. cubeMendelian) cube

#### **Details**

Imperial model sampling is currently only supported for one-node dynamics: a single node with mosquitoes parameterized by the distribution of human states. These nodes are set using the node\_list parameter. Mosquito-only node equilibrium calls equilibrium\_lifeycle, which follows one of two models: classic logistic dynamics or the Lotka-Volterra competition model. This is determined by the parameter log\_dd, and it changes elements of the return list: K is returned for logistic dynamics, or gamma is returned for Lotka-Volterra dynamics. This is parameterized with the NF parameter to define the adult female numbers. This parameter only needs to be supplied if there are mosquito-only nodes.

For human and mosquito nodes, this function calculates the number of SEI mosquitoes in each state.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The initial population genotype ratios are set by supplying the pop\_ratio\_Aq, pop\_ratio\_F, and pop\_ratio\_M values. The default value is NULL, and the function will use the wild-type alleles provided in the cube object. However, one can supply several different objects to set the initial genotype ratios. All genotypes provided must exist in the cube (this is checked by the function). If a single, named vector is provided, then all patches will be initialized with the same ratios. If a matrix is provided, with the number of columns (and column names) giving the initial genotypes,

and a row for each patch, each patch can be set to a different initial ratio. The three parameters do not need to match each other.

The params argument supplies all of the ecological and epidemiological parameters necessary to calculate equilibrium values. This is used to set the initial population distribution and during the simulation to maintain equilibrium. This params must include the following named parameters, noted as being the same as lifecycle parameters, or new for the epidemiological equilibrium

### • (Lifecycle parameters)

- qE: inverse of mean duration of egg stage
- nE: shape parameter of Erlang-distributed egg stage
- qL: inverse of mean duration of larval stage
- nL: shape parameter of Erlang-distributed larval stage
- qP: inverse of mean duration of pupal stage
- nP: shape parameter of Erlang-distributed pupal stage
- muE: egg mortality
- muL: density-independent larvae mortality
- muP: pupae mortality
- muF: adult female mortality, supplied from Imperial equilibrium function
- muM: adult male mortality, supplied from Imperial equilibrium function
- beta: egg-laying rate, daily
- nu: mating rate of unmated females

## • (Epidemiological parameters)

- NH: number of humans, can be a vector
- FOIv: force of infection on mosquitoes, supplied from Imperial equilibrium function
- Iv\_eq: per-capita proportion of infectious mosquitoes The return list contains all of the parameters necessary later in the simulations.

#### Value

a vector of the equilibrium number of females in each SEI stage

equilibrium\_SEI\_SEIR Calculate Equilibrium for Mosquito SEI - Human SEIR Model

### **Description**

Given prevalence of disease in humans (modeled as an SEIR: Susceptible-Latent-Infected-Recovered process with birth and death) and entomological parameters of transmission, this function calculates the quasi-stationary distribution of adult female mosquitoes across SEI (Susceptible-Exposed-Infectious) stages, allowing for Erlang distributed E stage.

#### Usage

```
equilibrium_SEI_SEIR(
  params,
  node_list = "b",
  NF = NULL,
  phi = 0.5,
  NH = NULL,
  log_dd = TRUE,
  spn_P,
  pop_ratio_Aq = NULL,
  pop_ratio_F = NULL,
  pop_ratio_H = C(1, 0, 0, 0),
  cube
)
```

### **Arguments**

params	a named list of parameters (see details)
node_list	a character vector specifying what type of nodes to create; $(m = a \text{ node with only mosquitoes}, h = a \text{ node with only humans}, b = a \text{ node with both humans and mosquitoes})$
NF	vector of female mosquitoes at equilibrium, for mosquito-only nodes
phi	sex ratio of mosquitoes at emergence
NH	vector of humans at equilibrium, for human-only nodes
log_dd	Boolean: TRUE implies logistic density dependence, FALSE implies Lotka-Volterra model
spn_P	the set of places (P) (see details)
pop_ratio_Aq	May be empty; if not, a named vector or matrix. (see details)
pop_ratio_F	May be empty; if not, a named vector or matrix. (see details)
pop_ratio_M	May be empty; if not, a named vector or matrix. (see details)
pop_ratio_H	Prevalence in human-only nodes, default is all susceptible
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

This function handles 3 types of nodes: Human only, mosquito only, and nodes with both. These nodes are set using the node\_list parameter. Mosquito-only node equilibrium calls equilibrium\_lifeycle, which follows one of two models: classic logistic dynamics or the Lotka-Volterra competition model. This is determined by the parameter log\_dd, and it changes elements of the return list: K is returned for logistic dynamics, or gamma is returned for Lotka-Volterra dynamics. This is parameterized with the NF parameter to define the adult female numbers. This parameter only needs to be supplied if there are mosquito-only nodes.

Human-only nodes don't require any equilibrium calculations. These nodes use the NH and pop\_ratio\_H to set adult human populations and infection rates in nodes. These two parameters only need to be

supplied if there are human-only nodes. pop\_ratio\_H needs to be a matrix with the number of rows equal to the number of human-only patches, and 4 columns. The columns are assumed to be fractions of the population in "S", "E", "I", or "R" states, and every row must sum to 1.

For human and mosquito nodes, this function calls make\_Q\_SEI to construct the infinitesimal generator matrix which is used to solve for the quasi-stationary (stochastic) or equilibrium (deterministic) distribution of mosquitoes over stages. Parameters are provided by params.

For information on the method used to solve this distribution, see section "3.1.3 Nonsingularity of the Subintensity Matrix" of:

• Bladt, Mogens, and Bo Friis Nielsen. Matrix-exponential distributions in applied probability. Vol. 81. New York: Springer, 2017.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The initial population genotype ratios are set by supplying the pop\_ratio\_Aq, pop\_ratio\_F, and pop\_ratio\_M values. The default value is NULL, and the function will use the wild-type alleles provided in the cube object. However, one can supply several different objects to set the initial genotype ratios. All genotypes provided must exist in the cube (this is checked by the function). If a single, named vector is provided, then all patches will be initialized with the same ratios. If a matrix is provided, with the number of columns (and column names) giving the initial genotypes, and a row for each patch, each patch can be set to a different initial ratio. The three parameters do not need to match each other.

The params argument supplies all of the ecological and epidemiological parameters necessary to calculate equilibrium values. This is used to set the initial population distribution and during the simulation to maintain equilibrium. This params must include the following named parameters, noted as being the same as lifecycle parameters, or new for the epidemiological equilibrium

## • (Lifecycle parameters)

- qE: inverse of mean duration of egg stage
- nE: shape parameter of Erlang-distributed egg stage
- qL: inverse of mean duration of larval stage
- nL: shape parameter of Erlang-distributed larval stage
- qP: inverse of mean duration of pupal stage
- nP: shape parameter of Erlang-distributed pupal stage
- muE: egg mortality
- muL: density-independent larvae mortality
- muP: pupae mortality
- muF: adult female mortality
- muM: adult male mortality
- beta: egg-laying rate, daily
- nu: mating rate of unmated females

#### • (Epidemiological parameters)

- NH: number of humans, can be a vector
- X: SEIR prevalence in humans, can be a vector of length 4 for 1 node, or a matrix for many nodes

- NFX: number of female mosquitoes, only required if any prevalence (X) is zero
- b: mosquito to human transmission efficiency, can be a vector
- c: human to mosquito transmission efficiency, can be a vector
- r: rate of recovery in humans (1/duration of infectiousness)
- muH: death rate of humans (1/avg lifespan)
- f: rate of blood feeding
- Q: human blood index
- qEIP: related to scale parameter of Gamma distributed EIP (1/qEIP is mean length of EIP)
- nEIP: shape parameter of Gamma distributed EIP
- delta: inverse duration of the latent stage (E)

The return list contains all of the parameters necessary later in the simulations.

For equilibrium without epidemiological parameters, see equilibrium\_lifeycle. For equilibrium without latent humans (SIS dynamics), see equilibrium\_SEI\_SIS.

#### Value

a vector of the equilibrium number of females in each SEI stage

equilibrium\_SEI\_SIS Calculate Equilibrium for Mosquito SEI - Human SIS Model

## **Description**

Given prevalence of disease in humans (modeled as an SIS: Susceptible-Infected-Susceptible process with birth and death) and entomological parameters of transmission, this function calculates the quasi-stationary distribution of adult female mosquitoes across SEI (Susceptible-Exposed-Infectious) stages, allowing for Erlang distributed E stage.

### Usage

```
equilibrium_SEI_SIS(
  params,
  node_list = "b",
  NF = NULL,
  phi = 0.5,
  NH = NULL,
  log_dd = TRUE,
  spn_P,
  pop_ratio_Aq = NULL,
  pop_ratio_F = NULL,
  pop_ratio_H = 1,
  cube
)
```

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### Arguments

params a named list of parameters (see details)

node\_list a character vector specifying what type of nodes to create; (m = a node with

only mosquitoes, h = a node with only humans, b = a node with both humans

and mosquitoes)

NF vector of female mosquitoes at equilibrium, for mosquito-only nodes

phi sex ratio of mosquitoes at emergence

NH vector of humans at equilibrium, for human-only nodes

log\_dd Boolean: TRUE implies logistic density dependence, FALSE implies Lotka-

Volterra model

spn\_P the set of places (P) (see details)

pop\_ratio\_Aq May be empty; if not, a named vector or matrix. (see details)
pop\_ratio\_F May be empty; if not, a named vector or matrix. (see details)
pop\_ratio\_M May be empty; if not, a named vector or matrix. (see details)

pop\_ratio\_H Prevalence in human-only nodes

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

This function handles 3 types of nodes: Human only, mosquito only, and nodes with both. These nodes are set using the node\_list parameter. Mosquito-only node equilibrium calls equilibrium\_lifeycle, which follows one of two models: classic logistic dynamics or the Lotka-Volterra competition model. This is determined by the parameter log\_dd, and it changes elements of the return list: K is returned for logistic dynamics, or gamma is returned for Lotka-Volterra dynamics. This is parameterized with the NF parameter to define the adult female numbers. This parameter only needs to be supplied if there are mosquito-only nodes.

Human-only nodes don't require any equilibrium calculations. These nodes use the NH and pop\_ratio\_H to set adult human populations and infection rates in nodes. These two parameters only need to be supplied if there are human-only nodes.

For human and mosquito nodes, this function calls make\_Q\_SEI to construct the infinitesimal generator matrix which is used to solve for the quasi-stationary (stochastic) or equilibrium (deterministic) distribution of mosquitoes over stages. Parameters are provided by params.

