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#####
#### INSTRUCTIONS
#####

## In the comments below, cp refers to the change point, w/ is "with", w/o
is "without".

## Data set should have these column names: response, months (in order),
and off_set.

## The researcher should provide the column numbers for the variables that
will be included as predictors in the data set.

## The researcher can provide where to look for the change points: first
and last.

#####

##### MAIN FUNCTION #####
#####

#####

## p_ncp = total prior prob for models w/o cp
## p_cp = total prior prob for models w/ cp
## input_column has the column numbers of the predictors chosen by the
researcher
## first and last are the first and last months to look for a change point,
respectively
## cp_loc is the month of vaccine introduction or any other introduction
of the intervention of interest

model_fit = function(input_column, dataset, first, last, cp_loc)
{

#####
## Create models:
#####

if(sum(input_column[1:length(input_column)] == 0) >= 1 )
  n_var = 0
else
  n_var = length(input_column)

## Add 2 more variables: cp as an intercept and cp as slope, cp ones will
be the last two variables.

n_var = n_var + 2

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var.mat <- matrix(NA, 2^(n_var*3), n_var)
for (i in 1:n_var)
    var.mat[, i] <- sample(c(0, 1), 2^(n_var*3), replace = T)

var.mat <- unique(var.mat)

## Models w/ cp as intercept and cp as slope are excluded. A model can either
have cp as intercept or cp as slope.

cancel.out = rep(0, dim(var.mat)[1])

for(i in 1:dim(var.mat)[1])
{
    if(var.mat[i, n_var] == 1 & var.mat[i, (n_var - 1)] == 1
)
        cancel.out[i] = i
}
var.mat = var.mat[cancel.out == 0, ]

model.matrix <- matrix(NA, length(var.mat[, 1]), length(var.mat[1,
])))

for (i in 1:length(var.mat[, 1])) for (j in 1:length(var.mat[1, ]))
    model.matrix[i, j] = ifelse(var.mat[i, j] == 1, paste(' + vart',
', ', j, ']', sep = ''), '')

## Models w/o any seasonality
models = c()

model.head1 <- "as.integer(response) ~ 1+offset(log(off_set))"
for (i in 1:length(var.mat[, 1]))
    models[i] <- paste(c(model.head1, model.matrix[i, 1:n_var]),
collapse = "")

## Models w/ 12-month seasonality

model.head2 <- "as.integer(response) ~
1+offset(log(off_set))+sin12+cos12"
for (i in 1:length(var.mat[, 1]) )
    models[(i + length(var.mat[, 1]))] <- paste(c(model.head2,
model.matrix[i, 1:n_var]), collapse = "")

## Models w/ 6-month seasonality

model.head3 <- "as.integer(response) ~ 1+offset(log(off_set))+sin6+cos6"

for (i in 1:length(var.mat[, 1]) )
    models[(i + 2 * length(var.mat[, 1]))] <- paste(c(model.head3,

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model.matrix[i, 1:n_var]), collapse = "")

## Models with both 12-month and 6-month seasonality

    model.head4 <- "as.integer(response) ~
1+offset(log(off_set))+sin12+cos12+sin6+cos6"

    for (i in 1:length(var.mat[, 1]) )
        models[(i + 3 * length(var.mat[, 1]))] <- paste(c(model.head4,
model.matrix[i, 1:n_var]), collapse = "")

#####
### MODEL FIT: We are fitting three types of change point models: no change
models, change in mean (aka change in intercept) and change in slope (aka
change in slope)
#####

n_var = n_var - 2

off_set = dataset$off_set

## Models w/ cp: cp_int -> changepoint as intercept, cp_slp->
changepoint as slope

cp_int = seq(1, length(var.mat[, 1]))[var.mat[, (n_var + 1)] == 1]
cp_int_models = c(cp_int, cp_int + length(var.mat[, 1]), cp_int +
2*length(var.mat[, 1]), cp_int + 3*length(var.mat[, 1]))

cp_slp = seq(1, length(var.mat[, 1]))[var.mat[, (n_var + 2)] == 1]
cp_slp_models = c(cp_slp, cp_slp + length(var.mat[, 1]), cp_slp +
2*length(var.mat[, 1]), cp_slp + 3*length(var.mat[, 1]))

cp_models = c(cp_int_models, cp_slp_models)

## Models w/o change point

n_cp_models = seq(1, length(models))[-c(cp_models)]

## Collect all variables besides seasonality into one place: input_column
are the column numbers of the predictors in the dataset provided by the
researcher.

