

# Package ‘netReg’

April 15, 2020

**Type** Package

**Title** Network-Regularized Regression Models

**Version** 1.10.0

**Date** 2018-04-28

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**Description** netReg fits linear regression models using network-penalization.

Graph prior knowledge, in the form of biological networks, is being incorporated into the loss function of the linear model.

The networks describe

biological relationships such as co-regulation or dependency of the same transcription factors/metabolites/etc. yielding a part sparse and part smooth solution for coefficient profiles.

**URL** <https://github.com/dirmeier/netReg>

**BugReports** <https://github.com/dirmeier/netReg/issues>

**Depends** R(>= 3.4)

**biocViews** Software, StatisticalMethod, Regression, FeatureExtraction, Network, GraphAndNetwork

**License** GPL-3 | BSL-1.0 + file LICENSE

**Encoding** UTF-8

**Suggests** testthat, knitr, rmarkdown, BiocStyle, lintr, lassoshooting

**VignetteBuilder** knitr

**RoxygenNote** 6.0.1

**SystemRequirements** C++11

**LinkingTo** Rcpp, RcppArmadillo

**Imports** Rcpp, stats

**NeedsCompilation** yes

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

**git\_branch** RELEASE\_3\_10

**git\_last\_commit** 8a63f73

**git\_last\_commit\_date** 2019-10-29

**Date/Publication** 2020-04-14

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netReg-package	<i>netReg</i>
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### Description

*netReg* is a package for generalized linear regression that includes prior graphs in the models objective function.

### Details

*netReg* uses *Armadillo*, *OpenBLAS*, *BLAS* and *LAPACK* for fast matrix computations and *Dlib* for constrained derivate-free optimization.

### Author(s)

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### References

Dirmeier, Simon and Fuchs, Christiane and Mueller, Nikola S and Theis, Fabian J (2018), netReg: Network-regularized linear models for biological association studies.

*Bioinformatics*

Friedman J., Hastie T., Hoefling H. and Tibshirani R. (2007), Pathwise coordinate optimization.

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*Journal of Statistical Software*

Fu W. J. (1998), Penalized Regression: The Bridge Versus the Lasso.

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*Bioinformatics*

Powell M.J.D. (2009), The BOBYQA algorithm for bound constrained optimization without derivatives.

[http://www.damtp.cam.ac.uk/user/na/NA\\_papers/NA2009\\_06.pdf](http://www.damtp.cam.ac.uk/user/na/NA_papers/NA2009_06.pdf)

cv.edgenet

*Find the optimal shrinkage parameters for edgenet***Description**

Finds the optimal shrinkage parameters using cross-validation for edgenet. We use the BOBYQA algorithm to find the optimal regularization parameters and coordinate descent in order to minimize the objective function of the linear model.

**Usage**

```
cv.edgenet(X, Y, G.X = NULL, G.Y = NULL, lambda = NULL, psigx = NULL,
  psigy = NULL, thresh = 1e-05, maxit = 1e+05, family = c("gaussian"),
  optim.epsilon = 0.001, optim.maxit = 10000, nfolds = 10)
```

**Arguments**

X	input matrix, of dimension (n x p) where n is the number of observations and p is the number of covariables. Each row is an observation vector.
Y	output matrix, of dimension (n x q) where n is the number of observations and q is the number of response variables. Each row is an observation vector.
G.X	non-negative affinity matrix for n, of dimensions (p x p) where p is the number of covariables X. Providing a graph G.X will optimize the regularization parameter <code>psi.gx</code> . If this is not desired just set G.X to NULL.
G.Y	non-negative affinity matrix for n, of dimensions (q x q) where q is the number of responses Y. Providing a graph G.Y will optimize the regularization parameter <code>psygy</code> . If this is not desired just set G.Y to NULL.
lambda	numerical shrinkage parameter for LASSO. Per default this parameter is set to NULL which means that lambda is going to be estimated using cross-validation. If any numerical value for lambda is set, estimation of the optimal parameter will <i>not</i> be conducted.
psigx	numerical shrinkage parameter for graph-regularization of G.X. Per default this parameter is set to NULL which means that psigx is going to be estimated in the cross-validation. If any numerical value for psigx is set, estimation of the optimal parameter will <i>not</i> be conducted.
psigy	numerical shrinkage parameter for graph-regularization of G.Y. Per default this parameter is set to NULL which means that psigy is going to be estimated in the cross-validation. If any numerical value for psigy is set, estimation of the optimal parameter will <i>not</i> be conducted.
thresh	numerical threshold for coordinate descent
maxit	maximum number of iterations for the coordinate descent (integer)
family	family of response, e.g. <i>gaussian</i>
optim.epsilon	numerical threshold criterion for the optimization to stop. Usually 1e-3 is a good choice.
optim.maxit	the maximum number of iterations for the optimization (integer). Usually 1e4 is a good choice.
nfolds	the number of folds to be used - default is 10 (minimum 3, maximum <code>nrow(X)</code> ).

