## Package 'BMRMM'

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**Title** An Implementation of the Bayesian Markov (Renewal) Mixed Models **Version** 0.0.3

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Description The Bayesian Markov renewal mixed models take sequentially observed categorical data with continuous duration times, being either state duration or inter-state duration. These models comprehensively analyze the stochastic dynamics of both state transitions and duration times under the influence of multiple exogenous factors and random individual effect. The default setting flexibly models the transition probabilities using Dirichlet mixtures and the duration times using gamma mixtures. It also provides the flexibility of modeling the categorical sequences using Bayesian Markov mixed models alone, either ignoring the duration times altogether or dividing duration time into multiples of an additional category in the sequence by a user-specific unit. The package allows extensive inference of the state transition probabilities and the duration times as well as relevant plots and graphs. It also includes a synthetic data set to demonstrate the desired format of input data set and the utility of various functions. Methods for Bayesian Markov renewal mixed models are as described in: Abhra Sarkar et al., (2018) <doi:10.1080/01621459.2018.1423986> and Yutong Wu et al., (2022) <doi:10.1093/biostatistics/kxac050>.

**Imports** fields, graphics, grDevices, gsubfn, logOfGamma, MCMCpack, mhsmm, multicool, pracma, stats, utils

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<b>BMRMM</b>																																								2
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2 BMRMM

Index		16
	get_tp_local_test_results	14
	get_tp_diagnostic_plots	
	get_mixture_probs_by_cov	12
	get_mixture_params	11
	get_lpml_waic_scores	10
	get_histogram_with_post_mean	9
	get_histogram_by_component	9
	get_global_test_results	8
	get_estimated_post_mean_and_sd	6
	get_duration_diagnostic_plots	6
	foxp2_sm	5
	toxp2	5

**BMRMM** 

Bayesian Markov Renewal Mixed Models (BMRMMs)

## **Description**

Provides inference results of both transition probabilities and duration times using BMRMMs.

## Usage

```
BMRMM(
  data,
  num_cov,
  switch_off_random = FALSE,
  switch_off_fixed = FALSE,
  trans_cov_index = 1:num_cov,
  duration_type = "Continuous",
  duration_cov_index = 1:num_cov,
  duration_excl_prev_state = FALSE,
  duration_unit = NULL,
  duration_num_comp = 4,
  duration_init_shape = rep(1, duration_num_comp),
  duration_init_rate = rep(1, duration_num_comp),
  simsize = 10000,
  burnin = simsize/2
)
```

## **Arguments**

data a data frame containing – individual ID, covariate values, previous state, current state, duration times (if applicable), in that order num\_cov total number of covariates provided in 'data' switch\_off\_random

TRUE if only population-level effects are considered, default is FALSE

BMRMM 3

switch\_off\_fixed

TRUE if only individual-level effects are considered, default is FALSE

trans\_cov\_index

indices of covariates that are used for transition probabilities, default is all of the covariates

duration\_type one of 'None', 'Discrete', 'Continuous', default is 'Continuous' duration\_cov\_index

indices of covariates that are used for duration times, default is all of the covariates plus the previous state

duration\_excl\_prev\_state

TRUE if the previous state is excluded from duration times inference, default is FALSE

duration\_unit the discretization of duration times, only used when 'duration\_type' is 'Discrete' duration\_num\_comp

number of gamma mixture components for duration times, default is 4

duration\_init\_shape

initialization of mixture gamma shape parameters, default is a vector of 1 of size 'duration\_num\_comp'

duration\_init\_rate

initialization of mixture gamma rate parameters, default is a vector of 1 of size 'duration\_num\_comp'

simsize total number of MCMC iterations, default is 10000

burnin number of burn-ins for the MCMC iterations, default is simsize/2

#### **Details**

Returns list(trans\_results, duration\_results)

(1) Users have the option to ignore duration times or model duration times as a discrete or continuous variable via defining 'duration\_type':

"None": ignores duration times

"Continuous": treat duration times as a continuous variable (DEFAULT)