dataset$vart = matrix(NA, nrow = dim(dataset)[1], ncol = (n_var + 2))
{
    if(n_var != 0)
    {
        for(tr in 1:n_var)
            dataset$vart[, tr] = dataset[, input_column[tr]]
        colnames(dataset$vart) =
c(names(dataset)[input_column], "change point as an intercept", "change"

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    point as a slope")
    }
    else
    {
        colnames(dataset$vart) = c("change point as an
intercept", "change point as a slope")
    }
}

# Response

response = dataset$response

### We will need this while distributing the beta estimates to the right
column in "betas" matrix defined below.

names.to.check <- c("(Intercept)","sin6","cos6","sin12","cos12")
for (i in 1:(n_var + 2))
    names.to.check[5 + i] = paste('vart', '[, ', i, ']', sep = '')

### Seasonality terms

dataset$sin12 = sin(2 * 3.14159 * dataset$months/12)
dataset$cos12 = cos(2 * 3.14159 * dataset$months/12)
dataset$sin6 = sin(2 * 3.14159 * dataset$months/6)
dataset$cos6 = cos(2 * 3.14159 * dataset$months/6)

### bics-> collects BIC values, betas -> collects regression coefficient
estimates, sigma.sq -> collects estimated standard errors of the regression
coefficients

### models_inorder -> keeps the order of the models fit

bics = rep(NA, (length(n_cp_models) + (last - first + 1) *
length(cp_models)))
betas_random = matrix(NA, length(bics), ncol =
length(dataset$months))
betas = matrix(0, length(bics), (length(names.to.check)))
sigma.sq = matrix(0, length(bics), (length(names.to.check)))
models_inorder = rep(NA, length(bics))
fitted_gam = matrix(0, ncol = length(bics), nrow = dim(dataset)[1])
fitted_lme = matrix(0, ncol = length(bics), nrow = dim(dataset)[1])
se_fitted_gam = matrix(0, ncol = length(bics), nrow = dim(dataset)[1])
)

## The same model is fit at different change points, so we need to keep
track of for which change point is fit: useful information for calculating
IRR

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change_points = rep(NA, length(bics))

## We are keeping which models have cp as intercept and cp as slope, we
will need this while calculating IRR

cp_slp_m = rep(0, length(bics))
cp_int_m = rep(0, length(bics))
assign("dataset", dataset, envir=globalenv())

## Models w/o cp:

aa = 1
for(i in n_cp_models)
{
    result =
suppressWarnings(try(gamm4(as.formula(models[i]), random=~(1|months),
family = "poisson", data=dataset), silent = TRUE))

    if (class(result)[1] == "try-error" ||
sum(is.na(coef(result$gam))[1:length(coef(result$gam))] == TRUE) >= 1)
    {
        bics[aa] = -100
        models_inorder[aa] = models[i]
        change_points[aa] = 0
        aa = aa + 1
    }
    else
    {

## This part assigns each coefficient to the right column in betas matrix
        similars = rep(0, length(names.to.check))
        for(k in 1:length(names.to.check))
        {
            for(j in 1:length(coef(result$gam)))
            {
                if(names.to.check[k] ==
attributes(coef(result$gam))$names[j])
                    similars[k] = j
            }
        }

        betas[aa, which(similars!=0)] = coef(result$gam)
        sigma.sq[aa, which(similars!=0)] =
diag(vcov(result$gam))
        betas_random[aa, ] = coef(result$mer)$months[,1]
        bics[aa] = BIC(result$mer)
        models_inorder[aa] = models[i]
        change_points[aa] = 0
        fitted_gam[,aa] = fitted(result$gam)
        se_fitted_gam[, aa] = predict(result$gam, se.fit =
TRUE)$se.fit
        fitted_lme[, aa] = fitted(result$mer)
    }
}

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        aa = aa + 1
    }
}

## Models w/ cp:

for(kk in first:last)
{
    dataset$vart[, (n_var + 1)] = (dataset$months>=
dataset$months[kk]) * 1
    dataset$vart[, (n_var + 2)] = (dataset$months>=
dataset$months[kk]) * (dataset$months - dataset$months[kk])

    assign("dataset", dataset, envir=globalenv())
}

## Intercept models

for(i in cp_int_models)
{
    result =
suppressWarnings(try(gamm4(as.formula(models[i]), random=~(1|months),
family = "poisson", data=dataset), silent = TRUE))

    if (class(result)[1] == "try-error" ||
sum(is.na(coef(result$gam))[1:length(coef(result$gam))]==TRUE)>=1 )
    {
        bics[aa] = -100
        models_inorder[aa] = models[i]
        change_points[aa] = kk
        cp_int_m[aa] = 1
        aa = aa + 1

