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# PairwiseCorrelations-IndependentVariables.R
# Computes Pearson and Spearman correlations and corresponding p-values between all possible pairs of variables
# that were considered as possible independent variables for our main multivariate regression model
# and as possible independent variables for the multivariate regressions in the sensitivity analyses
# Output written to spreadsheet PairwiseCorrelations-IndependentVariables.csv

fulldataset = read.csv(file="fulldataset.csv")

indepvariables =
fulldataset[,c("people_fully_vaccinated_per_hundred", "gdp_per_capita", "cardiovasc_death_rate", "diabetes_prevalence", "life_expectancy",
              "human_development_index", "Stringency", "aged_65_older", "masks.avg", "Obesity.rate", "Gini", "urban_density", "total_tests_per_thousa
nd")]

var1 = character()
var2 = character()
cor_pearson = numeric()
pval_pearson = numeric()
cor_spearman = numeric()
pval_spearman = numeric()

line = 1
for(i in 1:(ncol(indepvariables)-1))
{
  for(j in (i+1):ncol(indepvariables))
  {
    resultpearson = cor.test(indepvariables[,i], indepvariables[,j], method="pearson")
    resultspearman = suppressWarnings(cor.test(indepvariables[,i], indepvariables[,j], method="spearman"))
    var1[line] = names(indepvariables)[i]
    var2[line] = names(indepvariables)[j]
    cor_pearson[line] = resultpearson$estimate
    pval_pearson[line] = resultpearson$p.val
    cor_spearman[line] = resultspearman$estimate
    pval_spearman[line] = resultspearman$p.val
    line = line+1
  }
}

output =
data.frame(var1=var1, var2=var2, cor_pearson=cor_pearson, pval_pearson=pval_pearson, cor_spearman=cor_spearman, pval_spearman=pval_spearman)
output = output[order(abs(output$cor_pearson), decreasing=TRUE),]
write.csv(x=output, file="PairwiseCorrelations-IndependentVariables.csv", row.names=FALSE)

# CorrelationsWithPLAA.R
# Computes Pearson and Spearman correlations and corresponding p-values between the dependent variable PLAA and all variables
# that were considered as possible independent variables for our main multivariate regression model
# and as possible independent variables for the multivariate regressions in the sensitivity analyses
# cardiovascular death rate and life expectancy were substituted by cardlife, which is their principal component
# Output written to spreadsheet CorrelationsWithPLAA.csv

fulldataset = read.csv(file="fulldataset.csv")

# computes principal component of cardiovasc_death_rate and life_expectancy
prcomp_cardlife = prcomp(fulldataset[,c("cardiovasc_death_rate", "life_expectancy")], scale=TRUE)
prcomp1_cardlife = prcomp_cardlife$x[,1] # extracts the principal component
coef_cardlife = prcomp_cardlife$rot[,1] # coefficients used for calculating the principal component
prop1_cardlife = prcomp_cardlife$sdev[1]^2/length(prcomp_cardlife$sdev) # proportion of total variance contained in the principal
component

indepvariables = fulldataset[,c("people_fully_vaccinated_per_hundred", "gdp_per_capita", "diabetes_prevalence",
                              "human_development_index", "Stringency", "aged_65_older", "masks.avg", "Obesity.rate", "Gini", "urban_density", "total_tests_per_thousa
nd")]
indepvariables$cardlife = prcomp1_cardlife

PLAA = fulldataset$Per.Levitt.Age.Adjusted

variable = colnames(indepvariables)
cor_pearson = numeric()
pval_pearson = numeric()
cor_spearman = numeric()
pval_spearman = numeric()

for(i in 1:ncol(indepvariables))
{
  resultpearson = cor.test(PLAA, indepvariables[,i], method="pearson")
  resultspearman = suppressWarnings(cor.test(PLAA, indepvariables[,i], method="spearman"))
  cor_pearson[i] = resultpearson$estimate
  pval_pearson[i] = resultpearson$p.val
  cor_spearman[i] = resultspearman$estimate
  pval_spearman[i] = resultspearman$p.val
}

output =
data.frame(variable=variable, cor_pearson=cor_pearson, pval_pearson=pval_pearson, cor_spearman=cor_spearman, pval_spearman=pval_spearman)
write.csv(x=output, file="CorrelationsWithPLAA.csv", row.names=FALSE)

