

Package ‘EBEN’

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Type Package

Title Empirical Bayesian Elastic Net

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Maintainer Anhui Huang <anhuihuang@gmail.com>

Description Provides the Empirical Bayesian Elastic Net for handling multicollinearity in generalized linear regression models. As a special case of the 'EBglmnet' package (also available on CRAN), this package encourages a grouping effects to select relevant variables and estimate the corresponding non-zero effects.

License GPL

Depends R (>= 2.10)

NeedsCompilation yes

Repository CRAN

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Author Anhui Huang [aut, cre]

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EBEN-package

*Empirical Bayesian Elastic Net (EBEN)***Description**

Fast EBEN algorithms.

EBEN implements a normal and generalized gamma hierarchical priors.

(**) Two parameters (alpha, lambda) are equivalent with elastic net priors.

(**) When parameter alpha = 1, it is equivalent with EBlasso-NE (normal + exponential)

Two models are available for both methods:

(**) General linear regression model.

(**) Logistic regression model.

Multi-collinearity:

(**) for group of high correlated or collinear variables: EBEN identifies the group of variables estimates their effects together.

(**) group of variables can be selected together.

*Epistasis (two-way interactions) can be included for all models/priors

*model implemented with memory efficient c code.

*LAPACK/BLAS are used for most linear algebra computations.

Details

Package: EBEN
 Type: Package
 Version: 5.2
 Date: 2015-10-06
 License: gpl

Author(s)

Anhui Huang

References

key algorithms:

Cai, X., Huang, A., and Xu, S. (2011). Fast empirical Bayesian LASSO for multiple quantitative trait locus mapping. *BMC Bioinformatics* 12, 211.

Huang A, Xu S, Cai X. (2013). Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. *BMC genetics* 14(1):5.

Huang, A., Xu, S., and Cai, X. (2014). Empirical Bayesian elastic net for multiple quantitative trait locus mapping. *Heredity* 10.1038/hdy.2014.79

Other publications:

Huang, A., E. Martin, et al. (2014). "Detecting genetic interactions in pathway-based genome-wide

association studies." *Genet Epidemiol* 38(4): 300-309.
Huang, A., S. Xu, et al. (2014). "Whole-genome quantitative trait locus mapping reveals major role of epistasis on yield of rice." *PLoS ONE* 9(1): e87330.
Huang, A. (2014). "Sparse model learning for inferring genotype and phenotype associations." Ph.D Dissertation. University of Miami(1186).

BASIS

An Example Data File for the Gauss Model

Description

This is a 1000x481 sample feature matrix

Usage

```
data(BASIS)
```

Format

The format is: int [1:1000, 1:481] 0 -1 0 0 1 0 1 0 1 0 ...

Details

The data was simulated on a 2400cM chromosome, each column corresponded to an even spaced QTL

Source

Huang, A., Xu, S., and Cai, X. (2014). Empirical Bayesian elastic net for multiple quantitative trait locus mapping. *Heredity* 10.1038/hdy.2014.79

Examples

```
data(BASIS)
```

`BASISbinomial`*An Example Data File for the Binomial Model*

Description

This is a 500x481 sample feature matrix

Usage

```
data(BASISbinomial)
```

Format

The format is: int [1:500, 1:481] 0 -1 0 0 0 0 -1 -1 0 1 ...

Details

The data was simulated on a 2400cM chromosome, each column corresponded to an even spaced QTL

Source

Huang A, Xu S, Cai X: Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. BMC genetics 2013, 14(1):5.

