

Package ‘informedSen’

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Type Package

Title Sensitivity Analysis Informed by a Test for Bias

Version 1.0.7

Author Paul R Rosenbaum

Maintainer Paul R Rosenbaum <rosenbaum@wharton.upenn.edu>

Description After testing for biased treatment assignment in an observational study using an unaffected outcome, the sensitivity analysis is constrained to be compatible with that test. The package uses the optimization software gurobi obtainable from <<https://www.gurobi.com/>>, together with its associated R package, also called gurobi; see: <https://www.gurobi.com/documentation/7.0/refman/installing_the_r_package.html>. The method is a substantial computational and practical enhancement of a concept introduced in Rosenbaum (1992) Detecting bias with confidence in observational studies *Biometrika*, 79(2), 367-374 <[doi:10.1093/biomet/79.2.367](https://doi.org/10.1093/biomet/79.2.367)>.

License GPL-2

Encoding UTF-8

LazyData true

Imports sensitivitymult, stats

Enhances gurobi

Depends R (>= 3.5.0)

NeedsCompilation no

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informedSen-package *Sensitivity Analysis Informed by a Test for Bias*

Description

After testing for biased treatment assignment in an observational study using an unaffected outcome, the sensitivity analysis is constrained to be compatible with that test. The package uses the optimization software `gurobi` obtainable from <https://www.gurobi.com/>, together with its associated R package, also called `gurobi`; see: https://www.gurobi.com/documentation/7.0/refman/installing_the_r_package.html. The method is a substantial computational and practical enhancement of a concept introduced in Rosenbaum (1992) Detecting bias with confidence in observational studies *Biometrika*, 79(2), 367-374 <doi:10.1093/biomet/79.2.367>.

Details

The DESCRIPTION file:

```
Package:      informedSen
Type:         Package
Title:        Sensitivity Analysis Informed by a Test for Bias
Version:      1.0.7
Author:       Paul R Rosenbaum
Maintainer:   Paul R Rosenbaum <rosenbaum@wharton.upenn.edu>
Description:  After testing for biased treatment assignment in an observational study using an unaffected outcome, the sensit
License:      GPL-2
Encoding:     UTF-8
LazyData:    true
Imports:      sensitivitymult, stats
Enhances:    gurobi
Depends:      R (>= 3.5.0)
```

Index of help topics:

```
HDL           Light Daily Alcohol and HDL Cholesterol Levels
informedSen-package  Sensitivity Analysis Informed by a Test for
                    Bias
informedsen    Sensitivity Analysis Informed by a Test for
                    Unmeasured Bias
senmscores     Computes M-scores for M-tests.
```

The package performs a sensitivity analysis within a confidence set provided by a test for unmeasured bias. The method is a substantial computational and practical enhancement of a concept introduced in Rosenbaum (1992) Detecting bias with confidence in observational studies. *Biometrika*, 79(2), 367-374. <doi:10.1093/biomet/79.2.367> The main function in the package is `informedsen`. The package uses the optimization software `gurobi` obtainable from <https://www.gurobi.com/>, together with its associated R package, also called `gurobi`; see: https://www.gurobi.com/documentation/7.0/refman/installing_

See the example in `informedsen` for discussion about obtaining `gurobi` and its associated local R-package.

Author(s)

Paul R Rosenbaum

Maintainer: Paul R Rosenbaum <rosenbaum@wharton.upenn.edu>

References

- Rosenbaum, P. R. (1984). From association to causation in observational studies: The role of tests of strongly ignorable treatment assignment. *Journal of the American Statistical Association* 79, 41-48. <doi:10.1080/01621459.1984.10477060>
- Rosenbaum, P. R. (1989a). On permutation tests for hidden biases in observational studies. *The Annals of Statistics* 17, 643-653. <doi:10.1214/aos/1176347131>
- Rosenbaum, P. R. (1989b). The role of known effects in observational studies. *Biometrics* 45, 557-569. <doi:10.2307/2531497>
- Rosenbaum, P. R. (1992). Detecting bias with confidence in observational studies. *Biometrika*, 79(2), 367-374. <doi:10.1093/biomet/79.2.367>
- Rosenbaum, P. R. (2007) Sensitivity analysis for m-estimates, tests and confidence intervals in matched observational studies. *Biometrics*, 2007, 63, 456-464. <doi:10.1111/j.1541-0420.2006.00717.x>
- Rosenbaum, P. R. (2021). Sensitivity analyses informed by tests for bias in observational studies. Manuscript. This manuscript describes and illustrates the new computational tools that make feasible the method in Rosenbaum (1992). The example in the package is the example in this manuscript.

