

# Package ‘cfma’

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**Title** Causal Functional Mediation Analysis

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**Description** Performs causal functional mediation analysis (CFMA) for functional treatment, functional mediator, and functional outcome. This package includes two functional mediation model types: (1) a concurrent mediation model and (2) a historical influence mediation model. See Zhao et al. (2018), Functional Mediation Analysis with an Application to Functional Magnetic Resonance Imaging Data, <[arXiv:1805.06923](#)> for details.

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cfma-package

*Causal Functional Mediation Analysis*

---

### Description

cfma package performs causal functional mediation analysis (CFMA) for functional treatment, functional mediator, and functional outcome. This package includes two functional mediation model type: (1) a concurrent mediation model and (2) a historical influence mediation model.

### Details

Package: cfma  
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### Author(s)

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---

env.concurrent

*Simulated data from the concurrent mediation model*

---

### Description

"env.concurrent" is an R environment containing the data generated from a concurrent mediation model.

### Usage

```
data("env.concurrent")
```

### Format

An R environment

Z a  $n \times T$  data matrix, treatment trajectory of  $n$  subjects for  $T$  time points.

M a  $n \times T$  data matrix, mediator trajectory of  $n$  subjects for  $T$  time points.

$Y$  a  $n \times T$  data matrix, outcome trajectory of  $n$  subjects for  $T$  time points.

$\alpha$  a length  $T$  vector model coefficient.

$\beta$  a length  $T$  vector model coefficient.

$\gamma$  a length  $T$  vector model coefficient.

### Details

The data was generated from the concurrent mediation model

$$M(t) = Z(t)\alpha(t) + \epsilon_1(t),$$

$$R(t) = Z(t)\gamma(t) + M(t)\beta(t) + \epsilon_2(t).$$

$Z(t)$  is the convolution of hemodynamic response function (HRF) and event onsets.

### Examples

```
data(env.concurrent)
Z<-get("Z",env.concurrent)
M<-get("M",env.concurrent)
Y<-get("Y",env.concurrent)
```

---

env.historical

*Simulated data from the historical influence mediation model*

---

### Description

"env.historical" is an R environment containing the data generated from a historical influence mediation model.

### Usage

```
data("env.historical")
```

### Format

An R environment

$Z$  a  $n \times T$  data matrix, treatment trajectory of  $n$  subjects for  $T$  time points.

$M$  a  $n \times T$  data matrix, mediator trajectory of  $n$  subjects for  $T$  time points.

$Y$  a  $n \times T$  data matrix, outcome trajectory of  $n$  subjects for  $T$  time points.

$\alpha$  a  $T \times T$  matrix model coefficient.

$\beta$  a  $T \times T$  matrix model coefficient.

$\gamma$  a  $T \times T$  matrix model coefficient.

## Details

The data was generated from the historical influence mediation model

$$M(t) = \int_{\Omega_t^1} Z(s)\alpha(s,t)ds + \epsilon_1(t),$$

$$Y(t) = \int_{\Omega_t^2} Z(s)\gamma(s,t)ds + \int_{\Omega_t^3} M(s)\beta(s,t)ds + \epsilon_2(t),$$

where  $\alpha(s, t)$ ,  $\beta(s, t)$ ,  $\gamma(s, t)$  are coefficient curves;  $\Omega_t^j = [(t - \delta_j) \vee 0, t]$  for  $j = 1, 2, 3$ .  $Z(t)$  is the convolution of hemodynamic response function (HRF) and event onsets.

## Examples

```
data(env.historical)
Z<-get("Z",env.historical)
M<-get("M",env.historical)
Y<-get("Y",env.historical)
```

---

FMA.concurrent

*Functional mediation analysis under concurrent regression model*

---

## Description

This function performs functional mediation regression under the concurrent model with given tuning parameter.