Examples

```
data(BASISbinomial)
```

`EBlasticNet.Binomial` *The EB Elastic Net Algorithm for Binomial Model with Normal-Gamma(NG) Prior Distribution*

Description

Generalized linear regression, normal-Gxponential (NG) hierarchical prior for regression coefficients

Usage

```
EBlasticNet.Binomial(BASIS, Target, lambda, alpha, Epis = FALSE, verbose = 0)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Class label of each individual, TAKES VALUES OF 0 OR 1
lambda	Hyperparameter controls degree of shrinkage; can be obtained via Cross Validation; $\lambda > 0$
alpha	Hyperparameter controls degree of shrinkage; can be obtained via Cross Validation; $0 < \alpha < 1$
Epis	TRUE or FALSE for including two-way interactions
verbose	0 or 1; 1: display message; 0 no message

Details

If Epis=TRUE, the program adds two-way interaction of $K*(K-1)/2$ more columns to BASIS

Value

weight	the none-zero regression coefficients: col1,col2 are the indices of the bases(main if equal); col3: coefficient value; col4: posterior variance; col5: t-value; col6: p-value
logLikelihood	log likelihood from the final regression coefficients
WaldScore	Wald Score
Intercept	Intercept
lambda	the hyperparameter; same as input lambda
alpha	the hyperparameter; same as input alpha

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang A, Xu S, Cai X: Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. BMC genetics 2013, 14(1):5.

Examples

```
library(EBEN)
data(BASISbinomial)
data(yBinomial)
#reduce sample size to speed up the running time
n = 50;
k = 100;
N = length(yBinomial);
set = sample(N,n);
```

```
BASIS = BASISbinomial[set,1:k];
y = yBinomial[set];
output = EBelasticNet.Binomial(BASIS, y, lambda = 0.1, alpha = 0.5, Epis = FALSE, verbose = 5)
```

EBlasticNet.BinomialCV

Cross Validation (CV) Function to Determine Hyperparameter of the EB_Elastic Net Algorithm for Binomial Model with Normal-Gamma (NG) Prior Distribution

Description

Hyperparameter controls degree of shrinkage, and is obtained via Cross Validation (CV). This program calculates the maximum lambda that allows one non-zero basis; and performs a search down to $0.001 * \lambda_{\max}$ at even steps. (20 steps)

Usage

```
EBlasticNet.BinomialCV(BASIS, Target, nFolds, foldId, Epis = FALSE, verbose = 0)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Class label of each individual, TAKES VALUES OF 0 OR 1
nFolds	number of n-fold cv
Epis	TRUE or FALSE for including two-way interactions
foldId	random assign samples to different folds
verbose	from 0 to 5; larger verbose displays more messages

Details

If Epis=TRUE, the program adds two-way interaction $K*(K-1)/2$ more columns to BASIS

Value

CrossValidation	col1: hyperparameter; col2: loglikelihood mean; standard ERROR of nfold mean log likelihood
Lmabda_optimal	the optimal hyperparameter as computed
Alpha_optimal	the optimal hyperparameter as computed

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang A, Xu S, Cai X: Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. *BMC genetics* 2013, 14(1):5.

Examples

```
## not run
library(EBEN)
data(BASISbinomial)
data(yBinomial)
#reduce sample size to speed up the running time
n = 50;
k = 100;
N = length(yBinomial);
set.seed(1)
set = sample(N,n);
BASIS = BASISbinomial[set,1:k];
y = yBinomial[set];
nFolds = 3
## Not run:
CV = EBlasticNet.BinomialLCV(BASIS, y, nFolds = 3,Epis = FALSE)

## End(Not run)
```

EBlasticNet.Gaussian *The EB Elastic Net Algorithm for Gaussian Model*

Description

General linear regression, normal-Gamma (NG) hierarchical prior for regression coefficients

Usage

```
EBlasticNet.Gaussian(BASIS, Target, lambda, alpha,Epis = FALSE,verbose = 0)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Response each individual
lambda	Hyperparameter controls degree of shrinkage; can be obtained via Cross Validation; $\lambda > 0$
alpha	Hyperparameter controls degree of shrinkage; can be obtained via Cross Validation; $0 < \alpha < 1$
Epis	TRUE or FALSE for including two-way interactions
verbose	0 or 1; 1: display message; 0 no message

Details

If Epis=TRUE, the program adds two-way interaction of $K*(K-1)/2$ more columns to BASIS

Value

weight	the none-zero regression coefficients: col1,col2 are the indices of the bases(main if equal); col3: coefficient value; col4: posterior variance; col5: t-value; col6: p-value
WaldScore	Wald Score
Intercept	Intercept
lambda	the hyperparameter; same as input lambda
alpha	the hyperparameter; same as input alpha

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang, A., Xu, S., and Cai, X. (2014). Empirical Bayesian elastic net for multiple quantitative trait locus mapping. *Heredity* 10.1038/hdy.2014.79

Examples

