

# Package ‘ExpressionCellNet’

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**Type** Package

**Title** Network-Based Analysis of Gene Expression Perturbations

**Version** 0.1.1

**Description** Network-centric framework for integrative analysis of high-throughput gene expression data using user-supplied gene-gene interaction graphs. Constructs seed-centered multi-generation networks constrained by expression correlations and simulates expression perturbation scenarios via regression-based prediction (van Dam, 2018).

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AnalyzeNetwork	<i>Analyze network topology and identify hub genes</i>
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### Description

Computes degree, closeness, betweenness, and eigenvector centrality and stores: (i) Network Analysis table and (ii) Hub Genes table (top 10)

### Usage

```
AnalyzeNetwork(ExpCellNetObj)
```

### Arguments

ExpCellNetObj An ExpressionCellNet object.

### Value

Updated ExpressionCellNet object with "Network Analysis" and "Hub Genes".

### Examples

```
data("example_data")
obj <- AnalyzeNetwork(example_data$expcellnetobj)
```

---

**BuildNetwork***Construct a gene interaction network centered on a seed gene*

---

**Description**

Builds a multi-generation network starting from a seed gene using curated interactions and correlation filtering across generations.

**Usage**

```
BuildNetwork(  
  ExpCellNetObj,  
  Gene = "Gene1",  
  Generation = 3,  
  CorrelationThreshold = c(0.7, 0.7, 0.7)  
)
```

**Arguments**

**ExpCellNetObj** An ExpressionCellNet object containing expression and interaction data.

**Gene** Character; seed gene name (generation 0).

**Generation** Integer; number of generations to expand from the seed gene.

**CorrelationThreshold** Numeric vector; per-generation correlation thresholds.

**Value**

Updated ExpressionCellNet object with added components such as "Network" and "Generations".

**Examples**

```
data("example_data")  
obj <- BuildNetwork(  
  example_data$expcellnetobj,  
  Gene = "Gene1",  
  Generation = 3,  
  CorrelationThreshold = c(0.65, 0.70, 0.75)  
)
```

---

CountMatrixNormalization

*Normalize expression count matrix*

---

### Description

Filters low-expression genes (optional), applies CPM normalization (optional), and log2 transformation (optional). Stores results inside the ExpressionCellNet object.

### Usage

```
CountMatrixNormalization(  
  ExpCellNetObj,  
  CheckExpression = TRUE,  
  CPM = TRUE,  
  LogTransform = FALSE  
)
```

### Arguments

ExpCellNetObj An ExpressionCellNet object.

CheckExpression Logical; if TRUE, low-expression genes are filtered.

CPM Logical; if TRUE, counts are converted to counts-per-million.

LogTransform Logical; if TRUE, log2 transform is applied (typically after CPM).

### Value

Updated ExpressionCellNet object containing normalized matrix.

### Examples

```
data("example_data")  
obj <- CountMatrixNormalization(  
  example_data$expcellnetobj,  
  CheckExpression = TRUE,  
  CPM = TRUE,  
  LogTransform = TRUE  
)
```

---

createExpCellNetObj     *Create an ExpressionCellNet object*

---

**Description**

Wraps the core inputs (count matrix, annotation table, and interaction database) into a single ExpressionCellNet object used by downstream functions.

**Usage**

```
createExpCellNetObj(count_matrix, annotations, intraction_db)
```

**Arguments**

count\_matrix     A gene-by-sample expression count matrix (rows: genes, cols: samples).  
annotations     A data.frame containing gene annotations/mapping information.  
intraction\_db     A data.frame of gene-gene interactions (two columns).

**Value**

An ExpressionCellNet object (list) containing core data components.

**Examples**

```
data("example_data")  
obj <- createExpCellNetObj(  
  count_matrix = example_data$counts,  
  annotations = example_data$annot_df,  
  intraction_db = example_data$interactions  
)
```

---

example\_data     *Example dataset for ExpressionCellNet*

---

**Description**

Simulated example dataset containing four objects: counts, annot\_df, interactions, and expcellnetobj. All data are mock / artificial for testing.

