# Package 'FMM'

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Type Package

Title Rhythmic Patterns Modeling by FMM Models

Version 0.4.0

## **Description**

Provides a collection of functions to fit and explore single, multi-component and restricted Frequency Modulated Moebius (FMM) models. 'FMM' is a nonlinear parametric regression model capable of fitting non-sinusoidal shapes in rhythmic patterns. Details about the mathematical formulation of 'FMM' models can be found in Rueda et al. (2019) <doi:10.1038/s41598-019-54569-1>.

URL https://github.com/FMMGroupVa/FMM-base-R-package

 $\pmb{BugReports} \ \ \text{https://github.com/FMMGroupVa/FMM-base-R-package/issues}$ 

License GPL (>= 2)

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# **Description**

Provides a collection of functions to fit and explore single, multi-component and restricted Frequency Modulated Moebius (FMM) models. 'FMM' is a nonlinear parametric regression model capable of fitting non-sinusoidal shapes in rhythmic patterns.

## Details

Package: FMM
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For a complete list of functions with individual help pages, use library(help = "FMM").

## Author(s)

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#### References

Rueda C, Larriba Y, Peddada SD (2019). Frequency Modulated Moebius Model Accurately Predicts Rhythmic Signals in Biological and Physical Sciences. *Scientific reports*, **9** (1), 18701. https://www.nature.com/articles/s41598-019-54569-1

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ecgData

Fifth annotated beat (lead MLII) from patient sel100 in 'QT database'

## **Description**

Voltage electric activity data (mV) of the fifth annotated heartbeat for patient sel100 in 'QT database'. 200 samples were collected along 0.8 seconds with a sampling frequency of 250 Hz. Data records correspond to samples from 151248 to 151049 and can be dowload as text files from 'Physionet' website. Annotated beats were manually revised by experts including R peaks and fiducial marks.

#### Usage

data(ecgData)

#### **Format**

A numeric vector.

#### **Source**

'QT database' from 'Physionet' <a href="https://archive.physionet.org/cgi-bin/atm/ATM">https://archive.physionet.org/cgi-bin/atm/ATM</a>

#### References

Goldberger A, Amaral L, Glass L, Hausdorff J et al. (2000). A qrs detection and r point recognition method for wearable single-lead ecg devices. *Circulation*, **101** (23), E215-20. https://www.mdpi.com/1424-8220/17/9/1969

Laguna P, Mark RG, Goldberg A, Moody GB (1997). A database for evaluation of algorithms for measurement of qt and other waveform intervals in the ecg. *Computers in cardiology 1997*, 673-676. https://ieeexplore.ieee.org/document/648140

## **Examples**

data(ecgData)
str(ecgData)

extractWaves

Individual contribution to the fitted values of each FMM wave

#### **Description**

extractWaves extracts individual contribution to the fitted values of each FMM wave.

## Usage

extractWaves(objFMM)

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#### **Arguments**

objFMM Object of class 'FMM'.

#### Value

Individual contribution to the fitted values of each FMM wave. It is a list object with as many elements as FMM components have been fitted.

#### **Examples**

fitFMM

Fitting FMM models

# **Description**

fitFMM() is used to fit FMM models. The only required argument to fit FMM models is the input data. By default it is assumed that time points, corresponding to a single time period, are equally spaced from 0 to  $2\pi$ .

# Usage

```
fitFMM(
  vData,
  nPeriods = 1,
  timePoints = NULL,
  nback = 1,
  maxiter = nback,
  betaOmegaRestrictions = 1:nback,
  stopFunction = alwaysFalse,
  omegaMin = 1e-04,
  omegaMax = 0.9999,
  lengthAlphaGrid = 48,
  lengthOmegaGrid = 24,
  omegaGrid = NULL,
```