```
library(EBEN)
data(BASIS)
data(y)
n = 50;
k = 100;
BASIS = BASIS[1:n,1:k];
y = y[1:n];
Blup = EBlasticNet.Gaussian(BASIS, y, lambda = 0.0072, alpha = 0.95, Epis = FALSE, verbose = 0)
betas = Blup$weight
betas
```

EBlasticNet.GaussianCV

Cross Validation (CV) Function to Determine Hyperparameters of the EBEN Algorithm for Gaussian Model

Description

Hyperparameter controls degree of shrinkage, and is obtained via Cross Validation (CV). This program calculates the maximum lambda that allows one non-zero basis; and performs a search down to $0.0001 * \text{lambda_max}$ at even steps. (20 steps)

Usage

```
EBlasticNet.GaussianCV(BASIS, Target, nFolds, foldId, Epis = FALSE, verbose = 0)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Response each individual
nFolds	number of n-fold cv
Epis	TRUE or FALSE for including two-way interactions
foldId	random assign samples to different folds
verbose	from 0 to 5; larger verbose displays more messages

Details

If Epis=TRUE, the program adds two-way interaction $K*(K-1)/2$ more columns to BASIS

Value

CrossValidation	col1: hyperparameter; col2: loglikelihood mean; standard ERROR of nfold mean log likelihood
Lmabda_optimal	the optimal hyperparameter as computed
Alpha_optimal	the optimal hyperparameter as computed

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang, A., Xu, S., and Cai, X. (2013). Empirical Bayesian elastic net for multiple quantitative trait locus mapping. submitted.

Examples

```
library(EBEN)
data(BASIS)
data(y)
#reduce sample size to speed up the running time
n = 50;
k = 100;
BASIS = BASIS[1:n, 1:k];
```

```

y = y[1:n];
## Not run:
CV = EBlasticNet.GaussianCV(BASIS, y, nFolds = 3,Epis = FALSE)

## End(Not run)

```

EBlassoNEG.Binomial *The EBlasso Algorithm for Binomial Model with Normal-Exponential-Gamma (NEG) Prior Distribution*

Description

Generalized linear regression, normal-exponential-gamma (NEG) hierarchical prior for regression coefficients

Usage

```
EBlassoNEG.Binomial(BASIS, Target, a_gamma, b_gamma, Epis, verbose, group)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Class label of each individual, TAKES VALUES OF 0 OR 1
a_gamma	Hyperparameters control degree of shrinkage; can be obtained via Cross Validation; a_gamma >= -1
b_gamma	Hyperparameters control degree of shrinkage; can be obtained via Cross Validation; b_gamma > 0
Epis	TRUE or FALSE for including two-way interactions
verbose	0 or 1; 1: display message; 0 no message
group	0 or 1; 0: No group effect; 1 two-way interaction grouped. Only valid when Epis = TRUE

Details

If Epis=TRUE, the program adds two-way interaction $K*(K-1)/2$ more columns to BASIS

Value

weight	the none-zero regression coefficients: col1,col2 are the indices of the bases(main if equal); col3: coefficient value; col4: posterior variance; col5: t-value; col6: p-value
logLikelihood	log likelihood with the final regression coefficients

WaldScore	Wald Score
Intercept	Intercept
a_gamma	the hyperparameter; same as input
b_gamma	the hyperparameter; same as input

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang, A., Xu, S., and Cai, X.(2012). Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. BMC Genetics. Submitted

Examples

