# Package 'MGDrivE'

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Type Package

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Version 1.0
Description MGDrivE is a model designed to be a reliable testbed where various gene drive interventions for mosquito-borne diseases control.  It is being developed to accommodate the use of various mosquito-specific gene drive systems within a population dynamics framework that allows migration of individuals between patches in landscape.
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R topics documented:
accumulate_ADMnew_Patch

calcAverageGenerationTime	
calcDensityDependentDeathRate	
calcLarvalPopEquilibrium	
calcLarvalStageMortalityRate	. 9
calcMemoryWindow	. 10
calcPopulationGrowthRate	
calc_ExpKernel	
calc_GammaKernel	. 11
calc_haversine	
calc_HurdleExpKernel	. 12
calc_LognormalKernel	
close_allConnections_Network	
createNamedPopMatrix	. 13
createNamedPopVector	. 13
cube2csv	. 13
cubeModifiers	. 14
Cube_Homing1RA	. 14
Cube_HomingDrive	
Cube_KillerRescue	
Cube_MEDEA	
 Cube_Mendelian	. 18
Cube_oneLocusTA	
Cube_ReciprocalTranslocations	
Cube_RIDL	. 20
Cube_twoLocusTA	
Cube_Wolbachia	
eraseDirectory	
generateReleaseVector	
get_ADMdly_Patch	
get_ADMnew_Patch	23
get_ADM_Patch	
get_AdPopEQ_Network	
get_AF1dly_Patch	
get_AF1new_Patch	
get_AF1_Patch	
get_alpha_Network	
get_aiphia_Network	
get_conADM_Network	
get_conAF1 Network	25
get_conAr-1_Network	25
get_driveCube_genotype_Network	25
get_driveCube_genotype_network	26
get_driveCube_index_Network	26
•	26
get_EGG_Patch	
get_eta_Network	. 27 . 27
get_femaleMigration_Patch	
get_genotypesID_Network	. 27
get_genotypesN_Network	27
get_g_Network	28
get_LARdly_Patch	28
get_LAR_Patch	. 28
get_Leq_Network	. 28

get_maleMigration_Patch
get_migrationFemaleRow_Network
get_migrationFemale_Network
get_migrationMaleRow_Network
get_migrationMale_Network
get_moveVar_Network
get_muAd_Network
get_muAq_Network
get_NetworkPointer_Patch
get_nPatch_Network
get_omega_Network
get_patches_Network
get_patchID_Patch
get_patchReleases_Network
get_patch_Network
get_phi_Network
get_PUPdly_Patch
get_PUP_Patch
get_releaseType_Network
get_Rm_Network
get_simTime_Network
get_s_Network
get_tau_Network
get_thetaAq_Network
get_timeAq_Network
get_tNow_Network
get_wildType_Network
get_windowSize_Network
get_xiF_Network
get_xiM_Network
ggCol_utility
initPopMatrixArray
initPopVectorArray
initStagesDurations
kernels
MGDrivE
MGDrivE-Cube
MGDrivE-Model
MGDrivE.Setup
moveMatAll2
moveMatCascade3
moveMatDiag
moveMatDiagOneCity
moveMatDie
moveMatIndependent3
moveMatMixedSpil
moveMatTaleOfTwoCities
moveMatTriDiagonal
moveMatTriple
Network
Network.Parameters
normalise

4 R topics documented:

oneDay_admPupating_deterministic_Patch	
oneDay_admPupating_stochastic_Patch	
oneDay_admSurvival_deterministic_Patch	
oneDay_admSurvival_stochastic_Patch	
oneDay_af1Mating_deterministic_Patch	55
oneDay_af1Mating_stochastic_Patch	55
oneDay_af1Pupation_deterministic_Patch	56
oneDay_af1Pupation_stochastic_Patch	56
oneDay_af1Survival_deterministic_Patch	56
oneDay_af1Survival_stochastic_Patch	57
oneDay_calcCumulativeLarvalDensityDependentFactor_Patch	57
oneDay_calcCumulativePupaDensityDependentFactor_Patch	57
oneDay_calcLarvalDensityDependentFactor_Patch	58
oneDay_eggsFract2_deterministic_Patch	58
oneDay_eggsFract2_stochastic_Patch	58
oneDay_femaleReleases_Patch	59
oneDay_hatchingFract_deterministic_Patch	59
oneDay_hatchingFract_stochastic_Patch	59
oneDay_initOutput_Patch	59
oneDay_larHatching_deterministic_Patch	60
oneDay_larHatching_stochastic_Patch	60
oneDay_larPupating_deterministic_Patch	
oneDay_larPupating_stochastic_Patch	
oneDay_larSurvival_deterministic_Patch	
oneDay_larSurvival_stochastic_Patch	
oneDay_maleReleases_Patch	
oneDay_migrationIn_Patch	
oneDay_migrationOut_deterministic_Patch	62
oneDay_migrationOut_stochastic_Patch	
oneDay_Migration_Network	
oneDay_Network	
oneDay_numMaleFemale_deterministic_Patch	
oneDay_numMaleFemale_stochastic_Patch	
oneDay_ovipositG1_deterministic_Patch	
oneDay_ovipositG1_stochastic_Patch	
oneDay_PopDynamics_Patch	
oneDay_updatePopulation_Patch	
oneDay_writeOutput_Patch	65
oneRun_Network	65
Patch	66
primePopMatrixArray	69
primePopVectorArray	69
quantileC	70
rDirichlet	70
Release_basicRepeatedReleases	71
reset_Network	71
reset_Patch	72
retrieveOutput	72
set_ADMnew_Patch	72
set_AF1new_Patch	73
set_migrationFemale_Network	73
set_migrationMale_Network	73

accumu	late_ADMnew_Patch	5	
	set_NetworkPointer_Patch shiftAndUpdatePopVector splitOutput SymCubeC turnStochasticityOnOrOff	74 74 75	
Index		<b>76</b>	
accumulate_ADMnew_Patch  Accumulate ADMnew			

# Description

Accumulate new ADM males

# Usage

```
accumulate_ADMnew_Patch(count)
```

### **Arguments**

count vector of new ADM males

# Description

Aggregate over male mate genotype to convert female matrix output into vector output.

# Usage

```
aggregateFemales(directory, genotypes, remove = FALSE, multiCore = FALSE)
```

# **Arguments**

directory	Directory where output was written to; must not end in path seperator
genotypes	Character vector of possible genotypes; found in driveCube\$genotypesID
remove	Boolean flag to remove original (unaggregated) file
multiCore	Write output using multiple cores? Default is FALSE

6 AnalyzeQuantiles

	AnalyzeQuantiles	Summary Statistics for Stochastic MGDrivE
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# 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. (order does not matter)

### Usage

```
AnalyzeQuantiles(readDirectory, writeDirectory, mean = TRUE,
   quantiles = NULL)
```

### **Arguments**

 ${\tt readDirectory} \quad {\tt Directory} \ \ {\tt to} \ {\tt find} \ {\tt repetition} \ {\tt folders} \ {\tt in}$ 

writeDirectory Directory to write output

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

# **Details**

Given the readDirectory, this function assumes the follow file structure:

- readDirectory
  - repetition 1
    - \* patch 1
    - \* patch 2
    - \* patch 3
  - repetition 2
    - \* patch 1
    - \* patch 2
    - \* patch 3
  - repetition 3
  - repetition 4
  - **–** ...

### Value

Writes output to files in writeDirectory

 ${\tt calcAquaticStagesSurvivalProbability}$ 

Calculate Survival Probability of entire Aquatic Stage Life-cycle

# Description

Calculate vector of survival probabilities for each stage of aquatic lifecycle.

#### Usage

```
calcAquaticStagesSurvivalProbability(eggSurvivalProbability,
larvaSurvivalProbability, pupaSurvivalProbability)
```

#### **Arguments**

```
eggSurvivalProbability
see calcAquaticStageSurvivalProbability
larvaSurvivalProbability
see calcAquaticStageSurvivalProbability
pupaSurvivalProbability
see calcAquaticStageSurvivalProbability
```

calcAquaticStageSurvivalProbability

Calculate Aquatic Stage Surival Probability

# Description

Calculate  $\theta_{st}$ , density-independent survival probability, given by:

$$\theta_{st} = (1 - \mu_{st})^{T_{st}}$$

### Usage

 ${\tt calcAquaticStageSurvivalProbability(mortalityRate, stageDuration)}$ 

### **Arguments**

mortalityRate daily mortality probability,  $\mu_{st}$  stageDuration duration of aquatic stage,  $T^{st}$ 

calcAverageGenerationTime

Calculate Average Generation Time

#### **Description**

Calculate g, average generation time, given by:

$$g = T_e + T_l + T_p + \frac{1}{\mu_{ad}}$$

#### Usage

calcAverageGenerationTime(stagesDuration, adultMortality)

### **Arguments**

stagesDuration vector of lengths of aquatic stages,  $T_e, T_l, T_p$  adultMortality adult mortality rate,  $\mu_{ad}$ 

 ${\tt calcDensityDependentDeathRate}$ 

Calculate Density-dependent Larval Mortality

#### **Description**

Calculate  $\alpha$ , the strength of density-dependent mortality during the larval stage, given by:

$$\alpha = \left(\frac{1/2 * \beta_k * \theta_e * Ad_{eq}}{R_m - 1}\right) * \left(\frac{1 - (\theta_l/R_m)}{1 - (\theta_l/R_m)^{1/T_l}}\right)$$

### Usage

calcDensityDependentDeathRate(fertility, thetaAq, tAq, adultPopSizeEquilibrium, populationGrowthRate)

# Arguments

fertility number of eggs per oviposition for wild-type females,  $\beta_k$ 

thetaAq vector of density-independent survival probabilities of aquatic stages,  $\theta_e, \theta_l$ 

tAq vector of lengths of aquatic stages,  $T_e, T_l, T_p$ 

adultPopSizeEquilibrium

adult population size at equilbrium,  $Ad_{eq}$ 

populationGrowthRate

population growth in absence of density-dependent mortality  $R_m$ 

calcLarvalPopEquilibrium

Calculate Equilibrium Larval Population

### **Description**

Equilibrium larval population to sustain population.

#### Usage

calcLarvalPopEquilibrium(alpha, Rm)

### **Arguments**

alpha see calcDensityDependentDeathRate

Rm see calcPopulationGrowthRate

calcLarvalStageMortalityRate

Calculate Larval Stage Mortality Rate

# Description

Calculate  $\mu_l$ , the larval mortality, given by

$$\mu_l = 1 - \left(\frac{R_m * \mu_{ad}}{1/2 * \beta_k * (1 - \mu_m)}\right)^{\frac{1}{T_e + T_l + T_p}}$$

### Usage

# Arguments

 ${\tt generationPopGrowthRate}$ 

see calcPopulationGrowthRate

adultMortality adult mortality rate,  $\mu_{ad}$ 

fertility number of eggs per oviposition for wild-type females,  $\beta_k$ 

aquaticStagesDuration

vector of lengths of aquatic stages,  $T_e, T_l, T_p$ 

Calculate Memory Window

# Description

Calculates the necessary window of population history required for the model to work

# Usage

```
calcMemoryWindow(stagesDuration)
```

# Arguments

egg	length of egg stage (days)
larva	length of larval stage (days)
pupa	length of pupal stage (days)

 ${\tt calcPopulationGrowthRate}$ 

Calculate Population Growth Rate

# Description

Calculate  $R_m$ , population growth in absence of density-dependent mortality, given by:

$$(r_m)^g$$

### Usage

calcPopulationGrowthRate(dailyPopGrowthRate, averageGenerationTime)

# **Arguments**

```
dailyPopGrowthRate \mbox{daily population growth rate}, r_m \mbox{averageGenerationTime} \\ \mbox{see calcAverageGenerationTime}
```

calc\_ExpKernel 11

calc_ExpKernel	Calculate Exponential Stochastic Matrix
----------------	---

### **Description**

Given a distance matrix from calc\_haversine, calculate a stochastic matrix where one step movement probabilities follow an exponential density.

#### Usage

```
calc_ExpKernel(distMat, r)
```

### **Arguments**

distMat distance matrix from calc\_haversine
r rate parameter of Exponential distribution

calc\_GammaKernel Calculate Gamma Stochastic Matrix

#### **Description**

Given a distance matrix from calc\_haversine, calculate a stochastic matrix where one step movement probabilities follow a gamma density.

### Usage

```
calc_GammaKernel(distMat, shape, rate)
```

#### **Arguments**

distMat distance matrix from calc\_haversine
shape shape parameter of GammaDist distribution
rate rate parameter of GammaDist distribution

calc\_haversine Calculate Haversine Distance

### **Description**

Calculate the great-circle distance (Haversine distance) between sets of longitude - latitude points, see https://en.wikipedia.org/wiki/Haversine\_formula

# Usage

```
calc_haversine(longlats)
```

# Arguments

longlats numeric matrix where first column is vector of longitudes and second column is

vector of latitudes

calc\_HurdleExpKernel Calculate Hurdle Exponential Stochastic Matrix

### **Description**

Given a distance matrix from calc\_haversine, calculate a stochastic matrix where one step movement probabilities follow an zero-truncated exponential density with a point mass at zero.