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```
numReps = 1,
showProgress = FALSE,
showTime = FALSE,
parallelize = FALSE,
restrExactSolution = FALSE)
```

## **Arguments**

vData A numeric vector containing the data to be fitted a FMM model.

nPeriods A numeric value specifying the number of periods at which vData is observed.

timePoints A numeric vector containing the time points at which each data of one single

period is observed. The default value is NULL, in which case they are equally

spaced in range  $[0, 2\pi]$ . It must be between 0 and  $2\pi$ .

nback Number of FMM components to be fitted. Its default value is 1.

maxiter Maximum number of iterations for the backfitting algorithm. By default, it is

set at nback.

betaOmegaRestrictions

An integer vector of length nback indicating which FMM waves are constrained to have equal beta and omega parameters. For example, c(1,1,1,2,2) indicates that beta1=beta2=beta3 and beta4=beta5 as well as omega1=omega2=omega3 and omega4=omega5. In brief, some waves are restricted to have the same shape. Its default value is the sequence 1:nback to fit the FMM model without restrictions on shape parameters (beta and omega).

stopFunction Function to check the convergence criterion for the backfitting algorithm (see

Details).

Details).

omegaMin Lower bound for omega parameter and  $0 < omega_{Min} < omega_{Max} < 1$ . By

default, omegaMin = 0.0001.

omegaMax Upper bound for omega parameter and  $0 < omega_{Min} < omega_{Max} < 1$ . By

default, omegaMin = 0.9999.

lengthAlphaGrid

Precision of the grid of alpha in the search of the best model. If it is increased, more possible values of alpha will be considered, resulting in an increasing in the computation time too. By default, it is set to 48 possible values of alpha,

equally spaced between 0 and  $2\pi$ .

lengthOmegaGrid

Precision of the grid of omega in the search of the best model. If it is increased, more possible values of omega will be considered, resulting in an increasing in the computation time too. By default it is set to 24 possible values of omega.

omegaGrid Set of initial omega values in the search of the best model. By default, lengthOmegaGrid

equally spaced values between omegaMin and omegaMax in a logarithmic way.

numReps Number of times (alpha, omega) parameters are refined. Deprecated for non

restricted models.

showProgress TRUE to display a progress indicator on the console.

showTime TRUE to display execution time on the console.

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parallelize

TRUE to use parallelized procedure to fit restricted FMM model. Its default value is FALSE. When it is TRUE, the number of cores to be used is equal to 12, or if the machine has less, the number of cores - 1.

restrExactSolution

FALSE to use an aproximated algorithm to fit the model (default). If TRUE is specified, an nearly exact solution is computed.

#### **Details**

Data will be collected over nPeriods periods. When nPeriods > 1 the fitting is carried out by averaging the data collected at each time point across all considered periods. The model is fitting to summarized data. timePoints is a n-length numeric vector where n is the number of different time points per period.

The stopFunction argument can either be the functions alwaysFalse or R2 included in the package or user-defined functions that have the same arguments. The included functions serve for the following:

- alwaysFalse(), its default value, which returns FALSE to force maxiter iterations; and
- R2(vData, pred, prevPred, difMax = 0.001), a function that computes the difference between the explained variability in two consecutive iterations returning TRUE when the convergence criterion is reached. To calculate the explained variability difference, the data and the fitted values from the current and previous iteration are passed as arguments vData, pred and prevPred, respectively. The convergence criterion is fulfilled when the explained variability difference is less than the argument difMax (by default 0.001).

#### Value

An S4 object of class 'FMM' with information about the fitted model. The object contains the following slots:

- **@timePoints** The time points as specified by the input argument. It is a numeric vector containing the time points at which each data of one single period is observed.
- **@data** The data as specified by the input argument. It is a numeric vector containing the data to be fitted a FMM model. Data could be collected over multiple periods.
- @summarizedData When the data has more than one period, a numeric vector containing data averaging the data at each time point across all considered periods.
- @nPeriods A numeric value containing the number of periods in data as specified by the input argument.
- @fittedValues A numeric vector of the fitted values by the FMM model.
- @**M** A numeric value of the estimated intercept parameter M.
- **@A** A numeric value or vector of the estimated FMM wave amplitude parameter(s) A.
- **@alpha** A numeric value or vector of the estimated FMM wave phase translation parameter(s)  $\alpha$ .
- **@beta** A numeric value or vector of the estimated FMM wave skewness parameter(s)  $\beta$ .
- **@omega** A numeric value or vector of the estimated FMM wave kurtosis parameter(s)  $\omega$ .
- **@SSE** A numeric value of the sum of the residual squares values.
- **@R2** A numeric vector specifying the explained variance by each of the fitted FMM components.
- **@nIter** A numeric value specifying the number of iterations of the fitting algorithm.

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#### References

Rueda C, Larriba Y, Peddada SD (2019). Frequency Modulated Moebius Model Accurately Predicts Rhythmic Signals in Biological and Physical Sciences. *Scientific reports*, **9** (1), 18701. https://www.nature.com/articles/s41598-019-54569-1

# **Examples**

FMM-class

FMM Class Representation

## **Description**

Class representation for an S4 object of class 'FMM'.

#### Value

The S4 object of class 'FMM' contains the following slots:

@timePoints The time points as specified by the input argument. It is a numeric vector con-

taining the time points at which each data of one single period is observed.

edata The data as specified by the input argument. It is a numeric vector containing the

data to be fitted a FMM model. Data could be collected over multiple periods.

@summarizedData

When the data has more than one period, a numeric vector containing data

averaging the data at each time point across all considered periods.

@nPeriods A numeric value containing the number of periods in data as specified by the

input argument.

@fittedValues A numeric vector of the fitted values by the FMM model.

 ${\sf QM}$  A numeric value of the estimated intercept parameter M.

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| @alphaA numeric value or vector of the estimated FMM wave phase translation parameter(s) $\alpha$ .@betaA numeric value or vector of the estimated FMM wave skewness parameter(s) $\beta$ .@omegaA numeric value or vector of the estimated FMM wave kurtosis parameter(s) $\omega$ .@SSEA numeric value of the sum of the residual squares values.@R2A numeric vector specifying the explained variance by each of the fitted FMM components.@nIterA numeric value specifying the number of iterations of the fitting algorithm. | @A     | A numeric value or vector of the estimated FMM wave amplitude parameter(s) A.                    |
|---|--------|--|
| $\beta.$ @omega A numeric value or vector of the estimated FMM wave kurtosis parameter(s) $\omega.$ @SSE A numeric value of the sum of the residual squares values.  @R2 A numeric vector specifying the explained variance by each of the fitted FMM components.   | @alpha | 1  |
| QSSE A numeric value of the sum of the residual squares values.  A numeric vector specifying the explained variance by each of the fitted FMM components.   | @beta  |  |
| QR2 A numeric vector specifying the explained variance by each of the fitted FMM components.  | @omega | A numeric value or vector of the estimated FMM wave kurtosis parameter(s) $\boldsymbol{\omega}.$ |
| components.   | @SSE   | A numeric value of the sum of the residual squares values.                                       |
| @nIter A numeric value specifying the number of iterations of the fitting algorithm.  | @R2    |  |
|   | @nIter | A numeric value specifying the number of iterations of the fitting algorithm.                    |

# **Examples**