For information on the method used to solve this distribution, see section "3.1.3 Nonsingularity of the Subintensity Matrix" of:

• Bladt, Mogens, and Bo Friis Nielsen. Matrix-exponential distributions in applied probability. Vol. 81. New York: Springer, 2017.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The initial population genotype ratios are set by supplying the pop\_ratio\_Aq, pop\_ratio\_F, and pop\_ratio\_M values. The default value is NULL, and the function will use the wild-type alleles provided in the cube object. However, one can supply several different objects to set the initial genotype ratios. All genotypes provided must exist in the cube (this is checked by the function).

If a single, named vector is provided, then all patches will be initialized with the same ratios. If a matrix is provided, with the number of columns (and column names) giving the initial genotypes, and a row for each patch, each patch can be set to a different initial ratio. The three parameters do not need to match each other.

The params argument supplies all of the ecological and epidemiological parameters necessary to calculate equilibrium values. This is used to set the initial population distribution and during the simulation to maintain equilibrium. This params must include the following named parameters, noted as being the same as lifecycle parameters, or new for the epidemiological equilibrium

### • (Lifecycle parameters)

- qE: inverse of mean duration of egg stage
- nE: shape parameter of Erlang-distributed egg stage
- qL: inverse of mean duration of larval stage
- nL: shape parameter of Erlang-distributed larval stage
- qP: inverse of mean duration of pupal stage
- nP: shape parameter of Erlang-distributed pupal stage
- muE: egg mortality
- muL: density-independent larvae mortality
- muP: pupae mortality
- muF: adult female mortality
- muM: adult male mortality
- beta: egg-laying rate, daily
- nu: mating rate of unmated females

## • (Epidemiological parameters)

- NH: number of humans, can be a vector
- X: prevalence in humans, can be a vector
- NFX: number of female mosquitoes, only required if any prevalence (X) is zero
- b: mosquito to human transmission efficiency, can be a vector
- c: human to mosquito transmission efficiency, can be a vector
- r: rate of recovery in humans (1/duration of infectiousness)
- muH: death rate of humans (1/avg lifespan)
- f: rate of blood feeding
- Q: human blood index
- qEIP: related to scale parameter of Gamma distributed EIP (1/qEIP is mean length of EIP)
- nEIP: shape parameter of Gamma distributed EIP

The return list contains all of the parameters necessary later in the simulations.

For equilibrium without epidemiological parameters, see equilibrium\_lifeycle. For equilibrium with latent humans (SEIR dynamics), see equilibrium\_SEI\_SEIR.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

### Value

a vector of the equilibrium number of females in each SEI stage

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get\_shape

Calculate Erlang shape parameter

## **Description**

Calculate Erlang shape parameter

## Usage

```
get_shape(cv, q)
```

## **Arguments**

cv coefficient of variation (CV) between mean and standard deviation of dwell

times, smaller values of CV correspond to distributions less dispersed around

their mean and larger value to more dispersed distributions.

q inverse of mean dwell time

#### Value

integer value representing the coefficient of variation in Erlang-distributed life stages.

human\_Imperial\_ODE

ODE describing the age-structured Imperial model used in decoupled sampling, which will pass in values of  $I_V$  and return the human states for usein the mosquito portion of the model

## **Description**

ODE describing the age-structured Imperial model used in decoupled sampling, which will pass in values of I\_V and return the human states for usein the mosquito portion of the model

## Usage

```
human_Imperial_ODE(t, state, parameters)
```

## **Arguments**

t starting time of simulation state distributon of disease states

parameters parameter set

#### Value

matrix of disease states after integration

## **Description**

 $model\_param\_list\_create$  creates list of model parameters to be used within equilibrium\_init\_create

#### Usage

```
imperial_model_param_list_create(
 eta = 1/(21 * 365),
  rho = 0.85,
  a0 = 2920,
  sigma2 = 1.67,
 max_age = 100 * 365,
 rA = 1/195,
 rT = 0.2,
  rD = 0.2,
  rU = 1/110.299,
  rP = 1/15,
  dE = 12,
  delayGam = 12.5,
  cD = 0.0676909,
  cT = 0.322 * cD,
  cU = 0.006203,
  gamma1 = 1.82425,
  d1 = 0.160527,
  dID = 3650,
  ID0 = 1.577533,
  kD = 0.476614,
  uD = 9.44512,
  aD = 8001.99,
  fD0 = 0.007055,
  gammaD = 4.8183,
  alphaA = 0.75735,
  alphaU = 0.185624,
  b0 = 0.590076,
  b1 = 0.5,
  dB = 3650,
  IB0 = 43.8787,
  kB = 2.15506,
  uB = 7.19919,
  theta0 = 0.0749886,
  theta1 = 0.0001191,
  iv0 = 1.09629,
  kv = 2.00048,
```

```
av = 2493.41,
  gammaV = 2.91282,
  fvS = 0.141195,
  pctMort = 0.215,
  phi0 = 0.791666,
  phi1 = 0.000737,
  dCA = 10950,
  IC0 = 18.02366,
  kC = 2.36949,
  uCA = 6.06349,
 PM = 0.774368,
  dCM = 67.6952,
  dVM = 76.8365,
  dVA = 30 * 365,
 PVM = 0.195768,
  uVA = 11.4321,
  tau1 = 0.69,
  tau2 = 2.31,
 muF = 0.132,
 nEIP = 3,
 qEIP = 1/10,
 Q0 = 0.92,
 DY = 365,
  thetaB = 0.89,
  thetaI = 0.97,
  r_{llin} = 0.56,
  s_{llin} = 0.03,
 r_{irs} = 0.6,
  s_{irs} = 0,
 qE = 1/3,
 nE = 2,
 qL = 1/7,
 nL = 3,
 qP = 1/1,
  nP = 2,
 muE = 0.05,
 muL = 0.15,
 muP = 0.05,
 muM = 0.132,
 eps = 58.9,
 nu = 1/(4/24),
 NH = 1000,
)
```

#### **Arguments**

Death rate for expoential population distribtuion, i.e. 1/Mean Population Age. Default = 0.0001305

rho Age-dependent biting parameter. Default = 0.85 a0 Age-dependent biting parameter. Default = 2920

sigma2 Variance of the log heterogeneity in biting rates. Default = 1.67

max\_age Maximum age in days. Default = 100\*365

rA Rate of leaving asymptomatic infection. Default = 0.00512821

rT Rate of leaving treatment. Default = 0.2 rD Rate of leaving clinical disease. Default = 0.2

rU Rate of recovering from subpatent infection. Default = 0.00906627

rP Rate of leaving prophylaxis. Default = 0.06666667

dE Latent period of human infection. Default = 12

delayGam Lag from parasites to infectious gametocytes. Default = 12.5

cD Untreated disease contribution to infectiousness. Default = 0.0676909
 cT Treated disease contribution to infectiousness. Default = 0.322 \* cD
 cU Subpatent disease contribution to infectiousness. Default = 0.006203

gamma1 Parameter for infectiousness of state A. Default = 1.82425

d1 Minimum probability due to maximum immunity. Default = 0.160527

dID Inverse of decay rate. Default = 3650

ID0 Scale parameter. Default = 1.577533

kD Shape parameter. Default = 0.476614

uD Duration in which immunity is not boosted. Default = 9.44512

aD Scale parameter relating age to immunity. Default = 8001.99

fD0 Time-scale at which immunity changes with age. Default = 0.007055

gammaD Shape parameter relating age to immunity. Default = 4.8183

alphaA PCR detection probability parameters state A. Default = 0.757

alphaU PCR detection probability parameters state U. Default = 0.186

b0 Maximum probability due to no immunity. Default = 0.590076

b1 Maximum relative reduction due to immunity. Default = 0.5

dB Inverse of decay rate. Default = 3650

IB0 Scale parameter. Default = 43.8787

kB Shape parameter. Default = 2.15506

uB Duration in which immunity is not boosted. Default = 7.19919

theta0 Maximum probability of severe infection due to no immunity. Default = 0.0749886

theta1 Maximum reduction due to to immunity. Default = 0.0001191

iv0 Scale parameter. Default = 1.09629 kv Shape parameter. Default = 2.00048

av Age-dependent modifier. Default = 2493.41 gammaV Age-dependent modifier. Default = 2.91282

fvS	Age-dependent modifier. Default = $0.141195$
pctMort	Percentage of severe cases that die. Default = $0.215$
phi0	Maximum probability due to no immunity. Default = 0.791666
phi1	Maximum relative reduction due to immunity. Default = $0.000737$
dCA	Inverse of decay rate. Default = 10950
IC0	Scale parameter. Default = 18.02366
kC	Shape parameter. Default = $2.36949$
uCA	Duration in which immunity is not boosted. Default = 6.06349
PM	New-born immunity relative to mother's. Default = $0.774368$
dCM	Inverse of decay rate of maternal immunity. Default = 67.6952
dVM	Inverse of decay rate. Default = 76.8365
dVA	Inverse of decay rate. Default = 30 * 365
PVM	New-born immunity to severe disease relative to mothers. Default = $0.195768$
uVA	Duration in which immunity to severe disease is not boosted. Default = 11.4321
tau1	Duration of host seeking, assumed to be constant between species. Default = 0.69
tau2	Duration of mosquito resting after feed. Default = 2.31
muF	Daily mortality of adult mosquitos. Default = $0.132$
nEIP	Number of Erlang-distributed EIP compartments. Default = 6
qEIP	Inverse of the mean duration of the EIP. Default = 1/10 (days)
Q0	Anthrophagy probability. Default = 0.92
DY	number of days in a year. Default = 365
thetaB	proportion of bites on a person in bed. Default = 0.89
thetaI	proportion of bites on a person outdoors. Default = 0.97
r_llin	probability of repeating a feeding attempt due to LLINs. Default = 0.56
s_llin	probability of feeding and surviving in presence of LLINs. Default = $0.03$
r_irs	probability of repeating a feeding attempt due to IRS. Default = $0.60$
s_irs	probability of feeding and surviving in presence of IRS. Default = $0$
qE	mosquito egg lifecycle parameter. Default = 1/3
nE	mosquito egg lifecycle parameter. Default = 2
qL	mosquito larval lifecycle parameter. Default = 1/7
nL	mosquito larval lifecycle parameter. Default = 3
qP	mosquito pupae lifecycle parameter. Default = 1/1
nP	mosquito pupae lifecycle parameter. Default = 2
muE	death rate of egg stage. Default = $0.05$
muL	death rate of larval stage. Default = $0.15$
muP	death rate of pupae stage. Default = $0.05$

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eps eggs laid per day. Default = 0.132

eps eggs laid per day. Default = 58.9

nu mosquito lifecycle parameter. Default = 1/(4/24

NH number of humans. Default = 1000

... Any other parameters needed for non-standard model. If they share the same name as any of the defined parameters model\_param\_list\_create will stop.