#### Usage

```
calc_HurdleExpKernel(distMat, r, pi)
```

### **Arguments**

distMat distance matrix from calc\_haversine
r rate parameter of Exponential distribution

pi point mass at zero

calc\_LognormalKernel Calculate Lognormal Stochastic Matrix

# Description

Given a distance matrix from calc\_haversine, calculate a stochastic matrix where one step movement probabilities follow a lognormal density.

### Usage

```
calc_LognormalKernel(distMat, meanlog, sdlog)
```

#### **Arguments**

distMat distance matrix from calc\_haversine
meanlog log mean of Lognormal distribution

sdlog log standard deviation of Lognormal distribution

close\_allConnections\_Network

Close all Output Connections

### **Description**

Close private\$conADM and private\$conAF1

```
close_allConnections_Network()
```

createNamedPopMatrix Create a Named Matrix

# Description

Create a named matrix of 0s

### Usage

```
createNamedPopMatrix(genotypesID)
```

#### **Arguments**

genotypesID character vector of possible genotypes

### **Description**

Create a named vector of 0s

#### Usage

```
createNamedPopVector(genotypesID)
```

# Arguments

genotypesID character vector of possible genotypes

cube2csv Export a Cube to .csv

# Description

Export a cube as multiple .csv files (one for each genotype; slices of z-axis).

# Usage

```
cube2csv(cube, directory, digits = 3)
```

# Arguments

cube a cube object (see MGDrivE-Cube for options)

directory directory to write .csv files to

digits number of significant digits to retain in .csv output

14 Cube\_Homing1RA

cubeModifiers $G\epsilon$	enerate and Modify Default Genotype-specific Parameters
---------------------------	---

### **Description**

This is an internal function for cubes.

# Usage

```
cubeModifiers(gtype, eta = NULL, phi = NULL, omega = NULL, xiF = NULL,
  xiM = NULL, s = NULL)
```

### **Arguments**

gtype	character vector of genotypes
eta	genotype-specific mating fitness
phi	genotype-specific sex ratio at emergence
omega	genotype-specific multiplicative modifier of adult mortality
xiF	genotype-specific female pupatory success
xiM	genotype-specific male pupatory success
S	genotype-specific fractional reduction(increase) in fertility

# Description

This function creates an inheritance cube to model a homing gene drive (such as a CRISPR-Cas9 system) that creates 1 type of resistance allele. It assumes no sex-specific inheritance patterns and the construct is on an autosome.

# Usage

```
Cube_Homing1RA(e = 1, p = 0, eta = NULL, phi = NULL, omega = NULL,
    xiF = NULL, xiM = NULL, s = NULL)
```

### **Arguments**

е	Homing rate
p	Resistance allele generation rate
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

Cube\_HomingDrive 15

#### Value

Named list(inheritance cube, viability mask, genotypes ID, genotypes number, wild-type allele, mating fitness, sex ratio, adult mortality modifier, female pupatory success, male pupatory success, fertility modifier, release genotype)

Cube_HomingDrive	Inheritance Cube: CRISPR (Clustered Regularly Interspaced SWort
	Palindromic Repeats) witW 2 Resistance Allele

# Description

This is a sex-specific version of the original cube. It assumes that the construct is on an autosome and there can be different male/female homing rates

# Usage

```
Cube_HomingDrive(eM = 1, eF = 1, rM = 0, bM = 0, rF = 0, bF = 0,
  eta = NULL, phi = NULL, omega = NULL, xiF = NULL, xiM = NULL,
  s = NULL)
```

### **Arguments**

eM	Male homing rate
eF	Female homing rate
rM	Male no-cost resistance generation rate
bM	Male detrimental resistance generation rate
rF	Female no-cost resistance generation rate
bF	Female detrimental resistance generation rate
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
s	Genotype-specific fractional reduction(increase) in fertility

### Value

16 Cube\_KillerRescue

cube KillerRescue Inneritance Cube: Killer-Rescue Syste	Cube KillerRescue	Inheritance Cube: Killer-Rescue System
---	-------------------	--

# Description

This function creates an inheritance cube to model a Killer-Rescue system. Killer-Resuce is a 2-locus system: one locus has a toxin and the other locus contains the antidote. The loci are assumed independent and are non-homing.

This drive has 3 alleles at locus 1 and 2 alleles and locus 2:

- Locus 1
  - T: Wild-type allele
  - K: "Killer" toxin allele
  - R: Rroken toxin allele
- Locus 2
  - W: Wild-type allele
  - A: Antidote allele

### Usage

```
Cube_KillerRescue(eR = 0, Keff = 1, Aeff = 1, eta = NULL, phi = NULL,
  omega = NULL, xiF = NULL, xiM = NULL, s = NULL)
```

# Arguments

eR	Conversion of K allele to R allele, a basal mutation rate
Keff	Toxin efficacy
Aeff	Antidote efficacy
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

#### Value

Cube\_MEDEA 17

Cube_MEDEA	Inheritance Cube: MEDEA (Maternal Effect Dominant Embryonic Arrest)	
	,	

# Description

This function creates an inheritance cube to model a MEDEA drive system. This system was first discovered in flour beetles. It biases inheritance by expressing a maternal toxin such that offspring die unless they express a zygotic antidote.

This drive has 3 alleles at 1 locus:

• W: Wild-type allele

• M: MEDEA allele

• R: Resistance allele

# Usage

```
Cube_MEDEA(rM = 0, rW = 0, Teff = 1, eta = NULL, phi = NULL,
  omega = NULL, xiF = NULL, xiM = NULL, s = NULL)
```

### **Arguments**

rM	Breakdown of MEDEA allele, no homing/toxin/antidote, M -> R conversion
rW	De novo resistance generation, W -> R conversion
Teff	Efficacy of the toxin
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

### Value

18 Cube\_oneLocusTA

ian Inheritance Cube: Mendelian
---------------------------------

# Description

This function creates a Mendelian Inheritance Cube. It only handles simple, alphabetic genotypes. The default is 3 alleles at 1 locus, but this can be extended to however many alleles one is interested in, but only at 1 locus.

#### Usage

```
Cube_Mendelian(gtype = c("AA", "Aa", "aa"), eta = NULL, phi = NULL, omega = NULL, xiF = NULL, xiM = NULL, s = NULL)
```

#### **Arguments**

gtype	Vector of genotypes, with the wild-type in the first position
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

### Value

Named list containing the inheritance cube, transition matrix, genotypes, wild-type allele, and all genotype-specific parameters.

### **Description**

This function creates a 1 locus maternal-toxin/zygotic-antidote system. This is similar to the construct called UDmel. There is no resistance generation in this model. This drive has 3 alleles at 1 locus:

- A: Maternal-toxin 1, zygotic-antidote 2
- B: Maternal-toxin 2, zygotic-antidote 1
- W: Wild-type allele

```
Cube_oneLocusTA(TAEfficacy = 1, TBEfficacy = 1, eta = NULL, phi = NULL,
  omega = NULL, xiF = NULL, xiM = NULL, s = NULL)
```

#### **Arguments**

TAEfficacy	Maternal toxin A efficacy
TBEfficacy	Maternal toxin B efficacy
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

### Value

Named list containing the inheritance cube, transition matrix, genotypes, wild-type allele, and all genotype-specific parameters.

Cube\_ReciprocalTranslocations

Inheritance Cube: Reciprocal Translocation

# Description

This function creates an inheritance cube to model a reciprocal translocation. This technology was the original form of underdominant system. It involves 2 chromosomes, each with two alleles. This drive has 4 alleles at 2 loci:

- a: Wild-type at locus A
- A: Translocation at locus A
- b: Wile-type at locus B
- B: Translocation at locus B

#### Usage

### **Arguments**

eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

#### Value

20 Cube\_twoLocusTA

Cube_RIDL Inheritance Cube: RIDL (Release of Insects with Dominant Lethality	Cube_RIDL	Inheritance Cube:	RIDL (Release of Insects	with Dominant Lethality
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# Description

This function creates a RIDL system RIDL (Release of Insects with Dominant Lethality), is a form of SIT. Created by Oxitec, this is based on a positive feedback loop using the toxic tTAV gene, contolled under lab conditions by the TetO promoter. This has 2 alleles at 1 locus

- W: Wild-type allele
- R: OX513 RIDL allele

### Usage

```
Cube_RIDL(eta = NULL, phi = NULL, omega = NULL, xiF = NULL,
    xiM = NULL, s = NULL)
```

### **Arguments**

eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

### Value

Named list containing the inheritance cube, transition matrix, genotypes, wild-type allele, and all genotype-specific parameters.

# Description

This function creates a 2 locus maternal-toxin/zygotic-antidote system. This is similar to the construct called UDmel. There is no resistance generation in this model.

This drive has 2 unlinked alleles, 1 allele each at 2 loci:

- A: Maternal-toxin 1, zygotic-antidote 2
- a: Wild-type at locus A
- B: Maternal-toxin 2, zygotic-antidote 1
- b: Wild-type at locus B

```
Cube_twoLocusTA(TAEfficacy = 1, TBEfficacy = 1, eta = NULL, phi = NULL,
  omega = NULL, xiF = NULL, xiM = NULL, s = NULL)
```

Cube\_Wolbachia 21

#### **Arguments**

TAEfficacy	Maternal toxin A efficacy
TBEfficacy	Maternal toxin B efficacy
eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

#### Value

Named list containing the inheritance cube, transition matrix, genotypes, wild-type allele, and all genotype-specific parameters.

Cube\_Wolbachia Inheritance Cube: Wolbachia

# Description

This function creates an inheritance cube to model a Wolbachia infection. Wobachia is a parasite that can infect mosquitoes. It biases its inheritance through cytoplasmic incompatibility. This drive has 2 alleles at 1 locus:

- W: has Wolbachia
- w: does not have Wolbachia

#### Usage

```
Cube_Wolbachia(eta = NULL, phi = NULL, omega = NULL, xiF = NULL,
  xiM = NULL, s = NULL)
```

### **Arguments**

eta	Genotype-specific mating fitness
phi	Genotype-specific sex ratio at emergence
omega	Genotype-specific multiplicative modifier of adult mortality
xiF	Genotype-specific female pupatory success
xiM	Genotype-specific male pupatory success
S	Genotype-specific fractional reduction(increase) in fertility

#### **Details**

Cytoplasmic Incompatibility:

- male W cross female w -> all offspring die (complete penetrance)
- male w cross female W -> all offspring inherit Wolbachia

22 get\_ADMdly\_Patch

#### Value

Named list containing the inheritance cube, transition matrix, genotypes, wild-type allele, and all genotype-specific parameters.

eraseDirectory

Erase all files in a directory

### **Description**

Given a directory path, check it exists and if so delete all its contents.

#### Usage

```
eraseDirectory(directory)
```

#### **Arguments**

directory

directory whose contents will be deleted

generateReleaseVector Make List of Modified Mosquito Releases

### **Description**

Sets up a release schedule for a single patch, calls Release\_basicRepeatedReleases internally.

# Usage

```
generateReleaseVector(driveCube = driveCube,
  releasesParameters = releasesParameters, sex = "M")
```

### Arguments

 ${\tt driveCube}$ 

gene-drive cube

 ${\tt releases Parameters}$ 

A list containing the releasesStart, releasesNumber releasesInterval, and release-

Proportion named values.

sex

character in 'M','F'

get\_ADMdly\_Patch

Get ADMdly

# **Description**

Return adult male stage delay window population

```
get_ADMdly_Patch()
```

get\_ADMnew\_Patch 23

get\_ADMnew\_Patch

Get ADMnew

# Description

Return the new ADM females

# Usage

```
get_ADMnew_Patch()
```

get\_ADM\_Patch

Get ADM

# Description

Return adult male stage population

# Usage

```
get_ADM_Patch()
```

get\_AdPopEQ\_Network

Get AdPopEQ

# Description

Return equilibrium adult population

# Usage

```
get_AdPopEQ_Network(ix)
```

# Arguments

ix

index of patch

get\_AF1dly\_Patch

Get AF1dly

# Description

Return adult female stage delay window population

```
get_AF1dly_Patch()
```

24 get\_beta\_Network

get\_AF1new\_Patch

Get AF1new

# Description

Return the new AF1 females

# Usage

```
get_AF1new_Patch()
```

get\_AF1\_Patch

Get AF1

# Description

Return adult female stage population

# Usage

```
get_AF1_Patch()
```

get\_alpha\_Network

Get alpha

# Description

 $Return\ density\ dependent\ mortality,\ see\ {\tt calcDensityDependentDeathRate}$ 

# Usage

```
get_alpha_Network(ix)
```

# Arguments

ix

index of patch

get\_beta\_Network

Get beta

# Description

Return size of wild-type egg batch

```
get_beta_Network()
```

get\_conADM\_Network

get\_conADM\_Network

Get conADM

### **Description**

Return connection where adult male dynamics are written to

# Usage

```
get_conADM_Network()
```

get\_conAF1\_Network

Get conAF1

# Description

Return connection where adult female dynamics are written to

# Usage

```
get_conAF1_Network()
```

```
get_directory_Network Get directory
```

# Description

Return character string of directory being written to

# Usage

```
get_directory_Network()
```

```
get_driveCube_genotype_Network
```

 ${\it Get \ Element}(s) \ of \ Drive \ Cube \ by \ Genotype$ 

### **Description**

Return elements or slices of drive cube. If all NULL return entire cube.