```
## FMM class
getClass("FMM")
getSlots("FMM")
```

FMM-methods

Methods for objects of class 'FMM'

# Description

The methods for objects of class 'FMM' are:

```
coef coef method for S4 class 'FMM', summary method for S4 class 'FMM', fitted method for S4 class 'FMM', resid method for S4 class 'FMM'.
```

# Usage

```
## S4 coef method for signature 'FMM'
coef(object,...)
## S4 summary method for signature 'FMM'
summary(object,...)
## S4 fitted method for signature 'FMM'
fitted(object,...)
## S4 resid method for signature 'FMM'
resid(object,...)
```

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#### **Arguments**

```
object of class 'FMM'.
... additional arguments passed to the method.
```

#### Value

• The function coef() returns a list with two components:

A numeric value. The estimated intercept parameter M.

Wave A data. frame with the estimates of each FMM wave parameters. It is organised as one component per row.

• The function summary() displays relevant results of the fitting. When it is assigned, this function returns a list with the following components:

coef The list returns by the function coef().

peak.time A data.frame with the estimates of the peak and trough times of each wave. It is organised as one componer

peak.time A data.frame with the e resid The vector of residuals.

R. squared A numerical value with the R squared of each wave. The total R square is computed as the sum of the contrib

- The function fitted() returns a data. frame with two columns: timePoints and fittedValues.
- The function resid() returns a numeric vector with residuals of the model.