## Usage

```
FMA.concurrent(Z, M, Y, intercept = TRUE, basis = NULL, Ld2.basis = NULL,
  basis.type = c("fourier"), nbasis = 3, timeinv = c(0, 1), timegrids = NULL,
  lambda.m = 0.01, lambda.y = 0.01)
```

## Arguments

Z	a data matrix. Z is the treatment trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
M	a data matrix. M is the mediator trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
Y	a data matrix. Y is the outcome trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
intercept	a logic variable. Default is TRUE, an intercept term is included in the regression model.

basis	a data matrix. Basis function used in the functional data analysis. The number of columns is the number of basis function considered. If basis = NULL, Fourier basis functions will be generated.
Ld2.basis	a data matrix. The second derivative of the basis function. The number of columns is the number of basis function considered. If Ld2.basis = NULL, the second derivative of Fourier basis functions will be generated.
basis.type	a character of basis function type. Default is Fourier basis (basis.type = "fourier").
nbasis	an integer, the number of basis function included. If basis is provided, this argument will be ignored.
timeinv	a numeric vector of length two, the time interval considered in the analysis. Default is (0,1).
timegrids	a numeric vector of time grids of measurement. If timegrids = NULL, it is assumed the between measurement time interval is constant.
lambda.m	a numeric value of the tuning parameter in the mediator model.
lambda.y	a numeric value of the tuning parameter in the outcome model.

### Details

The concurrent mediation model is

$$M(t) = Z(t)\alpha(t) + \epsilon_1(t),$$

$$Y(t) = Z(t)\gamma(t) + M(t)\beta(t) + \epsilon_2(t),$$

where  $\alpha(t)$ ,  $\beta(t)$ ,  $\gamma(t)$  are coefficient curves. The model coefficient curves are estimated by minimizing the penalized  $L_2$ -loss.

### Value

basis	the basis functions used in the analysis.
M	a list of output for the mediator model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of M lambda: $\lambda$ value
Y	a list of output for the outcome model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of Y lambda: $\lambda$ value
IE	a list of output for the indirect effect comparing $Z_1(t) = 1$ versus $Z_0(t) = 0$ coefficients: the coefficient with respect to the basis function curve: the estimated causal curve
DE	a list of output for the direct effect comparing $Z_1(t) = 1$ versus $Z_0(t) = 0$ coefficients: the coefficient with respect to the basis function curve: the estimated causal curve

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 Brian Caffo, Johns Hopkins University, <bcaffo@gmail.com>

**References**

Zhao et al. (2017). *Functional Mediation Analysis with an Application to Functional Magnetic Resonance Imaging Data*. arXiv preprint arXiv:1805.06923.

**Examples**

```
#####
# Concurrent functional mediation model
data(env.concurrent)
Z<-get("Z",env.concurrent)
M<-get("M",env.concurrent)
Y<-get("Y",env.concurrent)

# consider Fourier basis
fit<-FMA.concurrent(Z,M,Y,intercept=FALSE,timeinv=c(0,300))

# estimate of alpha
plot(fit$M$curve[1,],type="l",lwd=5)
lines(get("alpha",env.concurrent),lty=2,lwd=2,col=2)

# estimate of gamma
plot(fit$Y$curve[1,],type="l",lwd=5)
lines(get("gamma",env.concurrent),lty=2,lwd=2,col=2)

# estimate of beta
plot(fit$Y$curve[2,],type="l",lwd=5)
lines(get("beta",env.concurrent),lty=2,lwd=2,col=2)

# estimate of causal curves
plot(fit$IE$curve,type="l",lwd=5)
plot(fit$DE$curve,type="l",lwd=5)
#####
```

---

FMA.concurrent.boot      *Functional mediation analysis under concurrent regression model  
with point-wise bootstrap confidence interval*

---

**Description**

This function performs functional mediation regression under the concurrent model with given tuning parameter. Point-wise confidence bands are obtained from bootstrap.

**Usage**

```
FMA.concurrent.boot(Z, M, Y, intercept = TRUE, basis = NULL, Ld2.basis = NULL,
  basis.type = c("fourier"), nbasis = 3, timeinv = c(0, 1), timegrids = NULL,
  lambda.m = 0.01, lambda.y = 0.01, sims = 1000, boot = TRUE,
  boot.ci.type = c("bca", "perc"), conf.level = 0.95, verbose = TRUE)
```

