Package 'mpersonalized'

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Description More about what it does (maybe more than one line) Use four spaces when indenting paragraphs within the Description.
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admm_optim lambda_estimate mpersonalized mpersonalized_cv plot.mp plot.mp_cv predict.mp predict.mp predict.mp_cv simulated_dataset
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 ${\it admm_optim} \qquad {\it ADMM\ Algorithm\ for\ Meta-analysis/Multiple\ Outcomes\ Personalized} \\ {\it Medicine}$

Usage

```
admm_optim(x, y, p, q, lambda1, lambda2, lambda3, abs.tol = 1e-05,
  rel.tol = 1e-05, maxit = 500L, rho = NULL)
```

lambda_estimate

Default Penalty Parameter Sequence if Not Given

Usage

```
lambda_estimate(modelXlist, modelYlist, penalty, unique_rule, alpha,
   num_lambda1, num_lambda2, num_unique_rule_lambda)
```

Arguments

modelXlist the Xlist in contrast framework after standardization modelYlist the Ylist in contrast framework after standardization

penalty penalty type

unique_rule whether a unique treatment rule is required

Value

estimated lambda for required penalty if not provided

mpersonalized

Meta-analysis/Multiple Outcomes for Personalized Medicine

Usage

```
mpersonalized(problem = c("meta-analysis", "multiple outcomes"), X, Trt,
   P = NULL, Xlist, Ylist, Trtlist, Plist = replicate(length(Xlist), NULL,
   simplify = FALSE), typelist = replicate(length(Xlist), "continuous",
   simplify = FALSE), penalty = c("none", "lasso", "GL", "SGL", "fused",
   "lasso+fused", "GL+fused", "SGL+fused"), lambda1 = NULL, lambda2 = NULL,
   unique_rule_lambda = NULL, num_lambda1 = ifelse(!is.null(lambda1),
   length(lambda1), 10), num_lambda2 = ifelse(!is.null(lambda2),
   length(lambda2), 10),
   num_unique_rule_lambda = ifelse(!is.null(unique_rule_lambda),
   length(unique_rule_lambda), 50), alpha = NULL, unique_rule = FALSE,
   admm_control = NULL, contrast_builder_control = NULL)
```

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Arguments

typlelist

problem a character specifiy whether the user want to solve "meta-analysis" or "multiple outcomes" problem. For "meta-analysis" problem, the user should supply Xlist, Ylist, Trtlist and Plist. For "multiple outcomes" problem, the user should supply X, Ylist, Trt and P. Χ the covariate matrix that should be supplied when the problem is "multiple outcomes" with rows indicating subjects and columns indicating covariates. Trt the treatment vector that should be supplied when the problem is "multiple outcomes". It should be coded as 0 or 1. Ρ the propensity score vector when the problem is "multiple outcomes". If not supplied, then study is treated as randomzied trial and the propensity score is estimated as the proportion of 1's in Trt. Xlist a list object with kth element denoting the covariate matrix of study k. This should be supplied when the problem is "meta-analysis". Ylist When the problem is "meta-analysis", Ylist should be a list object with kth element denoting the response vector of study k. When the problem is "multiple outcomes", Ylist should be a list object with kth element denoting the kth Trtlist a list object with kth element denoting the treatment vector of study k (coded as 0 or 1). This should be supplied when the problem is "meta-analysis". Plist a list object with kthe element denoting the propensity score vector of study k. If not supplied, then each study is treated as randomized trial and the corresponding propensity score is estimated as the proportion of 1's in Trt. For different rules, the penalty could be "none", "lasso", "GL", "SGL", "fused", penalty "lasso+fused", "GL+fused", "SGL+fused". For unique rule, the penalty could be "none" or "lasso". lambda1 lambda1 supplied in the framework when different rules are used. lambda2 lambda2 supplied in the framework when different rules are used. unique_rule_lambda λ when unique rule is used. length of the lambda1 sequence and default to be 10 if lambda1 is not provided num_lambda1 num_lambda2 length of the lambda2 sequence and default to be 10 if lambda2 is not provided num_unique_rule_lambda length of the unique_rule_lambda sequence and default to be 50 if unique_rule_lambda is not provided alpha alpha in the framework when different rules are used. a logical value, whether a unique treatment rule is required unique_rule a list of parameters which control the admm algorithm. In admm_control, the admm_control following parameters can be supplied: abs.tol, absolute tolerance; rel.tol, relative tolerance; maxit, maximum number of iterations; rho, Lagrangian paramecontrast_builder_control a list of parameters which control the contrast building process. In contrast_builder_control, the following parameters could be supplied: response_model, this could be "lasso" or "linear"; contrast builder folds, the number of folds used in cross