**Usage**

```
data(example_data)
```

**Format**

An RData file containing:

**counts** Numeric matrix of simulated RNA-seq counts.

**annot\_df** Data frame with ENSEMBL IDs and Gene Symbols.

**interactions** Data frame representing gene–gene interactions.

**expcellnetobj** Example ExpressionCellNet object.

---

ExtractFeature	<i>Extract network-derived expression features</i>
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---

**Description**

Extracts a feature matrix from the normalized expression data by restricting genes to those present in the constructed interaction network. The resulting matrix is intended for downstream modeling and dimensionality reduction.

**Usage**

```
ExtractFeature(ExpCellNetObj)
```

**Arguments**

**ExpCellNetObj** An ExpressionCellNet object containing a normalized count matrix and a constructed network.

**Value**

A data.frame or matrix of network-derived expression features (genes  $\times$  samples), suitable for regression or multivariate analysis.

**Examples**

```
data("example_data")
features <- ExtractFeature(example_data$expcellnetobj)
```

---

FindPathway	<i>Find the shortest path between two genes in the network</i>
-------------	--

---

**Description**

Computes the shortest interaction path (as a sequence of gene names) between a start gene and an end gene using the network stored in the ExpressionCellNet object.

**Usage**

```
FindPathway(ExpCellNetObj, Start_gene, End_gene)
```

**Arguments**

ExpCellNetObj	An ExpressionCellNet object containing a constructed network.
Start_gene	Character; source gene symbol (must exist as a node in the network).
End_gene	Character; destination gene symbol (must exist as a node in the network).

**Value**

A character vector of gene symbols representing the shortest path.

**Examples**

```
data("example_data")
path <- FindPathway(example_data$expcellnetobj, Start_gene="Gene1", End_gene="Gene11")
path
```

---

MeanExpression	<i>Compute mean expression of a gene</i>
----------------	--

---

**Description**

Computes mean expression from raw or normalized matrix (default: normalized).

**Usage**

```
MeanExpression(ExpCellNetObj, Gene, Normalized = TRUE)
```

**Arguments**

ExpCellNetObj	An ExpressionCellNet object.
Gene	Character; target gene.
Normalized	Logical; use normalized matrix (TRUE) or raw (FALSE).

**Value**

Numeric mean expression value.

**Examples**

```
data("example_data")
MeanExpression(example_data$expcellnetobj , "Gene1")
```

---

NetworkPrediction	<i>Predict network-wide expression changes after perturbing a seed gene</i>
-------------------	---

---

**Description**

Fits linear regression models for seed-to-network genes and predicts shifts under a user-defined perturbation scenario. Stores multiple result components in the object.

**Usage**

```
NetworkPrediction(ExpCellNetObj, Gene, Expression)
```

**Arguments**

ExpCellNetObj	An ExpressionCellNet object.
Gene	Character; seed gene to perturb.
Expression	Numeric; imposed expression value under perturbation.

**Value**

Updated ExpressionCellNet object with regression models and results.

**Examples**

```
data("example_data")
obj <- NetworkPrediction(example_data$expcellnetobj, Gene="Gene1", Expression=0)
```

---

PCAforGenes	<i>PCA on network-derived features</i>
-------------	--

---

**Description**

Performs principal component analysis (PCA) on the network-restricted feature matrix returned by `ExtractFeature()`, after min-max scaling, and returns a ggplot of PC1 vs PC2.

**Usage**

```
PCAforGenes(ExpCellNetObj)
```

**Arguments**

`ExpCellNetObj` An `ExpressionCellNet` object containing "Network", "Generations", and a normalized count matrix.

**Value**

A ggplot object showing PCA embedding of genes (nodes).

**Examples**

```
data("example_data")
p <- PCAforGenes(example_data$expcellnetobj)
p
```

---

PlotCorrelation	<i>Plot correlation between two genes</i>
-----------------	---

---

**Description**

Plots correlation/regression between two genes (e.g., seed and distal gene).

**Usage**

```
PlotCorrelation(ExpCellNetObj, Gene1, Gene2)
```

**Arguments**

`ExpCellNetObj` An `ExpressionCellNet` object.  
`Gene1` Character; first gene.  
`Gene2` Character; second gene.

**Value**

Invisibly returns NULL (plot is produced).

**Examples**

```
data("example_data")
PlotCorrelation(example_data$expcellnetobj, "Gene1", "Gene11")
```

---

PlotCountMatrix	<i>Plot expression matrix distribution</i>
-----------------	--

---

**Description**

Visualizes the distribution of raw or normalized expression values across samples.