**Arguments**

Z	a data matrix. Z is the treatment trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
M	a data matrix. M is the mediator trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
Y	a data matrix. Y is the outcome trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
intercept	a logic variable. Default is TRUE, an intercept term is included in the regression model.
basis	a data matrix. Basis function used in the functional data analysis. The number of columns is the number of basis function considered. If basis = NULL, Fourier basis functions will be generated.
Ld2.basis	a data matrix. The second derivative of the basis function. The number of columns is the number of basis function considered. If Ld2.basis = NULL, the second derivative of Fourier basis functions will be generated.
basis.type	a character of basis function type. Default is Fourier basis (basis.type = "fourier").
nbasis	an integer, the number of basis function included. If basis is provided, this argument will be ignored.
timeinv	a numeric vector of length two, the time interval considered in the analysis. Default is (0,1).
timegrids	a numeric vector of time grids of measurement. If timegrids = NULL, it is assumed the between measurement time interval is constant.
lambda.m	a numeric value of the tuning parameter in the mediator model.
lambda.y	a numeric value of the tuning parameter in the outcome model.
sims	an integer indicating the number of simulations for inference.
boot	a logical value, indicating whether or not bootstrap should be used. Default is TRUE.
boot.ci.type	a character of confidence interval method. boot.ci.type = "bca" bias corrected confidence interval; boot.ci.type = "perc" percentile confidence interval.
conf.level	a number of significance level. Default is 0.95.
verbose	a logical value, indicating whether print out bootstrap replications.

## Details

The concurrent mediation model is

$$M(t) = Z(t)\alpha(t) + \epsilon_1(t),$$

$$Y(t) = Z(t)\gamma(t) + M(t)\beta(t) + \epsilon_2(t),$$

where  $\alpha(t)$ ,  $\beta(t)$ ,  $\gamma(t)$  are coefficient curves. The model coefficient curves are estimated by minimizing the penalized  $L_2$ -loss.

## Value

alpha	a list of output for $\alpha$ estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve
gamma	: a list of output for $\gamma$ estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve
beta	a list of output for $\beta$ estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve
IE	a list of output for indirect effect estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve
DE	a list of output for direct effect estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve

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## References

Zhao et al. (2017). *Functional Mediation Analysis with an Application to Functional Magnetic Resonance Imaging Data*. arXiv preprint arXiv:1805.06923.

## Examples

```
#####
# Concurrent functional mediation model
data(env.concurrent)
Z<-get("Z",env.concurrent)
M<-get("M",env.concurrent)
Y<-get("Y",env.concurrent)

# consider Fourier basis
fit.boot<-FMA.concurrent.boot(Z,M,Y,intercept=FALSE,timeinv=c(0,300))

#####
```

---

FMA.concurrent.CV      *Functional mediation analysis under concurrent regression model*

---

## Description

This function performs functional mediation regression under the concurrent model. Tuning parameter is chosen based on cross-validation.

## Usage

```
FMA.concurrent.CV(Z, M, Y, intercept = TRUE, basis = NULL, Ld2.basis = NULL,
  basis.type = c("fourier"), nbasis = 3, timeinv = c(0, 1), timegrids = NULL,
  lambda = NULL, nfold = 5)
```

## Arguments

Z	a data matrix. Z is the treatment trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
M	a data matrix. M is the mediator trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
Y	a data matrix. Y is the outcome trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
intercept	a logic variable. Default is TRUE, an intercept term is included in the regression model.
basis	a data matrix. Basis function used in the functional data analysis. The number of columns is the number of basis function considered. If basis = NULL, Fourier basis functions will be generated.

Ld2.basis	a data matrix. The second derivative of the basis function. The number of columns is the number of basis function considered. If Ld2.basis = NULL, the second derivative of Fourier basis functions will be generated.
basis.type	a character of basis function type. Default is Fourier basis (basis.type = "fourier").
nbasis	an integer, the number of basis function included. If basis is provided, this argument will be ignored.
timeinv	a numeric vector of length two, the time interval considered in the analysis. Default is (0,1).
timegrids	a numeric vector of time grids of measurement. If timegrids = NULL, it is assumed the between measurement time interval is constant.
lambda	a numeric vector of tuning parameter values.
nfolds	a number gives the number of folds in cross-validation.

### Details

The concurrent mediation model is

$$M(t) = Z(t)\alpha(t) + \epsilon_1(t),$$

$$Y(t) = Z(t)\gamma(t) + M(t)\beta(t) + \epsilon_2(t),$$

where  $\alpha(t)$ ,  $\beta(t)$ ,  $\gamma(t)$  are coefficient curves. The model coefficient curves are estimated by minimizing the penalized  $L_2$ -loss. Tuning parameter  $\lambda$  controls the smoothness of the estimated curves, and is chosen by cross-validation.

