

# Package ‘Tsphere’

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**Type** Package

**Title** Transposable Sphering for Large-Scale Inference with Correlated Data.

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**Description** Adjusts for correlations among the rows and columns via the Transposable Sphering Algorithm when conducting large-scale inference on the rows of a data matrix.

**License** GPL-2

**Depends** glasso, rms

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ACE

*Alternating Conditional Expectations*

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**Description**

Predicts values for scattered missing elements according to the matrix-variate normal distribution. Internal use.

**Usage**

```
ACE(x, sig, delt, sigi, delti, M, thr = 1e-04, maxit = 1000)
```

**Arguments**

x	Data matrix with scattered missing values. Missing values should be denoted with "NA".
sig	Row covariance matrix.
delt	Column covariance matrix.
sigi	Row precision matrix.
delti	Column precision matrix.
M	Mean matrix.
thr	Convergence threshold.
maxit	Maximum number of iterations.

**Details**

For internal use.

**Value**

x	Matrix of predicted values.
iter	Number of iterations until convergence.

**Author(s)**

Genevera I. Allen

**References**

G. I. Allen and R. Tibshirani, "Transposable regularized covariance models with an application to missing data imputation", *Annals of Applied Statistics*, 4:2, 764-790, 2010.

**See Also**

[CVcov](#)

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covTranspose11                      *Covariance Estimation.*

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### Description

Inverse row and column covariance estimation for the L1 penalized matrix-variate normal model.

### Usage

```
covTranspose11(xc, rhor, rhoc, row = TRUE, sigi.init = NULL,
delti.init = NULL, thr = 1e-04, maxit = 1000, trace = TRUE, thr.glasso
= 1e-04, maxit.glasso = 1000, pen.diag = TRUE)
```

### Arguments

xc	Centered data matrix.
rhor	Row regularization parameter.
rhoc	Column regularization parameter.
row	Logical. TRUE = Start with row covariance.
sigi.init	Initialization for the row precision matrix. (Optional).
delti.init	Initialization for the column precision matrix. (Optional).
thr	Convergence threshold.
maxit	Maximum number of iterations.
trace	Prints matrix-variate log-likelihood for each iteration.
thr.glasso	Convergence threshold for the graphical lasso.
maxit.glasso	Maximum number of iterations for the graphical lasso.
pen.diag	Logical. Indicates whether the diagonal should be penalized.

### Details

Estimates row and column precision matrix via L1 penalized Transposable Regularized Covariance Models.

### Value

Sigmahat	Estimated row covariance.
Deltahat	Estimated column covariance.
Sigmaihat	Estimated sparse row precision matrix.
Deltaihat	Estimated sparse column precision matrix.
loglike	Trace of the penalized log-likelihood at each iteration.

### Author(s)

Genevera I. Allen

**References**

G. I. Allen and R. Tibshirani, "Transposable regularized covariance models with an application to missing data imputation", *Annals of Applied Statistics*, 4:2, 764-790, 2010.

**See Also**

[TransSphere](#)

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 CVcov

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*Cross-Validation.*


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**Description**

Cross-Validation to estimate regularization parameters for sparse inverse covariance estimation.

**Usage**

```
CVcov(x, maxlam, minlam, steps, pmiss = 0.01, do = 2, trace = TRUE)
```

**Arguments**

x	Data matrix.
maxlam	Maximum regularization parameter.
minlam	Minimum regularization parameter.
steps	Number of regularization parameters to test.
pmiss	Percentage missing in each fold.
do	Number of folds. Note that for medium or large size data matrices, often one fold is sufficient.
trace	Logical. Output the penalized log-likelihood and MSE for each step and fold.

**Details**

For internal use.

**Value**

cvmat	Matrix of cross-validation mean squared errors.
optlam	Optimal value of the regularization parameter as estimated by cross-validation.
lams	Values of the regularization parameters tested.

**Author(s)**

Genevera I. Allen

**References**

G. I. Allen and R. Tibshirani, "Transposable regularized covariance models with an application to missing data imputation", *Annals of Applied Statistics*, 4:2, 764-790, 2010.

**See Also**

[covTranspose11](#), [TransSphere](#)

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meanTranspose	<i>Transposable model mean.</i>
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**Description**

Estimates row and column means.

**Usage**

```
meanTranspose(x, tol = 1e-06)
```

**Arguments**

x	Data matrix.
tol	Tolerance for iterative algorithm when data has missing values.

**Details**

Estimates the row and column means.

**Value**

x	Original data matrix.
xcen	Centered data matrix.
mu	Column mean.
nu	Row mean.
M	Mean matrix.

**Author(s)**

Genevera I. Allen

**References**

G. I. Allen and R. Tibshirani, "Transposable regularized covariance models with an application to missing data imputation", *Annals of Applied Statistics*, 4:2, 764-790, 2010.

**See Also**

[covTranspose11](#), [TransSphere](#)

**Examples**

```
x = matrix(rnorm(100*50),100,50)

#row and column centered data matrix
xc = meanTranspose(x)$xcen
```

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MNloglike

*Penalized Matrix-variate normal log-likelihood.*

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**Description**

Penalized Matrix-variate normal log-likelihood.