# MainRegression.R
# Computes the main regression and sensitivity analyses based on bootstrap and robust sandwich standard errors
# Computes all diagnostics for the main regression
# Outputs: MainRegression.txt; summary of main regression (coefficients and standardized coefficients, 95% confidence intervals and p-
values)
# Sensitivity.txt; sensitivity analyses of main regression using bootstrap and heterocedastic-consistent sandwich standard errors
# VariousDiagnostics.txt; multicollinearity check, skewness, excess kurtosis and Shapiro-Wilk p-value for standardized residuals, Breusch-
Pagan test for homocedasticity
# fittedresidual.jpg; scatter plot of fitted values versus residuals
# fittedstdresidual.jpg; scatter plot of fitted values versus standardized residuals
# scalelocation.jpg; scale-location plot (fitted values versus square root of absolute values of standardized residuals)
# qqnormal.jpg; qq plot of standardized residuals against normal distribution
# cooks_distance.jpg; bar plot of Cook's distances

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library(ggplot2)
library(car)
library(sandwich)
library(DescTools)
library(lmtest)
library(betaDelta)
library(betaSandwich)

bootreg = function(y,X,nrep=10^5,conf.level=0.95){
  # computes multivariable linear regression coefficients and standardized coefficients
  # plus corresponding confidence intervals and p-values using bootstrap
  # y = dependent variable;
  # X = independent variables (should be a matrix and should not include a column of 1's for intercept, which is automatically included)
  # nrep = number of bootstraps; conf.level = confidence level for confidence intervals
  # returns a list with fields coef (coefficients), ICLwr, ICupr (confidence interval)
  # std_coef (standardized coefficients), std_IClwr, std_ICupr (confidence interval), pval (p-value)

  sestdcoefs = function(y,X,coefs,sigma2){
    # auxiliary function for function bootreg
    # computes standardized regression coefficients (stdcoef) and corresponding standard errors (se)
    # X should not include a column of 1's for intercept
    # coefs = (nonstandardized) coefficients for regression excluding intercept
    # sigma2 = estimate of residual variance

    X = as.matrix(X)
    SX = var(X)
    SYy = apply(X,2,function(u) cov(u,y))
    SY2 = var(y)
    n = length(y)
    stdcoef = coefs*sqrt(diag(SX))/sqrt(SY2)
    SXinv = solve(SX)
    se2 = diag(SX)*diag(SXinv)*sigma2/((n-3)*SY2) + coefs^2*(diag(SX)*as.vector(coefs%*%SX%*coefs)-diag(SX)*sigma2-SYy^2)/((n-
3)*SY2^2)
    return(list(stdcoef=stdcoef,se=sqrt(se2)))
  }

  oneboot = function(y,X,maincoefs,mainstdcoefs){
    # auxiliary function for function bootreg
    # does one iteration of bootstrap, returns "t-values" for coefficients and standardized coefficients
    # where "t-value" equals the coefficient from bootstrapped sample minus coefficient from main data divided by estimate of standard error
    # a column of 1's must be included as first column of X for intercept

    linreg = function(y,X,maincoefs,mainstdcoefs){
      # does the main calculation for function oneboot

      covunsc = solve(t(X)%*%X)
      coefs = covunsc%*%t(X)%*%y
      resid = y-X%*%coefs
      sigma2 = sum(resid^2)/(nrow(X)-ncol(X))
      se = sqrt(diag(covunsc))*sqrt(sigma2)
      tval = (coefs-maincoefs)/se
      calcstdcoefs = sestdcoefs(y,X[, -1],coefs[-1],sigma2)
      stdtval = (calcstdcoefs$stdcoef-mainstdcoefs)/calcstdcoefs$se
      return(c(tval, stdtval))
    }

    samp = sample.int(nrow(X),nrow(X),replace=TRUE) # bootstrap sample
    return(linreg(y[samp],X[samp,],maincoefs,mainstdcoefs))
  }