# Usage

```
get_driveCube_genotype_Network(fG = NULL, mG = NULL, oG = NULL)
```

# Arguments

fG	female genotype
mG	male genotype
oG	offspring genotype

get\_EGG\_Patch

```
{\tt get\_driveCube\_index\_Network}
```

Get Element(s) of Drive Cube by Index

# Description

Return elements or slices of drive cube. If all NULL return entire cube.

### Usage

```
get_driveCube_index_Network(fG = NULL, mG = NULL, oG = NULL)
```

# **Arguments**

fG	female genotype index
mG	male genotype index
oG	offspring genotype index

get\_EGGdly\_Patch

 $Get\ EGGdly$ 

# Description

Return egg stage delay window population

# Usage

```
get_EGGdly_Patch()
```

 ${\tt get\_EGG\_Patch}$ 

Get EGG

# Description

Return egg stage population

```
get_EGG_Patch()
```

get\_eta\_Network 27

get\_eta\_Network

Get eta

# Description

Return genotype-specific mating fitness

# Usage

```
get_eta_Network()
```

```
get_femaleMigration_Patch
```

Get maleMigration

# Description

Return outbound males (nGenotypes X nGenotypes X nPatch array)

# Usage

```
get_femaleMigration_Patch()
```

```
{\tt get\_genotypesID\_Network}
```

Get genotypesID

# Description

Return character vector of possible genotypes

# Usage

```
get_genotypesID_Network()
```

```
get_genotypesN_Network
```

Get genotypesN

# Description

Return number of possible genotypes

```
get_genotypesN_Network()
```

28 get\_Leq\_Network

get\_g\_Network

Get g

# Description

 $Return\ average\ generation\ time,\ see\ {\tt calcAverageGenerationTime}$ 

# Usage

```
get_g_Network()
```

get\_LARdly\_Patch

Get LARdly

# Description

Return larval stage delay window population

# Usage

```
get_LARdly_Patch()
```

get\_LAR\_Patch

Get LAR

# Description

Return larval stage population

# Usage

```
get_LAR_Patch()
```

get\_Leq\_Network

Get Leq

# Description

 $Return\ equilibrium\ larval\ population,\ see\ {\tt calcLarvalPopEquilibrium}$ 

# Usage

```
get_Leq_Network(ix)
```

# Arguments

ix

index of patch

```
get_maleMigration_Patch
```

 $Get\ male Migration$ 

# Description

Return outbound males (nGenotypes X nPatch integer matrix)

# Usage

```
get_maleMigration_Patch()
```

```
{\tt get\_migrationFemaleRow\_Network}
```

Get Row of Female Migration Matrix

# Description

Return a matrix object (does not drop dimensions)

# Usage

```
get_migrationFemaleRow_Network(ix)
```

### **Arguments**

ix index of row

```
get_migrationFemale_Network
```

Get Female Migration Matrix

# Description

Return a matrix object

```
get_migrationFemale_Network()
```

30 get\_muAd\_Network

```
get_migrationMaleRow_Network
```

Get Row of Male Migration Matrix

# Description

Return a matrix object (does not drop dimensions)

# Usage

```
get_migrationMaleRow_Network(ix)
```

### **Arguments**

ix

index of row

```
get_migrationMale_Network
```

Get Male Migration Matrix

# Description

Return a matrix object

# Usage

```
get_migrationMale_Network()
```

```
get_moveVar_Network
```

Get moveVar

# Description

Return numeric variance in Dirchlet-Multinomial movement

# Usage

```
get_moveVar_Network()
```

 ${\tt get\_muAd\_Network}$ 

Get muAd

# Description

Return adult mortality

```
get_muAd_Network()
```

get\_muAq\_Network 31

get\_muAq\_Network

Get muAq

# Description

Return larval mortality, see calcLarvalStageMortalityRate

### Usage

```
get_muAq_Network()
```

```
get_NetworkPointer_Patch
```

Get Network Pointer

# Description

Return a reference to the enclosing Network object

# Usage

```
get_NetworkPointer_Patch()
```

get\_nPatch\_Network

Get nPatch

# Description

Return number of patches

# Usage

```
get_nPatch_Network()
```

get\_omega\_Network

Get omega

### **Description**

Return genotype-specific multiplicative modifier of adult mortality

```
get_omega_Network()
```

# Description

Return a list of Patch objects

# Usage

```
get_patches_Network()
```

get\_patchID\_Patch

Get patchID

# Description

Return the ID of this patch

# Usage

```
get_patchID_Patch()
```

get\_patchReleases\_Network

Get Patch Release Schedule

# Description

Return the release schedule for a patch for male or female

# Usage

```
get_patchReleases_Network(ix, sex = "M")
```

# Arguments

ix index of patch

sex character in 'M', 'F'

get\_patch\_Network 33

get\_patch\_Network

Get Patch

# Description

Return a Patch object

# Usage

```
get_patch_Network(ix)
```

# **Arguments**

ix

integer id of patch to return

 ${\tt get\_phi\_Network}$ 

Get phi

# Description

Return genotype-specific sex ratio at emergence

# Usage

```
get_phi_Network()
```

get\_PUPdly\_Patch

Get PUP

# Description

Return pupae stage delay window population

# Usage

```
get_PUPdly_Patch()
```

get\_PUP\_Patch

Get PUP

# Description

Return pupae stage population

```
get_PUP_Patch()
```

34 get\_s\_Network

```
{\tt get\_releaseType\_Network}
```

Get releaseType

# Description

Return genotype of release

# Usage

```
get_releaseType_Network()
```

get\_Rm\_Network

Get Rm

# Description

Return population growth rate, see calcPopulationGrowthRate

# Usage

```
get_Rm_Network()
```

get\_simTime\_Network

Get simTime

# Description

Return maximum time to run simulation

# Usage

```
get_simTime_Network()
```

get\_s\_Network

Get s

### **Description**

Return genotype-specific fractional reduction(increase) in fertility

```
get_s_Network()
```

get\_tau\_Network 35

get\_tau\_Network

Get Viability Mash (tau)

# Description

Return matrix

# Usage

```
get_tau_Network()
```

 ${\tt get\_thetaAq\_Network}$ 

Get thetaAq

# Description

Return aquatic stage survival probability, see calcAquaticStagesSurvivalProbability and calcAquaticStageSurvivalProbability

# Usage

```
get_thetaAq_Network(stage)
```

# Arguments

stage

character in 'E', 'L', 'P'

 ${\tt get\_timeAq\_Network}$ 

Get timeAq

# Description

Return duration of aquatic stages, see initStagesDurations

# Usage

```
get_timeAq_Network(stage = NULL)
```

# Arguments

stage

character in 'E', 'L', 'P'; if NULL return total duration

36 get\_xiF\_Network

get\_tNow\_Network

Get tNow

# Description

Return current simulation time

### Usage

```
get_tNow_Network()
```

```
get_wildType_Network Get wildType
```

# Description

Return wild-type genotype

# Usage

```
get_wildType_Network()
```

```
get_windowSize_Network
```

Get windowSize

# Description

Return memory window size, see calcMemoryWindow

# Usage

```
get_windowSize_Network()
```

```
get_xiF_Network
```

Get xiF

### **Description**

Return genotype-specific female pupatory success

```
get_xiF_Network()
```

get\_xiM\_Network 37

# Description

Return genotype-specific male pupatory success

## Usage

```
get_xiM_Network()
```

ggCol\_utility

Utility to Imitate ggplot2 Colors

## **Description**

Sample at equally spaced intervals along the color wheel

# Usage

```
ggCol_utility(n, alpha = 1)
```

# Arguments

n number of colors alpha transparency

init Pop Matrix Array

Create a population array of matrices

# Description

Creates an array for the population history to be stored. The length of the array is equal to the window required for the model to run (in our specific case it is equal to the sum of aquatic stages lengths).

#### Usage

```
initPopMatrixArray(genotypesID, memoryWindow)
```

# Arguments

genotypesID character vector of possible genotypes

memoryWindow integer size of list structure

38 initStagesDurations

initPopVectorArray Create a population array of vectors

# **Description**

Creates an array for the population history to be stored. The length of the array is equal to the window required for the model to run (in our specific case it is equal to the sum of aquatic stages lengths).

# Usage

```
initPopVectorArray(genotypesID, memoryWindow)
```

# **Arguments**

genotypesID character vector of possible genotypes

memoryWindow integer size of list structure

# Description

Initialises the vector that holds the duration of each aquatic stage

# Usage

```
initStagesDurations(egg = 1, larva = 14, pupa = 1)
```

## **Arguments**

egg	length of egg stage (days)
larva	length of larval stage (days)
pupa	length of pupal stage (days)

kernels 39

kernels

Kernels Parameters

#### **Description**

A named list containing maximum likelihood fitted parameter values from mosquito dispersal estimates.

#### Usage

data(kernels)

#### **Format**

named list with 5 elements:

lnorm\_mean log mean of log-normal density
lnorm\_sd log standard deviation of log-normal density
gamma\_shape shape parameter of gamma density
gamma\_sd rate parameter of gamma density
exp\_rate rate parameter of exponential density

MGDrivE

MGDrivE: Mosquito Gene Drive Explorer

# Description

MGDrivE: Mosquito Gene Drive Explorer

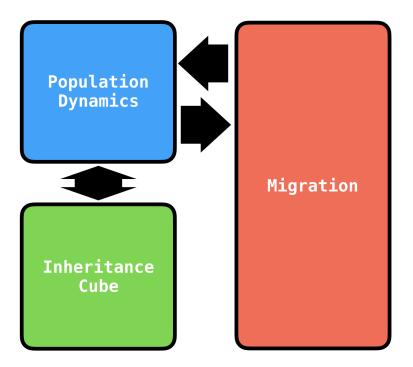
#### Introduction

Recent developments of CRISPR-Cas9 based homing endonuclease gene drive systems for the suppression or replacement of mosquito populations have generated much interest in their use for control of mosquito-borne diseases (such as dengue, malaria, chikungunya and Zika). This is because genetic control of pathogen transmission may complement or even substitute traditional vectorcontrol interventions, which have had limited success in bringing the spread of these diseases to a halt. Despite excitement for the use of gene drives for mosquito control, current modeling efforts have analyzed only a handful of these new approaches (usually studying just one per framework). Moreover, these models usually consider well-mixed populations with no explicit spatial dynamics. To this end, we are developing MGDrivE (Mosquito Gene DRIVe Explorer), in cooperation with the 'UCI Malaria Elimination Initiative', as a flexible modeling framework to evaluate a variety of drive systems in spatial networks of mosquito populations. This framework provides a reliable testbed to evaluate and optimize the efficacy of gene drive mosquito releases. What separates MGDrivE from other models is the incorporation of mathematical and computational mechanisms to simulate a wide array of inheritance-based technologies within the same, coherent set of equations. We do this by treating the population dynamics, genetic inheritance operations, and migration between habitats as separate processes coupled together through the use of mathematical tensor operations. This way we can conveniently swap inheritance patterns whilst still making use of the same set of population dynamics equations. This is a crucial advantage of our system, as it allows other research groups to test their ideas without developing new models and without the need to spend time adapting other frameworks to suit their needs.

40 MGDrivE

#### **Brief Description**

MGDrivE is based on the idea that we can decouple the genotype inheritance process from the population dynamics equations. This allows the system to be treated and developed in three semi-indpendent modules that come together to form the system. The way this is done will be described later in this document but a reference diagram is shown here.



## **Previous Work**

The original version of this model was based on work by (Deredec et al. 2011; Hancock and Godfray 2007) and adapted to accommodate CRISPR homing dynamics in a previous publication by our team (Marshall et al. 2017). As it was described, we extended this framework to be able to handle a variable number of genotypes, and migration across spatial scenarios. We did this by adapting the equations to work in a tensor-oriented manner, where each genotype can have different processes affecting their particular strain (death rates, mating fitness, sex-ratio bias, et cetera).

#### **Notation and Conventions**

Before beginning the full description of the model we will define some of the conventions we followed for the notation of the written description of the system.