**Usage**

```
PlotCountMatrix(ExpCellNetObj, Matrix = "Normalized")
```

**Arguments**

ExpCellNetObj An ExpressionCellNet object.  
Matrix Character; which matrix to plot (e.g., "RAW" or "Normalized").

**Value**

A plot (base R) is produced as a side effect.

**Examples**

```
data("example_data")
PlotCountMatrix(example_data$expcellnetobj, Matrix = "RAW")
PlotCountMatrix(example_data$expcellnetobj, Matrix = "Normalized")
```

---

PlotNetworkPrediction *Plot network prediction summary*

---

**Description**

Creates a summary plot of predicted percent changes across network genes (bar plot).

**Usage**

```
PlotNetworkPrediction(ExpCellNetObj)
```

**Arguments**

ExpCellNetObj An ExpressionCellNet object after NetworkPrediction().

**Value**

Invisibly returns NULL (plot is produced).

**Examples**

```
data("example_data")
PlotNetworkPrediction(example_data$expcellnetobj)
```

---

PlotPathwayPrediction *Plot pathway-restricted prediction to a target gene*

---

**Description**

Summarizes modeled expression changes along the seed-to-target pathway.

**Usage**

```
PlotPathwayPrediction(ExpCellNetObj, target_gene)
```

**Arguments**

ExpCellNetObj An ExpressionCellNet object after NetworkPrediction().  
target\_gene Character; destination gene to focus on.

**Value**

Invisibly returns NULL (plot is produced).

**Examples**

```
data("example_data")
obj <- NetworkPrediction(example_data$expcellnetobj, Gene="Gene1", Expression=0)
PlotPathwayPrediction(obj, "Gene11")
```

---

RegenerateTrainTest     *Regenerate train/test split*

---

**Description**

Re-creates the internal random 80/20 train-test split without stratification.

**Usage**

```
RegenerateTrainTest(ExpCellNetObj)
```

**Arguments**

ExpCellNetObj     An ExpressionCellNet object.

**Value**

Updated ExpressionCellNet object with regenerated split.

**Examples**

```
data("example_data")
obj <- RegenerateTrainTest(example_data$expcellnetobj)
```

---

ShowHubGenes             *Show hub genes on the network*

---

**Description**

Highlights hub genes on the igraph network visualization.

**Usage**

```
ShowHubGenes(ExpCellNetObj)
```

**Arguments**

ExpCellNetObj     An ExpressionCellNet object after AnalyzeNetwork().

**Value**

Invisibly returns NULL (plot is produced).

**Examples**

```
data("example_data")
ShowHubGenes(example_data$expcellnetobj)
```

---

ShowNetwork

*Visualize the constructed network*

---

**Description**

Plots the current network stored in the ExpressionCellNet object.

**Usage**

```
ShowNetwork(ExpCellNetObj)
```

**Arguments**

ExpCellNetObj An ExpressionCellNet object containing a "Network" edge table.

**Value**

A plot (igraph/base) is produced as a side effect.

**Examples**

```
data("example_data")
ShowNetwork(example_data$expcellnetobj)
```

---

ShowPathway

*Visualize a pathway (highlighted subgraph)*

---

**Description**

Highlights the nodes/edges of a provided pathway on the full network visualization.

**Usage**

```
ShowPathway(ExpCellNetObj, Pathway)
```

**Arguments**

ExpCellNetObj An ExpressionCellNet object containing a constructed network.  
Pathway A character vector of genes representing a path (e.g., output of FindPathway()).

**Value**

A plot (igraph/base) is produced as a side effect.

**Examples**

```
data("example_data")  
p <- FindPathway(example_data$expcellnetobj, Start_gene="Gene1", End_gene="Gene11")  
ShowPathway(example_data$expcellnetobj, Pathway = p)
```

---

UMAPforGenes

*UMAP on network-derived features*

---

**Description**

Computes a 2D UMAP embedding of the network-restricted feature matrix returned by ExtractFeature() (after min-max scaling) and returns a ggplot visualization.

**Usage**

```
UMAPforGenes(ExpCellNetObj)
```

**Arguments**

ExpCellNetObj An ExpressionCellNet object containing "Network", "Generations", and a normalized count matrix.

**Value**

A ggplot object showing UMAP embedding of genes (nodes).

**Examples**

```
data("example_data")  
u <- UMAPforGenes(example_data$expcellnetobj)  
u
```

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