Package 'qountstat'

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Type Package

Title Statistical Analysis of Count Data and Quantal Data

Version 0.1.1

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Description Methods for statistical analysis of count data and quantal data.

For the analysis of count data an implementation of the Closure Principle Computational Approach Test (``CPCAT") is provided (Lehmann, R et al. (2016) <doi:10.1007/s00477-015-1079-4>), as well as an implementation of a ``Dunnett GLM" approach using a Quasi-Poisson regression (Hothorn, L, Kluxen, F (2020) <doi:10.1101/2020.01.15.907881>).
For the analysis of quantal data an implementation of the Closure Principle Fisher–Freeman–Halton test (``CPFISH") is provided (Lehmann, R et al. (2018) <doi:10.1007/s00477-017-1392-1>). P-values and no/lowest observed (adverse) effect concentration values are calculated.
All implemented methods include further functions to evaluate the power and the minimum detectable difference using a bootstrapping approach.

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Encoding UTF-8

LazyData true

Imports multcomp

Suggests knitr, rmarkdown, testthat

RoxygenNote 7.3.2

Depends R (>= 2.10)

NeedsCompilation no

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CP.hypotheses CP hypotheses

Description

Function to generate hypotheses for the Closure Principle concept using 0/1 contrast matrices

Usage

```
CP.hypotheses(n, treatment.names = NULL)
```

Arguments

n Number of groups

treatment.names

Optional vector with names of treatment groups

Value

Contrast matrix (all 0/1 combinations)

CPCAT

CPCAT

Description

When conducting statistical tests with multiple treatments, such as a control group and increasing concentrations of a test substance, ANOVA and parametric post-hoc tests (e.g. Dunnett's test) are commonly used. However, these tests require the assumptions of homogeneous variances and normally distributed data. For count data (e.g. counts of animals), these assumptions are typically violated, as the data are usually Poisson-distributed. Additionally, multiple testing using post-hoc tests can lead to alpha-inflation. To address these issues, CPCAT was proposed by Lehmann et al. (2016). CPCAT has two components. The first is the Closure Principle (CP) developed by Bretz et al. (2010), which aims to eliminate alpha-inflation. CP applies a stepwise approach to identify at which concentration effects begin to occur. The second part of CPCAT is the actual significance test, CAT (Computational Approach Test; introduced by Chang et al., 2010), which uses a test based on the Poisson distribution rather than a parametric test based on normal distribution assumptions. For details on the structure of the input data, please refer to the dataset 'Daphnia.counts' provided alongside this package.

Usage

```
CPCAT(
  groups,
  counts,
  control.name = NULL,
  bootstrap.runs = 10000,
  hampel.threshold = 5,
  use.fixed.random.seed = NULL,
  get.contrasts.and.p.values = FALSE,
  show.output = TRUE
)
```

Arguments

groups	Group vector	
counts	Vector with count data	
control.name	Character string with control group name (optional)	
bootstrap.runs	Number of bootstrap runs	
hampel.threshold		
	Threshold for Hampel identifier (measure for over-/underdispersion)	
use.fixed.random.seed		
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
get.contrasts.and.p.values		
	Get each row of the contrast matrices evaluated	
show.output	Show/hide output	

Value

R object with results and information from CPCAT calculations

References

Bretz, F.; Hothorn, T.; Westfall, P. (2010): Multiple comparisons using R. 1st Edition, Chapman and Hall/CRC, New York

Chang, C.-H.; Pal, N.; Lin, J.-J. (2010): A Note on Comparing Several Poisson Means. In: Commun. Stat. Simul. Comput., 2010, 39(8), p. 1605-1627, https://doi.org/10.1080/03610918.2010.508860

Lehmann, R.; Bachmann, J.; Maletzki, D.; Polleichtner, C.; Ratte, H.; Ratte, M. (2016): A new approach to overcome shortcomings with multiple testing of reproduction data in ecotoxicology. In: Stochastic Environmental Research and Risk Assessment, 2016, 30(3), p. 871-882, https://doi.org/10.1007/s00477-015-1079-4