  # computes coefficients, standardized coefficients, standard errors and t-values for main data
  reg = summary(lm(y~X))
  maincoefs = reg$coefficients[,1]
  mainse = reg$coefficients[,2]
  maintvals = reg$coefficients[,3]
  calcstdcoefs = sestdcoefs(y,X,maincoefs[-1],reg$sigma^2)
  mainstdcoefs = calcstdcoefs$stdcoef
  mainsestd = calcstdcoefs$se

  X = cbind(1,X)
  # computes bootstrapped "t-values"; first ncol(X) lines of samptvals for regular coefficients (includes intercept)
  # last ncol(X)-1 lines of samptvals for standardized coefficients (excludes intercept)
  samptvals = matrix(replicate(nrep,oneboot(y,X,maincoefs,mainstdcoefs)),nrow=2*ncol(X)-1)

  pvals = sapply(1:ncol(X),function(i) sum(abs(samptvals[i,]) >= abs(maintvals[i]))/nrep)
  q = apply(samptvals,1,function(u) c(quantile(u,(1-conf.level)/2,na.rm=TRUE),quantile(u,(1+conf.level)/2,na.rm=TRUE))) #
quantiles from bootstraps
  return(list(coef=maincoefs,IClwr=maincoefs-mainse*q[2,1:ncol(X)],ICupr=maincoefs-
mainse*q[1,1:ncol(X)],std_coef=c(NA,mainstdcoefs),
std_IClwr=c(NA,mainstdcoefs-mainsestd*q[2,(ncol(X)+1):(2*ncol(X)-1)]),std_ICupr=c(NA,mainstdcoefs-mainsestd*q[1,
(ncol(X)+1):(2*ncol(X)-1)]),pval=pvals)
  }

  bootregression = function(resultreg,nrep=10^5,conf.level=0.95){
    # like function bootreg, but instead of receiving y and X as input
    # receives resultreg, which is the output of an lm command (regression must include intercept)

    return(bootreg(resultreg$model[,1],as.matrix(resultreg$model[, -1]),nrep,conf.level))
  }

  HCregression = function(resultreg,conf.level=0.95){
    # computes multivariable linear regression coefficients and standardized coefficients
    # plus corresponding confidence intervals and p-values using heterocedastic-consistent sandwich standard errors (method HC3)
    # resultreg = output of an lm command; conf.level = confidence level for confidence intervals
    # returns a list with fields coef (coefficients), ICLwr, ICupr (confidence interval)
    # std_coef (standardized coefficients), std_IClwr, std_ICupr (confidence interval), pval (p-value)

    se = sqrt(diag(vcovHC(resultreg,type="HC3")))
    pval = 2*pt(abs(resultreg$coefficients)/se,df=resultreg$df, lower.tail=FALSE)
  }

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radius = se*qt((1+conf.level)/2,df=resultreg$df)
stdreg = BetaHC(resultreg,type="hc3")
stdcoef = c(NA,coef(stdreg))
IC = confint(stdreg,level=conf.level)
return(list(coef=resultreg$coefficients,IClwr=resultreg$coefficients-radius,ICupr=resultreg$coefficients+radius,
            std_coef=stdcoef,std_IClwr=c(NA,IC[,1]),std_ICupr=c(NA,IC[,2]),pval=pval))
}

stdresiduals = function(resultreg){
# computes standardized regression residuals
# resultreg = output of an lm command

h = lm.influence(resultreg)$shat
return(resultreg$residuals/(sqrt(1-h)*summary(resultreg)$sigma))
}

checkmulticol = function(resultreg){
# multicollinearity check
# computes R squared for the regression of each independent variable against all others
# resultreg = output of an lm command (regression must include intercept)