validation when response_model = "lasso".

a list object with kth element denoting the type of response corresponding to the

kth element in the list Ylist. Each element should be "continuous" or "binary".

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Details

Assume the total number of studies is K. This function is aimed to solve meta-analysis/multiple outcomes problems for personalized medicine based on the following framework:

$$\min_{g_1,\dots,g_K} \frac{1}{2} \sum_{k=1}^K \sum_{i=1}^{n_k} \frac{|\hat{C}_k(X_i)|}{\sum_{i=1}^{n_k} |\hat{C}_k(X_i)|} \left[1\{\hat{C}_k(X_i) > 0\} - g_k(X_i) \right]^2 + h(g_1,\dots,g_K)$$

Here the regularization function h is of the form of a sum of sparse group lasso and fused lasso penalty

$$h = (1 - \alpha)\lambda_1 \sqrt{q} \sum_{j=1}^p \|\beta_j\|_2 + \alpha \lambda_1 \sum_{j=1}^p \|\beta_j\|_1 + \lambda_2 \sum_{j=1}^p \sum_{1 \le a < b \le K} |\beta_{ja} - \beta_{jb}|$$

where
$$\boldsymbol{\beta_j} = (\beta_{j1}, \dots, \beta_{jK})$$

If we would like a unique rule to be obtained, we let $g_1 = \ldots = g_K$ and solve the following question instead

$$\min_{g} \frac{1}{2} \sum_{k=1}^{K} \sum_{i=1}^{n_k} \frac{|\hat{C}_k(X_i)|}{\sum_{i=1}^{n_k} |\hat{C}_k(X_i)|} \left[1\{\hat{C}_k(X_i) > 0\} - g(X_i) \right]^2 + h(g_1, \dots, g_K) + \lambda_{uni} \|\beta\|_1$$

If we want different rules, by setting $\lambda_1, \lambda_2, \alpha$ differently, different penalties can be obtained.

- If $\lambda_1, \lambda_2 \neq 0$ and $\alpha \neq 0$ or 1, the penalty is "SGL+fused".
- If $\lambda_1,\lambda_2\neq 0$ and $\alpha=0$, the penalty is "GL+fused".
- If $\lambda_1, \lambda_2 \neq 0$ and $\alpha = 1$, the penalty is "lasso+fused".
- If $\lambda_1 = 0, \lambda_2 \neq 0$, the penalty is "fused".
- If $\lambda_1 \neq 0, \lambda_2 = 0$ and $\alpha \neq 0$ or 1, the penalty is "SGL".
- If $\lambda_1 \neq 0, \lambda_2 = 0$ and $\alpha = 0$, the penalty is "GL".
- If $\lambda_1 \neq 0, \lambda_2 = 0$ and $\alpha = 1$, the penalty is "lasso".
- If $\lambda_1, \lambda_2 = 0$, there is no penalty.

If we want unique rule,

- If $\lambda \neq 0$, the penalty is "lasso".
- If $\lambda = 0$, there is no penalty.

