

# Package ‘glmaag’

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**Title** Adaptive LASSO and Network Regularized Generalized Linear Models

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**Author** Kaiqiao Li [aut, cre],  
Pei Fen Kuan [aut],  
Xuefeng Wang [aut]

**Maintainer** Kaiqiao Li <kaiqiao.li@stonybrook.edu>

**Description** Efficient procedures for adaptive LASSO and network regularized for Gaussian, logistic, and Cox model. Provides network estimation procedure (combination of methods proposed by Ucar, et. al (2007) <doi:10.1093/bioinformatics/btm423> and Meinshausen and Buhlmann (2006) <doi:10.1214/009053606000000281>), cross validation and stability selection proposed by Meinshausen and Buhlmann (2010) <doi:10.1111/j.1467-9868.2010.00740.x> and Liu, Roeder and Wasserman (2010) <arXiv:1006.3316> methods. Interactive R app is available.

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---

coef.cv_glmaag	<i>Coefficients</i>
----------------	---------------------

---

**Description**

Get the coefficients estimated by the cv\_glmaag model

**Usage**

```
## S3 method for class 'cv_glmaag'
coef(object, type1se = T, ...)
```

**Arguments**

object	the estimated cv_glmaag model
type1se	whether or not used 1 SE error (default to be TRUE)
...	...

**Value**

estimated coefficient included intercept (Cox model does not return intercept)

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, -(1:3)]
cvwhich <- sample(rep(0:4, length.out = length(y)))
mod <- cv_glmaag(y, x, L0, cvwhich = cvwhich)
cc <- coef(mod)
```

---

coef.glmaag

*Coefficients for glmaag*


---

**Description**

Get coefficients for glmaag objects

**Usage**

```
## S3 method for class 'glmaag'
coef(object, lam1, lam2, ...)
```

**Arguments**

object	fitted glmaag object
lam1	lambda1 sequence need coefficients, must be within the fitted model
lam2	lambda2 sequence need coefficients, must be within the fitted model
...	...

**Value**

coefficients

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, -(1:3)]
mod <- glmaag(y, x, L0)
cc <- coef(mod)
```

---

coef.ss_glmaag	<i>Coefficients for ss_glmaag</i>
----------------	-----------------------------------

---

**Description**

Get the coefficients tuned by stability selection

**Usage**

```
## S3 method for class 'ss_glmaag'
coef(object, ...)
```

**Arguments**

object	the model estimated via stability selection
...	...

**Value**

the optimal coefficients get from stability selection including intercept (except for Cox)

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, 4:6]
mod <- ss_glmaag(y, x, L0[seq_len(3), seq_len(3)], nsam = 3)
cc <- coef(mod)
```

---

cv_glmaag	<i>Cross validation for glmaag</i>
-----------	------------------------------------

---

**Description**

Do k-fold cross-validation for glmaag

**Usage**

