

# Package ‘CNORode’

November 27, 2024

**Type** Package

**Title** ODE add-on to CellNOptR

**Version** 1.49.0

**Date** 2022-03-22

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**Description** Logic based ordinary differential equation (ODE) add-on to CellNOptR.

**License** GPL-2

**LazyLoad** yes

**Depends** CellNOptR, genalg

**Enhances** doParallel, foreach

**Suggests** knitr, rmarkdown

**VignetteBuilder** knitr

**biocViews** ImmunoOncology, CellBasedAssays, CellBiology, Proteomics, Bioinformatics, TimeCourse

**RoxygenNote** 7.1.2

**git\_url** <https://git.bioconductor.org/packages/CNORode>

**git\_branch** devel

**git\_last\_commit** 851f051

**git\_last\_commit\_date** 2024-10-29

**Repository** Bioconductor 3.21

**Date/Publication** 2024-11-27

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## Contents

cnodata . . . . .	2
cnolist . . . . .	2
cnolistCNORodeExample . . . . .	3

CNORode . . . . .	3
createLBodeContPars . . . . .	4
crossvalidateODE . . . . .	5
defaultParametersGA . . . . .	6
defaultParametersSSm . . . . .	7
getLBodeContObjFunction . . . . .	8
getLBodeDataSim . . . . .	10
getLBodeMINLPObjFunction . . . . .	12
getLBodeModelSim . . . . .	13
getLBodeSimFunction . . . . .	15
getStates . . . . .	16
incidence2Adjacency . . . . .	17
indices . . . . .	17
minlpLBodeSSm . . . . .	18
model . . . . .	20
parEstimationLBode . . . . .	20
parEstimationLBodeGA . . . . .	22
parEstimationLBodeSSm . . . . .	24
pknmodel . . . . .	26
plotLBodeFitness . . . . .	27
plotLBodeModelSim . . . . .	29
runCNORode . . . . .	30
simdata2cnolist . . . . .	32
simulate . . . . .	33

**Index** **35**

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cnodata	<i>A cnodata from CellNoptR</i>
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**Description**

A cnodata from CellNoptR to use with provided examples

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cnolist	<i>A cnolist from CellNoptR</i>
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**Description**

A cnolist from CellNoptR to use with provided examples

---

cnolistCNORodeExample *A cnolist from CellNOptR*

---

### Description

A cnolist from CellNOptR to use with provided CNORode examples.

---

CNORode *Logic based ODE extension for CellNOptR*

---

### Description

This package is used for the simulation and fitting of logic based ODE models based on the Odefy approach.

### Details

Package: CNORode  
Type: Package  
Version: 1.2.0  
Date: 2012-03-14  
License: GPL-3  
LazyLoad: yes

### Author(s)

David Henriques, Thomas Cokelaer Maintainer: David Henriques <dhenriques@ebi.ac.uk>

### References

Dominik Wittmann, Jan Krumsiek, Julio S. Rodriguez, Douglas Lauffenburger, Steffen Klamt, and Fabian Theis. Transforming boolean models to continuous models: methodology and application to t-cell receptor signaling. *BMC Systems Biology*, 3(1):98+, September 2009.

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R. Serban and A. C. Hindmarsh, "CVODES: the Sensitivity-Enabled ODE Solver in SUNDIALS," Proceedings of IDETC/CIE 2005, Sept. 2005, Long Beach, CA. Also available as LLNL technical report UCRL-JP-200039.

C. Terfve, T. Cokelaer, A. MacNamara, D. Henriques, E. Goncalves, MK. Morris, M. van Iersel, DA Lauffenburger, J. Saez-Rodriguez. CellNOptR: a flexible toolkit to train protein signaling networks to data using multiple logic formalisms. BMC Systems Biology, 2012, 6:133:

### See Also

[CellNOptR](#), [parEstimationLBode](#), [getLBodeModelSim](#), [parEstimationLBode](#) [plotLBodeFitness](#).

---

createLBodeContPars     *Create a list with ODE parameter information needed to perform parameter estimation*

---

### Description

Creates a list with the continuous parameters to simulate the model, upper and lower bounds for the parameter estimation, parameters names, indices of the parameters and other information.

### Usage

```
createLBodeContPars(model, LB_n = 1, LB_k = 0.1, LB_tau = 0.01,
  UB_n = 5, UB_k = 0.9, UB_tau = 10, default_n = 3, default_k = 0.5,
  default_tau = 1, LB_in = c(), UB_in = c(), opt_n = TRUE, opt_k = TRUE,
  opt_tau = TRUE, random = FALSE)
```

### Arguments

model	The logic model to be simulated.
LB_n	A numeric value to be used as lower bound for all parameters of type n.
LB_k	A numeric value to be used as lower bound for all parameters of type k.
LB_tau	A numeric value to be used as lower bound for all parameters of type tau.
UB_n	A numeric value to be used as upper bound for all parameters of type n.
UB_k	A numeric value to be used as upper bound for all parameters of type k.
UB_tau	A numeric value to be used as upper bound for all parameters of type tau.
default_n	The default parameter to be used for every parameter of type n.
default_k	The default parameter to be used for every parameter of type k.
default_tau	The default parameter to be used for every parameter of type tau.
LB_in	An array with the the same length as ode_parameters\$parValues with lower bounds for each specific parameter.
UB_in	An array with the the same length as ode_parameters\$parValues with upper bounds for each specific parameter.

opt_n	Add all parameter n to the index of parameters to be fitted.
opt_k	Add all parameter k to the index of parameters to be fitted.
opt_tau	Add all parameter tau to the index of parameters to be fitted.
random	logical value that determines that a random solution is for the parameters to be optimized.

**Value**

parNames	An array containing the names of the parameters.
parValues	An array containing the values of the parameters, in the same order as the names.
index_opt_pars	An array containing the indexes for the parameters to be fitted.
index_n	An array containing the indexes of the parameters of type n.
index_k	An array containing the indexes of the parameters of type k.
index_tau	An array containing the indexes of the parameters of type tau.
LB	An array containing the lower bound for each parameter.
UB	An array containing the upper bound for each parameter.