You can either write any extra parameters you like individually, e.g. model\_param\_list\_create(extra1 = 1, extra2 = 2) and these parameters will appear appended to the returned list, or you can pass explicitly the ellipsis argument as a list created before, e.g.

model\_param\_list\_create(...=list(extra1 = 1, extra2 = 2))

#### Value

A named vector of all baseline parameters required by the Imperial malaria model.

This function creates all of the necessary parameters for the Imperial model. Parameters furnished by MGDrivE will be removed from this function. Adapted from: https://github.com/mrc-ide/deterministic-malaria-model/blob/master/R/model\_parameters.R

A newer version of the model also includes parameters for severe disease. See: https://github.com/mrc-ide/malariasimulation for details.

make\_Q\_Imperial Rate Matrix (Q) for Adult Mosquito SEI Dynamics

#### **Description**

Construct the infinitesimal generator matrix for (individual) adult female infection dynamics. Adult females follow SEI (Susceptible-Exposed-Infectious) style dynamics with a Gamma distributed EIP, with a mean duration 1/q and variance 1/nq^2 (following shape-scale parameterization, EIP ~ Gamma(n,1/nq)). This function only constructs the rate matrix for either a single mosquito or cohort that all emerged at the same time (the rate matrix for a population with emergence is infinite in dimension).

## Usage

```
make_Q_Imperial(q, n, mu, FOIv)
```

## Arguments

q related to scale parameter of Gamma distributed EIP (1/q is mean length of EIP)

n shape parameter of Gamma distributed EIP

mu mosquito mortality rate

FOIV equilibrium force of infection on mosquitos

#### Value

rate matrix for a single (emergence) cohort of SEI mosquito

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make\_Q\_SEI

Rate Matrix (Q) for Adult Mosquito SEI Dynamics

## **Description**

Construct the infinitesimal generator matrix for (individual) adult female infection dynamics. Adult females follow SEI (Susceptible-Exposed-Infectious) style dynamics with a Gamma distributed EIP, with a mean duration 1/q and variance 1/nq^2 (following shape-scale parameterization, EIP ~ Gamma(n,1/nq)). This function only constructs the rate matrix for either a single mosquito or cohort that all emerged at the same time (the rate matrix for a population with emergence is infinite in dimension).

#### Usage

```
make_Q_SEI(q, n, mu, c, a, x)
```

## **Arguments**

q		related to scale parameter of Gamma distributed EIP (1/q is mean length of EIP)
n		shape parameter of Gamma distributed EIP
m	u	mosquito mortality rate
С		human to mosquito transmission efficiency
а		human biting rate
Х		prevalence of disease in humans

## Value

rate matrix for a single (emergence) cohort of SEI mosquito

movement\_prob2rate

Convert Stochastic Matrix to Rate Matrix

## Description

Given a stochastic matrix, return the rate matrix (infinitesimal generator) that would generate it when exponentiated over the interval of unit time.

## Usage

```
movement_prob2rate(tau)
```

### **Arguments**

tau

a row normalized stochastic matrix

36 mu\_ts

#### **Details**

Warning: if the matrix provided has diagonal-only rows (i.e., the location is independent), the rate matrix will return 0 in that row, as there is no movement rate that can generate that scenario.

#### Value

a list with two elements: gamma negative diagonal of the rate matrix, mat matrix of row normalized off-diagonal elements

## **Examples**

```
# generate random matrix for example
# This represents a 3-node landscape, with random movement between nodes
moveMat <- matrix(data = runif(n = 9), nrow = 3, ncol = 3)
moveMat <- moveMat/rowSums(moveMat)

moveRate <- movement_prob2rate(tau = moveMat)</pre>
```

 $mu_ts$ 

Mosquito Death Rates, Comoros Islands

#### **Description**

This is a matrix containing estimated mosquito death rates from the Comoros Islands, between Mozambique and Madagascar. It provides hourly death rates over the course of one year.

#### Usage

```
data(mu_ts)
```

## Format

matrix with 3 named columns and 8760 rows:

Grande\_Comore Hourly death rates for main island

Moheli Hourly death rates for second island

Anjouan Hourly death rates for smallest island

```
sim_trajectory_base_CSV
```

Simulate Trajectory From one SPN Model

# Description

This is an internal function to sim\_trajectory\_CSV. It does the actual sampling once all of the functions have been checked and setup.

# Usage

```
sim_trajectory_base_CSV(
   x0,
   times,
   stepFun,
   folders,
   stage,
   events0 = NULL,
   batch = NULL,
   sout = NULL,
   verbose = TRUE
)
```

# Arguments

x0	the initial marking of the SPN (initial state)
times	sequence of sampling times
stepFun	a sampling function
folders	vector of folders to write output
stage	vector of life-stages to print
events0	a data.frame of events (uses the same format as required in package deSolve for consistency, see events for more information)
batch	a list of batch migration events, created from ${\tt batch\_migration},$ may be set to NULL if not used
Sout	an optional matrix to track event firings
verbose	print a progress bar?

# Value

no return, prints .csv files into provided folders

# Description

This is an internal function to sim\_trajectory\_CSV. It does the actual sampling once all of the functions have been checked and setup.

# Usage

```
sim_trajectory_base_CSV_decoupled(
    x0,
    h0,
    SPN_P,
    theta,
    times,
    stepFun,
    events0 = NULL,
    batch = NULL,
    Sout = NULL,
    verbose = TRUE,
    human_ode = "Imperial",
    cube = NULL,
    folders = folders
)
```

x0	the initial marking of the SPN (initial state)
h0	initial human state distribution
SPN_P	stochastic petri net, places
theta	parameters
times	sequence of sampling times
stepFun	a sampling function
events0	a data.frame of events (uses the same format as required in package deSolve for consistency, see events for more information)
batch	a list of batch migration events, created from batch_migration, may be set to NULL if not used
Sout	an optional matrix to track event firings
verbose	print a progress bar?
human_ode	ode function used for human states
cube	inheritance cube
folders	vector of folders to write output

sim\_trajectory\_base\_R

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# Value

no return, prints .csv files into provided folders

```
\verb|sim_trajectory_base_R| \textit{Simulate Trajectory From one SPN Model}|
```

# Description

This is an internal function to sim\_trajectory\_R. It does the actual sampling once all of the functions have been checked and setup.

# Usage

```
sim_trajectory_base_R(
    x0,
    times,
    num_reps,
    stepFun,
    events = NULL,
    batch = NULL,
    Sout = NULL,
    verbose = TRUE
)
```

# Arguments

x0	the initial marking of the SPN (initial state)
times	sequence of sampling times
num_reps	number of repetitions to run
stepFun	a sampling function
events	a data.frame of events (uses the same format as required in package deSolve for consistency, see events for more information)
batch	a list of batch migration events, created from ${\tt batch\_migration},$ may be set to NULL if not used
Sout	an optional matrix to track event firings
verbose	print a progress bar?

# Value

matrix of sampled values

```
\begin{tabular}{ll} sim\_trajectory\_base\_R\_decoupled\_Imperial\\ Simulate\ Trajectory\ From\ one\ SPN\ Model\ using\ Imperial\ Malaria\\ model \end{tabular}
```

# Description

This is an internal function to sim\_trajectory\_R. It does the actual sampling once all of the functions have been checked and setup.

# Usage

```
sim_trajectory_base_R_decoupled_Imperial(
    x0,
    h0,
    SPN_P,
    theta,
    times,
    num_reps,
    stepFun,
    events = NULL,
    batch = NULL,
    verbose = TRUE,
    cube = NULL
)
```

x0	the initial marking of the SPN (initial state)
h0	initial human state distribution
SPN_P	stochastic petri net, places
theta	parameters
times	sequence of sampling times
num_reps	number of repetitions to run
stepFun	a sampling function
events	a data.frame of events (uses the same format as required in package deSolve for consistency, see events for more information)
batch	a list of batch migration events, created from ${\tt batch\_migration},$ may be set to NULL if not used
Sout	an optional matrix to track event firings
verbose	print a progress bar?
cube	inheritance cube

# Value

matrix of sampled values

```
sim_trajectory_base_R_decoupled_SIS

Simulate Trajectory From one SPN Model using Human SIS model
```

# **Description**

This is an internal function to sim\_trajectory\_R. It does the actual sampling once all of the functions have been checked and setup.

# Usage

```
sim_trajectory_base_R_decoupled_SIS(
    x0,
    h0,
    SPN_P,
    theta,
    times,
    num_reps,
    stepFun,
    events = NULL,
    batch = NULL,
    verbose = TRUE,
    cube = NULL
)
```

x0	the initial marking of the SPN (initial state)
h0	initial human state distribution
SPN_P	stochastic petri net, places
theta	parameters
times	sequence of sampling times
num_reps	number of repetitions to run
stepFun	a sampling function
events	a data.frame of events (uses the same format as required in package deSolve for consistency, see events for more information)
batch	a list of batch migration events, created from batch_migration, may be set to NULL if not used
Sout	an optional matrix to track event firings
verbose	print a progress bar?
cube	inheritance cube

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# Value

matrix of sampled values

sim\_trajectory\_CSV Simulate Trajectory From a SPN Model

# Description

This function provides a unified interface to the various simulation algorithms for SPN, returning output sampled at a lattice of time points to the user, and handling various exogenous events that may occur during the simulation (such as release of adult mosquitoes).

# Usage

```
sim_trajectory_CSV(
  χ0,
  tmax,
  dt = 1,
 dt_stoch = 0.1,
  folders = "./",
 stage = c("M", "F"),
  S,
 hazards,
 Sout = NULL,
  sampler = "tau",
 method = "lsoda",
 events = NULL,
 batch = NULL,
  verbose = TRUE,
)
```

x0	the initial marking of the SPN (initial state, M0)
tmax	the final time to end simulation
dt	the time-step at which to return output ( $\mathbf{not}$ the time-step of the sampling algorithm)
dt_stoch	time-step used for approximation of hazards
folders	vector of folders to write output
stage	life-stages to print. Any combination of: "E", "L", "P"," M", "U", "F", "H"
S	a stoichiometry Matrix-class object
hazards	list of hazard functions
Sout	an optional matrix to track event firings

sim\_trajectory\_CSV 43

sampler determines sampling algorithm, one of; "ode", "tau", "cle", or "dm"

method if sampler is "ode", the solver to use, from deSolve

events a data. frame of events

batch a list of batch migration events, created from batch\_migration, may be set

to NULL if not used

verbose print a progress bar?

... further named arguments passed to the step function

#### Details

dt\_stoch is used by the Poisson Time-Step (step\_PTS) and Chemical Langevin (step\_CLE) methods to approximate the hazards. A smaller dt\_stoch provides a better approximation, but will take longer to run.

The stoichiometry matrix (S) is generated in spn\_S.

The list of hazards (hazards) come from spn\_hazards.