```
cv_glmaag(y, x, L, nfolds = 5, cvwhich, foldseed, stratify = T,
  gam = 1, tune = F, est = T, lam1, lam2, dfmax, w0, adaptl1 = T,
  adaptl2 = T, pind, intercept = T, standardize = T,
  maxiter = 10000, cri = 0.001, fam = "Gaussian", measdev = T,
  type1se = T, parallel = F)
```

**Arguments**

y	outcome
x	predictors matrix
L	Laplacian matrix for the first network
nfolds	number of folds used in cross validation, default to be five
cvwhich	fold assignment, start from zero, if missing do random cross validation
foldseed	the random seed for cross validation design
stratify	whether to do stratified cross validation for Logistic or Cox model, default to be TRUE
gam	The power of weights of L1 penalty, default to be ones
tune	whether to tune the input network with estimated network or identity matrix, ignored if no input network
est	when there is no input network whether to use estimated network or identity matrix (elastic net) or mixed the network with estimated network or identity matrix, default to be estimated network
lam1	The tuning parameters for L1 penalty. If not defined, searched by default
lam2	The tuning parameters for quadratic penalty. If not defined, searched by default
dfmax	maximum number of parameters allowed in the model, default to be $p/2$
w0	Weights for L1 penalty. If not defined, estimated via quadratic penalized regression
adaptl1	whether to adapt the L1 penalty, default to be TRUE
adaptl2	whether to adapt the sign for quadratic penalty, default to be TRUE
pind	indicator vector whether to put L1 penalty on the feature, 1 means penalized while 0 means not penalized, default to be all ones (all penalized)
intercept	whether to include intercept. Ignore for Cox regression
standardize	whether to standardize predictors
maxiter	maximum number of iterations, default to be 500
cri	stoppint criterion, default to be 0.001
fam	family for the outcome, can be "Gaussian", "Logistic", and "Cox"
measdev	Whether to use deviance to tune, default to be deviance. If not, use mean absolute error, area under ROC curve, or concordance index for Gaussian, Logistic, and Cox
type1se	whether to use one standard error or maximum rule, default to be one standard error rule
parallel	whether to do parallel computing at each fold, need to set up parallel first, default to be FALSE

**Value**

input	input predictor matrix
inputweight	estimated weights if mixing network
lambda1	lambda1 path that has been searched
lambda1	lambda1 path that has been searched
lambda1_max	selected lambda1 based on maximum rule
lambda2_max	selected lambda2 based on maximum rule
lambda1_1se	selected lambda1 based on one standard error rule
lambda2_1se	selected lambda2 based on one standard error rule
cvm	the mean cross validation accuracy
cv1se	the standard error of cross validation accuracy
cvn	the mean number of parameter estimated among folds
n_max	number of selected features based on maximum rule
n_1se	number of selected features based on one standard error rule
intercept_max	estimated intercept based on maximum rule
intercept_1se	estimated intercept based on one standard error rule
coef_max	estimated coefficients based on maximum rule
coef_1se	estimated coefficients based on one standard error rule
fam	family of outcome
measure	measure in cross validation

**Examples**

```

data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, -(1:3)]
cvwhich <- sample(rep(0:4, length.out = length(y)))
mod <- cv_glmag(y, x, L0, cvwhich = cvwhich)

```

---

evaluate

*Evaluate prediction*

---

**Description**

Evaluate goodness of prediction.

**Usage**

```
evaluate(y_pre, y, cutpoint = 0.5, fam = "Gaussian")
```

**Arguments**

y_pre	predicted value
y	actual values (class for binary phenotype and Surv object for right censored phenotype)
cutpoint	cutpoints for binary phenotype, default to be 0.5
fam	family of the phenotype, can be "continuous", "binary", or "Cox"

**Value**

goodness of prediction

**Examples**

```
x <- rnorm(100)
y <- rnorm(100)
evaluate(x, y)
```

---

evaluate_plot	<i>Prediction visualization</i>
---------------	---------------------------------

---

**Description**

Sample plots for prediction evaluation (scatter plot for Gaussian, ROC curve for logistic, and Kaplan Meier curve for Cox)

**Usage**

```
evaluate_plot(y_pre, y_test, fam = "Gaussian", mod, y_train, cutp)
```

**Arguments**

y_pre	predicted value
y_test	actual value
fam	type of predicted outcome, can be "Gaussian" (default), "Logistic", and "Cox"
mod	fitted glmargraph model, must be available for Cox if cutpoint not provided
y_train	the training outcome to obtain
cutp	cutpoint for Cox model

**Value**

plots

**Examples**

```
x <- rnorm(100)
y <- x + rnorm(100)
evaluate_plot(x, y)
```

---

getcut	<i>Get optimal cut points for binary or right censored phenotype</i>
--------	--

---

**Description**

Obtain optimal cut point based on Youden index for binary phenotype and log rank test for right censored phenotype.

**Usage**

```
getcut(pre, act, fam = "Logistic")
```

**Arguments**

pre	predicted value
act	actual values (class for binary phenotype and Surv object for right censored phenotype)
fam	the family of the outcome, can be "Gaussian", "Logistic" or "Cox"

**Value**

optimal cut point

**Examples**