Examples

```
## Not run:
# To run these examples, you MUST have gurobi installed.
# The makers of gurobi provide free access to academics.
# Additionally, you must install the local R package gurobi
# that is provided by installing gurobi.
# The examples are from Rosenbaum (2021)
data(HDL)
shdl<-senmscores(HDL$hdl,HDL$z,HDL$mset)
smmerc<-senmscores(HDL$mmercury,HDL$z,HDL$mset)
sc<-cbind(shdl,smmerc)
informedsen(3.5, sc, HDL$z, HDL$mset, alpha = 0.05)
informedsen(3.4, sc, HDL$z, HDL$mset, alpha = c(0.04,0.01))

## End(Not run)
```

HDL

Light Daily Alcohol and HDL Cholesterol Levels

Description

An observational study of light daily alcohol consumption (1-3 drinks per day) versus little or no alcohol, and its possible effects on HDL cholesterol levels. The level of methylmercury is viewed as an unaffected outcome and used to test for biased treatment assignment. Data is from NHANES 2013/2014 and 2015/2016.

Usage

```
data("HDL")
```

Format

A data frame with 800 observations on the following 9 variables.

SEQN NHANES sequence number

nh Either 1314 for NHANES 2013/2014 or 1516 for NHANES 2015/2016

z Treatment indicator, z=1 for light daily alcohol or z=0 for little or no alcohol.

mset Matched set indicator, 1 to 200, for 200 matched sets, each containing one treated and three controls.

age Age in years. Used in matching.

female 1 for female, 0 for male. Used in matching.

education NHANES 1-5 education scale. 1 is <9th grade, 3 is high school, 5 is at least a BA degree.

hdl HDL cholesterol level

mmmercury Methylmercury level

Source

US National Health and Nutrition Examination Survey. Publicly available on-line.

References

LoConte, N. K., Brewster, A. M., Kaur, J. S., Merrill, J. K., and Alberg, A. J. (2018). Alcohol and cancer: a statement of the American Society of Clinical Oncology. *Journal of Clinical Oncology* 36, 83-93.

Rosenbaum, P. R. (2021). Sensitivity analyses informed by tests for bias in observational studies. Manuscript. See its data appendix.

Suh, I., Shaten, B. J., Cutler, J. A., and Kuller, L. H. (1992). Alcohol use and mortality from coronary heart disease: the role of high-density lipoprotein cholesterol. *Annals of Internal Medicine* 116, 881-887.

Examples

```
data(HDL)
boxplot(HDL$age~HDL$z) # ages are similar
boxplot(HDL$hdl~HDL$z) # hdl is higher
boxplot(HDL$mmercury~HDL$z,log="y") # methylmercury is higher
```

informedsen

Sensitivity Analysis Informed by a Test for Unmeasured Bias

Description

The function does a sensitivity analysis for one outcome informed or constrained by the results of a test for unmeasured bias based on another outcome known to be unaffected by the treatment. The package uses gurobi to solve a quadratically constrained quadratic program. To use the package, the gurobi solver must be installed. See the discussion about installing gurobi in the example below.

Usage

```
informedsen(gamma, sc, z, mset, alpha = 0.05)
```

Arguments

gamma	The sensitivity parameter. A number greater than or equal to 1.
sc	A matrix with N rows and at least two columns. The first column is the primary outcome, typically after scoring using senmscores. The remaining columns are unaffected outcomes used to test for bias, typically after scoring using senmscores.
z	A vector of length N whose N coordinates are 1 for treated, 0 for control.
mset	A vector of length N indicating the matched set. Each matched set contains one treated individual and the same fixed number of controls.
alpha	A vector with length equal to the number of columns of sc. The jth coordinate of alpha is the level of the test applied to the jth column of sc. If alpha is a scalar, it is repeated for every column of sc.

Value

result	Text indicating whether or not the test for bias rejects all biases of magnitude Gamma or less. If yes, then the conclusion is that you must increase Gamma to continue. If no, then the test on the primary outcome is conducted inside the confidence set defined by a test for bias. The text begins after gurobi prints its standard output for the underlying optimization problem.
optimization.problem	Reiterates the result above, where the word yes means the optimization problem is infeasible, and the word no means it is feasible. See the conclusion for a scientific interpretation of this aspect of the output.

conclusion	Text indicating the result of the test for effect on the primary outcome.
deviates	A vector of standardized deviates that might be compared with the standard Normal distribution. There is one deviate for each column of <code>sc</code> . If <code>sc</code> has column names, then the column names label the deviates. The deviates are computed at the treatment assignment probabilities, <code>theta</code> , that solve the constrained optimization problem.
alphas	A vector of two-sided levels used for the deviates, together with their total. The total is relevant if the Bonferroni inequality is used to ensure joint level of all the tests. The absolute deviates might be compared with <code>qnorm(1-alphas/2)</code> for a two-sided test.

Note

When `gurobi` is called, it produces extensive output. The output for `informedsen` appears at the end, after `gurobi` has produced its output. Most users will wish to skip to the end, for the output from `informedsen`, returning to the `gurobi` output only if needed.

`informedsen` checks that your input has the required form, and it will stop if there is a problem with your input. For instance, `informedsen` will stop if you supply a value of `gamma` that is less than one.