Several samplers are provided. The default is a Poisson Time-Step (step\_PTS) method. Other options are Gillespie's Direct Method (step\_DM) and a Chemical Langevin sampler (step\_CLE). Additionally, for convenience, an ODE "sampler" (step\_ODE) is provided for compatibility with other samplers. This function uses methods from deSolve.

If using the ode sampler, several methods are provided in the deSolve package, see ode. For inhomogeneous systems, consider using the "rk4" method to avoid excessive integration times.

Additionally, events objects must follow the format required by deSolve. This was done for consistency, see events for more information.

This function writes all output to .csv files. Each simulation is written to a folder element - the number of repetitions is the number of folders provided. What life-stages get recorded is specified by the stage parameter. All life-stages can be stored, or any subset thereof. Females are split by infection status, i.e. by "S", "E", or "I".

This function tracks state variables specified by argument stage by default; an optional argument Sout can be provided to track number of event firings each time step (for discrete stochastic simulations), or cumulative intensity (for continuous stochastic simulations), or the rate function of particular events for ODE simulation. The matrix must have number of columns equal to number of events in the system (the number of hazard functions), and a row for each tracking variable. If Sout is provided, it outputs an additional csv, "Tracking.csv". The function track\_hinf is provided, which builds a matrix to track human infection events.

To return simulations to R for further processing, see sim\_trajectory\_R.

#### Value

NULL - prints output to .csv files

## **Description**

This function provides a unified interface to the various simulation algorithms for SPN, returning output sampled at a lattice of time points to the user, and handling various exogenous events that may occur during the simulation (such as release of adult mosquitoes). This function is used in decoupled sampling, where the mosquito and human states are separated. This is used primarily when using the Imperial model of malaria transmission.

# Usage

```
sim_trajectory_CSV_decoupled(
  χ0,
 h0,
  inf_labels,
  tmax,
  dt = 1,
  dt_stoch = 0.1,
  folders = "./",
  S,
  hazards,
  SPN_P,
  theta,
  Sout = NULL,
  sampler = "tau",
 method = "lsoda",
  events = NULL,
 batch = NULL,
  verbose = TRUE,
  human_ode = "Imperial",
  cube = cube,
)
```

x0	the initial marking of the SPN (initial state, M0)
h0	the initial human state distribution
inf_labels	labels corresponding to female mosquito infection hazard
tmax	the final time to end simulation
dt	the time-step at which to return output ( <b>not</b> the time-step of the sampling algorithm)

dt\_stoch time-step used for approximation of hazards

folders vector of folders to write output

S a stoichiometry Matrix-class object

hazards list of hazard functions SPN\_P stochastic petri net places

theta parameters

Sout an optional matrix to track event firings

sampler determines sampling algorithm, one of; "ode", "tau", "cle", or "dm"

method if sampler is "ode", the solver to use, from deSolve

events a data. frame of events

batch a list of batch migration events, created from batch\_migration, may be set

to NULL if not used

verbose print a progress bar?
human\_ode human ode function
cube inheritance cube

... further named arguments passed to the step function

#### **Details**

dt\_stoch is used by the Poisson Time-Step (step\_PTS) and Chemical Langevin (step\_CLE) methods to approximate the hazards. A smaller dt\_stoch provides a better approximation, but will take longer to run.

The stoichiometry matrix (S) is generated in spn\_S.

The list of hazards (hazards) come from spn\_hazards.

Several samplers are provided. The default is a Poisson Time-Step (step\_PTS) method. Other options are Gillespie's Direct Method (step\_DM) and a Chemical Langevin sampler (step\_CLE). Additionally, for convenience, an ODE "sampler" (step\_ODE) is provided for compatibility with other samplers. This function uses methods from deSolve.

If using the ode sampler, several methods are provided in the deSolve package, see ode. For inhomogeneous systems, consider using the "rk4" method to avoid excessive integration times.

Additionally, events objects must follow the format required by deSolve. This was done for consistency, see events for more information.

This function writes all output to .csv files. Each simulation is written to a folder element - the number of repetitions is the number of folders provided. For now, only adult mosquito states, human states, clinical incidence, and pathogen prevalence are written to CSVs.

This function tracks state variables specified by argument stage by default; an optional argument Sout can be provided to track number of event firings each time step (for discrete stochastic simulations), or cumulative intensity (for continuous stochastic simulations), or the rate function of particular events for ODE simulation. The matrix must have number of columns equal to number of events in the system (the number of hazard functions), and a row for each tracking variable. If Sout is provided, it outputs an additional csv, "Tracking.csv". The function track\_hinf is provided, which builds a matrix to track human infection events.

To return simulations to R for further processing, see sim\_trajectory\_R.

sim\_trajectory\_R

# Value

NULL - prints output to .csv files

sim\_trajectory\_R

Simulate Trajectory From a SPN Model

# **Description**

This function provides a unified interface to the various simulation algorithms for SPN, returning output sampled at a lattice of time points to the user, and handling various exogenous events that may occur during the simulation (such as release of adult mosquitoes).

# Usage

```
sim_trajectory_R(
    x0,
    tmax,
    dt = 1,
    dt_stoch = 0.1,
    num_reps = 1,
    S,
    hazards,
    Sout = NULL,
    sampler = "tau",
    method = "lsoda",
    events = NULL,
    batch = NULL,
    verbose = TRUE,
    ...
)
```

x0	the initial marking of the SPN (initial state, M0)
tmax	the final time to end simulation (all simulations start at 0)
dt	the time-step at which to return output ( $\mathbf{not}$ the time-step of the sampling algorithm)
dt_stoch	time-step used for approximation of hazards
num_reps	number of repetitions to run, default is 1.
S	a stoichiometry Matrix-class object
hazards	list of hazard functions
Sout	an optional matrix to track event firings
sampler	determines sampling algorithm, one of; "ode", "tau", "cle", or "dm"
method	if sampler is "ode", the solver to use, from deSolve

events a data. frame of events, may be set to NULL if not used

batch a list of batch migration events, created from batch\_migration, may be set

to NULL if not used

verbose print a progress bar?

... further named arguments passed to the step function

#### **Details**

dt\_stoch is used by the Poisson Time-Step (step\_PTS) and Chemical Langevin (step\_CLE) methods to approximate the hazards. A smaller dt\_stoch provides a better approximation, but will take longer to run.

The stoichiometry matrix (S) is generated in spn\_S.

The list of hazards (hazards) come from spn\_hazards.

Several samplers are provided. The default is a Poisson Time-Step (step\_PTS) method. Other options are Gillespie's Direct Method (step\_DM) and a Chemical Langevin sampler (step\_CLE). Additionally, for convenience, an ODE "sampler" (step\_ODE) is provided for compatibility with other samplers. This function uses methods from deSolve.

If using the ode sampler, several methods are provided in the deSolve package, see ode. For inhomogeneous systems, consider using the "rk4" method to avoid excessive integration times.

Additionally, events objects must follow the format required by deSolve. This was done for consistency, see events for more information.

This function tracks state variables by default; an optional argument Sout can be provided to track number of event firings each time step (for discrete stochastic simulations), or cumulative intensity (for continuous stochastic simulations), or the rate function of particular events for ODE simulation. The matrix must have number of columns equal to number of events in the system (the number of hazard functions), and a row for each tracking variable. The function track\_hinf is provided, which builds a matrix to track human infection events.

To save output as .csv files, see sim\_trajectory\_CSV.

### Value

a list with 2 elements: "state" is the array of returned state values, and "events" will return events tracked with Sout if provided, otherwise is NULL

sim\_trajectory\_R\_decoupled

Simulate Trajectory From a SPN Model

## **Description**

This function provides a unified interface to the various simulation algorithms for SPN, returning output sampled at a lattice of time points to the user, and handling various exogenous events that may occur during the simulation (such as release of adult mosquitoes).

## Usage

```
sim_trajectory_R_decoupled(
  χ0,
  h0,
  tmax,
  inf_labels,
 dt = 1,
  dt_stoch = 0.1,
  num\_reps = 1,
  hazards,
  SPN_P,
  theta,
  Sout = NULL,
  sampler = "tau",
 method = "lsoda",
  events = NULL,
  batch = NULL,
  verbose = TRUE,
 human_ode = "Imperial",
  cube = cube,
)
```

# **Arguments**

x0 the initial marking of the SPN (initial state, M0) the initial human state distribution h0 the final time to end simulation (all simulations start at 0) tmax inf\_labels labels corresponding to female mosquito infection hazard dt the time-step at which to return output (not the time-step of the sampling algorithm) dt\_stoch time-step used for approximation of hazards number of repetitions to run, default is 1. num\_reps S a stoichiometry Matrix-class object list of hazard functions hazards SPN\_P stochastic petri net places theta parameters Sout an optional matrix to track event firings sampler determines sampling algorithm, one of; "ode", "tau", "cle", or "dm" method if sampler is "ode", the solver to use, from deSolve a data. frame of events, may be set to NULL if not used events a list of batch migration events, created from batch\_migration, may be set batch to NULL if not used

solve\_muAqua 49

verbose print a progress bar?
human\_ode human ode function
cube inheritance cube

... further named arguments passed to the step function

#### **Details**

dt\_stoch is used by the Poisson Time-Step (step\_PTS) and Chemical Langevin (step\_CLE) methods to approximate the hazards. A smaller dt\_stoch provides a better approximation, but will take longer to run.

The stoichiometry matrix (S) is generated in spn\_S.

The list of hazards (hazards) come from spn\_hazards.

Several samplers are provided. The default is a Poisson Time-Step (step\_PTS) method. Other options are Gillespie's Direct Method (step\_DM) and a Chemical Langevin sampler (step\_CLE). Additionally, for convenience, an ODE "sampler" (step\_ODE) is provided for compatibility with other samplers. This function uses methods from deSolve.

If using the ode sampler, several methods are provided in the deSolve package, see ode. For inhomogeneous systems, consider using the "rk4" method to avoid excessive integration times.

Additionally, events objects must follow the format required by deSolve. This was done for consistency, see events for more information.

This function tracks state variables by default; an optional argument Sout can be provided to track number of event firings each time step (for discrete stochastic simulations), or cumulative intensity (for continuous stochastic simulations), or the rate function of particular events for ODE simulation. The matrix must have number of columns equal to number of events in the system (the number of hazard functions), and a row for each tracking variable. The function track\_hinf is provided, which builds a matrix to track human infection events.

To save output as .csv files, see sim\_trajectory\_CSV.