Value

an S3 object of class "mp", which contains the information of the fitted model. It could be supplied to the predict function

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mpersonalized_cv	Meta-analysis/Multiple Outcomes for Personalized Medicine with Cross Validation	

Usage

```
mpersonalized_cv(problem = c("meta-analysis", "multiple outcomes"), X, Trt,
   P = NULL, Xlist, Ylist, Trtlist, Plist = replicate(length(Xlist), NULL,
   simplify = FALSE), typelist = replicate(length(Xlist), "continuous",
   simplify = FALSE), penalty = c("lasso", "GL", "SGL", "fused", "lasso+fused",
   "GL+fused", "SGL+fused"), lambda1 = NULL, lambda2 = NULL,
   unique_rule_lambda = NULL, num_lambda1 = ifelse(!is.null(lambda1),
   length(lambda1), 10), num_lambda2 = ifelse(!is.null(lambda2),
   length(lambda2), 10),
   num_unique_rule_lambda = ifelse(!is.null(unique_rule_lambda),
   length(unique_rule_lambda), 50), alpha = NULL, unique_rule = FALSE,
   cv_folds = 5, admm_control = NULL, contrast_builder_control = NULL)
```

Arguments

problem	a character specifiy whether you want to solve "meta-analysis" or "multiple outcomes" problem. For "meta-analysis" problem, the user should supply Xlist, Ylist, Trtlist and Plist. For "multiple outcomes" problem, the user should supply X, Ylist, Trt and P.	
X	the covariate matrix that should be supplied when the problem is "multiple out comes" with rows indicating subjects and columns indicating covariates.	
Trt	the treatment vector that should be supplied when the problem is "multiple outcomes". It should be coded as 0 or 1.	
Р	the propensity score vector when the problem is "multiple outcomes". If not supplied, then study is treated as randomzied trial and the propensity score is estimated as the proportion of 1's in Trt.	
Xlist	a list object with k th element denoting the covariate matrix of study k . This should be supplied when the problem is "meta-analysis".	
Ylist	When the problem is "meta-analysis", Ylist should be a list object with k th element denoting the response vector of study k . When the problem is "multiple outcomes", Ylist should be a list object with k th element denoting the k th outcome.	
Trtlist	a list object with k th element denoting the treatment vector of study k (coded as 0 or 1). This should be supplied when the problem is "meta-analysis".	
Plist	a list object with k the element denoting the propensity score vector of study k . If not supplied, then each study is treated as randomized trial and the corresponding propensity score is estimated as the proportion of 1's in Trt.	
penalty	For different rules, the penalty could be "none", "lasso", "GL", "SGL", "fused", "lasso+fused", "GL+fused", "SGL+fused". For unique rule, the penalty could be "none" or "lasso".	
lambda1	lambda1 supplied in the framework when different rules are used.	
lambda2	lambda2 supplied in the framework when different rules are used.	

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unique_rule_lambda

 λ when unique rule is used.

num_lambda1 length of the lambda1 sequence and default to be 10 if lambda1 is not provided num_lambda2 length of the lambda2 sequence and default to be 10 if lambda2 is not provided num_unique_rule_lambda.

num_unique_rule_lambda

length of the unique_rule_lambda sequence and default to be 50 if unique_rule_lambda

is not provided

alpha alpha in the framework when different rules are used.

unique_rule a logical value, whether a unique treatment rule is required cv_folds number of folds needed for cross-validation, default is 5

admm_control a list of parameters which control the admm algorithm. In admm_control, the

following parameters can be supplied: abs.tol, absolute tolerance; rel.tol, relative tolerance; maxit, maximum number of iterations; rho, Lagrangian parame-

ter.

contrast_builder_control

a list of parameters which control the contrast building process. In contrast_builder_control,

the following parameters could be supplied: response_model, this could be "lasso" or "linear"; contrast_builder_folds, the number of folds used in cross

validation when response_model = "lasso".

typlelist a list object with kth element denoting the type of response corresponding to the

kth element in the list Ylist. Each element should be "continuous" or "binary".

Details

This function implments mpersonalized but use cross validatation for the tuning of penalty parameter. The optimal penalty parameter is selected by minimizing

$$\sum_{i=1}^{n_k} \frac{|\hat{C}_k(X_i)|}{\sum_{i=1}^{n_k} |\hat{C}_k(X_i)|} \left[1\{\hat{C}_k(X_i) > 0\} - g_k(X_i) \right]^2$$

in the leave-out fold, where $\hat{C}_k(X_i)$ in the leave-out fold is separately estimated from the training set

Value

an S3 object of class "mp_cv", which contains the information of the model with the best fitted lambda. It can be supplied to the predict function.

plot.mp

Plot for a 'mp' Class Object.