```
## Generate example data:
fmm2.data <- generateFMM(0, rep(2, 2), c(1.5, 3.4), c(0.2, 2.3), c(0.1, 0.2),
                     plot = FALSE, outvalues = TRUE,
                     sigmaNoise = 0.5) # add a gaussian noise with sigma = 0.5
## Fit the FMM model with nback = 2 component
## fit is an object of S4 class 'FMM'
fit <- fitFMM(vData = fmm2.data$y,timePoints = fmm2.data$t,nback = 2,</pre>
              lengthAlphaGrid = 24,lengthOmegaGrid = 10)
## Extract coefficients of the model:
coef(fit)
## Summarize results:
summary(fit)
## Results on a list:
res <- summary(fit)</pre>
res$peak.time # fiducial points
## fitted values:
fit.values <- fitted(fit)</pre>
## residuals
res <- resid(fit)
```

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generateFMM

Simulating data from FMM models

# Description

generateFMM() simulates data from a FMM model defined by parameters M, A,  $\alpha$ ,  $\beta$  and  $\omega$ .

# Usage

```
generateFMM(
    M,
    A,
    alpha,
    beta,
    omega,
    from = 0,
    to = 2 * pi,
    length.out = 200,
    plot = TRUE,
    outvalues = TRUE,
    sigmaNoise = 0
)
```

# Arguments

| М          | A numeric vector which contains the value of the intercept parameter M.   |
|------------|---|
| A          | A positive numeric vector which contains the value of the FMM wave amplitude parameter A.   |
| alpha      | A numeric vector which contains the value of the FMM wave phase translation parameter $\boldsymbol{\alpha}.$                        |
| beta       | A numeric vector which contains the value of the FMM wave skewness parameter $\boldsymbol{\beta}.$                                  |
| omega      | A numeric vector which contains the value of the FMM wave kurtosis parameter $\omega.$ omega parameter must be between $0$ and $1.$ |
| from       | A numeric value which contains the initial time point of the simulated data. By default, it is $0$ .                                |
| to         | A numeric value which contains the final time point of the simulated data. By default, it is $2\pi$ .                               |
| length.out | A non-negative number wich contains the desired length of the simulation. By default, it is 100.                                    |
| plot       | A logical value indicating whether the simulated data should be drawn on a plot. By default, it is TRUE.                            |
| outvalues  | A logical value indicating whether the numerical simulation should be return. By default, it is TRUE.                               |
|            |   |

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sigmaNoise

A non-negative number which contains the standard deviation of the gaussian noise to be added. Its default value is zero equivalent to a simulation set-up without noise.

#### **Details**

To simulate a multicomponent FMM model, arguments A, alpha, beta and omega are vectors of length m, where m represents the number of FMM waves. With different lengths, the smaller vectors will be replicate until thay are the same length as the longest vector.

With sigmaNoise = s, s>0, the generateFMM function uses rnorm(length.out, 0, sigmaNoise) to create the normally distributed noise and adds it to the simulated values.

#### Value

When outvalues = TRUE a list of with the following components is returned:

input a list with the input parameters M, A, alpha, beta and omega.

t a numeric vector with the time points at each data is simulated.

y a numeric vector with the data simulated.

When plot = TRUE a scatter plot of y vs t is drawn.

#### References

Rueda C, Larriba Y, Peddada SD (2019). Frequency Modulated Moebius Model Accurately Predicts Rhythmic Signals in Biological and Physical Sciences. *Scientific reports*, **9** (1), 18701. https://www.nature.com/articles/s41598-019-54569-1

12 getFMMPeaks

| getFMMPeaks | Peak and trough times and signal values |  |
|-------------|---|--|

## **Description**

getFMMPeaks() is used to estimate peak and trough times and signal values at those times for each component of the model. These parameters result to be useful in multiple applications.

## Usage