"Discrete": treat duration times as a new state with discretization 'duration\_unit'. If 'Discrete' is used, duration times becomes a new state type. 'duration\_unit' must be specified if 'duration\_type' is 'Discrete'. For example, if an duration time entry is 20 and 'duration\_unit' is 5, then the model will add 4 consecutive new states. If an duration time entry is 23.33 and 'duration\_unit' is 5, then the model will still add 4 consecutive new states as the blocks are calculated with the floor operation

- (2) 'trans\_results' contains the results of the transition probabilities:
- "Num States": total number of states
- "Xexgns": covariates related to transition probabilities
- "dpreds": maximum level for each related covariate
- "MCMCparams": MCMC parameters: simsize, burnin and thinning factor
- "TP\_Exgns\_Post\_Mean": posterior mean of transition probabilities for different combinations of exogenous predictors

4 BMRMM

- "TP\_Exgns\_Post\_Std": posterior standard deviation of transition probabilities for different combinations of exogenous predictors

- "TP\_Anmls\_Post\_Mean": posterior mean of transition distribution components for different individuals
- "TP\_All\_Post\_Mean": posterior mean of transition distribution components for different combinations of exogenous predictors AND different individuals
- "TP\_Anmls\_Post\_Std": standard deviation of transition probabilities among different mice
- "TP\_Exgns\_Diffs\_Store": difference in posterior mean of transition probabilities for every pair of covariate levels given levels of the other covariates
- "TP Exgns All Itns": exogenous transition probabilities for every MCMC iteration
- "Clusters": number of clusters for each covariate for each MCMC iteration
- "Type": an identifier for results, which is "Transition Probabilities".
- (3) 'duration\_results' contains the results of the duration times ('duration\_results' is NULL if 'duration\_type' is 'None'):
- "K" <- number of gamma mixture components
- "Xexgns" <- covariates related to duration times
- "dpreds" <- maximum level for each related covariate
- "MCMCparams" <- MCMC parameters: simsize, burnin and thinning factor
- "Duration\_Times" <- input duration times
- "Comp\_Assign" <- mixture component assignment for each data point in the last MCMC iteration
- "Duration\_Exgns\_Store" <- posterior mean of mixture probabilities for different combinations of exogenous predictors of each MCMC iteration
- "Marginal\_Prob" <- estimated marginal mixture probabilities for each iteration
- "Shape\_Samples" <- estimated shape parameters for gamma mixtures for each iteration
- "Rate\_Samples" <- estimated rate parameters for gamma mixtures for each iteration
- "Clusters" <- number of clusters for each covariate for each MCMC iteration
- "Type" <- an identifier for results, which is "Inter-Syllable Intervals".

## Value

List of results for transition probabilities and durations times, list(trans\_results, duration\_results). See details.

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## **Examples**

```
# In the examples, we use a shorted version of the foxp2 dataset, foxp2_sm
```

# ignores duration times and only models transition probabilities using all three covariates
results <- BMRMM(foxp2\_sm,num\_cov=2,duration\_type='None',simsize=50)</pre>

```
\# models duration times as a continuous variable with 5 gamma mixture components,
```

# using covariate 1 and the previous state

results <- BMRMM(foxp2\_sm,num\_cov=2,trans\_cov\_index=c(1),'duration\_type'='Continuous',

duration\_cov\_index=c(1),duration\_num\_comp=5,simsize=50)

foxp2

Simulated FoxP2 Data Set.

## **Description**

A simulated data set of the original FoxP2 data set, which contains the sequences of syllables sung by male mice of different genotypes under various social contexts.

#### Usage

foxp2

#### **Format**

A data frame with 17391 rows and 6 variables:

Id Mouse Id

**Genotype** Genotype of the mouse, 1 = FoxP2 knocked out, 2 = wild type

**Context** Social context for the mouse, 1 = U (urine sample placed in the cage), 2 = L (living female mouse placed in the cage), 3 = A (an anesthetized female placed on the lid of the cage)

**Prev\_State** The previous syllable,  $\{1,2,3,4\} = \{d,m,s,u\}$ 

**Cur\_State** The current syllable,  $\{1,2,3,4\} = \{d,m,s,u\}$ 

**Transformed\_ISI** Modified inter-syllable interval times, log(original ISI + 1)

foxp2\_sm

Shortened Simulated FoxP2 Data Set.

## Description

A shortened version of the foxp2 data set for demonstrating R examples. See details of the foxp2 data set by calling ?foxp2.