**Author(s)**

David Henriques, Thomas Cokelaer

**Examples**

```
library(CNORode)
data("ToyCNOList",package="CNORode");
data("ToyModel",package="CNORode");
data("ToyIndices",package="CNORode");
ode_parameters=createLBodeContPars(model, opt_n=FALSE,default_n=2,
random=TRUE, LB_k=0.25, UB_k=0.8, LB_tau=0.01, UB_tau=10);
```

---

crossvalidateODE      *Crossvalidate ODE model*

---

**Description**

k-fold crossvalidation for logic ODE model

**Usage**

```
crossvalidateODE(
  CNOList,
  model,
  nfolds = 10,
  foldid = NULL,
  type = "datapoint",
```

```

parallel = FALSE,
ode_parameters = NULL,
paramsSSm = NULL,
method = "essm"
)

```

### Arguments

cnolist	Cnolist which contains all the experiments
model	a model prepared for the training
nfolds	number of folds - default is 10. Although nfolds can be as large as the sample size (leave-one-out CV), it is not recommended for large datasets.
foldid	an optional vector of values between '1' and 'nfold' identifying what fold each observation is in. If supplied, 'nfold' can be missing.
type	define the way to do the crossvalidation. The default is 'type="datapoint"', which assigns the data randomly into folds. The option 'type="experiment"' uses whole experiments for crossvalidation (all data corresponding to a cue combination). The 'type=observable' uses the subset of nodes across all experiments for crossvalidation.
parallel	use for parallel execution, requires the doParallel package
ode_parameters	list of fitted logic ODE parameter
paramsSSm	parameters for the SSm optimizer for running the optimization in crossvalidation
method	Selection of optimization method: only "ga" or "essm" arguments are accepted

### Details

Does a k-fold cross-validation for logic ODE CellNOpt models. In k-iterations a fraction of the data is eliminated from the CNolist. The model is trained on the remaining data and then the model predicts the held-out data. Then the prediction accuracy is reported for each iteration.

### See Also

[parEstimationLBode](#)

---

defaultParametersGA *Create default options to perform parameter estimation with a genetic algorithm.*

---

### Description

This function returns a list with several arguments for performing parameter estimation with the genetic algorithm from the package genalg.

### Usage

```
defaultParametersGA()
```

**Value**

mutationChance	NA
popSize	200
iters	100
elitism	NA
time	1
monitor	TRUE
verbose	0
transfer_function	3
reltol	1e-04
atol	0.001
maxStepSize	Inf
maxNumSteps	1e+05
maxErrTestsFails	50
nan_fac = 1	0

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [parEstimationLBode](#) [parEstimationLBodeGA](#)

---

defaultParametersSSm *Create default options to perform parameter estimation with scatter search meta-heuristic.*

---

**Description**

This function returns a list with several arguments for performing parameter estimation with scatter search meta-heuristic algorithm from the package `essR`.

**Usage**

```
defaultParametersSSm()
```

**Value**

maxeval	Inf
maxtime	100
ndiverse	NULL
dim_refset	NULL
local_solver	NULL
verbose	0
transfer_function	3
reltol	1e-04
atol	0.001
maxStepSize	Inf
maxNumSteps	1e+05
maxErrTestsFails	50
nan_fac	1
lambda_tau	0
lambda_k	0
bootstrap	0
SSpenalty_fac	0
SScontrolPenalty_fac	0
boot_seed	sample(1:10000,1)

**Author(s)**

David Henriques, Thomas Cokelaer, Federica Eduati

**See Also**

[CellNOptR](#) [parEstimationLBode](#) [parEstimationLBodeSSm](#)

---

getLBodeContObjFunction

*Returns the objective function to perform parameter estimation.*

---

**Description**

This function configures returns the objective function that can be used to evaluate the fitness of a logic based ODE model using a particular set of parameters. This function can be particularly useful if you are planing to couple a nonlinear optimization solver. The returned value of the objective function corresponds to the mean squared value normalized by the number of data points.



**Usage**

```
getLBodeContObjFunction(cnolist, model, ode_parameters, indices=NULL, time = 1,
  verbose = 0, transfer_function = 3, reltol = 1e-04, atol = 0.001, maxStepSize = Inf,
  maxNumSteps = 1e+05, maxErrTestsFails = 50, nan_fac = 1, lambda_tau=0, lambda_k=0,
  bootstrap=F, SSpenalty_fac=0, SScontrolPenalty_fac=0, boot_seed=sample(1:10000,1))
```

**Arguments**

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
time	An integer with the index of the time point to start the simulation. Default is 1.
verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum step size allowed to ODE solver.
maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.
nan_fac	A penalty for each data point the model is not able to simulate. We recommend higher than 0 and smaller than 1.
lambda_tau	Tunable regularisation parameters to penalise L1-norm of parameters tau and induce sparsity. We recommend testing values between 0 and 100 (in log scale) to find best compromise between good fit and sparse model. Default =0, corresponding to no regularisation.
lambda_k	Tunable regularisation parameters to penalise L1-norm of parameters k and induce sparsity. We recommend testing values between 0 and 100 (in log scale) to find best compromise between good fit and sparse model. Default =0, corresponding to no regularisation.
bootstrap	If set to TRUE performs random sampling with replacement of the measurements used in the optimisation (to be run multiple times to get bootstrapped distribution of parameters). Default =FALSE, no bootstrapping.
SSpenalty_fac	Penalty factor for penalising solutions which do not reach steady state. Default =0.
SScontrolPenalty_fac	Penalty factor for penalising solutions for which the control (unperturbed) condition (assumed to be first row) does not reach steady state. Default =0.
boot_seed	Seed used for random sampling if bootstrap=TRUE. Default chose random seed between 0 and 10000

### Details

Check [CellNOptR](#) for details about the cnoList and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

### Value

Returns a function to evaluate the model fitness. This function receives a vector containing both continuous parameters and integer values representing which reactions should be kept in the model.