Examples

Daphnia.counts # example data provided alongside the package

```
# Test CPCAT
CPCAT(groups = Daphnia.counts$Concentration,
    counts = Daphnia.counts$Number_Young,
    control.name = NULL,
    bootstrap.runs = 10000,
    use.fixed.random.seed = 123, #fixed seed for reproducible results
    get.contrasts.and.p.values = FALSE,
    show.output = TRUE)
```

CPCAT. bMDD

CPCAT bootstrap MDD (bMDD)

Description

The basic idea of the calculation of bootstrap MDD (bMDD) using the CPCAT approach is to shift the lambda parameter of Poisson distribution until there is a certain proportion of results significantly different from the control.

```
CPCAT.bMDD(
  groups,
  counts,
  control.name = NULL,
  alpha = 0.05,
  shift.step = -0.25,
  bootstrap.runs = 200,
  power = 0.8,
  max.iterations = 1000,
```

```
use.fixed.random.seed = NULL,
CPCAT.bootstrap.runs = 200,
show.progress = TRUE,
show.results = TRUE
```

)

	groups	Group vector
	counts	Vector with count data
	control.name	Character string with control group name (optional)
	alpha	Significance level
	shift.step	Step of shift (negative as a reduction is assumed)
	bootstrap.runs	Number of bootstrap runs
	power	Proportion of bootstrap.runs that return significant differences
	max.iterations	Max. number of iterations to not get stuck in the while loop
use.fixed.random.seed		
		Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.
CPCAT.bootstrap.runs		
		Bootstrap runs within CPCAT method
	show.progress	Show progress for each shift of lambda
	show.results	Show results

Value

Data frame with results from bMDD analysis

Examples

Daphnia.counts # example data provided alongside the package

```
# Test CPCAT bootstrap MDD
CPCAT.bMDD(groups = Daphnia.counts$Concentration,
    counts = Daphnia.counts$Number_Young,
    control.name = NULL,
    alpha = 0.05,
    shift.step = -1,# Caution: big step size for testing
    bootstrap.runs = 5,# Caution: low number of bootstrap runs for testing
    power = 0.8,
    max.iterations = 1000,
    use.fixed.random.seed = 123, #fixed seed for reproducible results
    CPCAT.bootstrap.runs = 10,
    show.progress = TRUE,
    show.results = TRUE)
```

CPCAT.Poisson.sub.test

CPCAT Poisson sub-test

Description

Helper function for CPCAT

Usage

CPCAT.Poisson.sub.test(dat, contrast, bootstrap.runs = 10000)

Arguments

dat	Data frame to be evaluated
contrast	Contrast matrix
bootstrap.runs	Number of bootstrap runs

Value

p-value for tested data and contrast

CPCAT.Poisson.test CPCAT Poisson test

Description

Helper function for CPCAT

Usage

```
CPCAT.Poisson.test(dat, contrastmatrix, bootstrap.runs = 10000)
```

Arguments

datData frame to be evaluatedcontrastmatrixContrast matrixbootstrap.runsNumber of bootstrap runs

Value

List of p-values for tested contrast matrices

CPCAT.power

CPCAT power

Description

The basic idea of CPCAT power calculations is to do parametric bootstrapping for each dose/concentration group and to evaluate the proportion of results significantly different from the control.

Usage

```
CPCAT.power(
  groups,
  counts,
  control.name = NULL,
  alpha = 0.05,
  bootstrap.runs = 200,
  use.fixed.random.seed = NULL,
  CPCAT.bootstrap.runs = 200,
  show.progress = TRUE,
  show.results = TRUE
)
```

Arguments

groups	Group vector	
counts	Vector with count data	
control.name	Character string with control group name (optional)	
alpha	Significance level	
bootstrap.runs	Number of bootstrap runs	
use.fixed.random.seed		
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
CPCAT.bootstrap.runs		
	Bootstrap runs within CPCAT method	
show.progress	Show progress for each shift of lambda	
show.results	Show results	