```
x <- rnorm(100)
y <- as.numeric(x + rlogis(100) > 0)
getcut(x, y)
```

---

getS	<i>Estimate standardized Laplacian matrix</i>
------	---

---

**Description**

Estimate standardized Laplacian matrix given data using gene co-expression network method

**Usage**

```
getS(x, sparse = T)
```

**Arguments**

x	data
sparse	estimate a sparse network or not, default to be T, but may be slow



**Value**

standardized laplacian matrix

**References**

Ucar D, Neuhaus I, Ross-MacDonald P, Tilford C, Parthasarathy S, et al. (2007) Construction of a reference gene association network from multiple profiling data: application to data analysis. *Bioinformatics* 23: 2716-2724.

Meinshausen, N., & Bühlmann, P. (2006). High-dimensional graphs and variable selection with the lasso. *The annals of statistics*, 1436-1462.

**Examples**

```
xx <- matrix(rnorm(12), 3, 4)
ss <- getS(xx, FALSE)
```

---

glmaag

*Fit glmaag model*

---

**Description**

Fit the glmaag model with given tuning parameters without cross validation or stability selection

**Usage**

```
glmaag(y, x, L, tune = F, est = T, gam = 1, lam1, lam2, nfolds = 5,
       dfmax, w0, adaptl1 = T, adaptl2 = T, pind, intercept = T,
       standardize = T, maxiter = 10000, cri = 0.001, fam = "Gaussian",
       measdev = T, typeise = T, parallel = F)
```

**Arguments**

y	outcome
x	predictors matrix
L	Laplacian matrix for the network
tune	whether to tune with an estimated network, default to be FALSE
est	whether to estimate a network from the data
gam	the parameter for l1 adaptive weight, default to be ones
lam1	The tuning parameters for L1 penalty. If not defined, searched by default
lam2	The tuning parameters for quadratic penalty. If not defined, searched by default
nfolds	number of folds used in cross validation to obtain network sign estimate and l1 weight estimate, default to be five
dfmax	maximum number of parameters allowed in the model, default to be p/2

w0	Weights for l1 penalty. If not defined, estimated via quadratic penalized regression
adaptl1	whether to adapt the l1 penalty, default to be TRUE
adaptl2	whether to adapt the sign for quadratic penalty, default to be TRUE
pin	indicator vector whether to put L1 penalty on the feature, 1 means penalized while 0 means not penalized, default to be all ones (all penalized)
intercept	whether to include intercept. Ignore for Cox regression
standardize	whether to standardize predictors
maxiter	maximum number of iterations, default to be 500
cri	stoppint criterion, default to be 0.001
fam	family for the outcome, can be "Gaussian", "Logistic", and "Cox"
measdev	Whether to use deviance to tune when estimate l1 weight and network sign, default to be deviance. If not, use mean absolute error, area under ROC curve, or concordance index for Gaussian, Logistic, and Cox
type1se	whether to use one standard error or maximum rule when estimate network sign and l1 weight, default to be one standard error rule
parallel	whether to do parallel computing at each lambda2, need to set up parallel first, default to be FALSE

### Value

input	input predictors
lambda1	l1 penalty parameter search sequence
lambda2	quadratic penalty parameter search sequence
ns	number of parameters selected given provided tuning parameter
coefs	coefficients estimated
intercept	intercepts estimated
loglik	log likelihood estimated
fam	family of the outcome

### Examples

```

data(sampledata)
data(L0)
y <- sampledata$Y_Gau
x <- sampledata[, -(1:3)]
mod <- glmaag(y, x, L0)

```

---

L0                      *sample network 0*

---

**Description**

A Laplacian matrix for the predictors

**Usage**

L0

**Format**

a matrix with 20 rows and 20 columns

---

L1                      *sample network 1*

---

**Description**

An alternative Laplacian matrix for the predictors

**Usage**

L1

**Format**

a matrix with 20 rows and 20 columns

---

laps                      *Standardized Laplacian matrix*

---

**Description**

Obtain standardized Laplacian matrix given adjacency matrix

**Usage**

laps(A)

**Arguments**

A                      adjacency matrix

**Value**

Laplacian matrix

**Examples**