### Author(s)

David Henriques, Thomas Cokelaer, Federica Eduati

### See Also

[CellNOptR](#) [createLBodeContPars](#)

### Examples

```
library(CNORode)
data("ToyCNOList", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");

ode_parameters=createLBodeContPars(model, random=TRUE);
minlp_obj_function=getLBodeContObjFunction(cnoListCNORodeExample, model, ode_parameters, indices);

x=ode_parameters$parValues;

f=minlp_obj_function(x);
```

---

getLBodeDataSim

*Simulate value signals a CNO list With Logic-Based ODEs.*

---

### Description

This function receives a set of inputs, namely the cnoList and the model and returns a list with the same size of the cnoList\$valueSignals.

### Usage

```
getLBodeDataSim(cnoList, model, ode_parameters = NULL, indices = NULL,
timeSignals=NULL, time = 1, verbose = 0, transfer_function = 3,
reltol = 1e-04, atol = 0.001, maxStepSize = Inf, maxNumSteps = 1e+05,
maxErrTestsFails = 50)
```

**Arguments**

<code>cnolist</code>	A list containing the experimental design and data.
<code>model</code>	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
<code>ode_parameters</code>	A list with the ODEs parameter information. Obtained with <code>makeParameterList</code> function.
<code>indices</code>	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
<code>timeSignals</code>	An array containing a different timeSignals. If you use this argument, it will also modify the dimensions from <code>valueSignals</code> .
<code>time</code>	An integer with the index of the time point to start the simulation. Default is 1.
<code>verbose</code>	A logical value that triggers a set of comments.
<code>transfer_function</code>	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
<code>reltol</code>	Relative Tolerance for numerical integration.
<code>atol</code>	Absolute tolerance for numerical integration.
<code>maxStepSize</code>	The maximum step size allowed to ODE solver.
<code>maxNumSteps</code>	The maximum number of internal steps between two points being sampled before the solver fails.
<code>maxErrTestsFails</code>	Specifies the maximum number of error test failures permitted in attempting one step.

**Details**

Check [CellNOptR](#) for details about the `cnolist` and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

**Value**

Returns a list with simulated data that has the same structure as the `cnolist$valueSignals`. One matrix for each time-point.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [parEstimationLBode](#) [parEstimationLBodeSSm](#)

**Examples**

```
library(CNORode)
data("ToyCNolist", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");
dataSimulation=getLBodeDataSim(cnolistCNORodeExample, model, indices=indices);
```

---

```
getLBodeMINLPObjFunction
```

*Get the objective function to evaluate the fitness of a given model structure and set of parameters.*

---

**Description**

This function configures returns the objective function that can be used to evaluate the fitness of a logic based ODE model using a particular set of parameters and model structure. This function can be particular useful if you are planing to couple a mixed integer nonlinear programming optimization solver. The returned value of the objective function corresponds to the mean squared value.

**Usage**

```
getLBodeMINLPObjFunction(cnolist, model, ode_parameters, indices=NULL, time = 1,
  verbose = 0, transfer_function = 3, reltol = 1e-04, atol = 0.001, maxStepSize = Inf,
  maxNumSteps = 1e+05, maxErrTestsFails = 50, nan_fac = 1)
```

**Arguments**

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createlBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <a href="#">indexFinder</a> function from <a href="#">CellNOptR</a> .
time	An integer with the index of the time point to start the simulation. Default is 1.
verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum step size allowed to ODE solver.
maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.

maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.
nan_fac	A penalty for each data point the model is not able to simulate. We recommend higher than 0 and smaller than 1.

**Details**

Check [CellNOptR](#) for details about the cnoList and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

**Value**

Returns a function to evaluate the model fitness. This function receives a continuous parameter vector.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [createLBodeContPars](#)

**Examples**

```
library(CNORode)
data("ToyCNolist", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");

ode_parameters=createLBodeContPars(model, random=TRUE);
minlp_obj_function=getLBodeMINLPobjFunction(cnoListCNORodeExample, model,ode_parameters,indices);

n_int_vars=dim(model$interMat)[2];
x_int=round(runif(n_int_vars))
x_cont=ode_parameters$parValues;
x=c(x_cont,x_int);
f=minlp_obj_function(x);
```

---

getLBodeModelSim

*Simulate the logic-based ODE model*

---

**Description**

This function simulates a logic-based ODE model and return a list with one matrix for each time point. The input species in the model are filled with NA values. If the simulation of a particular set of initial conditions fails the solver will fill the experience row with NA values.

**Usage**

```
getLBodeModelSim(cnolist, model, ode_parameters = NULL, indices = NULL, timeSignals=NULL,
time = 1, verbose = 0, transfer_function = 3, reltol = 1e-04, atol = 0.001, maxStepSize = Inf,
maxNumSteps = 1e+05, maxErrTestsFails = 50)
```

**Arguments**

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
timeSignals	An array containing a different timeSignals. If you use this argument, it will also modify the dimensions from valueSignals.
time	An integer with the index of the time point to start the simulation. Default is 1.
verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum number of internal steps between two points being sampled before the solver fails.
maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.

**Details**

Check [CellNOptR](#) for details about the cnolist and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

**Value**

Returns a list with simulated data with similar structure to `cnolist$valueSignals`. Contains one matrix for each time-point. Each matrix contains one row per experiment and one columns per model species.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [createLBodeContPars](#)

**Examples**

```
library(CNORode)
data('ToyCNolist',package='CNORode');
data('ToyModel',package='CNORode');
data('ToyIndices',package='CNORode');
modelSimulation=getLBodeModelSim(cnolistCNORodeExample, model,indices=indices);
```

---

getLBodeSimFunction     *Get a function to simulate a logic based ODE model.*

---

**Description**

This function is internally used by CNORode to configure the simulation function with default arguments.