X = as.matrix(resultreg$model[, -1])
if(ncol(X) == 1)
{
return(NULL)
}
R2 = numeric(ncol(X))
names(R2) = colnames(X)
for(i in 1:ncol(X))
{
y = X[,i]
X1 = X[, -i]
R2[i] = summary(lm(y~X1))$r.squared
}
return(R2)
}

predictedR2 = function(resultreg){
# computes predicted R squared
# resultreg = output of an lm command

h = lm.influence(resultreg)$shat
PRESS = sum((resultreg$residuals/(1-h))^2)
y = resultreg$model[,1]
TSS = sum((y-mean(y))^2)
return(1-PRESS/TSS)
}

summaryreg = function(resultreg,varnames=names(resultreg$coefficients)[-1],conf.level=0.95){
# computes multivariable linear regression coefficients and standardized coefficients
# plus corresponding confidence intervals and p-values
# resultreg = output of an lm command; varnames = names of independent variables (excluding intercept)
# conf.level = confidence level for confidence intervals
# returns a data frame with fields coef (coefficients), IClwr, ICupr (confidence interval)
# std_coef (standardized coefficients), std_IClwr, std_ICupr (confidence interval), pval (p-value)

coef = resultreg$coefficients
IClwr = coef-summary(resultreg)$coefficients[,2]*qt((1+conf.level)/2,df=resultreg$df)
ICupr = coef+summary(resultreg)$coefficients[,2]*qt((1+conf.level)/2,df=resultreg$df)
stdreg = BetaDelta(resultreg,type="mvn")
stdcoef = coef(stdreg)
stdIC = confint(stdreg,level=conf.level)
result = data.frame(coef=coef,IClwr=IClwr,ICupr=ICupr,
                    std_coef=c(NA,stdcoef),std_IClwr=c(NA,stdIC[,1]),std_ICupr=c(NA,stdIC[,2]),pval=summary(resultreg)$coefficients[,4])
rownames(result) = c("Intercept",varnames)
return(result)
}

output_summaryreg = function(resultreg,varnames=names(resultreg$coefficients)[-1],conf.level=0.95){
# generates text output for function summaryreg, includes adjusted and predicted R squared
# resultreg = output of an lm command; varnames = names of independent variables (excluding intercept)
# conf.level = confidence level for confidence intervals

ncols = getOption("width")
options(width=300)
output = capture.output(summaryreg(resultreg,varnames,conf.level))
output = c(output,"",paste("Adjusted R squared:",summary(resultreg)$adj.r.squared),paste("Predicted R
squared:",predictedR2(resultreg)))
options(width=ncols)
return(output)
}

output_bootregression = function(resultreg,varnames=names(resultreg$coefficients)[-1],nrep=10^5,conf.level=0.95){
# generates text output for function bootregression
# resultreg = output of an lm command (regression must include intercept); varnames = names of independent variables (excluding intercept)
# nrep = number of bootstraps; conf.level = confidence level for confidence intervals

result = data.frame(bootregression(resultreg,nrep,conf.level))
rownames(result) = c("Intercept",varnames)

ncols = getOption("width")
options(width=300)
output = capture.output(result)
options(width=ncols)
return(output)
}

output_HCregression = function(resultreg,varnames=names(resultreg$coefficients)[-1],conf.level=0.95){
# generates text output for function HCregression

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# resultreg = output of an lm command (regression must include intercept); varnames = names of independent variables (excluding intercept)
# conf.level = confidence level for confidence intervals

    result = data.frame(HCregression(resultreg, conf.level))
    rownames(result) = c("Intercept", varnames)

    ncols = getOption("width")
    options(width=300)
    output = capture.output(result)
    options(width=ncols)
    return(output)
}

plot_fittedresidual = function(resultreg, abbreviation=NA){
# generates plot of fitted values versus residuals
# resultreg = output of an lm command; abbreviation = labels for points

    if(is.na(abbreviation[1]))
    {
        abbreviation = rep("", nrow(resultreg$model))
    }
    dataplot = data.frame(labels=abbreviation, x=resultreg$fitted.values, y=resultreg$residuals)
    return(ggplot(dataplot, aes(x=x, y=y, label=labels))+geom_point()+geom_text(vjust=1.5)
        +labs(x="fitted values", y="residuals")+stat_smooth(formula=y~x, method="loess"))
}