```
a <- matrix(0, 2, 2)
la <- laps(a)
```

---

plot.cv\_glmaag            *Cross validation plot*

---

**Description**

plot cross validation performance paths

**Usage**

```
## S3 method for class 'cv_glmaag'
plot(x, col_count = 3, SE = T, ...)
```

**Arguments**

x	the cv_glmaag object
col_count	number of columns in the plots
SE	whether or not plot the standard error curves (when SE = TRUE)
...	...

**Value**

plot generated by the model

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, -(1:3)]
cvwhich <- sample(rep(0:4, length.out = length(y)))
mod <- cv_glmaag(y, x, L0, cvwhich = cvwhich)
gg <- plot(mod, SE = FALSE)
```

---

plot.glmaag	<i>Paths for glmaag object</i>
-------------	--------------------------------

---

**Description**

Generates coefficients, log likelihood, or number of parameters paths for glmaag models

**Usage**

```
## S3 method for class 'glmaag'
plot(x, col_count = 3, type = "coef", ...)
```

**Arguments**

x	glmaag object
col_count	number of columns shown in the plot (when type = 'coef')
type	can be "coef" (coefficients paths), "loglik" (log likelihood paths), or "n" (number of parameters paths)
...	...

**Value**

plots

**Examples**

```
data(sampledata)
data(L0)
y <- sampledata$Y_Gau
x <- sampledata[, -(1:3)]
mod <- glmaag(y, x, L0)
gg <- plot(mod, type = 'loglik')
```

---

plot.ss_glmaag	<i>Instability plot</i>
----------------	-------------------------

---

**Description**

Instability path plot

**Usage**

```
## S3 method for class 'ss_glmaag'
plot(x, ...)
```

**Arguments**

x                    the input ss\_glmagrph object  
 ...                    ...

**Value**

the instability path

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, 4:6]
mod <- ss_glmaag(y, x, L0[seq_len(3), seq_len(3)], nsam = 3)
gg <- plot(mod)
```

---

predict.cv\_glmaag      *Predict*

---

**Description**

Prediction for cv\_glmaag model

**Usage**

```
## S3 method for class 'cv_glmaag'
predict(object, x, type1se = T, type = "link", ...)
```

**Arguments**

object                the estimated cv\_glmaag model  
 x                      the new dataset for prediction, if omitted returns the training prediction  
 type1se                whether or not using the coefficients by one standard error rule, default to be TRUE  
 type                    can be either "link", or "response", link returns linear predicted score, For Gaussian model this option can be ignored, for logistic model "response" returns predicted probability, for Cox model "reponse" returns relative risk  
 ...                    ...

**Value**

the predicted value

**Examples**

```

data(sampledata)
data(L0)
y <- sampledata$Y_Gau
x <- sampledata[, -(1:3)]
cvwhich <- sample(rep(0:4, length.out = length(y)))
mod <- cv_glmaag(y, x, L0, cvwhich = cvwhich)
pp <- predict(mod)

```

---

predict.glmaag	<i>Prediction for glmaag</i>
----------------	------------------------------

---

**Description**

Prediction using glmaag model

**Usage**

```

## S3 method for class 'glmaag'
predict(object, x, lam1, lam2, type = "link",
        cutp = 0.5, ...)

```

**Arguments**

object	fitted glmaag object
x	The new dataset to be predicted, do training prediction if x is missing
lam1	lambda1 sequence for prediction, must be within the fitted model
lam2	lambda2 sequence for prediction, must be within the fitted model
type	type of prediction (can be "link", "reponse"), ignored for Gaussian model. "link" is the linear predicted score, "response" is the predicted probability for logistic model and relative risk for Cox model
cutp	the cut off value for binary outcome, default to be 0.5
...	...

**Value**

predicted values

**Examples**