**Usage**

```
getLBodeSimFunction(cnolist1, model1, adjMatrix1, indices1, odeParameters1,
time1 = 1, verbose1 = 0, transfer_function1 = 3, reltol1 = 1e-04, atol1 = 0.001,
maxStepSize1 = Inf, maxNumSteps1 = 1e+05, maxErrTestsFails1 = 50)
```

**Arguments**

cnolist1	A list containing the experimental design and data.
model1	The logic model to be simulated.
adjMatrix1	An adjacency matrix from the model.
indices1	Indices to map data in the model. Obtained with indexFinder function from CellNOptR.
odeParameters1	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
time1	An integer with the index of the time point to start the simulation. Default is 1.
verbose1	A logical value that triggers a set of comments.
transfer_function1	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol1	Relative Tolerance for numerical integration.
atol1	Absolute tolerance for numerical integration.
maxStepSize1	The maximum step size allowed to ODE solver.
maxNumSteps1	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails1	Specifies the maximum number of error test failures permitted in attempting one step.

**Value**

A function that returns a simulated model.

**Note**

This function is for CNORode internal use.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [CNORode](#)

---

getStates

*Find which species in the model are states.*

---

**Description**

Receives an adjacency matrix (model\$interMat from CellNOptR) and finds which species are states (i.e. not inputs).

**Usage**

```
getStates(adjacency)
```

**Arguments**

adjacency      An adjacency matrix from the model.

**Value**

A numeric vector with 0's for positions which are states and 1's for positions which are.

**Note**

For internal use of CNORode.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[incidence2Adjacency](#)



---

incidence2Adjacency     *Convert an incidence matrix into an adjacency matrix.*

---

**Description**

Convert the incidence matrix (model representation of CellNOptR) into an adjacency matrix. Denotes the inputs/output relationships.

**Usage**

```
incidence2Adjacency(model)
```

**Arguments**

model             Model from CellNOptR.

**Value**

Directed Adjacency matrix of size n\_species by n\_species.

**Note**

For internal use of CNORode.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#)

---

indices             *Indices that relate cnoList to model*

---

**Description**

A list with indices that relate the cnoList with the model from CellNOptR

---

minlpLBodeSSm	<i>Search for the best combination of continuous parameters and logic gates.</i>
---------------	--

---

### Description

This function uses `essR` to search for the best set of continuous parameters and model structure. The objective function is the same as the one provided by [getLBodeMINLPObjFunction](#).

### Usage

```
minlpLBodeSSm(cnolist, model, ode_parameters = NULL, int_x0=NULL, indices = NULL, maxeval = Inf,
maxtime = 100, ndiverse = NULL, dim_refset = NULL, local_solver = NULL, time = 1,
verbose = 0, transfer_function = 3, reltol = 1e-04, atol = 0.001, maxStepSize = Inf,
maxNumSteps = 1e+05, maxErrTestsFails = 50, nan_fac = 1)
```

### Arguments

<code>cnolist</code>	A list containing the experimental design and data.
<code>model</code>	The logic model to be simulated.
<code>ode_parameters</code>	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
<code>int_x0</code>	Vector with initial solution for integer parameters.
<code>indices</code>	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
<code>maxeval</code>	Maximum number of evaluation in the optimization procedure.
<code>maxtime</code>	Maximum number of evaluation spent in optimization procedure.
<code>ndiverse</code>	Duration of the optimisation procedure.
<code>dim_refset</code>	Number of diverse initial solutions.
<code>local_solver</code>	Local solver to be used in SSm.
<code>time</code>	An integer with the index of the time point to start the simulation. Default is 1.
<code>verbose</code>	A logical value that triggers a set of comments.
<code>transfer_function</code>	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and for normalized Hill function.
<code>reltol</code>	Relative Tolerance for numerical integration.
<code>atol</code>	Absolute tolerance for numerical integration.
<code>maxStepSize</code>	The maximum step size allowed to ODE solver.
<code>maxNumSteps</code>	The maximum number of internal steps between two points being sampled before the solver fails.
<code>maxErrTestsFails</code>	Specifies the maximum number of error test failures permitted in attempting one step.
<code>nan_fac</code>	A penalty for each data point the model is not able to simulate. We recommend higher than 0 and smaller than 1.

**Details**

Check [CellNOptR](#) for details about the cno1ist and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

**Value**

LB_n	A numeric value to be used as lower bound for all parameters of type n.
LB_k	A numeric value to be used as lower bound for all parameters of type k.
LB_tau	A numeric value to be used as lower bound for all parameters of type tau.
UB_n	A numeric value to be used as upper bound for all parameters of type n.
UB_k	A numeric value to be used as upper bound for all parameters of type k.
UB_tau	A numeric value to be used as upper bound for all parameters of type tau.
default_n	The default parameter to be used for every parameter of type n.
default_k	The default parameter to be used for every parameter of type k.
default_tau	The default parameter to be used for every parameter of type tau.
LB_in	An array with the the same length as ode_parameters\$parValues with lower bounds for each specific parameter.
UB_in	An array with the the same length as ode_parameters\$parValues with upper bounds for each specific parameter.
opt_n	Add all parameter n to the index of parameters to be fitted.
opt_k	Add all parameter k to the index of parameters to be fitted.
opt_tau	Add all parameter tau to the index of parameters to be fitted.
random	A logical value that determines that a random solution is for the parameters to be optimised.
model	The best fitting found model structure.
smm_results	A list containing the information provided by the nonlinear optimization solver.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR createLBodeContPars](#) `essR`

**Examples**

```
## Not run:
data("ToyCNO1ist", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");

ode_parameters=createLBodeContPars(model, random=TRUE);

#Visualize initial solution
```

```

simulatedData=plotLBodeFitness(cnolistCNORodeExample, model,ode_parameters,indices=indices)
ode_parameters=minlpLBodeSSm(cnolistCNORodeExample, model,ode_parameters);

model=ode_parameters$model;

#Visualize fitted solution
simulatedData=plotLBodeFitness(cnolistCNORodeExample, model,indices=indices);

## End(Not run)

```

---

model	<i>A model from CellNoptR</i>
-------	-------------------------------

---

### Description

A model from CellNoptR to use with provided examples

---

parEstimationLBode	<i>Perform parameter estimation using a genetic algorithm (package <code>genalg</code>) or ssm (if package <code>essm</code> available).</i>
--------------------	--

---

### Description

This function is an alias to the parEstimationLBode variants ([parEstimationLBodeGA](#) and [parEstimationLBodeSSm](#))

### Usage

```

parEstimationLBode(cnolist, model, method="ga", ode_parameters = NULL, indices = NULL,
  paramsGA=NULL, paramsSSm=NULL)

```

### Arguments

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
method	Only "ga" or "essm" arguments are accepted.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <code>indexFinder</code> function from CellNoptR.
paramsGA	A list of GA parameters. default is the list returned by <code>defaultParametersGA</code> .
paramsSSm	A list of SSm parameters. default is the list returned by <code>defaultParametersSSm</code> .