plot_fittedstdresidual = function(resultreg, abbreviation=NA){
# generates plot of fitted values versus standardized residuals
# resultreg = output of an lm command; abbreviation = labels for points

    if(is.na(abbreviation[1]))
    {
        abbreviation = rep("", nrow(resultreg$model))
    }
    dataplot = data.frame(labels=abbreviation, x=resultreg$fitted.values, y=stdresiduals(resultreg))
    return(ggplot(dataplot, aes(x=x, y=y, label=labels))+geom_point()+geom_text(vjust=1.5)
        +labs(x="fitted values", y="standardized residuals")+stat_smooth(formula=y~x, method="loess"))
}

plot_scalelocation = function(resultreg, abbreviation=NA){
# generates scale-location plot (fitted values versus square root of absolute values of standardized residuals)
# resultreg = output of an lm command; abbreviation = labels for points

    if(is.na(abbreviation[1]))
    {
        abbreviation = rep("", nrow(resultreg$model))
    }
    dataplot = data.frame(labels=abbreviation, x=resultreg$fitted.values, y=sqrt(abs(stdresiduals(resultreg))))
    return(ggplot(dataplot, aes(x=x, y=y, label=labels))+geom_point()+geom_text(vjust=1.5)
        +labs(x="fitted values", y="sqrt of abs value of standardized residuals")+stat_smooth(formula=y~x, method="loess"))
}

plot_qqnormal = function(resultreg){
# generates qq plot of standardized residuals against a normal distribution
# resultreg = output of an lm command

    dataplot = data.frame(y=stdresiduals(resultreg))
    return(ggplot(dataplot, aes(sample=y))+stat_qq()+stat_qq_line()+labs(x="theoretical quantile", y="standardized residuals"))
}

plot_cook = function(resultreg, abbreviation=NA){
# generates bar plot of Cook's distances
# resultreg = output of an lm command; abbreviation = labels for bars

    if(is.na(abbreviation[1]))
    {
        abbreviation = rep("", nrow(resultreg$model))
    }
    dataplot = data.frame(labels=abbreviation, y=cooks.distance(resultreg))
    return(ggplot(dataplot, aes(x=labels, y=y))+labs(x="countries", y="Cook's distance")+geom_bar(stat="identity"))
}

# beginning of main code

fulldataset = read.csv(file="fulldataset.csv")

# computes principal component of cardiovasc_death_rate and life_expectancy
prcomp_cardlife = prcomp(fulldataset[,c("cardiovasc_death_rate", "life_expectancy")], scale=TRUE)
prcomp1_cardlife = prcomp_cardlife$x[,1] # extracts the principal component
coef_cardlife = prcomp_cardlife$rot[,1] # coefficients used for calculating the principal component
prop1_cardlife = prcomp_cardlife$sdev[1]^2/length(prcomp_cardlife$sdev) # proportion of total variance contained in the principal component

# data for main regression
regdata = fulldataset[,c("people_fully_vaccinated_per_hundred", "human_development_index", "masks.avg", "Per.Levitt.Age.Adjusted")]
regdata$cardlife = prcomp1_cardlife

mainreg = lm(Per.Levitt.Age.Adjusted~people_fully_vaccinated_per_hundred+human_development_index+cardlife+masks.avg, data=regdata)

output = output_summaryreg(mainreg)
outputfile = file("MainRegression.txt")
writeLines(output, outputfile)
close(outputfile)

separator = paste(rep("=", 40), collapse="")

set.seed(123) # for reproducibility of bootstrap
output = c("Bootstrap:", output_bootstrapregression(mainreg), separator, "Heterocedastic-consistent sandwich standard
erros:", output_HCregression(mainreg))