### Value

basis	the basis functions used in the analysis.
M	a list of output for the mediator model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of M lambda: the chosen $\lambda$ value
Y	a list of output for the outcome model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of Y lambda: the chosen $\lambda$ value
IE	a list of output for the indirect effect comparing $Z_1(t) = 1$ versus $Z_0(t) = 0$ coefficients: the coefficient with respect to the basis function curve: the estimated causal curve
DE	a list of output for the direct effect comparing $Z_1(t) = 1$ versus $Z_0(t) = 0$ coefficients: the coefficient with respect to the basis function curve: the estimated causal curve

**Author(s)**

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**References**

Zhao et al. (2017). *Functional Mediation Analysis with an Application to Functional Magnetic Resonance Imaging Data*. arXiv preprint arXiv:1805.06923.

**Examples**

```
#####
# Concurrent functional mediation model
data(env.concurrent)
Z<-get("Z",env.concurrent)
M<-get("M",env.concurrent)
Y<-get("Y",env.concurrent)

# consider Fourier basis
fit<-FMA.concurrent.CV(Z,M,Y,intercept=FALSE,timeinv=c(0,300))

# estimate of alpha
plot(fit$M$curve[1,],type="l",lwd=5)
lines(get("alpha",env.concurrent),lty=2,lwd=2,col=2)

# estimate of gamma
plot(fit$Y$curve[1,],type="l",lwd=5)
lines(get("gamma",env.concurrent),lty=2,lwd=2,col=2)

# estimate of beta
plot(fit$Y$curve[2,],type="l",lwd=5)
lines(get("beta",env.concurrent),lty=2,lwd=2,col=2)

# estimate of causal curves
plot(fit$IE$curve,type="l",lwd=5)
plot(fit$DE$curve,type="l",lwd=5)

#####
```

**Description**

This function performs functional mediation regression under the historical influence model with given tuning parameter.

**Usage**

```
FMA.historical(Z, M, Y, delta.grid1 = 1, delta.grid2 = 1, delta.grid3 = 1,
  intercept = TRUE, basis1 = NULL, Ld2.basis1 = NULL, basis2 = NULL, Ld2.basis2 = NULL,
  basis.type = c("fourier"), nbasis1 = 3, nbasis2 = 3,
  timeinv = c(0, 1), timegrids = NULL,
  lambda1.m = 0.01, lambda2.m = 0.01, lambda1.y = 0.01, lambda2.y = 0.01)
```

**Arguments**

Z	a data matrix. Z is the treatment trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
M	a data matrix. M is the mediator trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
Y	a data matrix. Y is the outcome trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
delta.grid1	a number indicates the width of treatment-mediator time interval in the mediator model.
delta.grid2	a number indicates the width of treatment-outcome time interval in the outcome model.
delta.grid3	a number indicates the width of mediator-outcome time interval in the outcome model.
intercept	a logic variable. Default is TRUE, an intercept term is included in the regression model.
basis1	a data matrix. Basis function on the $s$ domain used in the functional data analysis. The number of columns is the number of basis function considered. If <code>basis = NULL</code> , Fourier basis functions will be generated.
Ld2.basis1	a data matrix. The second derivative of the basis function on the $s$ domain. The number of columns is the number of basis function considered. If <code>Ld2.basis = NULL</code> , the second derivative of Fourier basis functions will be generated.
basis2	a data matrix. Basis function on the $t$ domain used in the functional data analysis. The number of columns is the number of basis function considered. If <code>basis = NULL</code> , Fourier basis functions will be generated.
Ld2.basis2	a data matrix. The second derivative of the basis function on the $t$ domain. The number of columns is the number of basis function considered. If <code>Ld2.basis = NULL</code> , the second derivative of Fourier basis functions will be generated.
basis.type	a character of basis function type. Default is Fourier basis ( <code>basis.type = "fourier"</code> ).

nbasis1	an integer, the number of basis function on the $s$ domain included. If nbasis1 is provided, this argument will be ignored.
nbasis2	an integer, the number of basis function on the $t$ domain included. If nbasis2 is provided, this argument will be ignored.
timeinv	a numeric vector of length two, the time interval considered in the analysis. Default is (0,1).
timegrids	a numeric vector of time grids of measurement. If timegrids = NULL, it is assumed the between measurement time interval is constant.
lambda1.m	a numeric vector of tuning parameter values on the $s$ domain in the mediator model.
lambda2.m	a numeric vector of tuning parameter values on the $t$ domain in the mediator model.
lambda1.y	a numeric vector of tuning parameter values on the $s$ domain in the outcome model.
lambda2.y	a numeric vector of tuning parameter values on the $t$ domain in the outcome model.