**Value**

LB_n	A numeric value to be used as lower bound for all parameters of type n.
LB_k	A numeric value to be used as lower bound for all parameters of type k.
LB_tau	A numeric value to be used as lower bound for all parameters of type tau.
UB_n	A numeric value to be used as upper bound for all parameters of type n.
UB_k	A numeric value to be used as upper bound for all parameters of type k.
UB_tau	A numeric value to be used as upper bound for all parameters of type tau.
default_n	The default parameter to be used for every parameter of type n.
default_k	The default parameter to be used for every parameter of type k.
default_tau	The default parameter to be used for every parameter of type tau.
LB_in	An array with the the same length as ode_parameters\$parValues with lower bounds for each specific parameter.
UB_in	An array with the the same length as ode_parameters\$parValues with upper bounds for each specific parameter.
opt_n	Add all parameter n to the index of parameters to be fitted.
opt_k	Add all parameter k to the index of parameters to be fitted.
opt_tau	Add all parameter tau to the index of parameters to be fitted.
random	A logical value that determines that a random solution is for the parameters to be optimized.
res	A list containing the information provided by the solver.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR createLBodeContPars rbga](#)

**Examples**

```
data("ToyCN0list", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");

ode_parameters=createLBodeContPars(model, random=TRUE);
#Visualize initial solution
simulatedData=plotLBodeFitness(cnolistCNORodeExample, model, ode_parameters, indices=indices)
paramsGA = defaultParametersGA()
paramsGA$maxStepSize = 1
paramsGA$popSize = 10
paramsGA$iter = 10
paramsGA$transfer_function = 2

ode_parameters=parEstimationLBode(cnolistCNORodeExample, model, ode_parameters=ode_parameters,
```

```

paramsGA=paramsGA)
#Visualize fitted solution
simulatedData=plotLBodeFitness(cnolistCNORodeExample, model,ode_parameters,indices=indices)

```

---

parEstimationLBodeGA *Perform parameter estimation using a genetic algorithm (package genalg).*

---

## Description

This function uses a genetic algorithm (package genalg) to perform parameter estimation. The objective function is the same as the one provided by [getLBodeContObjFunction](#).

## Usage

```

parEstimationLBodeGA(cnolist, model, ode_parameters = NULL, indices = NULL, mutationChance = NA, popSi
elitism = NA, time = 1, monitor = TRUE, verbose = 0, transfer_function = 3, reltol = 1e-04,
atol = 0.001, maxStepSize = Inf, maxNumSteps = 1e+05, maxErrTestsFails = 50, nan_fac = 1)

```

## Arguments

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <a href="#">indexFinder</a> function from CellNOptR.
mutationChance	the chance that a gene in the chromosome mutates. By default $1/(size+1)$ . It affects the convergence rate and the probing of search space: a low chance results in quicker convergence, while a high chance increases the span of the search space.
popSize	the population size.
iters	the number of iterations.
elitism	the number of chromosomes that are kept into the next generation. By default is about 20% of the population size
time	An integer with the index of the time point to start the simulation. Default is 1.
monitor	If TRUE a plot will be generated to monitor the objective function
verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum step size allowed to ODE solver.

maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.
nan_fac	A penalty for each data point the model is not able to simulate. We recommend higher than 0 and smaller than 1.

**Value**

LB_n	A numeric value to be used as lower bound for all parameters of type n.
LB_k	A numeric value to be used as lower bound for all parameters of type k.
LB_tau	A numeric value to be used as lower bound for all parameters of type tau.
UB_n	A numeric value to be used as upper bound for all parameters of type n.
UB_k	A numeric value to be used as upper bound for all parameters of type k.
UB_tau	A numeric value to be used as upper bound for all parameters of type tau.
default_n	The default parameter to be used for every parameter of type n.
default_k	The default parameter to be used for every parameter of type k.
default_tau	The default parameter to be used for every parameter of type tau.
LB_in	An array with the the same length as ode_parameters\$parValues with lower bounds for each specific parameter.
UB_in	An array with the the same length as ode_parameters\$parValues with upper bounds for each specific parameter.
opt_n	Add all parameter n to the index of parameters to be fitted.
opt_k	Add all parameter k to the index of parameters to be fitted.
opt_tau	Add all parameter tau to the index of parameters to be fitted.
random	A logical value that determines that a random solution is for the parameters to be optimized.
res	A list containing the information provided by the nonlinear optimization solver (genalg).

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR createLBodeContPars rbg](#)

**Examples**

```

data("ToyCNolist", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");

ode_parameters=createLBodeContPars(model, random=TRUE);
#Visualize initial simulation
#simulatedData=plotLBodeFitness(cnolistCNORodeExample, model, ode_parameters, indices=indices)

ode_parameters=parEstimationLBodeGA(cnolistCNORodeExample, model, ode_parameters=ode_parameters,
indices=indices, maxStepSize=1, atol=1e-3, reltol=1e-5, transfer_function=2, popSize=10, iter=40);

#Visual solution after optimization
simulatedData=plotLBodeFitness(cnolistCNORodeExample, model, indices=indices, ode_parameters=ode_parameters);

```

---

parEstimationLBodeSSm *Perform parameter estimation using essR.*

---

**Description**

This function uses `essR` to perform parameter estimation. The objective function is the same as the one provided by [getLBodeContObjFunction](#).