- · Overlines are used to denote the dimension of a tensor
- Subscript brackets are used to indicate an element in time. For example:  $L_{[t-1]}$  is the larval population at time: t-1.
- Parentheses are used to indicate the parameter(s) of a function. For example:  $\overline{O(T_e+T_l)}$  represents the function O evaluated with the parameter:  $T_e+T_l$
- Matrices follow a 'row-first' indexing order (i: row, j: column)

MGDrivE-Cube 41

In the case of one dimensional tensors, each slot represents a genotype of the population. For example, the male population is stored in the following way:

$$\overline{Am} = \begin{pmatrix} g_1 \\ g_2 \\ g_3 \\ \vdots \\ g_n \end{pmatrix}_i$$

All the processes that affect mosquitoes in a genotype-specific way are defined and stored in this way within the framework.

There are two tensors of squared dimensionality in the model: the adult females matrix, and the genotype-specific viability mask. In the case of the former the rows represent the females' genotype, whilst the columns represent the genotype of the male they mated with:

$$\overline{\overline{Af}} = \begin{pmatrix}
g_{11} & g_{12} & g_{13} & \cdots & g_{1n} \\
g_{21} & g_{22} & g_{23} & \cdots & g_{2n} \\
g_{31} & g_{32} & g_{33} & \cdots & g_{3n} \\
\vdots & \vdots & \vdots & \ddots & \vdots \\
g_{n1} & g_{n2} & g_{n3} & \cdots & g_{nn}
\end{pmatrix}_{ij}$$

The genotype-specific viability mask, on the other hand, stores the mothers' genotype in the rows, and the potential eggs' genotype in the columns of the matrix.

#### References

Deredec A, Godfray HCJ, Burt A (2011). "Requirements for effective malaria control with homing endonuclease genes." *Proceedings of the National Academy of Sciences of the United States of America*, **108**(43), E874–80. ISSN 1091-6490, doi: 10.1073/pnas.1110717108, http://www.ncbi.nlm.nih.gov/pubmed/21976487{%}5Cnhttp://www.pubmedcentral.nih.gov/articlerender.fcgi?artid=PMC3203790.

Hancock PA, Godfray HCJ (2007). "Application of the lumped age-class technique to studying the dynamics of malaria-mosquito-human interactions." *Malaria journal*, **6**, 98. ISSN 1475-2875, doi: 10.1186/14752875698, http://www.pubmedcentral.nih.gov/articlerender.fcgi?artid=1971713{&}tool=pmcentrez{&}rendertype=abstract.

Marshall J, Buchman A, C. HMS, Akbari OS (2017). "Overcoming evolved resistance to population-suppressing homing-based gene drives." *Nature Scientific Reports*, 1–46. ISSN 2045-2322, doi: https://doi.org/10.1101/088427.

MGDrivE-Cube

MGDrivE: Inheritance Cube

#### **Description**

To model an arbitrary number of genotypes efficiently in the same mathematical framework we use a 3-dimensional array structure (cube) where each axis represents the following information:

- x: female adult mate genotype
- y: male adult mate genotype
- z: proportion of the offspring that inherits a given genotype (layer)

#### **Details**

The cube structure gives us the flexibility to apply tensor operations to the elements within our equations, so that we can calculate the stratified population dynamics rapidly; and within a readable, flexible computational framework. This becomes apparent when we define the equation we use for the computation of eggs laid at any given point in time:

$$\overline{O(T_x)} = \sum_{j=1}^{n} \left( \left( (\beta * \overline{s} * \overline{\overline{Af_{[t-T_x]}}}) * \overline{\overline{\overline{Ih}}} \right) * \Lambda \right)_{ij}^{\top}$$

In this equation, the matrix containing the number of mated adult females  $(\overline{Af})$  is multiplied element-wise with each one of the layers containing the eggs genotypes proportions expected from this cross  $(\overline{\overline{Ih}})$ . The resulting matrix is then multiplied by a binary 'viability mask'  $(\Lambda)$  that filters out female-parent to offspring genetic combinations that are not viable due to biological impediments (such as cytoplasmic incompatibility). The summation of the transposed resulting matrix returns us the total fraction of eggs resulting from all the male to female genotype crosses  $(\overline{O(T_x)})$ .

Note: For inheritance operations to be consistent within the framework the summation of each element in the z-axis (this is, the proportions of each one of the offspring's genotypes) must be equal to one.

#### **Drive-specific Cubes**

An inheritance cube in an array object that specifies inheritance probabilities (offspring genotype probability) stratified by male and female parent genotypes. MGDrivE provides the following cubes to model different gene drive systems:

- Cube\_oneLocusTA: 1 Locus Maternal-Toxin/Zygotic-Antidote System
- Cube\_twoLocusTA: 2 Locus Maternal-Toxin/Zygotic-Antidote System
- Cube\_Homing1RA: Homing Drive with 1 Resistance Allele
- Cube\_HomingDrive: CRISPR (Clustered Regularly Interspaced SWort Palindromic Repeats)
   witW 2 Resistance Allele
- Cube\_KillerRescue: Killer-Rescue System
- Cube\_MEDEA: MEDEA (Maternal Effect Dominant Embryonic Arrest)
- Cube\_ReciprocalTranslocations: Reciprocal Translocation
- Cube\_RIDL: RIDL (Release of Insects with Dominant Lethality)
- Cube\_Mendelian: Mendelian
- Cube\_Wolbachia: Wolbachia

## **Functions for Cubes**

There are several functions to operate on cube objects.

• cube2csv: Export slices of a cube to .csv format

MGDrivE-Model 43

MGDrivE-Model

MGDrivE: Model's Mathematical Description

#### **Description**

The original version of this model was based on work by (Deredec, Godfray, and Burt 2011; Hancock and Godfray 2007) and adapted to accommodate CRISPR homing dynamics in a previous publication by our team (Marshall, Buchman, C., and Akbari 2017). As it was described, we extended this framework to be able to handle a variable number of genotypes, and migration across spatial scenarios. We did this by adapting the equations to work in a tensor-oriented manner, where each genotype can have different processes affecting their particular strain (death rates, mating fitness, sex-ratio bias, etcetera).

#### **Inheritance Cube and Oviposition**

To allow the extension of the framework to an arbitrary number of genotypes we decided to transform traditional inheritance matrices into inheritance cubes where each of the axis represents the following information:

- x: female adult mate genotype
- y: male adult mate genotype
- z: proportion of the offspring that inherits a given genotype (slice)

The 'cube' structure gives us the flexibility to apply tensor operations to the elements within our equations, so that we can calculate the stratified population dynamics rapidly; and within a readable, flexible computational framework. This becomes apparent when we define the equation we use for the computation of eggs laid at any given point in time:

$$\overline{O(T_x)} = \sum_{j=1}^{n} \left( \left( (\beta * \overline{s} * \overline{\overline{Af_{[t-T_x]}}}) * \overline{\overline{\overline{Ih}}} \right) * \Lambda \right)_{ij}^{\top}$$

In this equation, the matrix containing the number of mated adult females  $(\overline{Af})$  is multiplied element-wise with each one of the slices containing the eggs genotypes proportions expected from this cross  $(\overline{Ih})$ . The resulting matrix is then multiplied by a binary 'viability mask'  $(\Lambda)$  that filters out female-parent to offspring genetic combinations that are not viable due to biological impediments (such as cytoplasmic incompatibility). The summation of the transposed resulting matrix returns us the total fraction of eggs resulting from all the male to female genotype crosses  $(O(T_x))$ .

Note: For inheritance operations to be consistent within the framework the summation of each element in the 'z' axis (this is, the proportions of each one of the offspring's genotypes) must be equal to one.

#### **Population Dynamics**

During the three aquatic stages, a density-independent mortality process takes place:

$$\theta_{st} = (1 - \mu_{st})^{T_{st}}$$

Along with a density dependent process dependent on the number of larvae in the environment:

$$F(L[t]) = \left(\frac{\alpha}{\alpha + \sum \overline{L[t]}}\right)^{1/T_l}$$

where  $\alpha$  represents the strength of the density-dependent process. This parameter is calculated with:

$$\alpha = \left(\frac{1/2 * \beta_k * \theta_e * Ad_{eq}}{R_m - 1}\right) * \left(\frac{1 - (\theta_l/R_m)}{1 - (\theta_l/R_m)^{1/T_l}}\right)$$

in which  $\beta_k$  is the species' fertility in the absence of gene-drives,  $Ad_{eq}$  is the adult mosquito population equilibrium size, and  $R_m$  is the population growth in the absence of density-dependent mortality. This population growth is calculated with the average generation time (g), the adult mortality rate  $(\mu_{ad})$ , and the daily population growth rate  $(r_m)$ :

$$g = T_e + T_l + T_p + \frac{1}{\mu_{ad}} R_m = (r_m)^g$$

**Larval Stages:** The computation of the larval stage in the population is crucial to the model because the density dependent processes necessary for equilibrium trajectories to be calculated occur here. This calculation is performed with the following equation:

$$D(\theta_{l}, T_{x}) = \begin{cases} \theta'_{l[0]} = \theta_{l} & i = 0\\ \theta'_{l[i+1]} = \theta'_{l[i]} * F(\overline{L_{[t-i-T_{x}]}}) & i \leq T_{l} \end{cases}$$

In addition to this, we need the larval mortality  $(\mu_l)$ :

$$\mu_l = 1 - \left(\frac{R_m * \mu_{ad}}{1/2 * \beta_k * (1 - \mu_m)}\right)^{\frac{1}{T_e + T_l + T_p}}$$

With these mortality processes, we are now able to calculate the larval population:

$$\overline{L_{[t]}} = \overline{L_{[t-1]}} * (1 - \mu_l) * F(\overline{L_{[t-1]}}) + \overline{O(T_e)} * \theta_e - \overline{O(T_e + T_l)} * \theta_e * D(\theta_l, 0)$$

where the first term accounts for larvae surviving one day to the other; the second therm accounts for the eggs that have hatched within the same period of time; and the last term computes the number of larvae that have transformed into pupae.

**Adult Stages:** We are ultimately interested in calculating how many adults of each genotype exist at any given point in time. For this, we first calculate the number of eggs that are laid and survive to the adult stages with the equation:

$$\overline{E'} = \overline{O(T_e + T_l + T_p)} \left( \overline{\xi_m} * (\theta_e * \theta_p) * (1 - \mu_{ad}) * D(\theta_l, T_p) \right)$$

With this information we can calculate the current number of male adults in the population by computing the following equation:

$$\overline{Am_{[t]}} = \overline{Am_{[t-1]}} * (1 - \mu_{ad}) * \overline{\omega_m} + (1 - \overline{\phi}) * \overline{E'} + \overline{\nu m_{[t-1]}}$$

in which the first term represents the number of males surviving from one day to the next; the second one, the fraction of males that survive to adulthood  $(\overline{E'})$  and emerge as males  $(1-\phi)$ ; the last one is used to add males into the population as part of gene-drive release campaigns. Female adult populations are calculated in a similar way:

$$\overline{\overline{Af_{[t]}}} = \overline{\overline{Af_{[t-1]}}} * (1 - \mu_{ad}) * \overline{\omega_f} + \left(\overline{\phi} * \overline{E'} + \overline{\nu f_{[t-1]}}\right)^\top * \left(\frac{\overline{\eta} * \overline{Am_{[t-1]}}}{\sum \overline{Am_{[t-1]}}}\right)$$

where we first compute the surviving female adults from one day to the next; and then we calculate the mating composition of the female fraction emerging from pupa stage. To do this, we obtain the surviving fraction of eggs that survive to adulthood  $(\overline{E'})$  and emerge as females  $(\phi)$ , we then add the new females added as a result of gene-drive releases  $(\overline{\nu f_{[t-1]}})$ . After doing this, we calculate the proportion of males that are allocated to each female genotype, taking into account their respective mating fitnesses  $(\overline{\eta})$  so that we can introduce the new adult females into the population pool.

MGDrivE-Model 45

#### **Gene Drive Releases and Effects**

As it was briefly mentioned before, we are including the option to release both male and/or female individuals into the populations. Another important thing to emphasize is that we allow flexible releases sizes and schedules. Our model handles releases internally as lists of populations compositions so, it is possible to have releases performed at irregular intervals and with different numbers of mosquito genetic compositions as long as no new genotypes are introduced (which have not been previously defined in the inheritance cube).

$$\overline{\nu} = \left\{ \begin{pmatrix} g_1 \\ g_2 \\ g_3 \\ \vdots \\ g_n \end{pmatrix}_{t=1}, \begin{pmatrix} g_1 \\ g_2 \\ g_3 \\ \vdots \\ g_n \end{pmatrix}_{t=2}, \dots, \begin{pmatrix} g_1 \\ g_2 \\ g_3 \\ \vdots \\ g_n \end{pmatrix}_{t=x} \right\}$$

So far, however, we have not described the way in which the effects of these gene-drives are included into the mosquito populations dynamics. This is done through the use of various modifiers included in the equations:

- $\overline{\omega}$ : Relative increase in mortality (zero being full mortality effects and one no mortality effect)
- $\overline{\phi}$ : Relative shift in the sex of the pupating mosquitoes (zero biases the sex ratio towards males, whilst 1 biases the ratio towards females).
- <del>\overline{\eta}</del>: Standardized mating fitness (zero being complete fitness ineptitude, and one being regular mating skills).
- $\overline{\beta}$ : Fecundity (average number of eggs laid).
- $\overline{\xi}$ : Pupation success (zero being full mortality and one full pupation success).