### Details

The historical influence mediation model is

$$M(t) = \int_{\Omega_t^1} Z(s)\alpha(s,t)ds + \epsilon_1(t),$$

$$Y(t) = \int_{\Omega_t^2} Z(s)\gamma(s,t)ds + \int_{\Omega_t^3} M(s)\beta(s,t)ds + \epsilon_2(t),$$

where  $\alpha(s,t)$ ,  $\beta(s,t)$ ,  $\gamma(s,t)$  are coefficient curves;  $\Omega_t^j = [(t - \delta_j) \vee 0, t]$  for  $j = 1, 2, 3$ . The model coefficient curves are estimated by minimizing the penalized  $L_2$ -loss.

### Value

basis1	the basis functions on the $s$ domain used in the analysis.
basis2	the basis functions on the $t$ domain used in the analysis.
M	a list of output for the mediator model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of M lambda1: the $\lambda$ value on the $s$ domain lambda2: the $\lambda$ value on the $t$ domain
Y	a list of output for the outcome model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of Y lambda1: the $\lambda$ value on the $s$ domain lambda2: the $\lambda$ value on the $t$ domain

- IE a list of output for the indirect effect comparing  $Z_1(t) = 1$  versus  $Z_0(t) = 0$  curve: the estimated causal curve
- DE a list of output for the direct effect comparing  $Z_1(t) = 1$  versus  $Z_0(t) = 0$  curve: the estimated causal curve

### Author(s)

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 Brian Caffo, Johns Hopkins University, <bcaffo@gmail.com>

### References

Zhao et al. (2017). *Functional Mediation Analysis with an Application to Functional Magnetic Resonance Imaging Data*. arXiv preprint arXiv:1805.06923.

### Examples

```
#####
# Historical influence functional mediation model
data(env.historical)
Z<-get("Z",env.historical)
M<-get("M",env.historical)
Y<-get("Y",env.historical)

# consider Fourier basis
fit<-FMA.historical(Z,M,Y,delta.grid1=3,delta.grid2=3,delta.grid3=3,
  intercept=FALSE,timeinv=c(0,300))

# estimate of causal curves
plot(fit$IE$curve,type="l",lwd=5)
plot(fit$DE$curve,type="l",lwd=5)
#####
```

---

FMA.historical.boot      *Functional mediation analysis under historical influence regression model with point-wise bootstrap confidence interval*

---

### Description

This function performs functional mediation regression under the historical influence model with given tuning parameter. Point-wise confidence bands are obtained from bootstrap.

**Usage**

```
FMA.historical.boot(Z, M, Y, delta.grid1 = 1, delta.grid2 = 1, delta.grid3 = 1,
  intercept = TRUE, basis1 = NULL, Ld2.basis1 = NULL, basis2 = NULL, Ld2.basis2 = NULL,
  basis.type = c("fourier"), nbasis1 = 3, nbasis2 = 3,
  timeinv = c(0, 1), timegrids = NULL,
  lambda1.m = 0.01, lambda2.m = 0.01, lambda1.y = 0.01, lambda2.y = 0.01,
  sims = 1000, boot = TRUE, boot.ci.type = c("bca", "perc"),
  conf.level = 0.95, verbose = TRUE)
```