```
getFMMPeaks(objFMM, timePointsIn2pi = TRUE)
```

## **Arguments**

```
objFMM Object of class 'FMM' timePointsIn2pi
```

TRUE to return peak and trough times in the  $[0,2\pi]$  interval. When timePointsIn2pi = FALSE the positions of peak and trough times are returned. Its default value is TRUE.

#### Value

A list with the following components is returned:

tpeakU a numeric vector with the time points at which the peak of each wave is esti-

mated.

tpeakL a numeric vector with the time points at which the trough of each wave is esti-

mated.

ZU a numeric vector with the estimated signal peak values of each wave.

ZL a numeric vector with the estimated signal trough values of each wave.

#### References

Rueda C, Larriba Y, Peddada SD (2019). Frequency Modulated Moebius Model Accurately Predicts Rhythmic Signals in Biological and Physical Sciences. *Scientific reports*, **9** (1), 18701. https://www.nature.com/articles/s41598-019-54569-1

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```
lengthAlphaGrid = 24,lengthOmegaGrid = 10)
getFMMPeaks(fit, timePointsIn2pi = TRUE) # times in the [0,2*pi] interval
```

getS4

General S4 Class Extractor Functions

# Description

A collection of functions to extract slots from S4 objects of class 'FMM'.

The extractor functions are:

| getTimePoints     | Extracts the timePoints slot from a S4 object of class 'FMM'.     |
|-------------------|---|
| getData           | Extracts the data slot from a S4 object of class 'FMM',           |
| getSummarizedData | Extracts the summarizedData slot from a S4 object of class 'FMM', |
| getNPeriods       | Extracts the nPeriods slot from a S4 object of class 'FMM',       |
| getFittedValues   | Extracts the fittedValues slot from a S4 object of class 'FMM',   |
| getM              | Extracts the M slot from a S4 object of class 'FMM',              |
| getA              | Extracts the A slot from a S4 object of class 'FMM',              |
| getAlpha          | Extracts the alpha slot from a S4 object of class 'FMM',          |
| getBeta           | Extracts the beta slot from a S4 object of class 'FMM',           |
| getOmega          | Extracts the omega slot from a S4 object of class 'FMM',          |
| getSSE            | Extracts the SSE slot from a S4 object of class 'FMM',            |
| getR2             | Extracts the R2 slot from a S4 object of class 'FMM',             |
| getNIter          | Extracts the nIter slot from a S4 object of class 'FMM',          |
|                   |   |

# Usage

```
getM(objFMM)
getOmega(objFMM)
getData(objFMM)
```

# Arguments

objFMM an object of class of class 'FMM'.

# Value

Return the content of the corresponding slot.

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mouseGeneExp

Iqgap2 gene expression data from mouse liver

# Description

Data from High-temporal resolution profiling of mouse liver. Samples were collected every hour for 48 hours from 3-5 mice per time point from liver. Samples were pooled and analyzed using Affymetrix arrays.

## Usage

data(mouseGeneExp)

#### **Format**

A numeric vector.

#### Source

'NCBI GEO', accession number GSE11923 <a href="https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE11923">https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE11923</a>

#### References

Hughes ME, DiTacchio L, Hayes KR, Vollmers C et al. Harmonics of circadian gene transcription in mammals. PLoS Genet 2009 Apr;5(4):e1000442. PMID: 19343201

#### **Examples**

data(mouseGeneExp)
str(mouseGeneExp)

neuronalAPTrain

Neuronal AP Train Data simulated with Hodgkin-Huxley model

# **Description**

Voltage data in mV simulated with Hodgkin Huxley model (parameters: C=1, gNa=232, gK=45, gL=0.215, vK=-12, vNa=115, vL=10.6, bar(alphaN)=0.95, bar(betaN)=1.3, bar(alphaM)=1, bar(betaM)=1.15, bar(alphaH)=1, bar(betaH)=1) and applied current of 4.5 microA 1 millisecond. The simulation has been done with a modified NeuroDynex Python module.

# Usage

data(neuronalAPTrain)

neuronalSpike 15

#### **Format**

A numeric vector.

#### Source

NeuroDynex Documentation, <a href="https://lcn-neurodynex-exercises.readthedocs.io/en/latest/#>

#### References

Wulfram Gerstner, Werner M. Kistler, Richard Naud, and Liam Paninski (2014). Neuronal Dynamics: From Single Neurons to Networks and Models of Cognition. ([Online Book](https://neuronaldynamics.epfl.ch/online/))

# **Examples**

```
data(neuronalAPTrain)
str(neuronalAPTrain)
```

neuronalSpike

Neuronal Spike Data simulated with Hodgkin-Huxley model

# **Description**

Voltage data in mV simulated with Hodgkin Huxley model (parameters: C=1, gNa=260, gK=30, gL=0.31, vK=-12, vNa=115, vL=10.6, bar(alphaN)=1.15, bar(betaN)=0.85, bar(alphaM)=0.9, bar(betaM)=1.3, bar(alphaH)=1, bar(betaH)=1) and applied current of 12 microA 1 millisecond. The simulation has been done with a modified NeuroDynex Python module.