#### Migration

To simulate migration within our framework we are considering patches (or nodes) of fully-mixed populations in a network structure. This allows us to handle mosquito movement across spatially-distributed populations with a transitions matrix, which is calculated with the tensor outer product of the genotypes populations tensors and the transitions matrix of the network as follows:

$$\overline{Am_{(t)}^i} = \sum \overline{A_m^j} \otimes \overline{\overline{\tau m_{[t-1]}}} \overline{\overline{Af_{(t)}^i}} = \sum \overline{\overline{A_f^j}} \otimes \overline{\overline{\tau f_{[t-1]}}}$$

In these equations the new population of the patch i is calculated by summing the migrating mosquitoes of all the j patches across the network defined by the transitions matrix  $\tau$ , which stores the mosquito migration probabilities from patch to patch. It is worth noting that the migration probabilities matrices can be different for males and females; and that there's no inherent need for them to be static (the migration probabilities may vary over time to accommodate wind changes due to seasonality).

# Parameters

This table compiles all the parameters required to run MGDrivE clustered in six categories:

- Life Stages: These deal with the structure of mosquito population.
- Bionomics: This set of parameters is related to the behavior of the specific mosquito species being modeled.
- Gene Drive: Genotype-specific vectors of parameters that affect how each gene-drive modifies the responses of populations to them.

- Releases: List of vectors that control the release of genetically-modified mosquitoes.
- Population: General mosquito-population parameters that control environmentally-determined variables.

• Network: Related to migration between nodes of population units

#### **Stochasticity**

*MGDrivE* allows stochasticity to be included in the dynamics of various processes; in an effort to simulate processes that affect various stages of mosquitoes lives. In the next section, we will describe all the stochastic processes that can be activated in the program. It should be noted that all of these can be turned on and off independently from one another as required by the researcher.

## Mosquito Biology: Oviposition

Stochastic egg laying by female/male pairs is separated into two steps: calculating the number of eggs laid by the females and then distributing laid eggs according to their genotypes. The number of eggs laid follows a Poisson distribution conditioned on the number of female/male pairs and the fertility of each female.

$$Poisson(\lambda = numFemales * Fertility)$$

Multinomial sampling, conditioned on the number of offspring and the relative viability of each genotype, determines the genotypes of the offspring.

$$Multinomial (numOffspring, p_1, p_2 \dots p_b) = \frac{numOffspring!}{p_1! p_2 \dots p_n} p_1^{n_1} p_2^{n_2} \dots p_n^{n_n}$$

#### **Sex Determination**

Sex of the offspring is determined by multinomial sampling. This is conditioned on the number of eggs that live to hatching and a probability of being female, allowing the user to design systems that skew the sex ratio of the offspring through reproductive mechanisms.

$$Multinomial(numHatchingEggs, p_{female}, p_{female})$$

**Mating** Stochastic mating is determined by multinomial sampling conditioned on the number of males and their fitness. It is assumed that females mate only once in their life, therefore each female will sample from the available males and be done, while the males are free to potentially mate with multiple females. The males' ability to mate is modulated with a fitness term, thereby allowing some genotypes to be less fit than others (as seen often with lab releases).

$$Multinomial(numFemales, p_1f_1, p_2f_2, \dots p_nf_n)$$

#### **Hatching**

Other Stochastic Processes All remaining stochastic processes (larval survival, hatching, pupating, surviving to adult hood) are determined by multinomial sampling conditioned on factors affecting the current life stage. These factors are determined empirically from mosquito population data.

**Migration:** Variance of stochastic movement (not used in diffusion model of migration). It affects the concentration of probability in the Dirchlet simplex, small values lead to high variance and large values lead to low variance.

MGDrivE.Setup 47

#### References

Deredec A, Godfray HCJ, Burt A (2011). "Requirements for effective malaria control with homing endonuclease genes." *Proceedings of the National Academy of Sciences of the United States of America*, **108**(43), E874–80. ISSN 1091-6490, doi: 10.1073/pnas.1110717108, http://www.ncbi.nlm.nih.gov/pubmed/21976487{%}5Cnhttp://www.pubmedcentral.nih.gov/articlerender.fcgi?artid=PMC3203790.

Hancock PA, Godfray HCJ (2007). "Application of the lumped age-class technique to studying the dynamics of malaria-mosquito-human interactions." *Malaria journal*, **6**, 98. ISSN 1475-2875, doi: 10.1186/14752875698, http://www.pubmedcentral.nih.gov/articlerender.fcgi?artid=1971713{&}tool=pmcentrez{&}rendertype=abstract.

Marshall J, Buchman A, C. HMS, Akbari OS (2017). "Overcoming evolved resistance to population-suppressing homing-based gene drives." *Nature Scientific Reports*, 1–46. ISSN 2045-2322, doi: https://doi.org/10.1101/088427.

MGDrivE.Setup

Setup MGDrivE

## **Description**

Initialize methods in Patch to run deterministic or stochastic simulations. This function must be called prior to any objects being created.

#### Usage

MGDrivE.Setup(stochasticityON = FALSE)

## **Arguments**

stochasticityON

enable/disable stochastic simulation.

moveMatAll2

Movement Matrix: All 2

## **Description**

A movement matrix for simulation with 3 patches.

#### Usage

data(moveMatAll2)

#### **Format**

A matrix with 3 rows and 3 columns:

Patches 1 and 3 are sources for patch 2 which is a sink.

moveMatCascade3

Movement Matrix: Cascade 3

# Description

A movement matrix for simulation with 3 patches.

# Usage

data(moveMatCascade3)

#### **Format**

A matrix with 3 rows and 3 columns:

Mosquitoes in patch 1 have equal probability to stay or move to 2; mosquitoes in patch 2 have equal probability to stay or move to 3; mosquitoes in patch 3 stay there.

moveMatDiag

Movement Matrix: Diagonal

## **Description**

A movement matrix for simulation with 3 patches.

## Usage

data(moveMatDiag)

# Format

A matrix with 3 rows and 3 columns:

3 independent patches.

moveMatDiagOneCity

Movement Matrix: Diagonal One City

# Description

A movement matrix for simulation with 1 patch.

# Usage

data(moveMatDiagOneCity)

#### **Format**

A matrix with 1 rows and 1 columns:

A 1 by 1 matrix with entry 1.

moveMatDie 49

moveMatDie

Movement Matrix: Die

## **Description**

A movement matrix for simulation with 3 patches.

## Usage

data(moveMatDie)

#### **Format**

A matrix with 3 rows and 3 columns:

All entries of matrix are 0 for testing that all mosquitoes will be killed.

moveMatIndependent3

Movement Matrix: Independent 3

## **Description**

A movement matrix for simulation with 3 patches.

# Usage

data(moveMatIndependent3)

#### **Format**

A matrix with 3 rows and 3 columns:

Mosquitoes in patch 1 stay with probability 0.975, move to patch 2 with probability 0.025, mosquitoes in patch 2 and 3 stay in their patches.

moveMatMixedSpil

Movement Matrix: Mixed Spill

## **Description**

A movement matrix for simulation with 3 patches.

## Usage

data(moveMatMixedSpil)

#### **Format**

A matrix with 3 rows and 3 columns:

Mosquitoes in patch 1 stay with probability 0.999, move to patch 2 with probability 0.001, mosquitoes in patch 2 and 3 stay in their patches.

50 moveMatTriple

moveMatTaleOfTwoCities

Movement Matrix: Tale of Two Cities

## **Description**

A movement matrix for simulation with 2 patches.

## Usage

data(moveMatTaleOfTwoCities)

#### **Format**

A matrix with 2 rows and 2 columns:

Mosquitoes do not move between the two patches.

moveMatTriDiagonal

Movement Matrix: Tri-diagonal

## **Description**

A movement matrix for simulation with 12 patches.

# Usage

data(moveMatTriDiagonal)

#### **Format**

A matrix with 12 rows and 12 columns:

Tri-diagonal matrix with approximately 0.985 probability on diagonal and rest of probability mass on k-1 and k+1 off-diagonal elements.

moveMatTriple

Movement Matrix: Triple

## **Description**

A movement matrix for simulation with 3 patches.

# Usage

data(moveMatTriple)

#### **Format**

A matrix with 3 rows and 3 columns:

All entries of matrix are 1 for testing that mosquitoes will be produced.

Network 51

Network

Network Class Definition

#### **Description**

A Network class object stores all the information for a simulation on a defined landscape.

## Usage

Network

#### **Format**

An R6Class generator object

#### Constructor

- networkParameters: see Network.Parameters
- driveCube: an inheritance cube, see MGDrivE-Cube for available cubes
- patchReleases: see Release\_basicRepeatedReleases for examples on how to set up release schedules
- migrationMale: a stochastic matrix whose dimensions conform to the number of patches
- migrationFemale: a stochastic matrix whose dimensions conform to the number of patches
- · directory: character string of output directory

#### Methods

- get\_moveVar: see get\_moveVar\_Network
- get\_timeAq: see get\_timeAq\_Network
- get\_thetaAq: see get\_thetaAq\_Network
- get\_windowSize: see get\_windowSize\_Network
- get\_beta: see get\_beta\_Network
- get\_muAd: see get\_muAd\_Network
- get\_rm: see get\_rm\_Network
- get\_AdPopEQ: see get\_AdPopEQ\_Network
- get\_g: see get\_g\_Network
- get\_Rm: see get\_Rm\_Network
- get\_muAq: see get\_muAq\_Network
- get\_alpha: see get\_alpha\_Network
- get\_Leq\_Network: see get\_Leq\_Network
- get\_driveCube\_genotype: see get\_driveCube\_genotype\_Network
- get\_driveCube\_index: see get\_driveCube\_index\_Network
- get\_tau: see get\_tau\_Network
- get\_genotypesID: see get\_genotypesID\_Network
- get\_genotypesN: see get\_genotypesN\_Network

52 Network

- get\_wildType: see get\_wildType\_Network
- get\_eta: see get\_eta\_Network
- get\_phi: see get\_phi\_Network
- get omega: see get\_omega\_Network
- get xiF: see get\_xiF\_Network
- get\_xiM: see get\_xiM\_Network
- get\_s: see get\_s\_Network
- get\_releaseType: see get\_releaseType\_Network
- get\_patch: see get\_patch\_Network
- get\_patches: see get\_patches\_Network
- get\_nPatch: see get\_nPatch\_Network
- get\_directory: see get\_directory\_Network
- get\_simTime: see get\_simTime\_Network
- get conADM: see get\_conADM\_Network
- get conAF1: see get\_conAF1\_Network
- close\_allConnections: see close\_allConnections\_Network
- get\_tNow: see get\_tNow\_Network
- get\_migrationMale: see get\_migrationMale\_Network
- get\_migrationMaleRow: see get\_migrationMaleRow\_Network
- set\_migrationMale: see set\_migrationMale\_Network
- get\_migrationFemale: see get\_migrationFemale\_Network
- get\_migrationFemaleRow: see get\_migrationFemaleRow\_Network
- set\_migrationFemale: see set\_migrationFemale\_Network
- get\_patchReleases: see get\_patchReleases\_Network
- oneDay\_Migration: see oneDay\_Migration\_Network
- reset: see reset\_Network
- oneRun: see oneRun\_Network
- oneDay: see oneDay\_Network

#### **Fields**

- parameters: see Network.Parameters
- patches: a list of Patch objects
- nPatch: number of patches
- simTime: maximum time of simulation
- driveCube: an inheritance cube
- tNow: current time of simulation (time starts at 2 because time 1 is the initial equilibrium state)
- runID: an identifier for the current simulation run, useful for Monte Carlo simulation
- directory: a character string of where to store output
- conADM: a connection to write male population dynamics out to
- conAF1: a connection to write female population dynamics out to
- migrationMale: a stochastic matrix whose dimensions conform to the number of patches
- migrationFemale: a stochastic matrix whose dimensions conform to the number of patches
- patchReleases: a list of release schedules for each patch

Network.Parameters 53

Network.Parameters Network Parameters

# Description

Generate parameters for simulation on a Network. Parameters average generation time g, population growth rate  $R_m$ , aquatic mortality  $\mu_{Aq}$ , and aquatic survival  $\theta_{Aq}$  are shared between patches and calculated by calcAverageGenerationTime, calcPopulationGrowthRate, calcLarvalStageMortalityRate, and calcAquaticStagesSurvivalProbability.