Value

Data frame with results from power analysis

Examples

Daphnia.counts # example data provided alongside the package

```
# Test CPCAT power
CPCAT.power(groups = Daphnia.counts$Concentration,
    counts = Daphnia.counts$Number_Young,
    control.name = NULL,
    alpha = 0.05,
    bootstrap.runs = 10,# Caution: low number of bootstrap runs for testing
    use.fixed.random.seed = 123, #fixed seed for reproducible results
    CPCAT.bootstrap.runs = 10,# Caution: low number of bootstrap runs for testing
    show.progress = TRUE,
    show.results = TRUE)
```

CPFISH

CPFISH

Description

For quantal data (e.g. survival data, '14 out of 20 animals died') e.g. in the form of a contingency table, CPFISH was proposed by Lehmann et al. (2018). Like CPCAT, CPFISH is based on the Closure Principle (CP), but instead of a bootstrapping approach, a Fisher test is performed for all sub-hypotheses to be analyzed. For details on the structure of the input table, please refer to the dataset 'CPFISH.contingency.table' provided alongside this package.

Usage

```
CPFISH(
   contingency.table,
   control.name = NULL,
   simulate.p.value = TRUE,
   use.fixed.random.seed = NULL,
   show.output = TRUE
)
```

Arguments

contingency.ta	ble	
	Matrix with observed data (e.g. survival counts, survival must be in first row)	
control.name	Character string with control group name (optional)	
simulate.p.value		
	Use simulated p-values in Fisher test or not	
use.fixed.rand	lom.seed	
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
show.output	Show/hide output	

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CPFISH.bMDD

Value

R object with results and information from CPFISH calculations

References

Lehmann, R.; Bachmann, J.; Karaoglan, B.; Lacker, J.; Polleichtner, C.; Ratte, H.; Ratte, M. (2018): An alternative approach to overcome shortcomings with multiple testing of binary data in ecotox-icology. In: Stochastic Environmental Research and Risk Assessment, 2018, 32(1), p. 213-222, https://doi.org/10.1007/s00477-017-1392-1

Examples

CPFISH.contingency.table # example data provided alongside the package

```
# Test CPFISH
CPFISH(contingency.table = CPFISH.contingency.table,
control.name = NULL,
simulate.p.value = TRUE,
use.fixed.random.seed = 123, #fixed seed for reproducible results
show.output = TRUE)
```

CPFISH.bMDD

```
CPFISH bootstrap MDD (bMDD)
```

Description

The basic idea of the calculation of bootstrap MDD (bMDD) using the CPCAT approach is to shift the probability of binomial distribution until there is a certain proportion of results significantly different from the control.

```
CPFISH.bMDD(
    contingency.table,
    control.name = NULL,
    alpha = 0.05,
    shift.step = -0.01,
    bootstrap.runs = 200,
    power = 0.8,
    max.iterations = 1000,
    simulate.p.value = TRUE,
    use.fixed.random.seed = NULL,
    show.progress = TRUE,
    show.results = TRUE
)
```

contingency.table		
	Matrix with observed data (e.g. survival counts, survival must be in first row)	
control.name	Character string with control group name (optional)	
alpha	Significance level	
shift.step	Step of shift (negative as a reduction is assumed)	
bootstrap.runs	Number of bootstrap runs (draw Poisson data n times)	
power	Proportion of bootstrap.runs that return significant differences	
max.iterations	Max. number of iterations to not get stuck in the while loop	
simulate.p.value		
	Use simulated p-values in Fisher test or not	
use.fixed.random.seed		
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
show.progress	Show progress for each shift of the probability	
show.results	Show results	