#### Value

a list with 2 elements: "state" is the array of returned state values, and "events" will return events tracked with Sout if provided, otherwise is NULL

solve\_muAqua Solve for Constant Aquatic Mortality

## **Description**

In MGDrivE, the model was typically solved at equilibrium assuming the density-independent mortality was constant over aquatic stages (eggs, larvae, pupae), given a daily growth rate,  $r_M$ . Given that growth rate, it solved for that mortality  $\mu_{Aqua}$  by relating it with  $R_M$ , the per-generation growth rate of the population, calculable from  $r_M$  and the mean duration of life stages. This function uses uniroot to solve for  $mu_{Aqua}$ .

split\_aggregate\_CSV

## Usage

```
solve_muAqua(params, rm)
```

# Arguments

params a named list of parameters rm the daily growth rate

#### **Details**

This function needs the following parameters in params:

- muF: adult female mortality
- beta: rate of egg laying
- phi: sex ratio at emergence
- qE: inverse of mean duration of egg stage
- nE: shape parameter of Erlang-distributed egg stage
- qL: inverse of mean duration of larval stage
- nL: shape parameter of Erlang-distributed larval stage
- qP: inverse of mean duration of pupal stage
- nP: shape parameter of Erlang-distributed pupal stage

### Value

location of the root, as provided from uniroot

# **Examples**

 $split\_aggregate\_CSV$ 

Split CSV output by Patch and Aggregate by Mate or Dwell-Stage

# Description

This function reads in the output files from sim\_trajectory\_CSV and splits them into smaller files. The files are output by patch, with the appropriate patch numbers for mosquitoes or humans, and specific stages are aggregated by a given metric.

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# Usage

```
split_aggregate_CSV(
  read_dir,
  write_dir = read_dir,
  stage = c("E", "L", "P", "M", "U", "FS", "FE", "FI", "H"),
  spn_P,
  tmax,
  dt,
  erlang = FALSE,
  sum_fem = FALSE,
  rem_file = FALSE,
  verbose = TRUE
)
```

# Arguments

read_dir	Directory where output was written to
write_dir	Directory to write output to. Default is read_dir
stage	Life stage to print, see details
spn_P	Places object, see details
tmax	The final time to end simulation
dt	The time-step at which to return output ( <b>not</b> the time-step of the sampling algorithm)
erlang	Boolean, default is FALSE, to return summaries by genotype
sum_fem	if TRUE, in addition to FS, FE, FI output by node and repetition, output an additional file F which sums over infection states (S,E,I). Does nothing if the simulation did not include epi dynamics.
rem_file	Remove original output? Default is FALSE
verbose	Chatty? Default is TRUE

# **Details**

Given the read\_dir, this function assumes the follow file structure:

```
    read_dir

            repetition 1
            * M.csv
            * FS.csv
            * ...

    repetition 2
    * M.csv
```

\* FS.csv

\* ...
- repetition 3
- ...

This function expects the write\_dir to be empty, and it sets up the same file structure as the read\_dir. For a 2-node simulation, the output will be organized similar to:

```
write_dir
repetition 1
* M_0001.csv
* M_0002.csv
* FS_0001.csv
* FS_0001.csv
* ...
repetition 2
* M_0001.csv
* M_0002.csv
* FS_0001.csv
* FS_0001.csv
* ...
repetition 3
...
```

stage defines which life-stages the function will analyze. These stages must be any combination of: "E", "L", "P", "M", "U", "FS", "FE", "FI", "H". These must come from the set of stages provided to sim\_trajectory\_CSV via the stage argument. It can be less than what was printed by the simulation, but any extra stages provided, but not printed, will throw a warning and then be ignored.

erlang defines how aquatic (eggs, larvae, and pupae) stages and adult females (only mated females) are aggregated. By default, erlang is FALSE, and all of these stages are summarized by genotype only, combining any Erlang-distributed dwell stages (for eggs, larvae, and pupae) or latent infection (for adult females) stages. If erlang is TRUE, summaries are returned by dwell stage or infection status, combining any genotype information.

Female summaries always combine over mate-genotype, so only female genotypes are returned.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

tmax, dt define the last sampling time, and each sampling time in-between.

For more details about using this function to process CSV output see: vignette("data-analysis", package = "MGDrivE2")

## Value

Writes output to files in write\_dir

```
split_aggregate_CSV_decoupled
```

Split CSV output for decoupled sampling with Imperial malaria model

# **Description**

This function reads in the output files from sim\_trajectory\_CSV and splits them into smaller files. The files are output by patch, with the appropriate patch numbers for mosquitoes or humans, and specific stages are aggregated by a given metric.

# Usage

```
split_aggregate_CSV_decoupled(
  read_dir,
  write_dir = read_dir,
  spn_P,
  tmax,
  dt,
  human_states,
  sum_fem = FALSE,
  rem_file = FALSE,
  verbose = TRUE,
  erlang = FALSE
)
```

# Arguments

read_dir	Directory when	re output was	written	to

write\_dir Directory to write output to. Default is read\_dir

spn\_P Places object, see details

tmax The final time to end simulation

dt The time-step at which to return output (**not** the time-step of the sampling algo-

rithm)

human\_states human state distribution

sum\_fem if TRUE, in addition to FS, FE, FI output by node and repetition, output an addi-

tional file F which sums over infection states (S,E,I). Does nothing if the simu-

lation did not include epi dynamics.

rem\_file Remove original output? Default is FALSE

verbose Chatty? Default is TRUE erlang erlang distributed states

## **Details**

Given the read\_dir, this function assumes the follow file structure:

```
    read_dir

            repetition 1
            * M.csv
            * FS.csv
            * ...

    repetition 2

            * M.csv
            * FS.csv
            * ...

    repetition 3

            ...
```

This function expects the write\_dir to be empty, and it sets up the same file structure as the read\_dir. For a 2-node simulation, the output will be organized similar to:

```
write_dir
repetition 1
* M_0001.csv
* M_0002.csv
* FS_0001.csv
* FS_0001.csv
* ...
repetition 2
* M_0001.csv
* M_0002.csv
* FS_0001.csv
* FS_0001.csv
* ...
repetition 3
- ...
```

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

tmax, dt define the last sampling time, and each sampling time in-between.

For more details about using this function to process CSV output see: vignette("data-analysis", package = "MGDrivE2")

spn\_hazards 55

# Value

Writes output to files in write\_dir

spn_hazards	spn_hazards	
-------------	-------------	--

# Description

Using the structural (topological) SPN model as well as parameters in the cube and params objects, generate a list (of length |v|) of hazards, each implemented as a function closure.

# Usage

```
spn_hazards(
  spn_P,
  spn_T,
  cube,
  params,
  type = "life",
  log_dd = TRUE,
  exact = TRUE,
  tol = 1e-12,
  verbose = TRUE
```

spn_P	the set of places (P) (see details)
spn_T	the set of transitions (T) (see details)
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
params	a named list of parameters (see details)
type	string indicating type of hazards, one of; "life", "SIS", or "SEIR"
log_dd	if TRUE, use logistic (carrying capacity) density dependent hazards, if FALSE use Lotka-Volterra density dependent hazards for larval mortality
exact	boolean, make exact (integer input) hazards? Default is TRUE
tol	if exact=FALSE, the value of hazard below which it is clipped to $\boldsymbol{0}$
verbose	display a progress bar when making hazards?

#### **Details**

If these hazards will be used in a continuous approximation algorithm, such as an ODE method (step\_ODE) or Gillespie's Direct Method (step\_DM), it is recommended to use exact=FALSE. If the hazards will be used in an integer state space method, such as tau-leaping (step\_PTS) or Chemical Langevin (step\_CLE) methods, it is recommended to use exact=TRUE.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The set of transitions (spn\_T) is generated from one of the following: spn\_T\_lifecycle\_node, spn\_T\_lifecycle\_network, spn\_T\_epiSIS\_node, spn\_T\_epiSIS\_network, spn\_T\_epiSEIR\_node, spn\_T\_epiSEIR\_network.

The params objected is generated from either equilibrium\_lifeycle or equilibrium\_SEI\_SIS; it is the "params" object in the return list. The equilibrium function used must match the type parameter.

The type parameter indicates what type of simulation is being run. It is one of: "life", "SIS", or "SEIR". This must match the params object supplied.

Use of this function is demonstrated in many vignettes, browseVignettes(package = "MGDrivE2")

## Value

list of length 2: hazards is a list of named closures for every state transition in the model, flag is a boolean indicating exact or approximate

spn\_hazards\_decoupled Make Hazards (Lambda) For a MGDrivE2: Node and Network Simulations

# Description

Using the structural (topological) SPN model as well as parameters in the cube and params objects, generate a list (of length |v|) of hazards, each implemented as a function closure.

#### **Usage**

```
spn_hazards_decoupled(
   spn_P,
   spn_T,
   cube,
   params,
   type = "SIS",
   log_dd = TRUE,
   exact = TRUE,
   tol = 1e-12,
   verbose = TRUE
)
```

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# **Arguments**

spn_P	the set of places (P) (see details)
spn_T	the set of transitions (T) (see details)
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
params	a named list of parameters (see details)
type	string indicating type of hazards, one of; "life", "SIS", "Imperial" or "SEIR"
log_dd	if TRUE, use logistic (carrying capacity) density dependent hazards, if FALSE use Lotka-Volterra density dependent hazards for larval mortality
exact	boolean, make exact (integer input) hazards? Default is TRUE
tol	if exact=FALSE, the value of hazard below which it is clipped to $\boldsymbol{0}$
verbose	display a progress bar when making hazards?

#### **Details**

If these hazards will be used in a continuous approximation algorithm, such as an ODE method (step\_ODE) or Gillespie's Direct Method (step\_DM), it is recommended to use exact=FALSE. If the hazards will be used in an integer state space method, such as tau-leaping (step\_PTS) or Chemical Langevin (step\_CLE) methods, it is recommended to use exact=TRUE.

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The set of transitions (spn\_T) is generated from one of the following: spn\_T\_lifecycle\_node, spn\_T\_lifecycle\_network, spn\_T\_epiSIS\_node, spn\_T\_epiSIS\_network, spn\_T\_epiSEIR\_node, spn\_T\_epiSEIR\_network.

The params objected is generated from either equilibrium\_lifeycle or equilibrium\_SEI\_SIS; it is the "params" object in the return list. The equilibrium function used must match the type parameter.

The type parameter indicates what type of simulation is being run. It is one of: "life", "SIS", or "SEIR". This must match the params object supplied.

Use of this function is demonstrated in many vignettes, browseVignettes(package = "MGDrivE2")

## Value

list of length 2: hazards is a list of named closures for every state transition in the model, flag is a boolean indicating exact or approximate

spn_Post	Make Post Matrix For a Petri Net	
----------	----------------------------------	--

### **Description**

Generate the Post (|v| by |u|) matrix for the SPN. This gives the edges from T to P (output arcs) in the bipartite network.

58 spn\_Pre

#### Usage

```
spn_Post(spn_P, spn_T)
```

## **Arguments**

```
spn_P set of places (P) (see details)
spn_T set of transitions (T) (see details)
```

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The set of transitions (spn\_T) is generated from one of the following: spn\_T\_lifecycle\_node, spn\_T\_lifecycle\_network, spn\_T\_epiSIS\_node, spn\_T\_epiSIS\_network, spn\_T\_epiSEIR\_node, spn\_T\_epiSEIR\_network.