Patch-specific parameters  $\alpha$  and  $L_{eq}$  are calculated for each patch by calcDensityDependentDeathRate and calcLarvalPopEquilibrium.

#### Usage

```
Network.Parameters(nPatch, simTime, parallel = FALSE, moveVar = 1000,
  tEgg = 1L, tLarva = 14L, tPupa = 1L, beta = 32, muAd = 0.123,
  popGrowth = 1.096, AdPopEQ, runID = 1L)
```

## **Arguments**

nPatch	number of Patch
simTime	maximum time to run simulation
parallel	append process id (see link[base]{Sys.getpid}) to output files for running in parallel
moveVar	variance of stochastic movement (not used in diffusion model of migration). It affects the concentration of probability in the Dirchlet simplex, small values lead to high variance and large values lead to low variance.
tEgg	length of egg stage
tLarva	length of larval instar stage
tPupa	length of pupal stage
beta	female egg batch size of wild-type
muAd	wild-type daily adult mortality (1/muAd is average wild-type lifespan)
popGrowth	daily population growth rate (used to calculate equilibrium)
AdPopEQ	vector of adult population size at equilibrium
runID	begin counting runs with this set of parameters from this value
normalise	Normalise a Numeric Vector

# Description

Normalise a numeric vector to sum to one

# Usage

```
normalise(vector)
```

# Arguments

vector numeric vector

oneDay\_admPupating\_deterministic\_Patch

Deterministc Adult Male Pupation

#### **Description**

Adult male emergence is calculated based on the number of male pupae multiplied by

$$\left(\overline{\xi_m} * (\theta_e * \theta_p) * (1 - \mu_{ad}) * D(\theta_l, T_p)\right)$$

, where  $\overline{\xi_m}$  is the genotype-specific pupation success probability.

## Usage

oneDay\_admPupating\_deterministic\_Patch()

## **Description**

Pupation from male pupae to adults is sampled from a binomial distribution for each genotype where the probability of pupation is given by

$$\left(\overline{\xi_m} * (\theta_e * \theta_p) * (1 - \mu_{ad}) * D(\theta_l, T_p)\right)$$

, where  $\overline{\xi_m}$  is the genotype-specific pupation success probability.

## Usage

oneDay\_admPupating\_stochastic\_Patch()

 $one {\it Day\_admSurvival\_deterministic\_Patch} \\ Deterministc~Adult~Male~Survival$ 

## **Description**

Daily adult male survival is calculated according to

$$\overline{\overline{Am_{[t-1]}}} * (1 - \mu_{ad}) * \overline{\omega_m}$$

, where  $\mu_{ad}$  is adult mortality rate and  $\overline{\omega_m}$  correponds to genotype-specific mortality effects.

#### Usage

oneDay\_admSurvival\_deterministic\_Patch()

 ${\it one Day\_adm Survival\_stochastic\_Patch} \\ {\it Stochastic Adult Male Survival}$ 

# Description

Daily adult male survival is sampled from a binomial distribution where survival probability is given by

$$(1-\mu_{ad})*\overline{\omega_m}$$

, where  $\mu_{ad}$  is adult mortality rate and  $\overline{\omega_m}$  correponds to genotype-specific mortality effects.

#### Usage

oneDay\_admSurvival\_stochastic\_Patch()

## **Description**

Mating is calculated as the outer product of newly emerging adult females and adult males, modulated by  $\overline{\eta}$ , genotype-specific male mating fitness.

## Usage

oneDay\_af1Mating\_deterministic\_Patch()

# **Description**

Mating for each newly emerging adult female genotype is sampled from a multinomial distribution with probabilities equal to the adult male population vector multiplied by  $\overline{\eta}$ , genotype-specific male mating fitness.

## Usage

oneDay\_af1Mating\_stochastic\_Patch()

oneDay\_af1Pupation\_deterministic\_Patch

Deterministc Adult Female Pupation

#### **Description**

Adult female emergence is calculated based on the number of female pupae multiplied by

$$\left(\overline{\xi_f} * (\theta_e * \theta_p) * (1 - \mu_{ad}) * D(\theta_l, T_p)\right)$$

, where  $\overline{\xi_f}$  is the genotype-specific pupation success probability.

## Usage

oneDay\_af1Pupation\_deterministic\_Patch()

oneDay\_af1Pupation\_stochastic\_Patch
Stochastic Adult Female Pupation

## **Description**

Pupation from female pupae to adults is sampled from a binomial distribution for each genotype where the probability of pupation is given by

$$\left(\overline{\xi_f} * (\theta_e * \theta_p) * (1 - \mu_{ad}) * D(\theta_l, T_p)\right)$$

, where  $\overline{\xi_f}$  is the genotype-specific pupation success probability.

## Usage

oneDay\_af1Pupation\_stochastic\_Patch()

oneDay\_af1Survival\_deterministic\_Patch

Deterministc Adult Female Survival

## **Description**

Daily adult female survival is calculated according to

$$\overline{\overline{Af_{[t-1]}}} * (1 - \mu_{ad}) * \overline{\omega_f}$$

, where  $\mu_{ad}$  is adult mortality rate and  $\overline{\omega_f}$  correponds to genotype-specific mortality effects.

#### Usage

oneDay\_af1Survival\_deterministic\_Patch()

## **Description**

Daily adult female survival is sampled from a binomial distribution where survival probability is given by

$$(1 - \mu_{ad}) * \overline{\omega_f}$$

, where  $\mu_{ad}$  is adult mortality rate and  $\overline{\omega_f}$  correponds to genotype-specific mortality effects.

#### Usage

oneDay\_af1Survival\_stochastic\_Patch()

oneDay\_calcCumulativeLarvalDensityDependentFactor\_Patch

Calculate Cumulative Larval density-dependent Mortality

## **Description**

Calculate

$$F(L[t]) = \left(\frac{\alpha}{\alpha + \sum \overline{L[t]}}\right)^{1/T_l}$$

, the cumulative effect of density-dependence on larval mortality over the entire duration of larval stage.

#### Usage

oneDay\_calcCumulativeLarvalDensityDependentFactor\_Patch()

oneDay\_calcCumulativePupaDensityDependentFactor\_Patch

Calculate Cumulative Pupae density-dependent Mortality

# Description

Calculate

$$D(\theta_{l}, T_{P}) = \begin{cases} \theta'_{l[0]} = \theta_{l} & i = 0\\ \theta'_{l[i+1]} = \theta'_{l[i]} * F(\overline{L_{[t-i-T_{P}]}}) & i \leq T_{P} \end{cases}$$

, the cumulative effect of density-dependence on larval mortality over the entire duration of larval and pupal stages.

#### Usage

oneDay\_calcCumulativePupaDensityDependentFactor\_Patch()

oneDay\_calcLarvalDensityDependentFactor\_Patch

Calculate Larval density-dependent Mortality

# Description

Calculate

$$D(\theta_{l}, T_{x}) = \begin{cases} \theta'_{l[0]} = \theta_{l} & i = 0\\ \theta'_{l[i+1]} = \theta'_{l[i]} * F(\overline{L_{[t-i-T_{x}]}}) & i \leq T_{l} \end{cases}$$

, the effect of density-dependence on larval mortality for one day.

## Usage

oneDay\_calcLarvalDensityDependentFactor\_Patch()

oneDay\_eggsFract2\_deterministic\_Patch

Deterministc Larval Pupation

## **Description**

Calculate the number of larvae that will pupate prior to calculating density-dependent effects in oneDay\_calcCumulativeLarvalDensityDependentFactor\_Patch.

# Usage

oneDay\_eggsFract2\_deterministic\_Patch()

oneDay\_eggsFract2\_stochastic\_Patch
Stochastic Larval Pupation

# Description

Calculate the number of larvae that will pupate prior to calculating density-dependent effects in oneDay\_calcCumulativeLarvalDensityDependentFactor\_Patch.

# Usage

oneDay\_eggsFract2\_stochastic\_Patch()

oneDay\_femaleReleases\_Patch

Release Female Mosquitoes in a Patch

#### **Description**

Based on this patch's release schedule, handle daily releases.

## Usage

```
oneDay_femaleReleases_Patch()
```

oneDay\_hatchingFract\_deterministic\_Patch

Deterministc Fraction of Eggs Maturing to Hatch

# **Description**

Calculate the fraction of hatching eggs accounting for delay given by:  $\overline{O(T_e)}$ 

## Usage

```
oneDay_hatchingFract_deterministic_Patch()
```

```
{\tt oneDay\_hatchingFract\_stochastic\_Patch}
```

Stochastic Fraction of Eggs Maturing to Hatch

## **Description**

Calculate the fraction of hatching eggs accounting for delay given by:  $\overline{O(T_e)}$  Stochasticity is introduced by assuming  $\overline{O(T_e)}$  defines the mean of a Poisson distributed number of eggs that can hatch.

#### Usage

```
oneDay_hatchingFract_stochastic_Patch()
```

```
oneDay_initOutput_Patch
```

Initialize Output from Focal Patch

## **Description**

Writes output to the text connections specified in the enclosing Network

# Usage

```
oneDay_initOutput_Patch()
```

oneDay\_larHatching\_deterministic\_Patch
Stochastic Egg Hatching to Larval Stage

#### **Description**

Calculate the eggs that have survived and hatched during a day given by:  $\overline{O(T_e)}*\theta_e$ 

## Usage

```
oneDay_larHatching_deterministic_Patch()
```

oneDay\_larHatching\_stochastic\_Patch
Stochastic Egg Hatching to Larval Stage

## **Description**

Calculate the eggs that have survived and hatched during a day given by:  $\overline{O(T_e)} * \theta_e$  The number of eggs that survive to hatch follows a binomial distribution.

## Usage

```
oneDay_larHatching_stochastic_Patch()
```

oneDay\_larPupating\_deterministic\_Patch

Deterministc Larval Pupation

## **Description**

Calculate the number of larvae that have transformed into pupae given by  $\overline{O(T_e+T_l)}*\theta_e*D(\theta_l,0)$ 

## Usage

```
oneDay_larPupating_deterministic_Patch()
```

oneDay\_larPupating\_stochastic\_Patch
Stochastic Larval Pupation

## **Description**

Calculate the number of larvae that have transformed into pupae given by  $\overline{O(T_e+T_l)}*\theta_e*D(\theta_l,0)$  This number follows a binomial distribution.

# Usage

```
oneDay_larPupating_stochastic_Patch()
```

 ${\tt one Day\_lar Survival\_deterministic\_Patch} \\ {\tt Deterministc\ Larval\ Survival}$ 

# Description

Calculate the number of larvae surviving from day to day, given by:

$$\overline{L_{[t-1]}} * (1-\mu_l) * F(\overline{L_{[t-1]}})$$

#### Usage

oneDay\_larSurvival\_deterministic\_Patch()

oneDay\_larSurvival\_stochastic\_Patch
Stochastic Larval Survival

# Description

Calculate the number of larvae surviving from day to day, given by:

$$\overline{L_{[t-1]}} * (1-\mu_l) * F(\overline{L_{[t-1]}})$$

Stochasticity is introduced by assuming  $(1-\mu_l)*F(\overline{L_{[t-1]}})$  defines binomial likelihood of survival for each genotype of larvae.

# Usage

oneDay\_larSurvival\_stochastic\_Patch()

oneDay\_maleReleases\_Patch

Release Male Mosquitoes in a Patch

# Description

Based on this patch's release schedule, handle daily releases.

## Usage

oneDay\_maleReleases\_Patch()

oneDay\_migrationIn\_Patch

Inbound Migration

## **Description**

Accumulate all inbound migration to this patch.

#### Usage

```
oneDay_migrationIn_Patch(maleIn, femaleIn)
```

## **Arguments**

maleIn vector of inbound migration femaleIn matrix of inbound migration

oneDay\_migrationOut\_deterministic\_Patch

Deterministic Oubound Migration from a Patch

## **Description**

Deterministic model of outbound migration of AF1new females from this patch, fills up the femaleMigration array.

#### Usage

```
oneDay_migrationOut_deterministic_Patch()
```

```
{\it one Day\_migration Out\_stochastic\_Patch} \\ {\it Stochastic~Oubound~Migration}
```

## **Description**

Stochastic model of migration of AF1new females from this patch, fills up the femaleMigration array. Migration is modeled as a Dirichlet-Multinomial process parameterized by moveVar multiplied by the row corresponding to this patch from the stochastic matrix. A Dirichlet distributed random variate is sampled from rdirichlet according to that parameter vector and then movement is sampled from rmultinom.

## Usage

```
oneDay_migrationOut_stochastic_Patch()
```

oneDay\_Migration\_Network

Inter-Patch Migration

## **Description**

Simulate migration between patches. See MGDrivE-Model, 'Migration' section for more details on how inter-patch migration is handled.