#### Usage

foxp2\_sm

#### **Format**

An object of class data. frame with 69 rows and 6 columns.

```
{\it get\_duration\_diagnostic\_plots} \\ {\it Duration\ Times\ Plots:\ MCMC\ Diagnostic\ Plots}
```

## Description

Provides the traceplots and autocorrelation plots for each gamma shape and rate parameter

#### Usage

```
get_duration_diagnostic_plots(results, comps = 1:results$K)
```

## Arguments

results of duration times, i.e., results results\_duration

comps a vector of the components to plot; default is 1:duration\_num\_comp

#### Value

No return value, called for plotting MCMC diagnostic plots for duration times.

## **Examples**

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=80)
# get diagnostic plots for all components
get_duration_diagnostic_plots(results$results_duration)
# get diagnostic plots for components 1,2
get_duration_diagnostic_plots(results$results_duration,comps=c(1,2))</pre>
```

```
{\tt get\_estimated\_post\_mean\_and\_sd}
```

Transition Probabilities: Posterior Mean and Standard Deviation

#### **Description**

Print and plot the posterior mean and standard deviation for transition probabilities from MCMC samples under given different combinations of covariate levels.

#### Usage

```
get_estimated_post_mean_and_sd(
  results,
  cov_labels = NULL,
  state_labels = 1:results$Num_States,
  cov_levels = NULL,
  decimal_pts = 2,
  include_plot = TRUE
)
```

#### **Arguments**

results	results of transition probabilities, i.e., results\$results_trans
cov_labels	a matrix such that row i represents the labels for covariate i; default labels for covariate i is $1:i$
state_labels	a vector of strings that represent the state labels; default is 1:Num_States
cov_levels	a matrix such that each row is a combination of covariate levels; default is all possible combinations of covariates
decimal_pts	specify the number of decimal points of the results; default is 2
include_plot	display plot if TRUE; default is TRUE

#### **Details**

For each row of 'cov\_levels', the function returns two matrices of size d0xd0 where d0 is the number of states: (1) the posterior mean and (2) the posterior standard deviation of transition probabilities, computed from MCMC samples after burn-ins and thinning. The default for 'cov\_levels' is all possible combinations of covariate levels.

## Value

No return value, called for printing and plotting posterior distribution of transition probabilities.

```
# Examples using the shortened version of the simulated Foxp2 data set, foxp2_sm

# get results for all combinations of covariate levels
results <- BMRMM(foxp2_sm,num_cov=2,duration_type='None',simsize=50)
get_estimated_post_mean_and_sd(results$results_trans)

# get results for covariate levels ("HET","U") and ("WT","U")
cov_labels <- matrix(c("HET","WT","","U","L","A"),nrow=2,byrow=TRUE)
cov_levels <- matrix(c(1,1,2,1),nrow=2,byrow=TRUE)
get_estimated_post_mean_and_sd(results$results_trans,cov_labels,cov_levels=cov_levels)</pre>
```

```
get_global_test_results
```

Global Test Results for Transition Probabilities and Duration Times

## **Description**

For each covariate, print (and plot) the percentage of the number of clusters for each covariate in stored MCMC iterations.

## Usage

```
get_global_test_results(results, decimal_pts = 2, include_plot = TRUE)
```

## **Arguments**

results either results\$results\_trans or results\$results\_duration

decimal\_pts specify the number of decimal points of the results; default is 2

include\_plot display plot if TRUE; default is TRUE

#### **Details**

The height of the bar represents the percentage of the number of the clusters within the stored MCMC samples.

 $Pr(\# \ clusters \ for \ covariate \ j ==1)$  is the probability for the null hypothesis that the covariate j is not significant for transition probabilities or duration times, depending on the result type.

#### Value

No return value, called for printing and plotting global test results.

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=50)
get_global_test_results(results$results_trans)
get_global_test_results(results$results_duration)</pre>
```

```
get_histogram_by_component
```

Duration Times Plots: Histogram of duration times by Component

## **Description**

Plots the histogram of each mixture component superimposed the gamma distribution with shape and rate parameters taken from the last MCMC iteration

## Usage

```
get_histogram_by_component(results, comp = 1:results$K)
```

## Arguments

results results of duration times, i.e., results\$results\_duration

comp a vector of the components to plot; default is 1:isi\_num\_comp

#### Value

No return value, called for plotting histogram for each mixture component.