    }
    else
    {
        similars = rep(0, length(names.to.check))
        for(k in 1:length(names.to.check))
        {
            for(j in 1:length(coef(result$gam)))
            {
                if(names.to.check[k] ==
attributes(coef(result$gam))$names[j])
                    similars[k] = j
            }
        }

        betas[aa, which(similars!=0)] = coef(result$gam)
        betas_random[aa, ] = coef(result$mer)$months[,1]

        sigma.sq[aa, which(similars!=0)] =
diag(vcov(result$gam))
    }
}

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        bics[aa] = BIC(result$mer)
        models_inorder[aa] = models[i]
        change_points[aa] = kk
        cp_int_m[aa] = 1
        fitted_gam[, aa] = fitted(result$gam)
        se_fitted_gam[, aa] = predict(result$gam, se.fit =
TRUE) $se.fit
        fitted_lme[, aa] = fitted(result$mer)
        aa = aa + 1
    }
}

## Slope models

for(i in cp_slp_models)
{
    result =
suppressWarnings(try(gamm4(as.formula(models[i]), random=~(1|months),
family = "poisson", data=dataset), silent = TRUE))

    if (class(result)[1] == "try-error" ||
sum(is.na(coef(result$gam)) [1:length(coef(result$gam)) ]==TRUE)>=1)
    {
        bics[aa] = -100
        models_inorder[aa] = models[i]
        cp_slp_m[aa] = 1
        change_points[aa] = kk
        aa = aa + 1
    }
else
{
    similars = rep(0, length(names.to.check))
    for(k in 1:length(names.to.check))
    {
        for(j in 1:length(coef(result$gam)))
        {
            if(names.to.check[k] ==
attributes(coef(result$gam)) $names[j])
                similars[k] = j
        }
    }
    betas[aa, which(similars!=0)] = coef(result$gam)
    betas_random[aa, ] = coef(result$mer)$months[,1]

    sigma.sq[aa, which(similars!=0)] =
diag(vcov(result$gam))
    bics[aa] = BIC(result$mer)
    models_inorder[aa] = models[i]
    change_points[aa] = kk
    cp_slp_m[aa] = 1
    fitted_gam[, aa] = fitted(result$gam)
}
}

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            se_fitted_gam[, aa] = predict(result$gam, se.fit =
TRUE) $se.fit
            fitted_lme[, aa] = fitted(result$mer)
aa = aa + 1

        }
    }
}

if(sum(bics== -100) == length(bics)) cat("Did you forget to load
package GAMM4?")

#####
### Calculate the posteriors
#####

p_ncp = 0.5
p_cp = 0.5

l1 = length(models_inorder[bics!= -100])
l2 = length(n_cp_models)

l3 = length(models_inorder)

l4 = length(n_cp_models[bics[1:length(n_cp_models)]!= -100])
priors = rep(0, l3)
priors[1:l2] = p_ncp/l4
priors[(l2 + 1): length(priors)] = p_cp/(l1-l4)

## Model fits that converge will have posteriors, rest won't.

index = seq(1:length(bics))[bics!= -100]

delta_bics = rep(0, l3)
delta_bics = (bics- min((bics[index])))

posteriors = rep(0, length(bics))
num_posterior = rep(0, length(bics))
for(i in index)
    num_posterior[i] = exp(-0.5 * delta_bics[i]) * priors[i]

posteriors[index] = num_posterior[index]/sum(num_posterior[index])

## Sum of posterior probabilities for each time point:

post_one = rep(0, last)

for(i in first:last)

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post_one[i] = sum(posteriors[change_points == i])

## Sum of posterior probabilities for change point models

posterior_cp = 1 - round(sum(posteriors[change_points == 0 &
bics != -100]), digits = 5)

#####
## COUNTERFACTUALS:
#####

sin12 = sin(2 * 3.14159 * dataset$months/12)
cos12 = cos(2 * 3.14159 * dataset$months/12)
sin6 = sin(2 * 3.14159 * dataset$months/6)
cos6 = cos(2 * 3.14159 * dataset$months/6)

n = dim(dataset)[1]

### IRR value will be stored in this vector:

IRR_fitted = rep(0, n)

## Matrix to store counterfactuals, n is the number of months, each column
## will correspond to a different model.

fitted_gam_ncp = matrix(NA, nrow = dim(fitted_gam)[1], ncol =
dim(fitted_gam)[2])

## Only focus on models that converged:

index = seq(1, length(bics))[bics != -100]