**Arguments**

Z	a data matrix. Z is the treatment trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
M	a data matrix. M is the mediator trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
Y	a data matrix. Y is the outcome trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
delta.grid1	a number indicates the width of treatment-mediator time interval in the mediator model.
delta.grid2	a number indicates the width of treatment-outcome time interval in the outcome model.
delta.grid3	a number indicates the width of mediator-outcome time interval in the outcome model.
intercept	a logic variable. Default is TRUE, an intercept term is included in the regression model.
basis1	a data matrix. Basis function on the $s$ domain used in the functional data analysis. The number of columns is the number of basis function considered. If <code>basis = NULL</code> , Fourier basis functions will be generated.
Ld2.basis1	a data matrix. The second derivative of the basis function on the $s$ domain. The number of columns is the number of basis function considered. If <code>Ld2.basis = NULL</code> , the second derivative of Fourier basis functions will be generated.
basis2	a data matrix. Basis function on the $t$ domain used in the functional data analysis. The number of columns is the number of basis function considered. If <code>basis = NULL</code> , Fourier basis functions will be generated.
Ld2.basis2	a data matrix. The second derivative of the basis function on the $t$ domain. The number of columns is the number of basis function considered. If <code>Ld2.basis = NULL</code> , the second derivative of Fourier basis functions will be generated.
basis.type	a character of basis function type. Default is Fourier basis ( <code>basis.type = "fourier"</code> ).
nbasis1	an integer, the number of basis function on the $s$ domain included. If <code>basis1</code> is provided, this argument will be ignored.
nbasis2	an integer, the number of basis function on the $t$ domain included. If <code>basis2</code> is provided, this argument will be ignored.

timeinv	a numeric vector of length two, the time interval considered in the analysis. Default is (0,1).
timegrids	a numeric vector of time grids of measurement. If timegrids = NULL, it is assumed the between measurement time interval is constant.
lambda1.m	a numeric vector of tuning parameter values on the $s$ domain in the mediator model.
lambda2.m	a numeric vector of tuning parameter values on the $t$ domain in the mediator model.
lambda1.y	a numeric vector of tuning parameter values on the $s$ domain in the outcome model.
lambda2.y	a numeric vector of tuning parameter values on the $t$ domain in the outcome model.
sims	an integer indicating the number of simulations for inference.
boot	a logical value, indicating whether or not bootstrap should be used. Default is TRUE.
boot.ci.type	a character of confidence interval method. boot.ci.type = "bca" bias corrected confidence interval; boot.ci.type = "perc" percentile confidence interval.
conf.level	a number of significance level. Default is 0.95.
verbose	a logical value, indicating whether print out bootstrap replications.

### Details

The historical influence mediation model is

$$M(t) = \int_{\Omega_t^1} Z(s)\alpha(s,t)ds + \epsilon_1(t),$$

$$Y(t) = \int_{\Omega_t^2} Z(s)\gamma(s,t)ds + \int_{\Omega_t^3} M(s)\beta(s,t)ds + \epsilon_2(t),$$

where  $\alpha(s,t)$ ,  $\beta(s,t)$ ,  $\gamma(s,t)$  are coefficient curves;  $\Omega_t^j = [(t - \delta_j) \vee 0, t]$  for  $j = 1, 2, 3$ . The model coefficient curves are estimated by minimizing the penalized  $L_2$ -loss.

### Value

alpha	a list of output for $\alpha$ estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve
gamma	a list of output for $\gamma$ estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve

beta	a list of output for $\beta$ estimate coefficients: the result of the coefficient estimates corresponding to the basis function curve: the point-wise estimate of the coefficient curve
IE	a list of output for indirect effect estimate curve: the point-wise estimate of the coefficient curve
DE	a list of output for direct effect estimate curve: the point-wise estimate of the coefficient curve

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 Brian Caffo, Johns Hopkins University, <bcaffo@gmail.com>

**References**

Zhao et al. (2017). *Functional Mediation Analysis with an Application to Functional Magnetic Resonance Imaging Data*. arXiv preprint arXiv:1805.06923.

**Examples**

```
#####
# Historical influence functional mediation model
data(env.historical)
Z<-get("Z",env.historical)
M<-get("M",env.historical)
Y<-get("Y",env.historical)

# consider Fourier basis
fit.boot<-FMA.historical.boot(Z,M,Y,delta.grid1=3,delta.grid2=3,delta.grid3=3,
  intercept=FALSE,timeinv=c(0,300))

#####
```

---

FMA.historical.CV

*Functional mediation analysis under historical influence model*


---

**Description**

This function performs functional mediation regression under the historical influence model. Tuning parameter is chosen based on cross-validation.

