

Package ‘fbstHWE’

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Type Package

Title FBST for Hardy-Weinberg Equilibrium Test

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Description

This package contains a straightforward method for the multiallelic Hardy-Weinberg Equilibrium (HWE) test. The core theory for the proposed method is given by the Full Bayesian Significance Test (FBST), an intuitive Bayesian approach which does not assign positive probabilities to zero measure sets when testing sharp hypotheses.

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fbstHWE-package *FBST for Hardy-Weinberg Equilibrium Test*

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Details

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Function: <fbstHWE>

Author(s)

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References

1. M.S.Lauretto, F.Nakano, S.R.Faria Jr, C.A.B.Pereira, J.M.Stern. A straightforward multiallelic significance test for the Hardy-Weinberg equilibrium law. *Genetics and Molecular Biology* 32(3), 619-625, 2009.
- 2.Pereira, C.A.B., Stern, J.M. Evidence and Credibility: Full Bayesian Significance Test for Precise Hypotheses. *Entropy* 1, 99-110, 1999.
- 3.C.A.B.Pereira, J.M.Stern, S. Wechsler. Can a significance test be genuinely Bayesian? *Bayesian Analysis* 3(1), 79-100, 2008.

fbstHWE

Full Bayesian Significance Test (FBST) for the Hardy-Weinberg Equilibrium

Description

Performs the test for Hardy-Weinberg Equilibrium (HWE) in a locus with several alleles A_1, A_2, \dots, A_k , $k \geq 2$, and returns the hypothesis evidence $Ev(H)$.

Usage

```
fbstHWE(X, delta = 0.005, simulate.p.value = FALSE, B = 200, verbose = FALSE)
```

Arguments

X vector of genotype counts in a lower triangular form:

$$X = c(\quad \quad \quad x_{11}, \\ x_{21}, x_{22}, \\ x_{31}, x_{32}, x_{33}, \\ \dots \\ x_{k1}, x_{k2}, \dots, x_{kk} \\ \text{where } k \text{ is the number of alleles}$$

X may also contain a square matrix with genotype counts.

delta	precision parameter $0 < \delta < 1$, it corresponds to the maximum length for the 95% confidence interval $ev(H)$
simulate.p.value	boolean parameter indicating whether to compute the Empirical p-value by Monte Carlo simulation. If FALSE, only asymptotic p-value is returned.
B	number of samples to be drawn for the Empirical p-value computation
verbose	verbose method. If TRUE, prints the estimated evidences and respective preci-

sions for simulated samples during the computation of empirical p-value computation

Details

Given k alleles A_1, A_2, \dots, A_k , $k \geq 2$, in a locus, and their genotype counts in lower triangular form $X = (x_{ij})$, $1 \leq j \leq i \leq k$, this function computes the evidence for the Hardy-Weinberg Equilibrium (HWE).

The parameter of interest, denoted by θ_{ij} , is the population relative frequencies of genotypes $A_i A_j$ ($1 \leq j \leq i \leq k$). Assuming a multinomial model for x and a Dirichlet(1,1,...,1) prior for θ , the posterior p.d.f is $f(\theta | x) = \text{Dirichlet}(x_{11}+1, x_{12}+1, \dots, x_{kk}+1)$.

Parameter space: $\text{THETA} = \theta = (\theta_{ij})$, $1 \leq j \leq i \leq k \mid 0 \leq \theta_{ij} \leq 1, \sum(\theta) = 1$

Parameter space under HWE: $H = \theta$ in $\text{THETA} \mid$ there exists a vector $p = (p_1, p_2, \dots, p_k)$, $0 \leq p_i \leq 1, \sum(p) = 1$, such that: $\theta_{ij} = (p_i)^2$ if $i=j$; $\theta_{ij} = 2 * p_i * p_j$ if $i < j$

The evidence measure against the hypothesis is computed in two steps: 1) compute $f_0 = \max_H f(\theta | x)$ where $f(\theta | x)$ denotes the posterior probability density function.

2) compute the integral $evb = \frac{\int_H f(\theta | x) d\theta}{\int_{\text{THETA}} f(\theta | x) d\theta}$ where $T = \theta$ in $\text{THETA} \mid f(\theta | x) > f_0$

The evidence measure supporting the hypothesis is: $evid = 1 - evb$

Value

<code>evid</code>	evidence of HWE hypothesis
<code>prec</code>	95% confidence interval length for <code>evid</code>
<code>p.value</code>	estimated asymptotic p-value
<code>emp.p.value</code>	empirical p-value (=NA if <code>simulate.p.value=FALSE</code>)

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References

1. M.S.Lauretto, F.Nakano, S.R.Faria Jr, C.A.B.Pereira, J.M.Stern. A straightforward multiallelic significance test for the Hardy-Weinberg equilibrium law. *Genetics and Molecular Biology* 32(3), 619-625, 2009.
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Examples

```
#####
# Example 1:
# Extracted from Louis and Dempster (1987),
# An exact test for Hardy-Weinberg and multiple alleles.
# Biometrics 43:805-811
#
X = c(
```

```
0,  
3, 1,  
5, 18, 1,  
3, 7, 5, 2)
```

```
EvFBST1 = fbstHWE(X, simulate.p.value=TRUE, B=100, verbose=TRUE)  
print(c(EvFBST1$evid, EvFBST1$delta, EvFBST1$p.value, EvFBST1$emp.p.value))
```