## Value

a matrix of type dgCMatrix-class

spn\_Pre

Make Pre Matrix For a Petri Net

## **Description**

Generate the Pre (|v| by |u|) matrix for the SPN. This gives the edges from P to T (input arcs) in the bipartite network.

#### **Usage**

```
spn_Pre(spn_P, spn_T)
```

# Arguments

```
spn_P set of places (P) (see details)
spn_T set of transitions (T) (see details)
```

## **Details**

```
The places (spn_P) object is generated from one of the following: spn_P_lifecycle_node, spn_P_lifecycle_network, spn_P_epiSIS_node, spn_P_epiSIS_network, spn_P_epiSEIR_node, or spn_P_epiSEIR_network.

The set of transitions (spn_T) is generated from one of the following: spn_T_lifecycle_node,
```

```
spn_T_lifecycle_network, spn_T_epiSIS_node, spn_T_epiSIS_network, spn_T_epiSEIR_node, spn_T_epiSEIR_network.
```

#### Value

```
a matrix of type dgCMatrix-class
```

spn\_P\_epiSEIR\_network Make Places (P) For a Network (SEI Mosquitoes - SEIR Humans)

## **Description**

This function makes the set of places (P) for a SPN model of a metapopulation network for simulation of coupled SEI-SEIR dynamics. It is the network version of spn\_P\_epiSEIR\_node.

## Usage

```
spn_P_epiSEIR_network(node_list, params, cube)
```

## **Arguments**

node\_list a character vector specifying what type of nodes to create; (m = a node with

only mosquitoes, h = a node with only humans, b = a node with both humans

and mosquitoes)

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SEIR

For examples of using this function, see: vignette("seir-dynamics", package = "MGDrivE2")

## Value

a list with two elements: ix contains labeled indices of the places by life stage and node, u is the character vector of places (P)

spn\_P\_epiSEIR\_node Make Places (P) For a Node (SEI Mosquitoes - SEIR Humans)

## **Description**

This function makes the set of places (P) for a SPN. It is used alone if our model is a single-node metapopulation for mosquito SEI and human SEIR dynamics; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, spn\_P\_epiSEIR\_network).

## Usage

```
spn_P_epiSEIR_node(params, cube)
```

## Arguments

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

## **Details**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SEIR

For examples of using this function, see: vignette("seir-dynamics", package = "MGDrivE2")

#### Value

a list with two elements: ix contains labeled indices of the places by life stage, u is the character vector of places (P)

## **Description**

This function makes the set of places (P) for a SPN model of a metapopulation network for simulation of coupled SEI-SIS dynamics. It is the network version of spn\_P\_epiSIS\_node.

#### **Usage**

```
spn_P_epiSIS_network(node_list, params, cube)
```

## **Arguments**

node\_list a character vector specifying what type of nodes to create; (m = a node\_id with

only mosquitoes, h = a node\_id with only humans, b = a node\_id with both

humans and mosquitoes)

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SIS

For examples of using this function, see: vignette("epi-network", package = "MGDrivE2")

# Value

a list with two elements: ix contains labeled indices of the places by life stage and node\_id, u is the character vector of places (P)

spn\_P\_epiSIS\_node 61

spn\_P\_epiSIS\_node

Make Places (P) For a Node (SEI Mosquitoes - SIS Humans)

#### **Description**

This function makes the set of places (P) for a SPN. It is used alone if our model is a single-node metapopulation for mosquito SEI and human SIS dynamics; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, see spn\_P\_epiSIS\_network).

#### Usage

```
spn_P_epiSIS_node(params, cube)
```

## **Arguments**

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

#### **Details**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SIS

For examples of using this function, see: vignette("epi-node", package = "MGDrivE2")

## Value

a list with two elements: ix contains labeled indices of the places by life stage, u is the character vector of places (P)

```
spn_P_epi_decoupled_node
```

Make Places (P) For a Node (SEI Mosquitoes). Note in the v2 epi module, we only use the SPN framework for the mosquito component of the model. The human component will be handled separately in the sampler, and will be formulated as an ODE. This function makes the set of places (P) for a SPN. It is used alone if our model is a single-node metapopulation for mosquito SEI and dynamics; This is used by both SIS and Imperial transmission models.

## **Description**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SIS

## Usage

```
spn_P_epi_decoupled_node(params, cube)
```

## **Arguments**

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

## **Details**

For examples of using this function, see: vignette("epi-node", package = "MGDrivE2")

#### Value

a list with two elements: ix contains labeled indices of the places by life stage, u is the character vector of places (P)

```
spn_P_lifecycle_network
```

Make Places (P) For a Network (Mosquitoes only)

## **Description**

This function makes the set of places (P) for a SPN model of a metapopulation network. It is the network version of spn\_P\_lifecycle\_node.

## Usage

```
spn_P_lifecycle_network(num_nodes, params, cube)
```

# Arguments

num\_nodes number of nodes in the network

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

# **Details**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, and nP parameters to be specified. For more details, see equilibrium\_lifeycle

For examples of using this function, see: vignette("lifecycle-network", package = "MGDrivE2")

## Value

a list with two elements: ix contains labeled indices of the places by life stage and node\_id, u is the character vector of places (P)

spn\_P\_lifecycle\_node 63

## Description

This function makes the set of places (P) for a SPN. It is used alone if our model is a single-node metapopulation for mosquito dynamics only; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, see spn\_P\_lifecycle\_network).

# Usage

```
spn_P_lifecycle_node(params, cube)
```

## **Arguments**

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

## **Details**

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, and nP parameters to be specified. For more details, see equilibrium\_lifeycle

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

## Value

a list with two elements: ix contains labeled indices of the places by life stage, u is the character vector of places (P)

spn\_S

Make stoichiometry Matrix For a Petri Net

## **Description**

Generate the stoichiometry (lul by lvl) matrix for the SPN. Each column gives the net effect of that transition firing upon the state space of the model. Internally, this creates a Pre (spn\_Pre) and Post (spn\_Post) matrix, and then calculates the final stoichiometry.

# Usage

```
spn_S(spn_P, spn_T)
```

## **Arguments**

```
spn_P set of places (P) (see details)
spn_T set of transitions (T) (see details)
```

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The set of transitions (spn\_T) is generated from one of the following: spn\_T\_lifecycle\_node, spn\_T\_lifecycle\_network, spn\_T\_epiSIS\_node, spn\_T\_epiSIS\_network, spn\_T\_epiSEIR\_node, spn\_T\_epiSEIR\_network.

#### Value

stoichiometry matrix representing the net effect of a transition in the SPN state model.

```
spn_T_epiSEIR_network Make Transitions (T) For a Network (SEI Mosquitoes - SEIR Humans)
```

# Description

This function makes the set of transitions (T) for a SPN model of a metapopulation network for simulation of coupled SEI-SEIR dynamics. It is the network version of spn\_T\_epiSEIR\_node.

## Usage

```
spn_T_epiSEIR_network(node_list, spn_P, params, cube, h_move, m_move)
```

node_list	a character vector specifying what type of nodes to create; $(m = a \text{ node with only mosquitoes}, h = a \text{ node with only humans}, b = a \text{ node with both humans and mosquitoes})$
spn_P	set of places produced by spn_P_epiSEIR_network
params	a named list of parameters (see details)
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
h_move	binary adjacency matrix indicating if movement of humans between nodes is possible or not
m_move	binary adjacency matrix indicating if movement of mosquitoes between nodes is possible or not

 $spn_TepiSEIR_node$  65

#### **Details**

This function takes the places produced from spn\_P\_epiSEIR\_network and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SEIR

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards.

For larger networks, this function may take some time to return, please be patient; the Petri Net modeling formalism trades additional computation time at model initialization for faster sampling of trajectories within a simulation.

Please note, the movement matrices (h\_move and m\_move) are NOT stochastic matrices, just binary matrices that say if i,j can exchange population. Diagonal elements must be FALSE, and both matrices are checked for validity; the function will stop with errors if the adjacency matrix specifies illegal movement rules (e.g.; mosquito movement from a "h" node to a "b" node)

For examples of using this function, see: vignette("seir-dynamics", package = "MGDrivE2")

#### Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions (T)

```
spn_T_epiSEIR_node
```

Make Transitions (T) For a Node (SEI Mosquitoes - SEIR Humans)

# **Description**

This function makes the set of transitions (T) for a SPN. It is used alone if our model is a single-node metapopulation of mosquito and human dynamics; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, see spn\_T\_epiSEIR\_network).

### Usage

```
spn_T_epiSEIR_node(spn_P, params, cube)
```

```
spn_P set of places produced by spn_P_epiSEIR_node

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
```

## **Details**

This function takes the places produced from spn\_P\_epiSEIR\_node and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SEIR

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards.

For examples of using this function, see: vignette("seir-dynamics", package = "MGDrivE2")

## Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions (T)

```
\verb|spn_T_epiSIS_network| & \textit{Make Transitions (T) For a Network (SEI Mosquitoes - SIS Humans)}|
```

## Description

This function makes the set of transitions (T) for a SPN model of a metapopulation network for simulation of coupled SEI-SIS dynamics. It is the network version of spn\_T\_epiSIS\_node.

# Usage

```
spn_T_epiSIS_network(node_list, spn_P, params, cube, h_move, m_move)
```

node_list	a character vector specifying what type of nodes to create; ( $m = a$ node with only mosquitoes, $h = a$ node with only humans, $b = a$ node with both humans and mosquitoes)
spn_P	set of places produced by spn_P_epiSIS_network
params	a named list of parameters (see details)
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
h_move	binary adjacency matrix indicating if movement of humans between nodes is possible or not
m_move	binary adjacency matrix indicating if movement of mosquitoes between nodes is possible or not

 $spn_TepiSIS_node$  67

#### **Details**

This function takes the places produced from spn\_P\_epiSIS\_network and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SIS

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards.

For larger networks, this function may take some time to return, please be patient; the Petri Net modeling formalism trades additional computation time at model initialization for faster sampling of trajectories within a simulation.

Please note, the movement matrices (h\_move and m\_move) are NOT stochastic matrices, just binary matrices that say if i,j can exchange population. Diagonal elements must be FALSE, and both matrices are checked for validity; the function will stop with errors if the adjacency matrix specifies illegal movement rules (e.g.; mosquito movement from a "h" node to a "b" node)

For examples of using this function, see: vignette("epi-network", package = "MGDrivE2")

#### Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions (T)

```
spn_T_epiSIS_node
```

*Make Transitions (T) For a Node (SEI Mosquitoes - SIS Humans)* 

# **Description**

This function makes the set of transitions (T) for a SPN. It is used alone if our model is a single-node metapopulation of mosquito and human dynamics; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, see spn\_T\_epiSIS\_network).