```

data(sampledata)
data(L0)
y <- sampledata$Y_Gau
x <- sampledata[, -(1:3)]
mod <- glmaag(y, x, L0)
pp <- predict(mod)

```

---

predict.ss\_glmaag      *Prediction via stability selection*

---

### Description

Predict using the model tuned by stability selection

### Usage

```
## S3 method for class 'ss_glmaag'
predict(object, x, type = "link", ...)
```

### Arguments

object	the ss_glmaag object
x	the new dataset to be predicted, do training prediction if x is missing
type	type of prediction (can be "link", or "reponse" ), ignored for Gaussian model. "link" is the linear predicted score, "response" is the predicted probability for logistic model and relative risk for Cox model
...	...

### Value

the predicted values

### Examples

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, 4:6]
mod <- ss_glmaag(y, x, L0[seq_len(3), seq_len(3)], nsam = 3)
pp <- predict(mod)
```

---

print.cv\_glmaag      *the results of the cross validation model*

---

### Description

print fitted information

### Usage

```
## S3 method for class 'cv_glmaag'
print(x, ...)
```



**Arguments**

x                    the fitted cv\_glmaag object  
 ...                    ...

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, -(1:3)]
cvwhich <- sample(rep(0:4, length.out = length(y)))
mod <- cv_glmaag(y, x, L0, cvwhich = cvwhich)
print(mod)
```

---

print.ss\_glmaag            *the results of the stability selection model*

---

**Description**

print fitted information

**Usage**

```
## S3 method for class 'ss_glmaag'
print(x, ...)
```

**Arguments**

x                    the fitted ss\_glmaag object  
 ...                    ...

**Examples**

```
data(sampleddata)
data(L0)
y <- sampleddata$Y_Gau
x <- sampleddata[, 4:6]
mod <- ss_glmaag(y, x, L0[seq_len(3), seq_len(3)], nsam = 3)
print(mod)
```

runtheExample      *Shiny app*

---

**Description**

Run the shiny app for web interactive using. You need to load data.table, dplyr, ggplot2, plotROC, and survminer beforehand. If you want to do parallel computing, you also need to register cores.

**Usage**

```
runtheExample(whetherrun)
```

**Arguments**

whetherrun      whether to run shiny app, default to be TRUE

**Examples**

```
runtheExample(FALSE)
```

---

sampledata      *Simulated data*

---

**Description**

A data set containing outcome for Gaussian, logistic, and Cox variables and 20 predictors

**Usage**

```
sampledata
```

**Format**

a data frame with 100 rows and 23 variables

ss\_glmaag

*Stability selection for glmaag***Description**

Do stability selection for glmaag

**Usage**