## **Examples**

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=50)
get_histogram_by_component(results$results_duration)</pre>
```

```
get_histogram_with_post_mean
```

Duration Times Plots: Histogram of duration times with Estimated Posterior Mean

## Description

Plots the histogram of Duration Times superimposed the posterior mean mixture gamma distribution

## Usage

```
get_histogram_with_post_mean(
  results,
  x_range = c(min(results$Duration_Times), max(results$Duration_Times)),
  breaks = (max(results$Duration_Times) - min(results$Duration_Times)) *
    length(results$Duration_Times)^(1/3)/2/IQR(results$Duration_Times)
)
```

## **Arguments**

results results of duration times, i.e., results\_duration

x\_range custom range for x-axis; default is (min(durations),max(durations))

breaks custom breaks for the histogram; default breaks follows the Freedman-Diaconis

rule, which would be  $(max(durations)-min(durations))*n^{(1/3)/2/IQR(durations)}$ 

#### Value

No return value, called for plotting histogram superimposed with the posterior distribution of mixture gamma.

#### **Examples**

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=50)
get_histogram_with_post_mean(results$results_duration,x_range=c(0,2),breaks=50)</pre>
```

## **Description**

Provides the LPML (Geisser and Eddy, 1979) and WAIC (Watanabe, 2010) scores of the Bayesian Markov renewal mixture models

#### Usage

```
get_lpml_waic_scores(results)
```

## **Arguments**

results results of duration times, i.e., results\$results duration

#### **Details**

The two scores can be used to compare different choices of isi\_num\_comp, i.e., the number of the mixture gamma components. Larger values of LPML and smaller values of WAIC indicate better model fits.

#### Value

No return value, called for printing LPML and WAIC scores for gamma mixture models.

get\_mixture\_params 11

#### References

Geisser, S. and Eddy, W. F. (1979). A predictive approach to model selection. Journal of the American Statistical Association, 74, 153–160.

Watanabe, S. (2010). Asymptotic equivalence of Bayes cross validation and widely applicable information criterion in singular learning theory. Journal of Machine Learning Research, 11, 3571–3594.

## **Examples**

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=50)
get_lpml_waic_scores(results$results_duration)</pre>
```

## **Description**

Print and plot the heat-map for the gamma mixture parameters for each component

## Usage

```
get_mixture_params(results, decimal_pts = 2, include_plot = FALSE)
```

## **Arguments**

results results of duration times, i.e., results\$results\_duration

decimal\_pts specify the number of decimal points of the results; default is 2

include\_plot display plot if TRUE; default is FALSE

#### **Details**

Print a matrix of size K\*2, where K is the number of components. Each row i represents the shape and rate parameter of component i.

#### Value

No return value, called for printing and plotting heatmaps for mixture parameters.

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=50)
get_mixture_params(results$results_duration)</pre>
```

```
get_mixture_probs_by_cov
```

Duration Times Plot: Heat-map of Mixture Probabilities for Different Covariates

## Description

Print and plot the heat-map of mixture probabilities for covariate levels

#### Usage

```
get_mixture_probs_by_cov(
  results,
  cov_labels = NULL,
  cov_index = 1:ncol(results$Xexgns),
  decimal_pts = 2,
  include_plot = FALSE
)
```

## **Arguments**

results	results of duration times, i.e., results\$results_duration
cov_labels	a matrix such that row i represents the labels for covariate $i$ ; default labels for covariate $i$ is $1$ : $i$
cov_index	a vector of indices of covariate to plot; default is 1:Num_Covariates
decimal_pts	specify the number of decimal points of the results; default is 2
include_plot	display plot if TRUE; default is FALSE

## **Details**

Print a matrix for each covariate. Each matrix has K rows, where K is the number of components. The number of columns of a matrix equals to the total number of levels of the covariate.