Value

Data frame with results from bMDD analysis

Examples

CPFISH.contingency.table # example data provided alongside the package

```
# Test CPFISH bootstrap MDD
CPFISH.bMDD(contingency.table = CPFISH.contingency.table,
    control.name = NULL,
    alpha = 0.05,
    shift.step = -0.1,# Caution: big step size for testing
    bootstrap.runs = 10,# Caution: low number of bootstrap runs for testing
    power = 0.8,
    max.iterations = 1000,
    simulate.p.value = TRUE,
    use.fixed.random.seed = 123, #fixed seed for reproducible results
    show.progress = TRUE,
    show.results = TRUE)
```

CPFISH.contingency.table CPFISH.contingency.table is a dataset to showcase analyses with CP-FISH

Description

CPFISH.contingency.table is a dataset to showcase analyses with CPFISH

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CPFISH.power

Usage

CPFISH.contingency.table

Format

An object of class matrix (inherits from array) with 2 rows and 3 columns.

CPFISH power

Source

The CPFISH data was artificially created.

Examples

data(CPFISH.contingency.table)

CPFISH.power

Description

The basic idea of CPFISH power calculations is to do parametric bootstrapping for each dose/concentration group and to evaluate the proportion of results significantly different from the control.

Usage

```
CPFISH.power(
   contingency.table,
   control.name = NULL,
   alpha = 0.05,
   bootstrap.runs = 200,
   simulate.p.value = TRUE,
   use.fixed.random.seed = NULL,
   show.progress = TRUE,
   show.results = TRUE
)
```

Arguments

contingency.table

	Matrix with observed data (e.g. survival counts, survival must be in first row)
control.name	Character string with control group name (optional)
alpha	Significance level
bootstrap.runs	Number of bootstrap runs (draw Poisson data n times)
simulate.p.value	
	Use simulated p-values in Fisher test or not

Daphnia.counts

use.fixed.random.seed Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set. show.progress Show progress for each shift of the probability show.results Show results

Value

Data frame with results from power analysis

Examples

CPFISH.contingency.table # example data provided alongside the package

```
# Test CPFISH power
CPFISH.power(contingency.table = CPFISH.contingency.table,
control.name = NULL,
alpha = 0.05,
bootstrap.runs = 10,# Caution: low number of bootstrap runs for testing
simulate.p.value = TRUE,
use.fixed.random.seed = 123, #fixed seed for reproducible results
show.progress = TRUE,
show.results = TRUE)
```

Daphnia.counts	Daphnia.counts is a dataset to showcase analyses with CPCAT and
	Dunnett.GLM

Description

Daphnia.counts is a dataset to showcase analyses with CPCAT and Dunnett.GLM

Usage

Daphnia.counts

Format

An object of class data. frame with 60 rows and 3 columns.

Source

The daphnia data was taken from Hothorn and Kluxen (2020).

Examples

data(Daphnia.counts)

Dunnett.GLM

Dunnett.GLM

Description

When conducting statistical tests with multiple treatments, such as a control group and increasing concentrations of a test substance, ANOVA and parametric post-hoc tests (e.g. Dunnett's test) are commonly used. However, these tests require the assumptions of homogeneous variances and normally distributed data. For count data (e.g. counts of animals), these assumptions are typically violated, as the data are usually Poisson-distributed. The Dunnett.GLM function is based on a GLM followed by a Dunnett test to the model estimates. It was implemented to serve as an alternative approach to CPCAT while using a Quasi-Poisson regression. The basic approach from Hothorn and Kluxen (2020) was adjusted to overcome methodological issues (see description of 'zero.treatment.action parameter'). For details on the structure of the input data, please refer to the dataset 'Daphnia.counts' provided alongside this package.