```
library(EBEN)
data(BASISbinomial)
data(yBinomial)
#reduce sample size to speed up the running time
n = 50;
k = 100;
BASIS = BASISbinomial[1:n,1:k];
y = yBinomial[1:n];
output = EBlassoNEG.Binomial(BASIS,y,0.1,0.1,Epis = FALSE)
```

EBlassoNEG.BinomialCV *Cross Validation (CV) Function to Determine Hyperparameters of the EBlasso Algorithm for Binomial Model with Normal-Exponential-Gamma (NEG) Prior Distribution*

Description

Hyperparameters control degree of shrinkage, and are obtained via Cross Validation. This program performs three steps of CV.

1st: a = b = 0.001, 0.01, 0.1, 1;

2nd: fix b= b1; a=[-0.5, -0.4, -0.3, -0.2, -0.1, -0.01, 0.01, 0.05, 0.1, 0.5, 1];

3rd: fix a = a2; b= 0.01 to 10 with a step size of one for b > 1 and a step size of one on the logarithmic scale for b < 1

In the 2nd step, a can take value from -1 and values in [-1, -0.5] can be added to the set in line 13 of this function (The smaller a is, the less shrinkage.)

Usage

```
EBlassoNEG.BinomialCV(BASIS, Target, nFolds, foldId, Epis, verbose, group)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Class label of each individual, TAKES VALUES OF 0 OR 1
nFolds	number of n-fold cv
foldId	random assign samples to different folds
Epis	TRUE or FALSE for including two-way interactions
verbose	from 0 to 5; larger verbose displays more messages
group	TRUE or FALSE; FALSE: No group effect; TRUE two-way interaction grouped. Only valid when Epis = TRUE

Details

If Epis=TRUE, the program adds two-way interaction $K*(K-1)/2$ more columns to BASIS

Note: Given the fact that degree of shrinkage is a monotonic function of (a,b),

The function implemented a 3-step search as described in Huang, A. 2014, for full grid search, user needs to modify the function accordingly.

Value

CrossValidation	col1: hyperparameters; col2: loglikelihood mean; standard ERROR of nfold mean log likelihood
a_optimal	the optimal hyperparameter as computed
b_optimal	the optimal hyperparameter as computed

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang A, Xu S, Cai X: Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. BMC genetics 2013, 14(1):5.

Huang, A., S. Xu, et al. Whole-genome quantitative trait locus mapping reveals major role of epistasis on yield of rice. PLoS ONE 2014, 9(1): e87330.

Examples

```
library(EBEN)
data(BASISbinomial)
data(yBinomial)
#reduce sample size to speed up the running time
n = 50;
k = 100;
BASIS = BASISbinomial[1:n,1:k];
y = yBinomial[1:n];
## Not run:
CV = EBlassoNEG.BinomialCV(BASIS, y, nFolds = 3,Epis = FALSE, verbose = 0)
```

```
## End(Not run)
```

EBlassoNEG.Gaussian *The EBlasso Algorithm for Gaussian Model with Normal-Exponential-Gamma (NEG) Prior Distribution*

Description

General linear regression, normal-exponential-gamma (NEG) hierarchical prior for regression coefficients

Usage

```
EBlassoNEG.Gaussian(BASIS, Target, a_gamma, b_gamma, Epis, verbose, group)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Response each individual
a_gamma	Hyperparameters control degree of shrinkage; can be obtained via Cross Validation
b_gamma	Hyperparameters control degree of shrinkage; can be obtained via Cross Validation
Epis	TRUE or FALSE for including two-way interactions
verbose	from 0 to 5; larger verbose displays more messages
group	0 or 1; 0: No group effect; 1 two-way interaction grouped. Only valid when Epis = TRUE

Details

If Epis=TURE, the program adds two-way interaction $K*(K-1)/2$ more columns to BASIS for memory efficient, the function pass `n_effect` to C. `n_effect > n_true` effects, which is a rough guess on how many variables will be selected by the function by providing a relative 'small' `n_effect`, the function will not allocate a large trunk of memory during computation.

Value

weight	the none-zero regression coefficients: col1,col2 are the indices of the bases(main if equal); col3: coefficient value; col4: posterior variance; col5: t-value; col6: p-value
--------	--

WaldScore	Wald Score
Intercept	Intercept
residVar	residual variance
a_gamma	the hyperparameter; same as input
b_gamma	the hyperparameter; same as input

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Cai, X., Huang, A., and Xu, S. (2011). Fast empirical Bayesian LASSO for multiple quantitative trait locus mapping. *BMC Bioinformatics* 12, 211.

Examples