#### Value

No return value, called for printing and plotting heatmaps for mixture probabilities for each covariate.

```
results <- BMRMM(foxp2_sm,num_cov=2,simsize=50)
get_mixture_probs_by_cov(results$results_duration)
get_mixture_probs_by_cov(results$results_duration,cov_index=c(1,2))</pre>
```

```
{\tt get\_tp\_diagnostic\_plots}
```

Transition Probabilities: MCMC Diagnostic Plots

## **Description**

Provides the traceplots and autocorrelation plots for each transition type under combinations of covariate levels

## Usage

```
get_tp_diagnostic_plots(
  results,
  from = 1:results$Num_States,
  to = 1:results$Num_States,
  cov_levels = NULL,
  cov_labels = NULL,
  state_labels = 1:results$Num_States
)
```

#### **Arguments**

results	results of transition probabilities, i.e., results\$results_trans
from	list of "from" states in target state transitions; default is all states
to	list of "to" states in target state transitions; default is all states
cov_levels	a matrix such that each row is a combination of covariate levels; default is all possible combinations of covariates
cov_labels	a matrix such that row i represents the labels for covariate i; default labels for covariate i is 1:i
state_labels	a vector of strings that represent the state labels; default is 1:Num_States

## Value

No return value, called for plotting MCMC diagnostic plots for transition probabilities.

```
# Examples using the shortened built-in simulated Foxp2 data set, foxp2_sm
# recall the covariate labels are: ("HET","WT") and ("U","L","A")
# recall the state labels are 'd', 'm', 's', 'u'

results <- BMRMM(foxp2_sm,num_cov=2,duration_type='None',simsize=50)
# results for all transition types for every combinations of covariates
get_tp_diagnostic_plots(results$results_trans)</pre>
```

```
get_tp_local_test_results
```

Transition Probabilities: Local Tests for a Specific Covariate

#### **Description**

Given some covariate 'cov' and the levels of other covariates, prints and plots the local test results for pairs of levels of 'cov', including the absolute differences in transition probabilities, and the probabilities for the null hypothesis.

## Usage

```
get_tp_local_test_results(
    results,
    cov,
    delta,
    all_cov_labels = NULL,
    state_labels = 1:results$Num_States,
    other_cov_levels = NULL,
    comp_pairs = NULL,
    decimal_pts = 2,
    include_plot = TRUE
)
```

## **Arguments**

results results of transition probabilities, i.e., results\$results\_trans the index of the covariate to perform local tests on cov delta threshold for the null hypothesis H\_0; see Details all\_cov\_labels a matrix such that row i represents the labels for covariate i; default labels for covariate i is 1:i state\_labels a vector of strings that represent the state labels; default is 1:Num\_States other\_cov\_levels a matrix such that each row is a combination of covariate levels excluding 'cov'; default is all possible combinations of covariates excluding 'cov' comp\_pairs a matrix such that each row is a pair of levels of 'cov' to compare decimal\_pts specify the number of decimal points of the results; default is 2 display plot if TRUE; default is TRUE include\_plot

#### **Details**

Fix a covariate i as the target of the local test.

The function provides two matrices of size d0xd0 where d0 is the number of states:

- (1) the mean of the absolute difference in all transition types for each pair of levels of covariate i;
- (2) the probability of the null hypothesis of the local test for all transition types.

Given a pair of levels of covariate i, say i\_1 and i\_2, and given the levels of other covariates, the null hypothesis is that the difference between i\_1 and i\_2 is not significant for transition probabilities. It is calculated as the percentage of the samples with absolute difference less than 'delta'.

'comp\_pairs' are user-specified pairs of 'cov' for the local test. Each row of 'comp\_pairs' is a pair of indices of 'cov'.

#### Value

No return value, called for printing and plotting local test results for transition probabilities.

# **Index**

```
* datasets
    foxp2, 5
    foxp2_sm, 5
BMRMM, 2
foxp2, 5
foxp2_sm, 5
get_duration_diagnostic_plots, 6
{\tt get\_estimated\_post\_mean\_and\_sd}, \\ 6
get_global_test_results, 8
get_histogram_by_component, 9
{\tt get\_histogram\_with\_post\_mean}, 9
{\tt get\_lpml\_waic\_scores}, 10
get_mixture_params, 11
get_mixture_probs_by_cov, 12
get_tp_diagnostic_plots, 13
{\tt get\_tp\_local\_test\_results}, 14
```