**Usage**

```

parEstimationLBodeSSm(cnolist, model, ode_parameters = NULL, indices = NULL,
maxeval = Inf, maxtime = 100, ndiverse = NULL, dim_refset = NULL, local_solver = NULL,
time = 1, verbose = 0, transfer_function = 3, reltol = 1e-04, atol = 0.001,
maxStepSize = Inf, maxNumSteps = 1e+05, maxErrTestsFails = 50, nan_fac = 1,
lambda_tau = 0, lambda_k = 0, bootstrap = FALSE, SSpenalty_fac = 0,
SScontrolPenalty_fac = 0, boot_seed = sample(1:10000,1))

```

**Arguments**

<code>cnolist</code>	A list containing the experimental design and data.
<code>model</code>	The logic model to be simulated.
<code>ode_parameters</code>	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
<code>indices</code>	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
<code>maxeval</code>	Maximum number of evaluation in the optimization procedure.
<code>maxtime</code>	Duration of the optimization procedure.
<code>ndiverse</code>	Number of diverse initial solutions.
<code>dim_refset</code>	Size of the reference set.
<code>local_solver</code>	Local solver to be used in SSm.
<code>time</code>	An integer with the index of the time point to start the simulation. Default is 1.



verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum step size allowed to ODE solver.
maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.
nan_fac	A penalty for each data point the model is not able to simulate. We recommend higher than 0 and smaller than 1.
lambda_tau	penalty parameter for node parameters (tau)
lambda_k	penalty parameter for edge parameters (k)
bootstrap	Boolean, default: FALSE. If the residuals should be bootstrapped.
SSpenalty_fac	Steady-state penalty: at the end of the simulation the model states should reach steady state. The steady state is measured by the sum of squares of the state derivatives.
SScontrolPenalty_fac	Steady-state penalty for a control experiment, the default is 0. The first condition should represent a control condition (no stimulus or inhibition). Then the model simulation is penalised if it deviates from the initial conditions. This is to make sure that the predicted dynamics is not due to the initial conditions, but because of the stimuli.
boot_seed	random seed used for the bootstrapping.

## Details

Check [CellNOptR](#) for details about the codelist and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

## Value

LB_n	A numeric value to be used as lower bound for all parameters of type n.
LB_k	A numeric value to be used as lower bound for all parameters of type k.
LB_tau	A numeric value to be used as lower bound for all parameters of type tau.
UB_n	A numeric value to be used as upper bound for all parameters of type n.
UB_k	A numeric value to be used as upper bound for all parameters of type k.
UB_tau	A numeric value to be used as upper bound for all parameters of type tau.
default_n	The default parameter to be used for every parameter of type n.
default_k	The default parameter to be used for every parameter of type k.

default_tau	The default parameter to be used for every parameter of type tau.
LB_in	An array with the the same length as ode_parameters\$parValues with lower bounds for each specific parameter.
UB_in	An array with the the same length as ode_parameters\$parValues with upper bounds for each specific parameter.
opt_n	Add all parameter n to the index of parameters to be fitted.
opt_k	Add all parameter k to the index of parameters to be fitted.
opt_tau	Add all parameter tau to the index of parameters to be fitted.
random	A logical value that determines that a random solution is for the parameters to be optimized.
smm_results	A list containing the information provided by the nonlinear optimization solver.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR createLBodeContPars](#)

**Examples**

```
## Not run:
data("ToyCN0list", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");

ode_parameters=createLBodeContPars(model, random=TRUE);

#Visualize intial simulation
simulatedData=plotLBodeFitness(cnolistCNORodeExample, model,ode_parameters,indices=indices)

ode_parameters=parEstimationLBodeSSm(cnolistCNORodeExample,model,ode_parameters,
indices=indices,maxtime=20,ndiverse=50,dim_refset=6);

#Visualize fitterd solution
simulatedData=plotLBodeFitness(cnolistCNORodeExample, model,indices=indices,ode_parameters=ode_parameters);

## End(Not run)
```

---

pknmodel

*A pknmodel from CellNOptR*

---

**Description**

A pknmodel from CellNOptR to use with provided examples

---

plotLBodeFitness      *Plot data against simulated values.*

---

### Description

Plots the simulated values with the logic-based ODE against the the data contained contained the data contained in the cnolist. The data values are represented with a black line and the simulated values with a blue line. Additionally this functions returns the the simulated values.

### Usage

```
plotLBodeFitness(cnolist, model, ode_parameters = NULL, indices = NULL,
  adjMatrix = NULL, time = 1, verbose = 0, transfer_function = 3, reltol = 1e-04,
  atol = 0.001, maxStepSize = Inf, maxNumSteps = 1e+05, maxErrTestsFails = 50,
  plot_index_signals = NULL, plot_index_experiments = NULL,
  plot_index_cues = NULL, colormap="heat", plotParams=list(margin=0.1, width=15, height=12,
  cmap_scale=1, cex=1.6, ymin=NULL)
)
```

### Arguments

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
adjMatrix	Model representation in the form of an adjacency matrix. When not provided will be automatically computed based in the model.
time	An integer with the index of the time point to start the simulation. Default is 1.
verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum step size allowed to ODE solver.
maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.

plot_index_signals	In case you only want to plot some signals, provide an integer vector with the indexes.
plot_index_experiments	In case you only want to plot some experiments, provide an integer vector with the indexes.
plot_index_cues	In case you only want to plot some cues, provide an integer vector with the indexes.
colormap	Uses the same colormap as in CellNOptR by default. If set to "green", it uses the deprecated colormap.
plotParams	additional parameters to refine the ploggin. See plotOptimResultsPan function in CellNOptR for more details.

### Details

Check [CellNOptR](#) for details about the cnoList and the model format. For more details in the configuration of the ODE solver check the CVODES manual.

### Value

Returns a list with simulated data that has the same structure as the cnoList\$valueSignals. One matrix for each time-point.