**Usage**

```
FMA.historical.CV(Z, M, Y, delta.grid1 = 1, delta.grid2 = 1, delta.grid3 = 1,
  intercept = TRUE, basis1 = NULL, Ld2.basis1 = NULL, basis2 = NULL, Ld2.basis2 = NULL,
  basis.type = c("fourier"), nbasis1 = 3, nbasis2 = 3,
  timeinv = c(0, 1), timegrids = NULL, lambda1 = NULL, lambda2 = NULL, nolds = 5)
```

**Arguments**

Z	a data matrix. Z is the treatment trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
M	a data matrix. M is the mediator trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
Y	a data matrix. Y is the outcome trajectory in the mediation analysis. The number of rows is the number of subjects, and the number of columns is the number of measured time points.
delta.grid1	a number indicates the width of treatment-mediator time interval in the mediator model.
delta.grid2	a number indicates the width of treatment-outcome time interval in the outcome model.
delta.grid3	a number indicates the width of mediator-outcome time interval in the outcome model.
intercept	a logic variable. Default is TRUE, an intercept term is included in the regression model.
basis1	a data matrix. Basis function on the $s$ domain used in the functional data analysis. The number of columns is the number of basis function considered. If basis = NULL, Fourier basis functions will be generated.
Ld2.basis1	a data matrix. The second derivative of the basis function on the $s$ domain. The number of columns is the number of basis function considered. If Ld2.basis = NULL, the second derivative of Fourier basis functions will be generated.
basis2	a data matrix. Basis function on the $t$ domain used in the functional data analysis. The number of columns is the number of basis function considered. If basis = NULL, Fourier basis functions will be generated.
Ld2.basis2	a data matrix. The second derivative of the basis function on the $t$ domain. The number of columns is the number of basis function considered. If Ld2.basis = NULL, the second derivative of Fourier basis functions will be generated.
basis.type	a character of basis function type. Default is Fourier basis (basis.type = "fourier").
nbasis1	an integer, the number of basis function on the $s$ domain included. If basis1 is provided, this argument will be ignored.
nbasis2	an integer, the number of basis function on the $t$ domain included. If basis2 is provided, this argument will be ignored.
timeinv	a numeric vector of length two, the time interval considered in the analysis. Default is (0,1).

timegrids	a numeric vector of time grids of measurement. If timegrids = NULL, it is assumed the between measurement time interval is constant.
lambda1	a numeric vector of tuning parameter values on the $s$ domain.
lambda2	a numeric vector of tuning parameter values on the $t$ domain.
nfolds	a number gives the number of folds in cross-validation.

### Details

The historical influence mediation model is

$$M(t) = \int_{\Omega_t^1} Z(s)\alpha(s,t)ds + \epsilon_1(t),$$

$$Y(t) = \int_{\Omega_t^2} Z(s)\gamma(s,t)ds + \int_{\Omega_t^3} M(s)\beta(s,t)ds + \epsilon_2(t),$$

where  $\alpha(s,t)$ ,  $\beta(s,t)$ ,  $\gamma(s,t)$  are coefficient curves;  $\Omega_t^j = [(t - \delta_j) \vee 0, t]$  for  $j = 1, 2, 3$ . The model coefficient curves are estimated by minimizing the penalized  $L_2$ -loss. Tuning parameter  $\lambda$  controls the smoothness of the estimated curves, and is chosen by cross-validation.

### Value

basis1	the basis functions on the $s$ domain used in the analysis.
basis2	the basis functions on the $t$ domain used in the analysis.
M	a list of output for the mediator model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of M lambda1: the chosen $\lambda$ value on the $s$ domain lambda2: the chosen $\lambda$ value on the $t$ domain
Y	a list of output for the outcome model coefficient: the estimated coefficient with respect to the basis function curve: the estimated coefficient curve fitted: the fitted value of Y lambda1: the chosen $\lambda$ value on the $s$ domain lambda2: the chosen $\lambda$ value on the $t$ domain
IE	a list of output for the indirect effect comparing $Z_1(t) = 1$ versus $Z_0(t) = 0$ curve: the estimated causal curve
DE	a list of output for the direct effect comparing $Z_1(t) = 1$ versus $Z_0(t) = 0$ curve: the estimated causal curve

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## References

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## Examples

```
#####  
# Historical influence functional mediation model  
data(env.historical)  
Z<-get("Z",env.historical)  
M<-get("M",env.historical)  
Y<-get("Y",env.historical)  
  
# consider Fourier basis  
fit<-FMA.historical.CV(Z,M,Y,delta.grid1=3,delta.grid2=3,delta.grid3=3,  
  intercept=FALSE,timeinv=c(0,300))  
  
#####
```

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