### Matrix for covariates, each model has a different value for change
## points, so we'll fill in the last two columns of X_1 separately for each
## model

X_1 = cbind(rep(1, n), sin12, cos12, sin6, cos6, dataset[, input_column])

cp_loc = cp_loc

for(j in index)
{

## For models with no change point we assigned counterfactual to be equal
## to the ones obtained from the model

if(change_points[j]==0)

fitted_gam_ncp[, j] = fitted_lme[, j]
}

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    else
    {

        cp1 = (dataset$months>= change_points[j]) * 1 ## change
in mean
        cp2 = (dataset$months>= change_points[j]) *
(dataset$months - change_points[j]) ## change in slope

        X_11= cbind(X_1,
cp1*(sum(posteriors[models_inorder==models_inorder[j] &
change_points<=cp_loc])/sum(posteriors[models_inorder==models_inorder[j]])), cp2*(sum(posteriors[models_inorder==models_inorder[j] &
change_points<=cp_loc])/sum(posteriors[models_inorder==models_inorder[j]])))

        fitted_gam_ncp[, j] = exp(betas[j, ] %*% t(X_11) +
betas_random[j, ])*(dataset$off_set)
    }
}

## Model_averaged predictors: weighted_fitted
## Weighted counterfactuals: weighted_fitted_ncp

weighted_fitted = rep(0, dim(dataset)[1])
weighted_fitted_ncp = rep(0, dim(dataset)[1])
weighted_fitted_se = rep(0, dim(dataset)[1])

for(i in 1:n)
{
    weighted_fitted_ncp[i] =
(fitted_gam_ncp[i,index])%*%posterior[index]
    weighted_fitted[i] =
(fitted_lme[i,index])%*%posterior[index]
    weighted_fitted_se[i] =
(se_fitted_gam[i,index])%*%posterior[index]
}

### IRR fitted vector has the IRR values, and IRR_CI_1 and IRR_CI_2 have
the components of our naive confidence interval estimate (Please see
Supplementary methods for details and comparison of this confidence
interval to bootstrap confidence intervals):

IRR_fitted = weighted_fitted/weighted_fitted_ncp
IRR_CI_1 =
exp(log(weighted_fitted)+1.96*weighted_fitted_se)/weighted_fitted_ncp
IRR_CI_2 =
exp(log(weighted_fitted)-1.96*weighted_fitted_se)/weighted_fitted_ncp

```

```

list(bics = bics, betas = betas, sigma.sq = sigma.sq, models_inorder =
models_inorder, change_points = change_points, n_cp_models = n_cp_models,
cp_int_m = cp_int_m, cp_slp_m = cp_slp_m, posteriors = posteriors,
posterior_ncp = 1-posterior_cp, fitted_gam = fitted_gam, fitted_lme =
fitted_lme, se_fitted_gam = se_fitted_gam, response= response,
betas_random = betas_random, IRR_fitted = IRR_fitted, IRR_CI_1 =
IRR_CI_1, IRR_CI_2 = IRR_CI_2, weighted_fitted = weighted_fitted,
weighted_fitted_ncp = weighted_fitted_ncp)

}

```

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#####
### CODE TO RUN:
#####

##packages required
require(gamm4)

## Read the data:

dataset = read.csv('location_of_data')

# Define, input column, first, last, and cp_loc and then run the function
#(The following numbers are specific to the dataset provided):

input_column = 0
first = 3
last = 129
cp_loc = 50
result = model_fit(input_column, dataset, first, last, cp_loc)

##Fitted values versus observed values:

dev.new(width = 9.5, height = 5.7)
plot(dataset$months, result$weighted_fitted, type="l", xlab = "months",
ylab = "", lwd = 2)
lines(dataset$months, dataset$response, col="coral", lwd = 2)
lines(dataset$months, result$weighted_fitted_ncp, col="cornflowerblue",
lwd = 2)
legend("topright", col = c("black", "coral", "cornflowerblue"),
c("Observed values", "Fitted values", "Counterfactual predictions"), lty =
c(1,1, 1), lwd = 2)

### Posterior probability of change point models:

print(paste("Probability of any change in the time series is", 1-
result$posterior_ncp))

###IRR with our naive confidence interval estimate:

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```
dev.new(width = 8.5, height = 4.7)
plot(dataset$months, result$IRR_fitted, type="l", ylim = c(0.6, 1.1), xlab
= "months", ylab = "IRR")
lines(dataset$months, result$IRR_CI_1, lty = 2)
lines(dataset$months, result$IRR_CI_2, lty = 2)
```