**Usage**

```
MNloglike(x, M, Sig, Delt, rhor, rhoc, qr = 2, qc = 2, Sigi = NULL,
Delti = NULL)
```

**Arguments**

x	Data matrix.
M	Mean matrix.
Sig	Row covariance matrix.
Delt	Column covariance matrix.
rhor	Row regularization parameter.
rhoc	Column regularization parameter.
qr	Row regularization norm. Either 1 or 2.
qc	Column regularization norm. Either 1 or 2.
Sigi	Row precision matrix. (optional).
Delti	Column precision matrix. (optional).

**Details**

For internal use.

**Value**

value	Log-likelihood.
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**Author(s)**

Genevera I. Allen

**References**

G. I. Allen and R. Tibshirani, "Transposable regularized covariance models with an application to missing data imputation", *Annals of Applied Statistics*, 4:2, 764-790, 2010.

**See Also**

[covTranspose11](#)

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TransSphere

*Transposable Sphering Algorithm for Large-Scale Inference.*

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**Description**

Applies the Transposable Sphering Algorithm to adjust for correlations among the rows and columns when conducting large-scale inference on the rows of a data matrix.

**Usage**

```
TransSphere(dat, y, fdr, minlam, maxlam = NULL)
```

**Arguments**

<code>dat</code>	Data matrix. Inference will be conducted on the rows and the matrix should be oriented in this manner. For example in gene expression data, the data matrix should be oriented as genes by samples.
<code>y</code>	A vector of group labels. Labels should be denoted as a numeric 1 or 2.
<code>fdr</code>	Desired False Discovery Rate to be controlled. Default is 0.1.
<code>minlam</code>	Minimum regularization parameter to test via cross-validation for sparse inverse covariance estimation. Default is 0.15. Note that small values of this parameter may result in numerical instabilities. It is recommended to keep this parameter at the default.
<code>maxlam</code>	Maximum regularization parameter to test via cross-validation for sparse inverse covariance estimation. Default is 0.25.

**Details**

The Transposable Sphering Algorithm adjusts for correlations among the rows and columns of a data matrix before conducting large-scale inference. Currently, this method is only written for two-sample problems. The data matrix is row and column centered and two-sample T-statistics are computed for each row. The Transposable Sphering method is applied to the top 500 rows corresponding to the largest absolute T-statistics. The matrix is decomposed into a signal matrix, corresponding to the two classes of interest, and a noise matrix. This noise matrix is sphered so that both the rows

and columns are approximately independent. Specifically, sparse inverse covariances of the rows and columns are estimated via Transposable Regularized Covariance Models and used to whiten the noise matrix. Cross-validation is used to estimate the regularization parameters controlling the amount of sparsity. The estimated signal matrix and sphered noise matrix are then added to form the sphered data matrix that is used to conduct large-scale inference. Test statistics are adjusted using central-matching, and the Benjamini-Hochberg step-up procedure is used to control the False Discovery Rate.

### Value

sig.rows	The indices of the statistically significant rows after controlling the False Discovery Rate at the value fdr.
t.stats	Sphered two-sample T-statistics.
p.vals	Sphered (unadjusted) p-values.
x.sphered	The sphered data matrix. Note that only the top 500 rows are used in the algorithm so this data matrix is has row dimension at most 500.

### Author(s)

Genevera I. Allen

### References

- G. I. Allen and R. Tibshirani, "Inference with Transposable Data: Modeling the Effects of Row and Column Correlations", To Appear in Journal of the Royal Statistical Society, Series B (Theory & Methods), 2011.
- G. I. Allen and R. Tibshirani, "Transposable regularized covariance models with an application to missing data imputation", Annals of Applied Statistics, 4:2, 764-790, 2010.

### Examples

```
#batch-effect simulation
n = 250
p = 50
y = c(rep(1,25),rep(2,25))
mu1true = c(rep(.5,25),rep(-.5,25),rep(0,n-50))
mu2true = c(rep(-.5,25),rep(.5,25),rep(0,n-50))
Smat = cbind(matrix(mu1true,n,p/2),matrix(mu2true,n,p/2))
mus = c(-.5,-.25,0,.25,.5)
Bmatsig = matrix(1,n,1) %*% t(rep(mus,each=10))
Bmat = Bmatsig + matrix(rnorm(n*p)*.75,n,p)
xxt = matrix(rnorm(2*n^2),n,2*n)
Sig = xxt %*% t(xxt)/(2*n); eSig = eigen(Sig);
xx = matrix(rnorm(n*p),n,p)
x.b = Smat + eSig$eigenvectors %*% diag(sqrt(eSig$values)) %*%
eSig$eigenvectors %*% xx + Bmat

#Transposable Sphering Algorithm
ans = TransSphere(x.b,y,fdr=.1,.15,.25)
```



```
#significant rows
ans$sig.rows

#true positive rate
sum(ans$sig.rows<=50)/50

#false positive rate
sum(ans$sig.rows>50)/200
```

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