Author(s)

Paul R. Rosenbaum

References

- Berger, R. L. and Boos, D. D. (1994). P-values maximized over a confidence set for the nuisance parameter. *Journal of the American Statistical Association*, 89, 1012-1016. <doi:10.1080/01621459.1994.10476836>
- Rosenbaum, P. R. (1984). From association to causation in observational studies: The role of tests of strongly ignorable treatment assignment. *Journal of the American Statistical Association* 79, 41-48. <doi:10.1080/01621459.1984.10477060>
- Rosenbaum, P. R. (1989a). On permutation tests for hidden biases in observational studies. *The Annals of Statistics* 17, 643-653. <doi:10.1214/aos/1176347131>
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- Rosenbaum, P. R. (1992). Detecting bias with confidence in observational studies. *Biometrika*, 79(2), 367-374. <doi:10.1093/biomet/79.2.367>
- Rosenbaum, P. R. (2007) Sensitivity analysis for m-estimates, tests and confidence intervals in matched observational studies. *Biometrics*, 2007, 63, 456-464. <doi:10.1111/j.1541-0420.2006.00717.x>
- Rosenbaum, P. R. (2021). Sensitivity analyses informed by tests for bias in observational studies. Manuscript.

Examples

```
## Not run:
# To run these examples, you MUST have gurobi installed.
# gurobi is available for free to academic faculty
# Search for the gurobi web page, and click the menu for Academia
```

```

# Search for "gurobi and R" to find gurobi's local R package
# connecting gurobi and R. You must install both gurobi
# and its local R package to run informedsen.
#
# The examples are from Rosenbaum (2021)
#
# gurobi generates output before the output from informedsen
# appears. In a first use, you might skip to the output
# from informedsen, which begins with text labeled result.
#
data(HDL)
shdl<-senmscores(HDL$hdl,HDL$z,HDL$mset)
smmerc<-senmscores(HDL$mercury,HDL$z,HDL$mset)
sc<-cbind(shdl,smmerc)

# A test within the confidence set
informedsen(3.5, sc, HDL$z, HDL$mset, alpha = 0.05)

# A test within the confidence set using
# the method of Berger and Boos (1994)
informedsen(3.4, sc, HDL$z, HDL$mset, alpha = c(0.04,0.01))

# An example in which the confidence set is empty
informedsen(1.25, sc, HDL$z, HDL$mset, alpha = .05)

## End(Not run)

```

senmscores

Computes M-scores for M-tests.

Description

Computes M-scores for an M-test with one outcome in 1-to-k matched sets, for fixed $k \geq 1$. For the one-sample problem or matched pairs, Maritz (1979) proposed robust tests and confidence intervals based on Huber's (1981) M-estimates. These tests are extended to matching with several controls in Rosenbaum (2007).

Usage

```
senmscores(y, z, mset, inner = 0, trim = 3, lambda = 1/2)
```

Arguments

y	A vector of length N for one outcome.
z	A vector whose N coordinates are 1 for treated, 0 for control.
mset	A vector of length N giving the matched set.
inner	See trim.

trim	The two values, inner and trim, define the M-statistic's psi-function. The psi-function is an odd function, $\psi(y) = -\psi(-y)$, so it suffices to define it for non-negative y . For nonnegative y , $\psi(y)$ equals 0 for y between 0 and inner, rises linearly from 0 to 1 for y between inner and trim, and equals 1 for y greater than trim. There are two requirements: inner must be nonnegative, and trim must be larger than inner.
lambda	A number strictly between 0 and 1. The M-scores are $\psi(y/s)$ where s is the lambda quantile of the within-set absolute pair differences.

Details

The choice of psi-function to increase insensitivity to unmeasured bias is discussed in Rosenbaum (2013), where the parameter inner is proposed.

Value

A vector of length N containing the M-scores.

Note

The function is essentially a wrapper for the mscorev function in the sensitivitymult package. It is easier to use senmscores when using the informedSen package.

Author(s)

Paul R. Rosenbaum

References

- Huber, P. (1981). Robust Statistics. NY: Wiley.
- Maritz, J. S. (1979). A note on exact robust confidence intervals for location. *Biometrika* 66, 163-170.
- Rosenbaum, P. R. (2007) Sensitivity analysis for m-estimates, tests and confidence intervals in matched observational studies. *Biometrics*, 2007, 63, 456-464. <doi:10.1111/j.1541-0420.2006.00717.x>
- Rosenbaum, P. R. (2013). Impact of multiple matched controls on design sensitivity in observational studies. *Biometrics* 69 118-127. (Introduces inner trimming.) <doi:10.1111/j.1541-0420.2012.01821.x>
- Rosenbaum, P. R. (2015). Two R packages for sensitivity analysis in observational studies. *Observational Studies*, v. 1. (Free on-line.)

Examples

```
data(HDL)
shd1<-senmscores(HDL$hd1, HDL$z, HDL$mset)
plot(HDL$hd1, shd1)
```


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