### Author(s)

David Henriques, Thomas Cokelaer

### See Also

[CellNOptR createLBodeContPars](#)

### Examples

```
library(CNORode)
data("ToyCNOList", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");
ode_parameters=createLBodeContPars(model, random=TRUE);
dataSimulation=plotLBodeFitness(cnoListCNORodeExample, model, indices=indices);
```

---

plotLBodeModelSim	<i>Simulate the model and plot the obtained with the different experimental conditions.</i>
-------------------	---

---

### Description

Plots the simulated values of the logic based ODE model. Only dynamic states are plotted, i.e. those that are not inputs. a blue line. Additionally this functions returns the the simulated values.

### Usage

```
plotLBodeModelSim(cnolist, model, ode_parameters = NULL, indices = NULL,
adjMatrix = NULL, timeSignals=NULL, time = 1, verbose = 0, transfer_function = 3,
reltol = 1e-04, atol = 0.001, maxStepSize = Inf, maxNumSteps = 1e+05,
maxErrTestsFails = 50, large = FALSE, nsplit = 4, show = TRUE)
```

### Arguments

cnolist	A list containing the experimental design and data.
model	The logic model to be simulated.
ode_parameters	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
indices	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
adjMatrix	Model representation in the form of an adjacency matrix. When not provided will be automatically computed based in the model.
timeSignals	An array containing a different timeSignals. If you use this argument, it will also modify the dimensions from valueSignals.
time	An integer with the index of the time point to start the simulation. Default is 1.
verbose	A logical value that triggers a set of comments.
transfer_function	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
reltol	Relative Tolerance for numerical integration.
atol	Absolute tolerance for numerical integration.
maxStepSize	The maximum step size allowed to ODE solver.
maxNumSteps	The maximum number of internal steps between two points being sampled before the solver fails.
maxErrTestsFails	Specifies the maximum number of error test failures permitted in attempting one step.
large	Boolean variable defining if the plot should split into several subplots.
nsplit	In case the large plot options is selected define how many subplots will exist. Default is 4.
show	Boolean variable defining if we should plot the CNolist object.

**Value**

Returns a list with simulated Model values. One matrix of size number of species by number of experimental conditions for each time-point.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [createLBodeContPars](#)

**Examples**

```
library(CNORode)
data("ToyCNOList", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");
modelSimulation=plotLBodeModelSim(cnolistCNORodeExample, model, indices=indices);
```

---

runCNORode

*runCNORode*

---

**Description**

A one-line wrapper of the CNORode pipeline

**Usage**

```
runCNORode(
  model,
  data,
  compression = TRUE,
  results_folder = "CNORode_results",
  cutNONC = TRUE,
  expansion = FALSE,
  LB_n = 1,
  LB_k = 0.1,
  LB_tau = 0.01,
  UB_n = 5,
  UB_k = 0.9,
  UB_tau = 10,
  default_n = 3,
  default_k = 0.5,
  default_tau = 1,
  opt_n = TRUE,
  opt_k = TRUE,
  opt_tau = TRUE,
```

```

    random = TRUE,
    maxeval = 1e+05,
    maxtime = 60,
    transfer_function = 3,
    nan_fac = 1,
    lambda_tau = 0,
    lambda_k = 0
)

```

### Arguments

model	A filename of prior knowledge network (PKN) in the SIF format
data	A measurement filename in the MIDAS format
compression	compress the prior knowledge network (TRUE), see <a href="#">preprocessing</a>
results_folder	results folder for the analysis.
cutNONC	cut non-observable non-controllable node from PKN (TRUE), see <a href="#">preprocessing</a>
expansion	expand OR gates in the PKN (FALSE), see <a href="#">preprocessing</a>
LB_n	lower bound on parameter n, see <a href="#">createLBodeContPars</a>
LB_k	lower bound on parameter k, see <a href="#">createLBodeContPars</a>
LB_tau	lower bound on parameter tau, see <a href="#">createLBodeContPars</a>
UB_n	upper bound on parameter n, see <a href="#">createLBodeContPars</a>
UB_k	upper bound on parameter k, see <a href="#">createLBodeContPars</a>
UB_tau	upper bound on parameter tau, see <a href="#">createLBodeContPars</a>
default_n	default value of parameter n, see <a href="#">createLBodeContPars</a>
default_k	default value of parameter k, see <a href="#">createLBodeContPars</a>
default_tau	default value of parameter tau, see <a href="#">createLBodeContPars</a>
opt_n	should parameter n be optimised, see <a href="#">createLBodeContPars</a>
opt_k	should parameter k be optimised, see <a href="#">createLBodeContPars</a>
opt_tau	should parameter tau be optimised, see <a href="#">createLBodeContPars</a>
random	initial parameter vector generation (TRUE: random, FALSE: half of the LB-UB)
maxeval	maximum number of function evaluations in the optimisation, see <a href="#">parEstimationLBodeSSm</a>
maxtime	maximum CPU time (in seconds) spent on optimisation before calling final refinement, see <a href="#">parEstimationLBodeSSm</a>
transfer_function	transfer function types represented by the edges, see <a href="#">parEstimationLBodeSSm</a>
nan_fac	penalty for NA simulations, see <a href="#">parEstimationLBodeSSm</a>
lambda_tau	regularisation penalty for tau parameters, see <a href="#">parEstimationLBodeSSm</a>
lambda_k	regularisation penalty for k parameters for optimisation, see <a href="#">parEstimationLBodeSSm</a>

## Examples

```
## Not run:
model = system.file("extdata", "ToyModelMMB_FeedbackAnd.sif", package="CNORode")
data = system.file("extdata", "ToyModelMMB_FeedbackAnd.csv", package="CNORode")
res = runCNORode(model,data,results_folder = "./results")

## End(Not run)
```

---

simdata2cnolist      *converts output of getLBodeModelSim to cnolist*

---

## Description

This function converts the simulated data returned by `getLBodeModelSim` into a valid CNolist data structure.