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outputfile = file("Sensitivity.txt")
writeLines(output,outputfile)
close(outputfile)

output = c("R2 of regression of each independent variable against all other independent
variables:",capture.output(checkmulticol(mainreg)))
output = c(output,separator,"Skewness for standardized
residuals:",capture.output(Skew(stdresiduals(mainreg),method=2,conf.level=0.95,ci.type="bca",R=10^5)))
output = c(output,separator,"Excess kurtosis for standardized
residuals:",capture.output(Kurt(stdresiduals(mainreg),method=2,conf.level=0.95,ci.type="bca",R=10^5)))
output = c(output,separator,"Shapiro-Wilk for standardized residuals:",capture.output(shapiro.test(stdresiduals(mainreg))))
output = c(output,separator,"Breusch-Pagan test for homocedasticity:",capture.output(bptest(mainreg,studentize=TRUE)))
outputfile = file("VariousDiagnostics.txt")
writeLines(output,outputfile)
close(outputfile)

plot_fittedresidual(mainreg,fulldataset$abbreviation)
ggsave(filename="fittedresidual.jpg",device="jpeg",width=14.23,height=6.77)

plot_fittedstdresidual(mainreg,fulldataset$abbreviation)
ggsave(filename="fittedstdresidual.jpg",device="jpeg",width=14.23,height=6.77)

plot_scalelocation(mainreg,fulldataset$abbreviation)
ggsave(filename="scalelocation.jpg",device="jpeg",width=14.23,height=6.77)

plot_qqnormal(mainreg)
ggsave(filename="qqnormal.jpg",device="jpeg",width=14.23,height=6.77)

plot_cook(mainreg,fulldataset$abbreviation)
ggsave(filename="cooks_distance.jpg",device="jpeg",width=14.23,height=6.77)

# OtherRegressions.R
# Computes all possible regressions obtained by adding at most 3 extra independent variables to the 4 independent variables considered in
the main regression
# computes coefficients, standardized coefficients, 95% confidence intervals and p-values
# Outputs: OtherRegressions.txt; text file with results of all regressions
# OtherRegressions.csv; spreadsheet with results of all regressions

library(ggplot2)
library(car)
library(betaDelta)

predictedR2 = function(resultreg){
# computes predicted R squared
# resultreg = output of an lm command

  h = lm.influence(resultreg)$shat
  PRESS = sum((resultreg$residuals/(1-h))^2)
  y = resultreg$model[,1]
  TSS = sum((y-mean(y))^2)
  return(1-PRESS/TSS)
}

summaryreg = function(resultreg,varnames=names(resultreg$coefficients)[-1],conf.level=0.95){
# computes multivariable linear regression coefficients and standardized coefficients
# plus corresponding confidence intervals and p-values
# resultreg = output of an lm command; varnames = names of independent variables (excluding intercept)
# conf.level = confidence level for confidence intervals
# returns a data frame with fields coef (coefficients), IC1lwr, ICupr (confidence interval)
# std_coef (standardized coefficients), std_IC1lwr, std_ICupr (confidence interval), pval (p-value)

  coef = resultreg$coefficients
  IC1lwr = coef-summary(resultreg)$coefficients[,2]*qt((1+conf.level)/2,df=resultreg$df)
  ICupr = coef+summary(resultreg)$coefficients[,2]*qt((1+conf.level)/2,df=resultreg$df)
  stdreg = BetaDelta(resultreg,type="mvn")
  stdcoef = coef(stdreg)
  stdIC = confint(stdreg,level=conf.level)
  result = data.frame(coef=coef,IC1lwr=IC1lwr,ICupr=ICupr,
    std_coef=c(NA,stdcoef),std_IC1lwr=c(NA,stdIC[,1]),std_ICupr=c(NA,stdIC[,2]),pval=summary(resultreg)$coefficients[,4])
  rownames(result) = c("Intercept",varnames)
  return(result)
}

output_summaryreg = function(resultreg,varnames=names(resultreg$coefficients)[-1],conf.level=0.95){
# generates text output for function summaryreg, includes adjusted and predicted R squared
# resultreg = output of an lm command; varnames = names of independent variables (excluding intercept)
# conf.level = confidence level for confidence intervals

  ncols = getOption("width")
  options(width=300)
  output = capture.output(summaryreg(resultreg,varnames,conf.level))
  output = c(output,"",paste("Adjusted R squared:",summary(resultreg)$adj.r.squared),paste("Predicted R
squared:",predictedR2(resultreg)))
  options(width=ncols)
  return(output)
}