## Usage

```
oneDay_Migration_Network()
```

oneDay\_Network

Run a Single Day on a Network

#### **Description**

Runs a single day of simulation on a Network object, handling population dynamics, migration, population update, and output.

## Usage

```
oneDay_Network()
```

```
oneDay_numMaleFemale_deterministic_Patch

Deterministc Sex Ratio
```

## **Description**

Calculate the number of males,  $(1 - \overline{\phi}) * \overline{E'}$  and females,  $\overline{\phi} * \overline{E'}$ 

## Usage

```
oneDay_numMaleFemale_deterministic_Patch()
```

```
oneDay_numMaleFemale_stochastic_Patch
Stochastic Sex Ratio
```

#### **Description**

Calculate the number of males,  $(1-\overline{\phi})*\overline{E'}$  and females,  $\overline{\phi}*\overline{E'}$  These counts are follow a binomial distribution.

# Usage

```
oneDay_numMaleFemale_stochastic_Patch()
```

 ${\it one Day\_oviposit G1\_deterministic\_Patch} \\ {\it Deterministc~Oviposition}$ 

# Description

Calculate the number of eggs oviposited by female mosquitoes following:

$$\overline{O(T_x)} = \sum_{j=1}^{n} \left( \left( (\beta * \overline{s} * \overline{\overline{Af_{[t-T_x]}}}) * \overline{\overline{\overline{Ih}}} \right) * \Lambda \right)_{ij}^{\top}$$

# Usage

oneDay\_ovipositG1\_deterministic\_Patch()

## **Description**

Calculate the number of eggs oviposited by female mosquitoes following:

$$\overline{O(T_x)} = \sum_{j=1}^n \left( \left( (\beta * \overline{s} * \overline{\overline{Af_{[t-T_x]}}}) * \overline{\overline{\overline{Ih}}} \right) * \Lambda \right)_{ij}^{\top}$$

The deterministic result for number of eggs is used as the mean of a Poisson-distributed number of actual eggs oviposited.

# Usage

oneDay\_ovipositG1\_stochastic\_Patch()

oneDay\_PopDynamics\_Patch

Daily Population Dynamics for a Patch

#### **Description**

Run population dynamics (not including migration) for this patch.

#### Usage

oneDay\_PopDynamics\_Patch()

## **Description**

Update larval and adult populations daily at end of time step, calls shiftAndUpdatePopVector

## Usage

```
oneDay_updatePopulation_Patch()
```

```
oneDay_writeOutput_Patch
```

Write Output from Focal Patch

## **Description**

Writes output to the text connections specified in the enclosing Network

## Usage

```
oneDay_writeOutput_Patch()
```

oneRun\_Network

Run Simulation

#### **Description**

Run a single simulation on this network.

## Usage

```
oneRun_Network(conADM = NULL, conAF1 = NULL)
```

## **Arguments**

conADM an optional connection to write male population dynamics to, if NULL use the

directory specified in the constructor of Network with the current runID ap-

pended to the file.

conAF1 an optional connection to write female population dynamics to, if NULL use

the directory specified in the constructor of Network with the current runID

appended to the file.

66 Patch

Patch

Patch Class Definition

#### **Description**

A Patch is a single well-mixed population that is the smallest unit of simulation for MGDrivE.

#### Usage

Patch

#### **Format**

An R6Class generator object

#### Constructor

- patchID: integer ID of this patch
- genotypesID: character vector of genotypes
- simTime: maximum time of simulation
- · windowSize: necessary memory window size for model
- EGGt0: initial egg population,  $L_{eq}$
- LARt0: initial larval population
- PUPt0: initial pupae population
- ADMt0: initial adult male population,  $Ad_{eq}$
- AF1t0: initial adult female population,  $Ad_{eq}$
- maleReleases: integer ID of this patch
- $\bullet \ \ female Releases: female \ release \ schedule \ for \ this \ patch, see \ Release\_basic Repeated Releases$

#### Methods

- get\_patchID: see get\_patchID\_Patch
- get\_AF1new: see get\_AF1new\_Patch
- set\_AF1new: see set\_AF1new\_Patch
- get\_ADMnew: see get\_ADMnew\_Patch
- set\_ADMnew: see set\_ADMnew\_Patch
- accumulate\_ADMnew: see accumulate\_ADMnew\_Patch
- get\_EGG: see get\_EGG\_Patch
- get\_LAR: see get\_LAR\_Patch
- get\_PUP: see get\_PUP\_Patch
- get\_ADM: see get\_ADM\_Patch
- get\_AF1: see get\_AF1\_Patch
- get\_EGGdly: see get\_EGGdly\_Patch
- get\_LARdly: see get\_LARdly\_Patch

Patch 67

- get\_PUPdly: see get\_PUPdly\_Patch
- get\_ADMdly: see get\_ADMdly\_Patch
- get\_AF1dly: see get\_AF1dly\_Patch
- get\_maleMigration: see get\_maleMigration\_Patch
- get\_femaleMigration: see get\_femaleMigration\_Patch
- set\_NetworkPointer: see set\_NetworkPointer\_Patch
- get\_NetworkPointer: see get\_NetworkPointer\_Patch
- reset: see reset\_Patch
- oneDay\_initOutput: see oneDay\_initOutput\_Patch
- oneDay\_writeOutput: see oneDay\_writeOutput\_Patch
- oneDay\_migrationIn: see oneDay\_migrationIn\_Patch
- oneDay\_maleReleases: see oneDay\_maleReleases\_Patch
- oneDay\_femaleReleases: see oneDay\_femaleReleases\_Patch
- oneDay\_PopDynamics: see oneDay\_PopDynamics\_Patch
- oneDay\_updatePopulation: see oneDay\_updatePopulation\_Patch
- oneDay\_calcLarvalDensityDependentFactor: see oneDay\_calcLarvalDensityDependentFactor\_Patch
- one buy\_cutebut varbensity begonden a detor. See one buy\_cutebut varbensity begonden a detor. I deter
- oneDay\_calcCumulativeLarvalDensityDependentFactor: see oneDay\_calcCumulativeLarvalDensityDependent
- oneDay\_calcCumulativePupaDensityDependentFactor: see oneDay\_calcCumulativePupaDensityDependentFactor
- $\bullet \ \ one Day\_migration Out: see \ one Day\_migration Out\_stochastic\_Patch \ or \ one Day\_migration Out\_determinist on the property of the pr$
- oneDay\_ovipositG1: see oneDay\_ovipositG1\_stochastic\_Patch or oneDay\_ovipositG1\_deterministic\_Pat
- $\bullet \ \ one Day\_lar Survival: see \ one Day\_lar Survival\_stochastic\_Patch \ or \ one Day\_lar Survival\_deterministic\_Patch \ or$
- oneDay\_hatchingFract: see oneDay\_hatchingFract\_stochastic\_Patch or oneDay\_hatchingFract\_deterministic\_
   oneDay\_larHatching: see oneDay\_larHatching\_stochastic\_Patch or oneDay\_larHatching\_deterministic\_
- oneDay\_eggsFract2: see oneDay\_eggsFract2\_stochastic\_Patch or oneDay\_eggsFract2\_deterministic\_Pat
- oneDay\_larPupating: see oneDay\_larPupating\_stochastic\_Patch or oneDay\_larPupating\_deterministic\_larPupating\_stochastic\_Patch or oneDay\_larPupating\_stochastic\_Patch or oneDay\_larPupating\_stochastic\_P
- oneDay\_numMaleFemale: see oneDay\_numMaleFemale\_stochastic\_Patch or oneDay\_numMaleFemale\_determ
- oneDay\_admSurvival: see oneDay\_admSurvival\_stochastic\_Patch or oneDay\_admSurvival\_deterministic\_
- $\bullet \ \ one Day\_adm Pupating: see \ one Day\_adm Pupating\_stochastic\_Patch \ or \ one Day\_adm Pupating\_deterministic\_Patch \ or$
- $\bullet\ one Day\_af1 Survival: see\ one Day\_af1 Survival\_stochastic\_Patch\ or\ one Day\_af1 Survival\_deterministic\_Patch\ or\ one Day\_af1 Survival$
- oneDay\_af1Pupation: see oneDay\_af1Pupation\_stochastic\_Patch or oneDay\_af1Pupation\_deterministic\_
- oneDay\_aflMating: see oneDay\_af1Mating\_stochastic\_Patch or oneDay\_af1Mating\_deterministic\_Patch

#### **Fields**

- patchID: integer ID of this patch
- EGGt0: vector of initial egg stage population
- LARt0: vector of initial larval stage population
- PUPt0: vector of initial pupae stage population
- ADMt0: vector of initial adult male stage population
- AF1t0: matrix of initial adult female stage population
- EGG: egg stage population

68 Patch

- LAR: larvae stage population
- PUP: pupae stage population
- ADM: adult male stage population
- AF1: adult female stage population
- EGGdly: delay egg stage population
- LARdly: delay larvae stage population
- PUPdly: delay pupae stage population
- ADMdly: delay adult male stage population
- AF1dly: delay adult female stage population
- LARnew: new larval population after difference equations; needed to store population prior to migration exchange
- ADMnew: new adult male population after difference equations; needed to store population prior to migration exchange
- AF1new: new adult female population after difference equations; needed to store population prior to migration exchange
- maleMigration: matrix of outbound migrating males of dimension nGenotypes X nPatch
- femaleMigration: array of outbound migrating females of dimension nGenotypes X nGenotypes X nPatch
- NetworkPointer: a reference to enclosing Network
- ovipositG1: new eggs after oviposition by mated female mosquitoes
- larSurvival: surviving larvae
- hatchingFract: fraction of larvae that hatch
- larPupating: fraction of larvae that undergo pupation
- numMaleFemale: number of male vs. female emerging imago stage adults
- admSurvival: number of surviving adult males
- admPupating: number of pupating imago stage adults that become males
- af1Survival: number of surviving adult females
- af1Pupation: number of pupating imago stage adults that become females
- maleMatrix: row of male migration matrix corresponding to migration from this patch
- femaleMatrix: row of female migration matrix corresponding to migration from this patch
- larDDMortal: larval mortality
- f: density dependent factor in larval mortality

primePopMatrixArray 69

primePopMatrixArray Create a primed population array of matrices

## **Description**

Primes an array for the population history to be stored. The length of the array is equal to the window required for the model to run (in our specific case it is equal to the sum of aquatic stages lengths).

# Usage

```
primePopMatrixArray(primingMatrix, memoryWindow)
```

## **Arguments**

```
primingMatrix a named matrix population
memoryWindow integer size of list structure
```

primePopVectorArray Create a primed population array of vectors

## **Description**

Primes an array for the population history to be stored. The length of the array is equal to the window required for the model to run (in our specific case it is equal to the sum of aquatic stages lengths).

# Usage

```
primePopVectorArray(primingVector, memoryWindow)
```

# Arguments

primingVector a named vector population
memoryWindow integer size of list structure

70 rDirichlet

quantileC

Quantiles Function

## **Description**

Calculate the given quantiles of a matrix.

#### Usage

```
quantileC(Trials, Probs)
```

#### **Arguments**

Trials Integer matrix to calculate quantiles over

Probs Vector of quantiles

#### **Details**

This function calculates the given quantiles over the rows of an integer matrix. It uses method 8 of the stat::quantiles() function. It gives the same result, to numerical accuracy, and is designed to handle matrix input. It is only designed to work on integer matrices!

#### Value

Numeric Matrix

# **Examples**

```
Trials <- matrix(data = sample(x=1:100, size = 150, replace = TRUE), nrow=15, ncol=10)
DrawSize <- c(.25, .5, .75)
quantileC(Trials, Probs)</pre>
```

rDirichlet

Dirichlet Distribution

# Description

Make a single draw from a dirichlet distribution with the shape parameter one. This replaces the MCMCpack rDirichlet function, which was wholly written in R.

#### Usage

```
rDirichlet(migrationPoint)
```

## **Arguments**

migrationPoint Vector of weights for draws. Must be positive.

Release\_basicRepeatedReleases

Make List of Modified Mosquito Releases

## **Description**

Sets up a release schedule for a single patch, returns a list to be used in oneDay\_maleReleases\_Patch or oneDay\_femaleReleases\_Patch.

## Usage

```
Release_basicRepeatedReleases(genotypes, releaseStart, releaseEnd,
  releaseInterval, releaseVector, sex = "M")
```

## **Arguments**

```
genotypes possible genotypes

releaseStart day releases start

releaseEnd day releases end

releaseInterval

interval between releases

releaseVector named character vector of release composition

sex character in 'M','F'
```

## **Examples**

```
patchReleases = replicate(n = 3,expr = {
    list(maleReleases = NULL,femaleReleases = NULL)
},simplify = FALSE)

patchReleases[[1]]$femaleReleases = Release_basicRepeatedReleases(genotypes = Cube_Homing1RA$genotypesID,releases[[1]]$maleReleases = Release_basicRepeatedReleases(genotypes = Cube_Homing1RA$genotypesID,releases[[1]]$maleReleases = Release_basicRepeatedReleases(genotypes = Cube_Homing1RA$genotypesID,releases[[1]]$maleReleases = Release_basicRepeatedReleases(genotypes = Cube_Homing1RA$genotypesID,releases[[1]]$maleReleases = Release_basicRepeatedReleases(genotypes = Cube_Homing1RA$genotypesID,releases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleReleases[[1]]$maleRelease
```

reset\_Network Reset Network

# Description

Reset a Network between runs, useful for Monte Carlo simulation. This calls reset\_Patch on each patch and resets tNow = 2 and increments the runID.