```
ss_glmaag(y, x, L, nfolds = 5, subn, nsam = 100, beta = 0.15,
  gam = 1, tune = F, est = T, lam1, lam2, w0, adaptl1 = T,
  adaptl2 = T, pind, intercept = T, standardize = T,
  maxiter = 10000, cri = 0.001, fam = "Gaussian", measdev = T,
  type1se = T, parallel = F)
```

**Arguments**

y	outcome
x	predictors matrix
L	Laplacian matrix for the first network
nfolds	number of folds used in cross validation to estimate the l1 weights or network tuning, default to be five
subn	number of samples in each subset, default to be n/2 if n<400 and 10sqrt(10) if n>400
nsam	number of subsets, default to be 100
beta	the cut off for instability score
gam	The power of weights of l1 penalty, default to be ones
tune	whether to tune the input network with estimated network or identity matrix, ignored if no input network
est	when there is no input network whether to use estimated network or identity matrix (elastic net) or mixed the network with estimated network or identity matrix, default to be estimated network
lam1	The tuning parameters for l1 penalty. If not defined, searched by default
lam2	The tuning parameters for quadratic penalty. If not defined, searched by default
w0	Weights for l1 penalty. If not defined, estimated via quadratic penalized regression
adaptl1	whether to adapt the l1 penalty, default to be TRUE
adaptl2	whether to adapt the sign for quadratic penalty, default to be TRUE
pind	indicator vector whether to put l1 penalty on the feature, 1 means penalized while 0 means not penalized, default to be all ones (all penalized)
intercept	whether to include intercept. Ignore for Cox regression
standardize	whether to standardize predictors

maxiter	maximum number of iterations, default to be 500
cri	stoppint criterion, default to be 0.001
fam	family for the outcome, can be "Gaussian", "Logistic", and "Cox"
measdev	Whether to use deviance to tune when estimate l1 weight and network sign, default to be deviance. If not, use mean absolute error, area under ROC curve, or concordance index for Gaussian, Logistic, and Cox
type1se	whether to use one standard error or maximum rule for l1 weight estimation and network sign, default to be one standard error rule
parallel	whether to do parallel computing at each subset, need to set up parallel first, default to be FALSE

### Value

input	input matrix for predictors
lambda1	searching sequence for l1 penalty parameters
lambda2	searching sequence for quadratic penalty parameters
lambda1_ss	optimal l1 parameter
lambda2_ss	optimal quadratic parameter
n_ss	number of parameters obtained by the optimal model
ssm	instability score paths
ssf	selection probability paths
intercept_ss	intercept estimated by the optimal model
coef_ss	coefficients estimated by the optimal model
fam	the family of the outcome

### References

Meinshausen, N., & Bühlmann, P. (2010). Stability selection. *Journal of the Royal Statistical Society: Series B (Statistical Methodology)*, 72(4), 417-473.

Liu, H., Roeder, K., & Wasserman, L. (2010). Stability approach to regularization selection (stars) for high dimensional graphical models. In *Advances in neural information processing systems* (pp. 1432-1440).

### Examples

```
data(sampledata)
data(L0)
y <- sampledata$Y_Gau
x <- sampledata[, 4:6]
mod <- ss_glmaag(y, x, L0[seq_len(3), seq_len(3)], nsam = 3)
```

---

tune_network	<i>tune two network</i>
--------------	-------------------------

---

### Description

Tune two network for better prediction.

### Usage

```
tune_network(y, x, L1, L2, adaptl2 = T, nfolds = 5, cvwhich, foldseed,
  stratify = T, lam0, bets, intercept = T, standardize = T,
  fam = "Gaussian", type1se = T, measdev = T, maxiter = 10000,
  cri = 0.001, parallel = F)
```

### Arguments

y	outcome
x	predictors matrix
L1	Laplacian matrix for the first network
L2	Laplacian matrix for the second network
adaptl2	whether to adapt the sign for quadratic penalty, default to be TRUE
nfolds	number of folds used in cross validation, default to be five
cvwhich	fold assignment, start from zero, if missing do random cross validation
foldseed	the random seed for cross validation design
stratify	whether to do stratified cross validation for Logistic or Cox model, default to be TRUE
lam0	The tuning parameters for quadratic penalty. If not defined, tuned by default
bets	The candidate weight for the first network, must be between 0 and 1, default to be 0, 0.1,..., 1
intercept	whether to include intercept. Ignore for Cox regression
standardize	whether to standardize predictors
fam	family for the outcome, can be "Gaussian", "Logistic", and "Cox"
type1se	whether to use one standard error or maximum rule, default to be one standard error rule
measdev	Whether to use deviance to tune, default to be deviance. If not, use mean absolute error, area under ROC curve, or concordance index for Gaussian, Logistic, and Cox
maxiter	maximum number of iterations, default to be 500
cri	stoppint criterion, default to be 0.001
parallel	whether to do parallel computing at each fold

**Value**

est	estimated mixed Laplacian matrix
weight	weights for the two Laplacian matrix

**Examples**

```
data(sampledata)
data(L0)
data(L1)
y <- sampledata$Y_Gau
x <- sampledata[, -(1:3)]
Ltune <- tune_network(y, x, L0, L1, adaptl2 = FALSE)
weight <- Ltune@weight
Lest <- Ltune@est
```

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