Package 'plasmut'

December 5, 2025

Type Package

Title Stratifying mutations observed in cell-free DNA and white blood
cells as germline, hematopoietic, or somatic

Version 1.9.0

Description A Bayesian method for quantifying the liklihood that a given plasma muta-
tion arises from clonal hematopoesis or the underlying tumor. It requires sequenc-
ing data of the mutation in plasma and white blood cells with the number of distinct and mu-
tant reads in both tissues. We implement a Monte Carlo importance sampling method to as-
sess the likelihood that a mutation arises from the tumor relative to non-tumor origin.

License Artistic-2.0

Encoding UTF-8

biocViews Bayesian, SomaticMutation, GermlineMutation, Sequencing

RoxygenNote 7.2.3

Imports tibble, stats, dplyr

Depends R (>= 4.3.0)

Suggests knitr, rmarkdown, tidyverse, ggrepel, magrittr, qpdf,
BiocStyle, biocViews, testthat (>= 3.0.0)

VignetteBuilder knitr

Config/testthat/edition 3

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Author Adith Arun [aut, cre],
Robert Scharpf [aut]

Maintainer Adith Arun <adith.3.arun@gmail.com>

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crcseq   *This data is an example dataset to show how to use the package*

Description

A cohort of metastatic colorectal cancer patients whose plasma and buffy coat were sequenced as part of the CAIRO5 trial. The cohort and analyses are described here: https://pubmed.ncbi.nlm.nih.gov/36534496/

Value

An example DNA sequencing dataset of matched plasma and wbc colorectal cancer samples crcseq

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importance_sampler   *Importance sampler to estimate marginal likelihoods and Bayes factors*

Description

Importance sampler to estimate marginal likelihoods and Bayes factors

Usage

```
importance_sampler(dat, params, save_montecarlo = TRUE)
```

Arguments

dat	data frame with observed mutant and total counts and the analyte (plasma or buffy coat) it was taken from and the identifiers on what the mutation is (e.g., KRASG12C) and pt id
params	list with ctc, ctdna and chip a and b beta parameters reflect beliefs on what fraction of fragments belong to each class; montecarlo.samples being the number of MC samples; prior weight is the prior.weight reflects how much importance sampling to implement, closer to zero means more importance density considered
save_montecarlo	
	save more indepth monte carlo results

Value

implement importance sampling for a data set to assess probability of tumor derived mutations from sequencing results

Examples

```
param.list <- list(ctc=list(a=1, b=9999),
                    ctdna=list(a=1, b=9),
                    chip=list(a=1, b=9),
                    montecarlo.samples=50e3,
                    prior.weight=0.1)
dat <- data.frame(y=c(4, 1),
              n=c(1000, 1000),
              analyte=c("plasma", "buffy coat"),
              mutation="mutA",
              sample_id="id1")
importance_sampler(dat, param.list)
```

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model_w	*Estimate the marginal likelihood that mutations in buffy coat and cfDNA reflect CH or correspond to germline mutations. If germline, the allele frequency should be 50 percent. The prior should be diffuse enough to handle CHIP mutations which are potentially way less than 50 percent*

Description

Estimate the marginal likelihood that mutations in buffy coat and cfDNA reflect CH or correspond to germline mutations. If germline, the allele frequency should be 50 percent. The prior should be diffuse enough to handle CHIP mutations which are potentially way less than 50 percent

Usage

```
model_w(dat, params)
```

Arguments

dat	tibble containing vectors yand n. y and n should be named
params	a list with named elements that must include the following a which is the prior expectation for number of CH or germline variants observed in the sequencing data b which is the prior expectation for number of fragments reflecting CH or germline

Value

list of samples, probability densities, and likelihood for non-tumor assumption

Examples

```
param.list <- list(ctc=list(a=1, b=9999),
                    ctdna=list(a=1, b=9),
                    chip=list(a=1, b=9),
                    montecarlo.samples=50e3,
                    prior.weight=0.1)
dat <- data.frame(y=c(4, 1),
              n=c(1000, 1000),
              analyte=c("plasma", "buffy coat"),
              mutation="mutA",
              sample_id="id1")
model_w(dat, param.list)
```

---

plasma_somatic	*Estimate the marginal likelihood that variants identified in cell-free DNA are derived from tumor cells (ctDNA-derived)*

Description

Estimate the marginal likelihood that variants identified in cell-free DNA are derived from tumor cells (ctDNA-derived)

Usage

```
plasma_somatic(dat, params)
```

Arguments

dat   tibble containing vectors 'y'and 'n'; 'y' and 'n' should be named

params   a list with named elements that must include the following: 'a': prior expectation for number of plasma somatic variants observed in the plasma sequencing data 'b': prior expectation for number of plasma fragments not containing variants

Value

generate importance samples for plasma somatic model

Examples

```
param.list <- list(ctc=list(a=1, b=9999),
                    ctdna=list(a=1, b=9),
                    chip=list(a=1, b=9),
                    montecarlo.samples=50e3,
                    prior.weight=0.1)
dat <- data.frame(y=c(4, 1),
              n=c(1000, 1000),
              analyte=c("plasma", "buffy coat"),
```

```
                   mutation="mutA",
                   sample_id="id1")
     plasma_somatic(dat, param.list)
```

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plasmut	*Bayesian models for estimating the origin of a sequenced DNA fragment*

Description

The plasmut package provides a Bayesian importance sampling based approach to estimate the liklihood of a mutation arising from clonal hematopoiesis or tumor

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wbc_somatic	*Estimate the marginal likelihood of observing somatic mutations from CTCs present in buffy coat p(y_w | theta_w, n_w, model_S) x p(theta_w| Model_S) theta_w	model_S ~ beta(1, 999) ## sequencing error or CTC*

Description

Estimate the marginal likelihood of observing somatic mutations from CTCs present in buffy coat p(y_w | theta_w, n_w, model_S) x p(theta_w| Model_S) theta_w | model_S ~ beta(1, 999) ## sequencing error or CTC

Usage

```
     wbc_somatic(dat, params)
```

Arguments

dat	tibble containing vectors 'y'and 'n'; 'y' and 'n' should be named
params	a list with named elements that must include the following: 'a': prior expectation for number of somatic variants observed in the WBC sequencing data (either by error or from a CTC) 'b': prior expectation for number of WBCs not containing the variant

Value

generate importance samples for wbc somatic model

Examples

```
param.list <- list(ctc=list(a=1, b=9999),
                    ctdna=list(a=1, b=9),
                    chip=list(a=1, b=9),
                    montecarlo.samples=50e3,
                    prior.weight=0.1)
dat <- data.frame(y=c(4, 1),
                  n=c(1000, 1000),
                  analyte=c("plasma", "buffy coat"),
                  mutation="mutA",
                  sample_id="id1")
wbc_somatic(dat, param.list)
```

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