#### Usage

data(neuronalSpike)

#### **Format**

A numeric vector.

#### **Source**

NeuroDynex Documentation, <a href="https://lcn-neurodynex-exercises.readthedocs.io/en/latest/#>

### References

Wulfram Gerstner, Werner M. Kistler, Richard Naud, and Liam Paninski (2014). Neuronal Dynamics: From Single Neurons to Networks and Models of Cognition. ([Online Book](https://neuronaldynamics.epfl.ch/online/))

```
data(neuronalSpike)
str(neuronalSpike)
```

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plotFMM

Plot fitted FMM models

## **Description**

plotFMM() is used to plot fitted FMM models. The function can either plot the fitted model against the data or each of the components of the model separately. Optionally 'ggplot2' can be used as graphic library.

# Usage

```
plotFMM(
  objFMM,
  components = FALSE,
  plotAlongPeriods = FALSE,
  use_ggplot2 = FALSE,
  legendInComponentsPlot = TRUE,
  textExtra = ""
)
```

## Arguments

objFMM Object of class FMM

components A logical value indicating if the centered wave components of the model should

be separately plotted (case where it is TRUE). If FALSE, the default, the fitted

FMM model along with the observed data is plotted.

plotAlongPeriods

A logical value indicating if more than one period should be plotted in the plots

by default. Its default value is FALSE.

use\_ggplot2 A logical value. If FALSE, the default, R base graphics are used. If TRUE,

'ggplot2' library is used as graphics engine.

legendInComponentsPlot

A logical value indicating whether the legend should be plotted in the compo-

nents plot. By defaults it is TRUE.

textExtra A character vector for extra text to be added to the titles of the plots.

#### **Details**

plotFMM() can generate two types of plots: the basic plot compares the fitted model against the original data while the components plot represents separately the centered waves of the model (if the argument components is TRUE).

The function is also capable of plotting multiple periods if the data has more than one, as is the case in many applications such as chronobiology. In this case, the argument plotAlongPeriods should be TRUE. In the case of components plots the value taken by the latter argument is ignored as they are plotted along just one period.

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While, by default, plots are created using base R graphics, 'ggplot2' can also be used for more aesthetic and customizable plots. Optional arguments legendInComponentsPlot and textExtra serve to control, respectively, whether a legend to the components plot should be added and adding extra text to the plot's title.

#### Value

None if base R graphics are used, a named ggplot2 list if 'ggplot2' is used.

```
# Simulates an scenario in which an FMM model is suitable,
res <- generateFMM(2,3,1.5,2.3,0.1,outvalues = TRUE,sigmaNoise = 0.3, plot=FALSE)
# then a FMM model is fitted to the data.
fit <- fitFMM(res$y, lengthAlphaGrid=20,lengthOmegaGrid=12)</pre>
plotFMM(fit)
# Components plot of FMM Model fitted to neuronal data with various optional aesthetics
data("neuronalSpike")
fittedFMM2<-fitFMM(neuronalSpike, nback=2,</pre>
                   lengthAlphaGrid = 24,lengthOmegaGrid = 10, numReps = 1)
plotFMM(fittedFMM2, components = TRUE)
plotFMM(fittedFMM2, components = TRUE,
        legendInComponentsPlot = FALSE,
        textExtra = "Neuronal Data")
# With ggplot2, customizable plots can be created,
library(ggplot2)
# standard plots
plotFMM(fittedFMM2, use_ggplot2 = TRUE)
# and components plots
plotFMM(fittedFMM2, components = TRUE, use_ggplot2 = TRUE)
# Plot of fitted model to more than one period.
data("mouseGeneExp")
fittedFMM2<-fitFMM(mouseGeneExp, nPeriods = 2,</pre>
                   lengthAlphaGrid = 20,lengthOmegaGrid = 10)
plotFMM(fittedFMM2, plotAlongPeriods = TRUE)
```

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