Usage

```
Dunnett.GLM(
  groups,
  counts,
  control.name = NULL,
  zero.treatment.action = "identity.link",
  show.output = TRUE
)
```

Arguments

groups	Group vector
counts	Vector with count data
control.name	Character string with control group name (optional)
zero.treatment.	action
	Method for dealing with treatments only containing zeros (use either "iden- tity.link" or " $log(x+1)$ "). The method is only used if the data set contains dose/concentration groups that exclusively contain zero values (since the basic method provides for a logarithmic transformation of the data averages, it would lead to incorrect results). To deal with this methodological shortcoming, two options were im- plemented. The 'identity.link' option: the 'identity' link is used in the GLM instead of the 'log' link, i.e. the data are no longer transformed. The 'log(x+1)' option: The 'log' link is retained and 1 is added to each count value at the start of the procedure so that the subsequent log-transformation can be carried out without any problems. Note that both options may slightly distort the results.
show.output	Show/hide output

Value

R object with results and information from Dunnett.GLM calculations

References

Hothorn, L.; Kluxen, F. (2020): Statistical analysis of no observed effect concentrations or levels in eco-toxicological assays with overdispersed count endpoints. In: bioRxiv, 2020, https://doi.org/10.1101/2020.01.15.907881

Examples

Daphnia.counts # example data provided alongside the package

```
# Test Dunnett.GLM with 'identity.link' option
Dunnett.GLM(groups = Daphnia.counts$Concentration,
    counts = Daphnia.counts$Number_Young,
    control.name = NULL,
    zero.treatment.action = "identity.link",
    show.output = TRUE)
# Test Dunnett.GLM with 'log(x+1)' option
Dunnett.GLM(groups = Daphnia.counts$Concentration,
    counts = Daphnia.counts$Number_Young,
    control.name = NULL,
    zero.treatment.action = "log(x+1)",
    show.output = TRUE)
```

Dunnett.GLM.bMDD Dunnett.GLM bootstrap MDD (bMDD)

Description

The basic idea of the calculation of bootstrap MDD (bMDD) using the Dunnett.GLM approach is to shift the lambda parameter of Poisson distribution until there is a certain proportion of results significantly different from the control.

```
Dunnett.GLM.bMDD(
  groups,
  counts,
  control.name = NULL,
  alpha = 0.05,
  shift.step = -0.25,
  bootstrap.runs = 200,
  power = 0.8,
  max.iterations = 1000,
  use.fixed.random.seed = NULL,
  Dunnett.GLM.zero.treatment.action = "log(x+1)",
  show.progress = TRUE,
  show.results = TRUE
)
```

groups	Group vector	
counts	Vector with count data	
control.name	Character string with control group name (optional)	
alpha	Significance level	
shift.step	Step of shift (negative as a reduction is assumed)	
bootstrap.runs	Number of bootstrap runs	
power	Proportion of bootstrap.runs that return significant differences	
max.iterations	Max. number of iterations to not get stuck in the while loop	
use.fixed.random.seed		
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
Dunnett.GLM.zero.treatment.action		
	Dunnett.GLM method to be used for treatments only containing zeros	
show.progress	Show progress for each shift of lambda	
show.results	Show results	

Value

Data frame with results from bMDD analysis

Examples

Daphnia.counts # example data provided alongside the package

```
# Test Dunnett.GLM bootstrap MDD
Dunnett.GLM.bMDD(groups = Daphnia.counts$Concentration,
counts = Daphnia.counts$Number_Young,
control.name = NULL,
alpha = 0.05,
shift.step = -1,# Caution: big step size for testing
bootstrap.runs = 5,# Caution: low number of bootstrap runs for testing
power = 0.8,
max.iterations = 1000,
use.fixed.random.seed = 123, #fixed seed for reproducible results
Dunnett.GLM.zero.treatment.action = "log(x+1)",
show.progress = TRUE,
show.results = TRUE)
```

Dunnett.GLM.power Dunnett.GLM power

Description

The basic idea of Dunnett.GLM power calculations is to do parametric bootstrapping for each dose/concentration group and to evaluate the proportion of results significantly different from the control.