**Value**

An object of class `cv.edgenet`

<code>call</code>	the call that produced the object
<code>lambda</code>	the estimated $(p \times q)$ -dimensional coefficient matrix $B.hat$
<code>psigx</code>	the estimated $(q \times 1)$ -dimensional vector of intercepts
<code>psigy</code>	the estimated $(q \times 1)$ -dimensional vector of intercepts

**References**

Dirmeier, Simon and Fuchs, Christiane and Mueller, Nikola S and Theis, Fabian J (2018), *netReg: Network-regularized linear models for biological association studies.*

Friedman J., Hastie T., Hoefling H. and Tibshirani R. (2007), Pathwise coordinate optimization. *The Annals of Applied Statistics*

Friedman J., Hastie T. and Tibshirani R. (2010), Regularization Paths for Generalized Linear Models via Coordinate Descent. *Journal of Statistical Software*

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Powell M.J.D. (2009), The BOBYQA algorithm for bound constrained optimization without derivatives. [http://www.damtp.cam.ac.uk/user/na/NA\\_papers/NA2009\\_06.pdf](http://www.damtp.cam.ac.uk/user/na/NA_papers/NA2009_06.pdf)

**Examples**

```
X <- matrix(rnorm(100*10), 100, 10)
b <- rnorm(10)
G.X <- matrix(rpois(10*10,1),10)
G.X <- t(G.X) + G.X
diag(G.X) <- 0

# fit a Gaussian model
Y <- X%*%b + rnorm(100)
cv.edge <- cv.edgenet(X=X, Y=Y, G.X=G.X, family="gaussian")
```

---

edgenet

*Fit a graph-regularized linear regression model using edge-based regularization.*

---

**Description**

Fit a graph-regularized linear regression model using edge-penalization. The coefficients are computed using graph-prior knowledge in the form of one/two affinity matrices. Graph-regularization is an extension to previously introduced regularization techniques, such as the LASSO. For that reason we are also using coordinate descent for minimization of the objective function of the linear model.

**Usage**

```
edgenet(X, Y, G.X = NULL, G.Y = NULL, lambda = 1, psigx = 1,
        psigy = 1, thresh = 1e-05, maxit = 1e+05, family = c("gaussian"))
```

**Arguments**

X	input matrix, of dimension (n x p) where n is the number of observations and p is the number of covariables. Each row is an observation vector.
Y	output matrix, of dimension (n x q) where n is the number of observations and q is the number of response variables. Each row is an observation vector.
G.X	non-negativ affinity matrix for n, of dimensions (p x p) where p is the number of covariables X
G.Y	non-negativ affinity matrix for n, of dimensions (q x q) where q is the number of responses Y
lambda	numerical shrinkage parameter for LASSO.
psigx	numerical shrinkage parameter for graph-regularization of G.X
psigy	numerical shrinkage parameter for graph-regularization of G.Y
thresh	numerical threshold for coordinate descent
maxit	maximum number of iterations for coordinate descent (integer)
family	family of response, e.g. <i>gaussian</i>

**Value**

An object of class edgenet

coefficients	the estimated (p x q)-dimensional coefficient matrix $\hat{B}$
intercept	the estimated (q x 1)-dimensional vector of intercepts
call	the call that produced the object
family	the family of the response

**References**

Dirmeier, Simon and Fuchs, Christiane and Mueller, Nikola S and Theis, Fabian J (2018), netReg: Network-regularized linear models for biological association studies.  
*Bioinformatics*

Friedman J., Hastie T., Hoefling H. and Tibshirani R. (2007), Pathwise coordinate optimization.  
*The Annals of Applied Statistics*

Friedman J., Hastie T. and Tibshirani R. (2010), Regularization Paths for Generalized Linear Models via Coordinate Descent.  
*Journal of Statistical Software*

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*Journal of Computational and Graphical Statistics*

Cheng W. and Wang W. (2014), Graph-regularized dual Lasso for robust eQTL mapping.  
*Bioinformatics*

**Examples**

```
X <- matrix(rnorm(100*10), 100, 10)
b <- rnorm(10)
G.X <- matrix(rpois(100,1), 10)
G.X <- t(G.X) + G.X
diag(G.X) <- 0

# fit a Gaussian model
Y <- X%*%b + rnorm(100)
fit <- edgenet(X=X, Y=Y, G.X=G.X, family="gaussian")
```

---

predict.gaussian.edgenet

*Predict method for gaussian edgenet fits*

---

**Description**

Predicts the estimated  $\hat{Y}$  values for a newdata design matrix  $X$  similar to the other predict methods, e.g. from glm and glmnet

**Usage**

```
## S3 method for class 'gaussian.edgenet'
predict(object, newdata = NULL, ...)
```

**Arguments**

object	a fitted object of class <i>gaussian.edgenet</i>
newdata	a new (m x p)-dimensional design matrix with a variable number of observations m, but a constant number of co-variables p
...	further arguments

**Value**

A (m x q)-dimensional matrix

**Examples**

```
## Not run:
X <- matrix(rnorm(100*10),100,10)
G.X <- matrix(rpois(10*10,1),10)
G.X <- t(G.X) + G.X
diag(G.X) <- 0

Y <- matrix(rnorm(100*10),100,10)
fit <- edgenet(X=X, Y=Y, G.X=G.X, family="gaussian")
pred <- predict(fit, X)

## End(Not run)
```

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yeast

*A sample yeast data set for regression*

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### **Description**

The yeast data set is a `list` containing three matrices that can be used as an example for using `netReg`. The data have been taken from the references listed below.

### **Usage**

```
data(yeast)
```

### **Format**

A `list` containing three matrices

### **Details**

- $X$  (112 x 500)-dimensional binary matrix of 500 genetic markers for 112 yeast samples
- $Y$  (112 x 231)-dimensional double matrix of 231 gene expression values for 112 yeast samples
- $GY$  (231 x 231)-dimensional adjacency matrix representing protein-protein interactions for 231 yeast genes

### **References**

Brem, Rachel B., et al. (2005), Genetic interactions between polymorphisms that affect gene expression in yeast.  
*Nature*

Storey, John D., Joshua M. Akey, and Leonid Kruglyak (2005), Multiple locus linkage analysis of genomewide expression in yeast.  
*PLoS Biology*

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