### Usage

```
spn_T_epiSIS_node(spn_P, params, cube)
```

```
spn_P set of places produced by spn_P_epiSIS_node

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
```

#### **Details**

This function takes the places produced from spn\_P\_epiSIS\_node and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SIS

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards.

For examples of using this function, see: vignette("epi-node", package = "MGDrivE2")

#### Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions T

```
spn_T_epi_decoupled_node

Make Transitions (T) For a Node (SEI Mosquitoes)
```

# **Description**

This function makes the set of transitions (T) for a SPN. It is used alone if our model is a single-node metapopulation of mosquito; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, see spn\_T\_epiSIS\_network).

#### **Usage**

```
spn_T_epi_decoupled_node(spn_P, params, cube)
```

# Arguments

spn\_P set of places produced by spn\_P\_epiSIS\_node\_decoupled function
params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

## **Details**

This function takes the places produced from spn\_P\_epiSIS\_node\_decoupled and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, nP, and nEIP parameters to be specified. For more details, see equilibrium\_SEI\_SIS

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards. This is used by both decoupled SIS and Imperial transmission model sampling. For examples of using this function, see: vignette("epi-node-decoupled", package = "MGDrivE2")

#### Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions (T)

```
spn_T_lifecycle_network

Make Transitions (T) For a Network (Mosquitoes only)
```

## **Description**

This function makes the set of transitions (T) for a SPN model of a metapopulation network. It is the network version of spn\_T\_lifecycle\_node.

#### **Usage**

```
spn_T_lifecycle_network(spn_P, params, cube, n = NULL, m_move = NULL)
```

## **Arguments**

spn_P	set of places produced by spn_P_lifecycle_network
params	a named list of parameters (see details)
cube	an inheritance cube from the MGDrivE package (e.g. cubeMendelian)
n	an integer giving the number of nodes
m_move	binary adjacency matrix indicating if movement of mosquitoes between nodes is possible or not

### Details

This function takes the places produced from spn\_P\_lifecycle\_network and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, and nP parameters to be specified. For more details, see equilibrium\_lifeycle

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards.

For larger networks, this function may take some time to return, please be patient; the Petri Net modeling formalism trades additional computation time at model initialization for faster sampling of trajectories within a simulation.

Please note, the movement matrix  $(m\_move)$  is NOT a stochastic matrices, just a binary matrix that say if i,j can exchange population. Diagonal elements must be FALSE.

At least one of the arguments n and m\_move must be provided. If both are provided n is ignored.

For examples of using this function, see: vignette("lifecycle-network", package = "MGDrivE2")

## Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions (T)

```
spn_T_lifecycle_node Make Transitions (T) For a Node (Mosquitoes only)
```

## **Description**

This function makes the set of transitions (T) for a SPN. It is used alone if our model is a single-node metapopulation for mosquito dynamics only; otherwise it is used as part of other functions to make SPN models with larger state spaces (metapopulation models, see spn\_T\_lifecycle\_network).

# Usage

```
spn_T_lifecycle_node(spn_P, params, cube)
```

## **Arguments**

spn\_P set of places produced by spn\_P\_lifecycle\_node

params a named list of parameters (see details)

cube an inheritance cube from the MGDrivE package (e.g. cubeMendelian)

## **Details**

This function takes the places produced from spn\_P\_lifecycle\_node and builds all possible transitions between subsets of those places.

The params argument supplies all of the ecological parameters necessary to calculate equilibrium values. This function requires the nE, nL, and nP parameters to be specified. For more details, see equilibrium\_lifeycle

While this function produces all structural information related to transitions, hazards are produced by a separate function, spn\_hazards.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

## Value

a list with two elements: T contains transitions packets as lists, v is the character vector of transitions (T)

step\_CLE 71

step_CLE Make Chemical Langevin (CLE) Sampler for a SPN model	
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# **Description**

Make a function closure to implement a chemical Langevin (continuous-state) approximation for a SPN.

# Usage

```
step\_CLE(S, Sout, haz, dt = 0.01, maxhaz = 1e+06)
```

## **Arguments**

S	a stoichiometry Matrix-class object
Sout	an optional matrix to track of event firings. In the continuous stochastic model this will be the approximate cumulative intensity of each event.
haz	a list of hazard functions
dt	time-step for Euler-Maruyama method used to solve the SDE system
maxhaz	maximum allowable hazard

# **Details**

The chemical Langevin approximation is a numerical simulation of a Fokker-Planck approximation to the Master equations (Kolmogorov Forwards Equations) governing the stochastic model; the CLE approximation is a second-order approximation that will get the correct mean and variance but higher order moments will be incorrect.

The design of step\_CLE is from: Wilkinson, D. J. (2011). Stochastic modeling for systems biology. CRC press

Elements of the N list come from two places: The stoichiometry matrix (S) is generated in spn\_S and the hazards (h) come from spn\_hazards.

```
For other samplers, see: step_PTS, step_DM, step_ODE
```

## Value

function closure for use in sim\_trajectory\_R or sim\_trajectory\_CSV

72 step\_ODE

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Make Gillespie's Direct Method (DM) Sampler for a SPN model

# **Description**

Make a function closure to implement Gillespie's Direct Method sampler for a SPN.

# Usage

```
step_DM(S, Sout, haz, maxhaz = 1e+06)
```

# Arguments

S a stoichiometry Matrix-class object
Sout an optional matrix to track of event firings

haz a list of hazard functions maxhaz maximum allowable hazard

#### **Details**

The direct method is an exact sampling algorithm; it simulates each event individually. Because of this it may be extremely slow for non-trivial population sizes, and thus should be used to debug and test rather than for serious Monte Carlo simulation.

The design of step\_DM is from: Wilkinson, D. J. (2011). Stochastic modeling for systems biology. CRC press

Elements of the N list come from two places: The stoichiometry matrix (S) is generated in spn\_S and the hazards (h) come from spn\_hazards.

For other samplers, see: step\_CLE, step\_PTS, step\_ODE

# Value

function closure for use in sim\_trajectory\_R or sim\_trajectory\_CSV

step_ODE	Make Mean-field Approximation (ODE) Numerical Integrator for a
	SPN Model

# **Description**

Make a function closure to implement a first order mean-field ODE approximation for a SPN.

## Usage

```
step_ODE(S, Sout, haz, method = "lsoda")
```

## **Arguments**

S a stoichiometry Matrix-class object

Sout an optional matrix to track of event firings. In the deterministic case it will return

the rate of that event at the end of the time step

haz a list of hazard functions

method a character giving the type of numerical integrator used, the default is "Isoda"

#### **Details**

This method is equivalent to considering the ODEs describing the time evolution of the mean trajectory (first moment) and setting all higher order moments which appear on the right hand side to zero.

The solvers used within can be found in the deSolve package, see ode. For inhomogeneous systems, consider using the "rk4" method to avoid excessive integration times.

The stoichiometry matrix (S) is generated in spn\_S.

The list of hazards (haz) come from spn\_hazards.

For other samplers, see: step\_CLE, step\_PTS, step\_DM

#### Value

function closure for use in sim\_trajectory\_R or sim\_trajectory\_CSV

step_ODE_decoupled	Make Mean-field Approximation (ODE) Numerical Integrator for a
	SPN Model for Decoupled Epi Dynamics

## Description

Make a function closure to implement a first order mean-field ODE approximation for a SPN.

# Usage

```
step_ODE_decoupled(S, Sout, haz, method = "lsoda", human_ode = "SIS")
```

## Arguments

S a stoichiometry Matrix-class ob	1ect
-----------------------------------	------

Sout an optional matrix to track of event firings. In the deterministic case it will return

the rate of that event at the end of the time step

haz a list of hazard functions

method a character giving the type of numerical integrator used, the default is "Isoda"

human\_ode ODE function used for human states

74 step\_PTS

## **Details**

This method is equivalent to considering the ODEs describing the time evolution of the mean trajectory (first moment) and setting all higher order moments which appear on the right hand side to zero.

The solvers used within can be found in the deSolve package, see ode. For inhomogeneous systems, consider using the "rk4" method to avoid excessive integration times.

The stoichiometry matrix (S) is generated in spn\_S.

The list of hazards (haz) come from spn\_hazards.

For other samplers, see: step\_CLE, step\_PTS, step\_DM

#### Value

function closure for use in sim\_trajectory\_R or sim\_trajectory\_CSV

step_PTS Make Poisson Time-Step (PTS) Sampler for a SPN Model
---

## **Description**

Make a function closure to implement a Poisson time-step (tau-leaping with fixed tau) sampler for a SPN.

## Usage

```
step_PTS(S, Sout, haz, dt = 0.01, maxhaz = 1e+06)
```

#### **Arguments**

S a stoichiometry Matrix-class object

Sout an optional matrix to track of event firings
haz a list of hazard functions

dt time-step for tau-leap method maxhaz maximum allowable hazard

#### **Details**

This sampling algorithm is based on representing a SPN as a set of competing Poisson processes; it thus uses an integer valued state space but approximates the number of events over dt.

The design of step\_PTS is from: Wilkinson, D. J. (2011). Stochastic modeling for systems biology. CRC press

Elements of the N list come from two places: The stoichiometry matrix (S) is generated in spn\_S and the hazards (h) come from spn\_hazards.

For other samplers, see: step\_CLE, step\_DM, step\_ODE

step\_PTS\_decoupled 75

# Value

function closure for use in sim\_trajectory\_R or sim\_trajectory\_CSV

step\_PTS\_decoupled Make Poisson Time-Step (PTS) Sampler for a SPN Model

# **Description**

Make a function closure to implement a Poisson time-step (tau-leaping with fixed tau) sampler for a SPN.

# Usage

```
step_PTS_decoupled(S, Sout, haz, dt = 0.01, maxhaz = 1e+06, human_ode = "SIS")
```

## Arguments

S a stoichiometry Matrix-class object

Sout an optional matrix to track of event firings

haz a list of hazard functions

dt time-step for tau-leap method
maxhaz maximum allowable hazard
human\_ode ode used for human states

## **Details**

This sampling algorithm is based on representing a SPN as a set of competing Poisson processes; it thus uses an integer valued state space but approximates the number of events over dt.

The design of step\_PTS is from: Wilkinson, D. J. (2011). Stochastic modeling for systems biology.  $CRC\ press$ 

Elements of the N list come from two places: The stoichiometry matrix (S) is generated in spn\_S and the hazards (h) come from spn\_hazards.

For other samplers, see: step\_CLE, step\_DM, step\_ODE

#### Value

function closure for use in sim\_trajectory\_R or sim\_trajectory\_CSV

summarize\_eggs\_geno Summarize Eggs by Genotype

## **Description**

This function summarizes egg stage by genotype. It calls base\_aquatic\_geno to do all of the work.