Usage

```
Dunnett.GLM.power(
  groups,
  counts,
  control.name = NULL,
  alpha = 0.05,
  bootstrap.runs = 200,
  use.fixed.random.seed = NULL,
  Dunnett.GLM.zero.treatment.action = "log(x+1)",
  show.progress = TRUE,
  show.results = TRUE
)
```

Arguments

groups	Group vector	
counts	Vector with count data	
control.name	Character string with control group name (optional)	
alpha	Significance level	
bootstrap.runs	Number of bootstrap runs	
use.fixed.random.seed		
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
Dunnett.GLM.zero.treatment.action		
	GLM.Dunnett method to be used for treatments only containing zeros	
show.progress	Show progress for each shift of lambda	
show.results	Show results	

Value

Data frame with results from power analysis

Multi.group.test.bMDD

Examples

Daphnia.counts # example data provided alongside the package

```
# Test Dunnett.GLM power
Dunnett.GLM.power(groups = Daphnia.counts$Concentration,
counts = Daphnia.counts$Number_Young,
control.name = NULL,
alpha = 0.05,
bootstrap.runs = 10,# Caution: low number of bootstrap runs for testing
use.fixed.random.seed = 123, #fixed seed for reproducible results
Dunnett.GLM.zero.treatment.action = "log(x+1)",
show.progress = TRUE,
show.results = TRUE)
```

Multi.group.test.bMDD Multi-group test bMDD

Description

Idea: shift lambda of Poisson distribution until there is a certain proportion of significant results

Usage

```
Multi.group.test.bMDD(
  groups,
  counts,
  control.name = NULL,
  alpha = 0.05,
  shift.step = -0.25,
  bootstrap.runs = 200,
  power = 0.8,
 max.iterations = 1000,
  use.fixed.random.seed = NULL,
  CPCAT.bootstrap.runs = 200,
  Dunnett.GLM.zero.treatment.action = "log(x+1)",
  show.progress = TRUE,
  show.results = TRUE,
  get.effect.and.power = FALSE,
  use.CMP.distribution = FALSE,
  CMP.dispersion.factor = 1,
  test = "CPCAT"
)
```

Arguments

groups	Group vector
counts	Vector with count data

control.name	Character string with control group name (optional)
alpha	Significance level
shift.step	Step of shift (negative as a reduction is assumed)
bootstrap.runs	Number of bootstrap runs
power	Proportion of bootstrap.runs that return significant differences
max.iterations	Max. number of iterations to not get stuck in the while loop
use.fixed.rando	om.seed
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.
CPCAT.bootstrap	o. runs
	Bootstrap runs within CPCAT method
Dunnett.GLM.zer	ro.treatment.action
	Dunnett.GLM method to be used for treatments only containing zeros
show.progress	Show progress for each shift of lambda
<pre>show.results get.effect.and.</pre>	
	Return effect size (percent of control) and power for each step (only for last treatment)
use.CMP.distrib	pution
	Use Conway-Maxwell-Poisson distribution for sampling
CMP.dispersion.	factor
	Dispersion parameter phi has to be sqrt(factor) to scale the variance by this factor
test	Either "CPCAT" or "GLM.Dunnett"

Value

Data frame with results from bMDD analysis

Multi.group.test.power

Multi-group test power

Description

Idea: Do parametric bootstrapping for each group and evaluate the proportion of significant results.

```
Multi.group.test.power(
  groups,
  counts,
  control.name = NULL,
  alpha = 0.05,
  bootstrap.runs = 200,
  use.fixed.random.seed = NULL,
```

```
CPCAT.bootstrap.runs = 200,
Dunnett.GLM.zero.treatment.action = "log(x+1)",
show.progress = TRUE,
show.results = TRUE,
test = "CPCAT"
)
```

groups	Group vector	
counts	Vector with count data	
control.name	Character string with control group name (optional)	
alpha	Significance level	
bootstrap.runs	Number of bootstrap runs	
use.fixed.random.seed		
	Use fixed seed, e.g. 123, for reproducible results. If NULL no seed is set.	
CPCAT.bootstrap.runs		
	Bootstrap runs within CPCAT method	
Dunnett.GLM.zero.treatment.action		
	GLM.Dunnett method to be used for treatments only containing zeros	
show.progress	Show progress for each shift of lambda	
show.results	Show results	
test	Either "CPCAT" or "GLM.Dunnett"	

Value

Data frame with results from power analysis

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