## Usage

```
simdata2cnolist(sim_data, cnolist, model)
```

## Arguments

<code>sim_data</code>	structure returned by <code>getLBodeModelSim</code>
<code>cnolist</code>	A list containing the experimental design and data.
<code>model</code>	The logic model to be simulated.

## Value

a CNolist

## Author(s)

Thomas Cokelaer

## See Also

[CellNOptR](#) [createLBodeContPars](#)

## Examples

```
data('ToyCNolist', package='CNORode');
data('ToyModel', package='CNORode');
data('ToyIndices', package='CNORode');
simdata = getLBodeModelSim(cnolistCNORodeExample, model, indices=indices)
cnolist = simdata2cnolist(simdata, cnolistCNORodeExample, model)

cnolist = simdata2cnolist(simdata, cnolistCNORodeExample, model)
```



---

simulate	<i>Simulate value signals a CNO list With Logic-Based ODEs.</i>
----------	---

---

### Description

This function receives a set of inputs, namely the `cnolist` and the `model` and returns a list with the same size of the `cnolist$valueSignals`.

### Usage

```
simulate(cnolist, model, ode_parameters=NULL, indices=NULL,
adjMatrix=NULL, time=1, verbose=0, transfer_function=3,
reltol=1e-04, atol=0.001, maxStepSize=Inf, maxNumSteps=1e+05,
maxErrTestsFails=50)
```

### Arguments

<code>cnolist</code>	A list containing the experimental design and data.
<code>model</code>	A list with the ODEs parameter information. Obtained with <a href="#">createLBodeContPars</a> .
<code>ode_parameters</code>	A list with the ODEs parameter information. Obtained with <code>makeParameterList</code> function.
<code>indices</code>	Indices to map data in the model. Obtained with <code>indexFinder</code> function from <code>CellNOptR</code> .
<code>adjMatrix</code>	The adjacency matrix. Recomputed if not provided
<code>time</code>	An integer with the index of the time point to start the simulation. Default is 1.
<code>verbose</code>	A logical value that triggers a set of comments.
<code>transfer_function</code>	The type of used transfer. Use 1 for no transfer function, 2 for Hill function and 3 for normalized Hill function.
<code>reltol</code>	Relative Tolerance for numerical integration.
<code>atol</code>	Absolute tolerance for numerical integration.
<code>maxStepSize</code>	The maximum step size allowed to ODE solver.
<code>maxNumSteps</code>	The maximum number of internal steps between two points being sampled before the solver fails.
<code>maxErrTestsFails</code>	Specifies the maximum number of error test failures permitted in attempting one step.

### Details

Check [CellNOptR](#) for details about the `cnolist` and the `model` format. For more details in the configuration of the ODE solver check the CVODES manual.

**Value**

Returns a list with simulated data that has the same structure as the `cnolist$valueSignals`. One matrix for each time-point.

**Author(s)**

David Henriques, Thomas Cokelaer

**See Also**

[CellNOptR](#) [parEstimationLBode](#) [parEstimationLBodeSSm](#)

**Examples**

```
library(CNORode)
data("ToyCNolist", package="CNORode");
data("ToyModel", package="CNORode");
data("ToyIndices", package="CNORode");
dataSimulation = simulate(cnolistCNORodeExample, model, indices=indices);
```

# Index

- \* **CNORode**
  - CNORode, [3](#)
- \* **CVODES**
  - getLBodeSimFunction, [15](#)
- \* **CellNOptR**
  - parEstimationLBode, [20](#)
  - parEstimationLBodeGA, [22](#)
- \* **SSm**
  - defaultParametersSSm, [7](#)
- \* **adjacency**
  - getStates, [16](#)
  - incidence2Adjacency, [17](#)
- \* **algorithm**
  - defaultParametersGA, [6](#)
  - parEstimationLBode, [20](#)
  - parEstimationLBodeGA, [22](#)
- \* **default**
  - defaultParametersGA, [6](#)
  - defaultParametersSSm, [7](#)
- \* **essR**
  - defaultParametersSSm, [7](#)
- \* **genetic**
  - defaultParametersGA, [6](#)
  - parEstimationLBode, [20](#)
  - parEstimationLBodeGA, [22](#)
- \* **incidence**
  - incidence2Adjacency, [17](#)
- \* **logic**
  - parEstimationLBode, [20](#)
  - parEstimationLBodeGA, [22](#)
- \* **matrix**
  - incidence2Adjacency, [17](#)
- \* **model**
  - parEstimationLBode, [20](#)
  - parEstimationLBodeGA, [22](#)
- \* **parameters**
  - defaultParametersGA, [6](#)
- \* **states**
  - getStates, [16](#)
- CellNOptR, [4, 7, 8, 10, 11, 13, 14, 16, 17, 19, 21, 23, 25, 26, 28, 30, 32–34](#)
- cnodata, [2](#)
- cnolist, [2](#)
- cnolistCNORodeExample, [3](#)
- CNORode, [3, 16](#)
- createLBodeContPars, [4, 9–15, 18–24, 26–33](#)
- crossvalidateODE, [5](#)
- defaultParametersGA, [6](#)
- defaultParametersSSm, [7](#)
- getLBodeContObjFunction, [8, 22, 24](#)
- getLBodeDataSim, [10](#)
- getLBodeMINLPObjFunction, [12, 18](#)
- getLBodeModelSim, [4, 13](#)
- getLBodeSimFunction, [15](#)
- getStates, [16](#)
- incidence2Adjacency, [16, 17](#)
- indices, [17](#)
- minlpLBodeSSm, [18](#)
- model, [20](#)
- parEstimationLBode, [4, 6–8, 11, 20, 34](#)
- parEstimationLBodeGA, [7, 20, 22](#)
- parEstimationLBodeSSm, [8, 11, 20, 24, 31, 34](#)
- pknmodel, [26](#)
- plotLBodeFitness, [4, 27](#)
- plotLBodeModelSim, [29](#)
- preprocessing, [31](#)
- rbga, [21, 23](#)
- runCNORode, [30](#)
- simdata2cnolist, [32](#)
- simulate, [33](#)