# Usage

```
summarize_eggs_geno(out, spn_P)
```

## **Arguments**

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, genotype, and value.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

## Value

a 3 to 5 column dataframe for plotting with ggplot2

```
summarize_eggs_stage Summarize Eggs by Erlang-Stage
```

## **Description**

This function summarizes egg stage by Erlang-stages. It calls base\_aquatic\_stage to do all of the work.

## Usage

```
summarize_eggs_stage(out, spn_P)
```

# Arguments

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

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## **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, Erlang-stage, and value.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

#### Value

a 3 to 5 column dataframe for plotting with ggplot2

 ${\it summarize\_females} \qquad {\it Summarize\ Adult\ Females\ (One\ Node\ or\ Metapopulation\ Network,} \\ {\it Lifecycle\ Model)}$ 

# **Description**

For MGDrivE2 simulations of mosquito lifecycle dynamics in a single node or metapopulation network, this function sums over the male mate genotype to get population trajectories of adult female mosquitoes by their genotype.

#### **Usage**

```
summarize_females(out, spn_P)
```

## **Arguments**

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

## **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node or spn\_P\_lifecycle\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, genotype, and value.

For examples of using this function, this or any vignette which visualizes output: vignette("lifecycle-node", package = "MGDrivE2")

#### Value

a 3 to 5 column dataframe for plotting with ggplot2

# **Description**

For MGDrivE2 simulations of mosquito epidemiological dynamics in a single node or metapopulation network, this function sums over the male mate genotype as well as EIP bins to get population trajectories of adult female mosquitoes by their genotype and (S,E,I) status.

## Usage

```
summarize_females_epi(out, spn_P)
```

## **Arguments**

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, inf, genotype, and value.

For examples of using this function, this or any vignette which simulates epi dynamics: vignette("epi-node", package = "MGDrivE2")

#### Value

a 4 to 6 column dataframe for plotting with ggplot2

```
summarize_humans_epiImperial
```

Summarize Humans for Imperial Model

## **Description**

The Imperial model outputs six human states for each age compartment. This function accepts the output matrix and the desired index of an age compartment and returns the trajectory of all human states in that given age compartment (default 1)

## **Usage**

```
summarize_humans_epiImperial(out, index = 1)
```

## **Arguments**

out the output of sim\_trajectory\_R

index the desired age compartment for which to pull trajectory

#### Value

dataframe for plotting with ggplot2

```
summarize_humans_epiSEIR
```

Summarize Humans (One Node or Metapopulation Network, SEI Mosquitoes - SEIR Humans)

# Description

For MGDrivE2 simulations of mosquito epidemiological dynamics in a node or network, this function summarizes human infection status, S, E, I, and R. It uses base\_summarize\_humans to do all of the work.

# Usage

```
summarize_humans_epiSEIR(out)
```

#### **Arguments**

out the output of sim\_trajectory\_R

#### **Details**

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, inf, genotype, and value.

For examples of using this function, see: vignette("seir-dynamics", package = "MGDrivE2")

# Value

a 4 to 6 column dataframe for plotting with ggplot2

summarize\_humans\_epiSIS

Summarize Humans (One Node or Metapopulation Network, SEI Mosquitoes - SIS Humans)

# **Description**

For MGDrivE2 simulations of mosquito epidemiological dynamics in a node or network, this function summarizes human infection status, S and I. It uses base\_summarize\_humans to do all of the work.

# Usage

```
summarize_humans_epiSIS(out)
```

## **Arguments**

out

the output of sim\_trajectory\_R

#### **Details**

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, inf, genotype, and value.

For examples of using this function, see: vignette("epi-node", package = "MGDrivE2")

## Value

a 4 to 6 column dataframe for plotting with ggplot2

```
summarize_larvae_geno Summarize Larvae by Genotype
```

## **Description**

This function summarizes larval stage by genotype. It calls base\_aquatic\_geno to do all of the work.

# Usage

```
summarize_larvae_geno(out, spn_P)
```

# Arguments

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, genotype, and value.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

#### Value

a 3 to 5 column dataframe for plotting with ggplot2

```
summarize_larvae_stage
```

Summarize Larval by Erlang-Stage

## **Description**

This function summarizes larval stage by Erlang-stages. It calls base\_aquatic\_stage to do all of the work.

#### Usage

```
summarize_larvae_stage(out, spn_P)
```

# **Arguments**

```
out the output of sim_trajectory_R spn_P the places of the SPN, see details
```

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, Erlang-stage, and value.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

## Value

a 3 to 5 column dataframe for plotting with ggplot2

summarize\_males

Summarize Adult Males (One Node or Metapopulation Network)

# **Description**

For MGDrivE2 simulations of mosquito lifecycle dynamics or human infection dynamics, in a node or metapopulation network, this function summarizes population trajectories of adult male mosquitoes by their genotype.

# Usage

```
summarize_males(out)
```

## **Arguments**

out

the output of sim\_trajectory\_R

#### **Details**

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, genotype, and value.

For examples of using this function, this or any vignette which visualizes output: vignette("lifecycle-node", package = "MGDrivE2")

#### Value

a 3 to 5 column dataframe for plotting with ggplot2

```
summarize_pupae_geno Summarize Pupal by Genotype
```

# Description

This function summarizes pupal stage by genotype. It calls base\_aquatic\_geno to do all of the work.

## Usage

```
summarize_pupae_geno(out, spn_P)
```

# Arguments

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, genotype, and value.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

## Value

a 3 to 5 column dataframe for plotting with ggplot2

summarize\_pupae\_stage Summarize Pupal by Erlang-Stage

# **Description**

This function summarizes pupal stage by Erlang-stages. It calls base\_aquatic\_stage to do all of the work.

#### Usage

```
summarize_pupae_stage(out, spn_P)
```

## **Arguments**

out the output of sim\_trajectory\_R spn\_P the places of the SPN, see details

#### **Details**

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

The return object depends on the data provided. If the simulation was only 1 node, then no node designation is returned. If only one repetition was performed, no rep designation is returned. Columns always returned include: time, Erlang-stage, and value.

For examples of using this function, see: vignette("lifecycle-node", package = "MGDrivE2")

#### Value

a 3 to 5 column dataframe for plotting with ggplot2

```
summarize_stats_CSV Summary Statistics for MGDrivE2
```

# **Description**

This function reads in all repetitions for each patch and calculates either the mean, quantiles, or both. User chooses the quantiles, up to 4 decimal places, and enters them as a vector. Quantiles are calculated empirically. (order does not matter)

# Usage

```
summarize_stats_CSV(
  read_dir,
  write_dir = read_dir,
  mean = TRUE,
  quantiles = NULL,
  spn_P,
  tmax,
  dt,
  rem_file = FALSE,
  verbose = TRUE
)
```

## **Arguments**

read_dir	Directory to find repetition folders in
write_dir	Directory to write output
mean	Boolean, calculate mean or not. Default is TRUE
quantiles	Vector of quantiles to calculate. Default is NULL
spn_P	Places object, see details
tmax	The final time to end simulation
dt	The time-step at which to return output ( <b>not</b> the time-step of the sampling algorithm)
rem_file	Remove original output? Default is FALSE
verbose	Chatty? Default is TRUE

## **Details**

Given the read\_dir, this function assumes the follow file structure:

```
read_dirrepetition 1
```

- \* M\_0001.csv
- \* M\_0002.csv
- \* FS\_0001.csv
- \* FS 0001.csv
- \* ...
- repetition 2
  - \* M\_0001.csv
  - \* M\_0002.csv
  - \* FS 0001.csv
  - \* FS\_0001.csv
  - \* ..
- repetition 3
- **–** ...

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

t0, tt, dt define the first sampling time, the last sampling time, and each sampling time in-between.

Output files are \*.csv and contain the mean or quantile in the file name, e.g. stage *Mean* (patch Num).csv and stage *Quantile* (quant Num)\_(patch Num).csv.

For more details about using this function to process CSV output see: vignette("data-analysis", package = "MGDrivE2")

# Value

Writes output to files in write dir

```
summarize_stats_CSV_decoupled
```

Summary Statistics for MGDrivE2 - Decoupled samples

# Description

This function reads in all repetitions for each patch and calculates either the mean, quantiles, or both. User chooses the quantiles, up to 4 decimal places, and enters them as a vector. Quantiles are calculated empirically. (order does not matter)

## Usage

```
summarize_stats_CSV_decoupled(
  read_dir,
  write_dir = read_dir,
  mean = TRUE,
  quantiles = NULL,
  spn_P,
  tmax,
  dt,
  human_states,
  rem_file = FALSE,
  verbose = TRUE
)
```

# Arguments

read\_dir Directory to find repetition folders in

write\_dir Directory to write output

mean Boolean, calculate mean or not. Default is TRUE quantiles Vector of quantiles to calculate. Default is NULL

spn\_P Places object, see details

tmax The final time to end simulation

dt The time-step at which to return output (**not** the time-step of the sampling algo-

rithm)

human\_states human state distribution

rem\_file Remove original output? Default is FALSE

verbose Chatty? Default is TRUE

# **Details**

Given the read\_dir, this function assumes the follow file structure:

```
• read_dir
```

- repetition 1
  - \* M\_0001.csv
  - \* M\_0002.csv
  - \* FS\_0001.csv
  - \* FS\_0001.csv
  - \* ...
- repetition 2
  - \* M\_0001.csv
  - \* M\_0002.csv

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```
* FS_0001.csv

* FS_0001.csv

* ...

- repetition 3

- ...
```

The places (spn\_P) object is generated from one of the following: spn\_P\_lifecycle\_node, spn\_P\_lifecycle\_network, spn\_P\_epiSIS\_node, spn\_P\_epiSIS\_network, spn\_P\_epiSEIR\_node, or spn\_P\_epiSEIR\_network.

t0, tt, dt define the first sampling time, the last sampling time, and each sampling time in-between.

Output files are \*.csv and contain the mean or quantile in the file name, e.g. stage *Mean* (patch Num).csv and stage *Quantile* (quant Num)\_(patch Num).csv.

For more details about using this function to process CSV output see: vignette("data-analysis", package = "MGDrivE2")

#### Value

Writes output to files in write\_dir

track\_hinf

Make tracking matrix for human infection events

# Description

Create a matrix object for tracking incidence in human population to be passed to either sim\_trajectory\_CSV or sim\_trajectory\_R.

#### **Usage**

```
track_hinf(spn_T, S)
```

# Arguments

spn\_T set of transitions
S stoichiometry matrix

#### **Details**

The returned matrix can be passed to the Sout argument of sim\_trajectory\_CSV or sim\_trajectory\_R.

# Value

```
a sparseMatrix object
```

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