# to setup for 3 patches but only release in the first with a defined release schedule:

#### Usage

```
reset_Network()
```

72 set\_ADMnew\_Patch

reset	Patch

Reset Patch to Initial Conditions

#### **Description**

Resets a patch to its initial configuration so that a new one does not have to be created and allocated in the network (for Monte Carlo simulation).

## Usage

```
reset_Patch()
```

retrieveOutput

Retrieve Output

## **Description**

Read in output from directory. The resulting object will be a nested list; outermost nesting dimension indexes runID, within runID elements are split by sex and innermost nesting is over patches.

## Usage

```
retrieveOutput(directory, genotypes)
```

# Arguments

directory directory where output was written to; must not end in path seperator genotypes character vector of possible genotypes; found in driveCube\$genotypesID

set\_ADMnew\_Patch

Set ADMnew

# Description

Set an element in new ADM males

#### Usage

```
set_ADMnew_Patch(count, genotype_M)
```

# Arguments

count number of mosquitoes genotype\_M genotype of male

set\_AF1new\_Patch 73

set\_AF1new\_Patch

Set AF1new

## **Description**

Set an element in new AF1 females

# Usage

```
set_AF1new_Patch(count, genotype_F, genotype_M)
```

# Arguments

count number of mosquitoes

genotype\_F genotype of female (row of matrix)
genotype\_M genotype of male (column of matrix)

set\_migrationFemale\_Network

Set Female Migration Matrix

## **Description**

Sets link{private\$migrationFemale} field

## Usage

```
set_migrationFemale_Network(migrationFemale)
```

# **Arguments**

migrationFemale

matrix object (rows must sum to one)

set\_migrationMale\_Network

Set Male Migration Matrix

# Description

Sets link{private\$migrationMale} field

# Usage

```
set_migrationMale_Network(migrationMale)
```

# Arguments

migrationMale matrix object (rows must sum to one)

74 splitOutput

```
set_NetworkPointer_Patch
```

Set Network Pointer

#### **Description**

Set a reference to the enclosing Network object

# Usage

```
set_NetworkPointer_Patch(NetworkPointer)
```

#### **Arguments**

NetworkPointer a Network object

```
{\it shift} And Update Pop Vector
```

Shift a Vector

## **Description**

Shift a population vector by one day and insert the new population. This was written to remove the dependency "binhf".

# Usage

```
shiftAndUpdatePopVector(popVector, newPop)
```

# **Arguments**

popVector	List of population vectors of length(Tegg+Tlarva+Tpupa)
newPop	Vector of length equal to the number of genotypes

splitOutput Split Output by Patch

# Description

Split output into multiple files by patches.

## Usage

```
splitOutput(directory, multiCore = FALSE)
```

# Arguments

directory Directory where output was written to; must not end in path seperator multiCore Write output using multiple cores? Default is FALSE

SymCubeC 75

SymCubeC

Make a Symmetric Cube

# **Description**

This function makes a lower-triangular cube symmetric over the z-axis. It was written to remove the dependency "Matrix". It is only used in building cubes.

# Usage

```
SymCubeC(lowerMat)
```

# Arguments

lowerMat

A lower-triangular matrix of depth 1 or more

turnStochasticityOnOrOff

Enable or Disable Stochastic Model

# Description

Set switches for deterministic or stochastic model

# Usage

```
turnStochasticityOnOrOff(on = TRUE)
```

## **Arguments**

on

enable/disable stochastic behaviour

# Index

*Topic <b>R6</b>	Cuba Haming1DA 14 42
Network, 51	Cube_Homing1RA, 14, 42
	Cube_HomingDrive, 15, 42
Patch, 66	Cube_KillerRescue, 16, 42
*Topic <b>class</b>	Cube_MEDEA, 17, 42
Network, 51	Cube_Mendelian, 18, 42
Patch, 66	Cube_oneLocusTA, 18, 42
*Topic datasets	Cube_ReciprocalTranslocations, 19, 42
kernels, 39	Cube_RIDL, 20, 42
moveMatAll2, 47	Cube_twoLocusTA, 20, 42
moveMatCascade3, 48	Cube_Wolbachia, 21, 42
moveMatDiag, 48	cubeModifiers, 14
moveMatDiagOneCity, 48	anacaDimactany 22
moveMatDie, 49	eraseDirectory, 22
moveMatIndependent3, 49	Exponential, 11, 12
moveMatMixedSpil, 49	GammaDist, 11
moveMatTaleOfTwoCities, 50	generateReleaseVector, 22
moveMatTriDiagonal, 50	get_ADM_Patch, 23, 66
moveMatTriple, 50	
1	get_ADMdly_Patch, 22, 67
accumulate_ADMnew_Patch, 5, 66	get_ADMnew_Patch, 23, 66
aggregateFemales, 5	get_AdPopEQ_Network, 23, 51
AnalyzeQuantiles, 6	get_AF1_Patch, 24, 66
1 F K 1 11	get_AF1dly_Patch, 23, 67
calc_ExpKernel, 11	get_AF1new_Patch, 24, 66
calc_GammaKernel, 11	get_alpha_Network, 24, 51
calc_haversine, <i>11</i> , 11, <i>12</i>	get_beta_Network, 24, 51
calc_HurdleExpKernel, 12	get_conADM_Network, 25, 52
calc_LognormalKernel, 12	get_conAF1_Network, 25, 52
calcAquaticStagesSurvivalProbability,	get_directory_Network, 25, 52
7, 35, 53	get_driveCube_genotype_Network, 25, 51
calcAquaticStageSurvivalProbability, 7,	get_driveCube_index_Network, 26, 51
7, 35	get_EGG_Patch, 26, 66
calcAverageGenerationTime, 8, 10, 28, 53	get_EGGdly_Patch, 26, 66
calcDensityDependentDeathRate, 8, 9, 24,	get_eta_Network, 27, 52
53	<pre>get_femaleMigration_Patch, 27, 67</pre>
calcLarvalPopEquilibrium, 9, 28, 53	$get_g_Network, 28, 51$
calcLarvalStageMortalityRate, 9, 31, 53	<pre>get_genotypesID_Network, 27, 51</pre>
calcMemoryWindow, 10, 36	<pre>get_genotypesN_Network, 27, 51</pre>
calcPopulationGrowthRate, $9$ , $10$ , $34$ , $53$	get_LAR_Patch, 28, 66
close_allConnections_Network, 12, 52	get_LARdly_Patch, 28, 66
connection, 25, 52, 65	get_Leq_Network, 28, 51
createNamedPopMatrix, 13	<pre>get_maleMigration_Patch, 29, 67</pre>
createNamedPopVector, 13	<pre>get_migrationFemale_Network, 29, 52</pre>
cube2csv, 13, <i>42</i>	<pre>get_migrationFemaleRow_Network, 29, 52</pre>

INDEX 77

<pre>get_migrationMale_Network, 30, 52</pre>	Network.Parameters, <i>51</i> , <i>52</i> , <i>53</i>
<pre>get_migrationMaleRow_Network, 30, 52</pre>	normalise, 53
get_moveVar_Network, 30, 51	
get_muAd_Network, 30, 51	<pre>oneDay_admPupating_deterministic_Patch,</pre>
get_muAq_Network, 31, 51	54, 67
get_NetworkPointer_Patch, 31, 67	<pre>oneDay_admPupating_stochastic_Patch,</pre>
get_nPatch_Network, 31, 52	54, 67
get_omega_Network, 31, 52	<pre>oneDay_admSurvival_deterministic_Patch,</pre>
get_patch_Network, 33, 52	54, 67
get_patches_Network, 32, 52	<pre>oneDay_admSurvival_stochastic_Patch,</pre>
get_patchID_Patch, 32, 66	55, 67
get_patchReleases_Network, 32, 52	<pre>oneDay_af1Mating_deterministic_Patch,</pre>
get_phi_Network, 33, 52	55, 67
get_PUP_Patch, 33, 66	<pre>oneDay_af1Mating_stochastic_Patch, 55,</pre>
get_PUPdly_Patch, 33, 67	67
get_releaseType_Network, 34, 52	<pre>oneDay_af1Pupation_deterministic_Patch,</pre>
get_Rm_Network, 34, 51	56, 67
get_rm_Network, 51	oneDay_af1Pupation_stochastic_Patch,
get_s_Network, 34, 52	56, 67
get_simTime_Network, 34, 52	<pre>oneDay_af1Survival_deterministic_Patch,</pre>
get_tau_Network, 35, 51	56, 67
get_thetaAq_Network, 35, 51	oneDay_af1Survival_stochastic_Patch,
get_timeAq_Network, 35, 51	57, 67
get_tNow_Network, 36, 52	oneDay_calcCumulativeLarvalDensityDependentFactor_Patc
get_wildType_Network, 36, 52	57, 58, 67
get_windowSize_Network, 36, 51	oneDay_calcCumulativePupaDensityDependentFactor_Patch,
get_xiF_Network, 36, 52	57, 67
get_xiM_Network, 30, 52 get_xiM_Network, 37, 52	oneDay_calcLarvalDensityDependentFactor_Patch,
ggCol_utility, 37	58, 67
ggcoi_utility, 37	oneDay_eggsFract2_deterministic_Patch,
<pre>initPopMatrixArray, 37</pre>	58, 67
initPopVectorArray, 38	oneDay_eggsFract2_stochastic_Patch, 58,
initStagesDurations, 35, 38	67
111111111111111111111111111111111111111	oneDay_femaleReleases_Patch, 59, 67, 71
kernels, 39	oneDay_hatchingFract_deterministic_Patch,
	59, 67
Lognormal, 12	oneDay_hatchingFract_stochastic_Patch,
MGDrivE, 39	59, 67
MGDrivE-Cube, 41	oneDay_initOutput_Patch, 59, 67
MGDrivE-Model, 43	<pre>oneDay_larHatching_deterministic_Patch,</pre>
MGDrivE.Setup, 47	60, 67
moveMatAll2, 47	oneDay_larHatching_stochastic_Patch,
moveMatCascade3, 48	60, <i>67</i>
moveMatDiag, 48	oneDay_larPupating_deterministic_Patch,
moveMatDiagOneCity, 48	60, 67
moveMatDie, 49	oneDay_larPupating_stochastic_Patch,
moveMatIndependent3, 49	60, 67
moveMatMixedSpil, 49	<pre>oneDay_larSurvival_deterministic_Patch,</pre>
${\tt moveMatTaleOfTwoCities}, {\tt 50}$	61, 67
moveMatTriDiagonal, 50	oneDay_larSurvival_stochastic_Patch,
<pre>moveMatTriple, 50</pre>	61, 67
	oneDay_maleReleases_Patch, 61, 67, 71
Network, 31, 51, 53, 59, 63, 65, 68, 71, 74	oneDay_Migration_Network, 52, 63

78 INDEX

```
oneDay_migrationIn_Patch, 62, 67
oneDay_migrationOut_deterministic_Patch,
        62, 67
oneDay_migrationOut_stochastic_Patch,
        62, 67
oneDay_Network, 52, 63
{\tt one Day\_num Male Female\_deterministic\_Patch},
oneDay_numMaleFemale_stochastic_Patch,
        63, 67
oneDay_ovipositG1_deterministic_Patch,
        64, 67
oneDay_ovipositG1_stochastic_Patch, 64,
        67
oneDay_PopDynamics_Patch, 64, 67
oneDay_updatePopulation_Patch, 65, 67
oneDay_writeOutput_Patch, 65, 67
oneRun_Network, 52, 65
Patch, 32, 33, 47, 52, 53, 66
primePopMatrixArray, 69
{\tt primePopVectorArray}, {\tt 69}
quantileC, 70
R6Class, 51, 66
rDirichlet, 70
rdirichlet, 62
Release_basicRepeatedReleases, 22, 51,
        66, 71
reset_Network, 52, 71
reset_Patch, 67, 71, 72
retrieveOutput, 72
rmultinom, 62
set_ADMnew_Patch, 66, 72
set_AF1new_Patch, 66, 73
set\_migrationFemale\_Network, 52, 73
set_migrationMale_Network, 52, 73
set_NetworkPointer_Patch, 67, 74
shiftAndUpdatePopVector, 65, 74
splitOutput, 74
SymCubeC, 75
turnStochasticityOnOrOff, 75
```