```
library(EBEN)
data(BASIS)
data(y)
n = 50;
k = 100;
BASIS = BASIS[1:n,1:k];
y = y[1:n];
output = EBlassoNEG.Gaussian(BASIS, y, a_gamma = 0.1, b_gamma = 0.1)
```

EBlassoNEG.GaussianCV *Cross Validation (CV) Function to Determine Hyperparameters of the EBlasso Algorithm for Gaussian Model with Normal-Exponential-Gamma (NEG) Prior Distribution*

Description

Hyperparameters control degree of shrinkage, and are obtained via Cross Validation. This program performs three steps of CV.

1st: $a = b = 0.001, 0.01, 0.1, 1$;

2nd: fix $b = b_1$; $a = [-0.5, -0.4, -0.3, -0.2, -0.1, -0.01, 0.01, 0.05, 0.1, 0.5, 1]$;

3rd: fix $a = a_2$; $b = 0.01$ to 10 with a step size of one for $b > 1$ and a step size of one on the logarithmic scale for $b < 1$

In the 2nd step, a can take value from -1 and values in $[-1, -0.5]$ can be added to the set in line 13 of this function (The smaller a is, the less shrinkage.)

Usage

```
EBlassoNEG.GaussianCV(BASIS, Target, nFolds, foldId, Epis, verbose, group)
```

Arguments

BASIS	sample matrix; rows correspond to samples, columns correspond to features
Target	Class label of each individual, TAKES VALUES OF 0 OR 1
nFolds	number of n-fold cv
foldId	random assign samples to different folds
Epis	TRUE or FALSE for including two-way interactions
verbose	from 0 to 5; larger verbose displays more messages
group	TRUE or FALSE; FALSE: No group effect; TRUE two-way interaction grouped. Only valid when Epis = TRUE

Details

If Epis= TRUE, the program adds two-way interaction $K*(K-1)/2$ more columns to BASIS

Note: Given the fact that degree of shrinkage is a monotonic function of (a,b),

The function implemented a 3-step search as described in Huang, A. 2014, for full

grid search, user needs to modify the function accordingly.

Value

CrossValidation	col1: hyperparameters; col2: loglikelihood mean; standard ERROR of nfold mean log likelihood
a_optimal	the optimal hyperparameter as computed
b_optimal	the optimal hyperparameter as computed

Author(s)

Anhui Huang; Dept of Electrical and Computer Engineering, Univ of Miami, Coral Gables, FL

References

Huang A, Xu S, Cai X: Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. BMC genetics 2013, 14(1):5.

Huang, A., S. Xu, et al. Whole-genome quantitative trait locus mapping reveals major role of epistasis on yield of rice. PLoS ONE 2014, 9(1): e87330.

Examples

```
library(EBEN)
data(BASIS)
data(y)
#reduce sample size to speed up the running time
n = 50;
k = 100;
BASIS = BASIS[1:n,1:k];
y = y[1:n];
## Not run:
CV = EBlassoNEG.GaussianCV(BASIS, y, nFolds = 3,Epis = FALSE)
```

```
## End(Not run)
```

y	<i>Sample Response Data for Gaussian Model</i>
---	--

Description

Corresponding to the response of BASIS

Usage

```
data(y)
```

Format

The format is: num [1:1000, 1] 113.5 97.1 116.6 96.7 105.5 ...

Source

Huang, A., Xu, S., and Cai, X. (2014). Empirical Bayesian elastic net for multiple quantitative trait locus mapping. *Heredity* 10.1038/hdy.2014.79

Examples

```
data(y)
```

yBinomial	<i>Sample Variable Data for Binomial Model</i>
-----------	--

Description

Corresponding to the class label of BASISbinomial

Usage

```
data(yBinomial)
```

Format

The format is: int [1:500, 1] 1 1 1 1 1 1 1 1 1 ...

Source

Huang A, Xu S, Cai X: Empirical Bayesian LASSO-logistic regression for multiple binary trait locus mapping. *BMC genetics* 2013, 14(1):5.

Examples

```
data(BASISbinomial)
```

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