Usage

```
## S3 method for class 'mp'
plot(mp, ind1, ind2, unique_ind)
```

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Arguments

mp the 'mp' class object returned by mpersonalzied in	
ind1	the index of the lambda1 if different rules are used
ind2 the index of the lambda2 if different rules are used	
unique_ind	the index of the unique_rule_lambda if an unique rule is used

Details

This function plots the results for estimated treatment effects. Depending on the received treatment and recommended treatment, the group means of the outcome are computed and the relations between them are plotted. This plot provides a sanity check of the treatment recommendation rule. By specifying the index of the penalty parameters, we can obtain the plots of the corresponding treatment recommendation rule.

Value

a list object with kth element denoting the plot of study k

plot.mp_cv	Plot for a 'mp' Class Object.

Usage

```
## S3 method for class 'mp_cv'
plot(mp_cv)
```

Arguments

mp_cv the 'mp_cv' class object returned by mpersonalzied_cv function

Details

This function plots the results for estimated treatment effects by using the estimated optimal treatment recommendation rule obtained from corss validation.

Value

a list object with kth element denoting the plot of study k

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Description

This function predict the benefit scores and optimal treatment for new patients

Usage

```
## S3 method for class 'mp'
predict(mp, newx = NULL, weight = NULL, overall_rec = TRUE)
```

Arguments

mp the fitted "mp" object returned by "mpersonalized"

news the covariate matrix of the new patients. If newx = NULL, then the prediction is

made for each data set based on the rule of that study/outcome.

weight a weight vector for the overall recommendation. If leave as NULL, a equally

weighted recommendation will be made.

overall_rec a logical value. If TRUE, an overall recommendation will be made weighted by

the "weight" parameter.

Value

a list of results with each element in the list corresponding to the prediciton by each different penalty parameter. Each element in this list contains:

treatment recommended treatment for each patient for each study/outcome. If overall_rec = TRUE,

the weighted overall recommended treatment will be computed as well. If the overall recommend treatment is equal to 0.5, it means the sum of weight is

equal for 0 and 1.

benefit_score $\ \ \$ the benefit score computed from g_1,\ldots,g_K

Description

This function predict the benefit scores and optimal treatment for new patients

Usage

```
## S3 method for class 'mp_cv'
predict(mp_cv, newx = NULL, weight = NULL,
    overall_rec = TRUE)
```

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Arguments

mp_cv the fitted "mp_cv" object returned from "mpersonalized_cv" function

newx the covariate matrix of the new patients. If newx = NULL, then the prediction is

made for each data set based on the rule of that study/outcome.

weight a weight vector for the overall recommendation. If leave as NULL, a equally

weighted recommendation will be made.

overall_rec a logical value. If TRUE, an overall recommendation will be made weighted by

the "weight" parameter.

Value

the prediciton by using the optimal penalty parameter selected by cross validation. It contains:

treatment recommended treatment for each patient for each study/outcome. If overall_rec = TRUE,

the weighted overall recommended treatment will be computed as well. If the overall recommend treatment is equal to 0.5, it means the sum of weight is

equal for 0 and 1.

benefit_score $\ \ \$ the benefit score computed from g_1,\ldots,g_K

Description

This function could generate a simulated dataset for the test and usage of this package.

Usage

```
simulated_dataset(n, sim_seed, problem = c("meta-analysis",
   "multiple outcomes"))
```

Arguments

n sample size

sim_seed the seed supplied to the generation

problem a character specifiy whether to generate simulated data set for "meta-analysis"

or "multiple outcomes" problem.

Value

A simulated dataset.

X1ist a list object with kth element denoting the covariate matrix of study k Y1ist a list object with kth element denoting the response vector of study k

Trtlist a list object with kth element denoting the treatment vector of study k (coded as

0 or 1)

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