# beginning of main code

fulldataset = read.csv(file="fulldataset.csv")

# computes principal component of cardiovasc_death_rate and life_expectancy
prcomp_cardlife = prcomp(fulldataset[,c("cardiovasc_death_rate","life_expectancy")],scale=TRUE)
prcomp1_cardlife = prcomp_cardlife$x[,1] # extracts the principal component
coef_cardlife = prcomp_cardlife$rot[,1] # coefficients used for calculating the principal component
prop1_cardlife = prcomp_cardlife$sdev[1]^2/length(prcomp_cardlife$sdev) # proportion of total variance contained in the principal
component

# four variables that must be included as independent variables in the sensitivity analyses regressions

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mustinclude = fulldataset[,c("masks.avg", "people_fully_vaccinated_per_hundred", "human_development_index")]
mustinclude$cardlife = prcomp1_cardlife
mustinclude = as.matrix(mustinclude)

# other variables that can be included as independent variables in the sensitivity analyses regressions - at most 3 at a time
others =
as.matrix(fulldataset[,c("Stringency", "total_tests_per_thousand", "aged_65_older", "urban_density", "diabetes_prevalence", "Obesity.rate", "Gini")])

PLAA = fulldataset$Per.Levitt.Age.Adjusted

# generates data frame corresponding to all possible subsets of size at most 3 from the "other" variables
possibilities = expand.grid(replicate(ncol(others),c(0,1),simplify=FALSE))
possibilities = possibilities[apply(possibilities,1,sum)<=3,]

outputsheet = matrix(nrow=nrow(possibilities), ncol=4+(ncol(mustinclude)+ncol(others))*7+2)
colnamesheet = c("Intercept", "Intercept_IClwr", "Intercept_ICupr", "Intercept_pval")
colnamesheet = c(colnamesheet,
  unlist(lapply(c(colnames(mustinclude), colnames(others)),
    function(x) sapply(c("", "IClwr", "ICupr", "_std", "_std_IClwr", "_std_ICupr", "_pval"), function(y) paste(x,y, sep="")))))
colnamesheet = c(colnamesheet, "adj_R2", "pred_R2")
colnames(outputsheet) = colnamesheet

separator = paste(rep("=", 40), collapse="")

legend_textfile = c("Legend:", "coef: coefficient value", "IClwr: lower extremity of 95% confidence interval for coefficient",
  "ICupr: upper extremity of 95% confidence interval for coefficient",
  "std_coef: standardized coefficient", "std_IClwr: lower extremity of 95% confidence interval for standardized coefficient",
  "std_ICupr: upper extremity of 95% confidence interval for standardized coefficient", "pval: p-value")

legend_sheet = c("Legend:", "Column with variable name (varname) alone is the coefficient",
  "varname_IClwr is the lower extremity of 95% confidence interval for the coefficient",
  "varname_ICupr is the upper extremity of 95% confidence interval for the coefficient",
  "varname_std is the standardized coefficient",
  "varname_std_IClwr is the lower extremity of the 95% confidence interval for the standardized coefficient",
  "varname_std_ICupr is the upper extremity of the 95% confidence interval for the standardized coefficient",
  "varname_pval is the p-value",
  "adj_R2 is the adjusted R squared",
  "pred_R2 is the predicted R squared")

outputtext = c()
for(i in 1:nrow(possibilities))
{
  X = cbind(mustinclude, others[, possibilities[i,]==1])
  regression = lm(PLAA~X)
  sumreg = summaryreg(regression)
  sheetline = unlist(sumreg[1,c("coef", "IClwr", "ICupr", "pval")])
  sheetline = c(sheetline, unlist(lapply(1:ncol(mustinclude), function(j) sumreg[j+1,])))
  k = ncol(mustinclude)+1
  for(j in 1:ncol(others))
  {
    if(possibilities[i,j] == 1)
    {
      k = k+1
      sheetline = c(sheetline, unlist(sumreg[k,]))
    } else
    {
      sheetline = c(sheetline, rep(NA, 7))
    }
  }
  sheetline = c(sheetline, summary(regression)$adj.r.squared, predictedR2(regression))
  outputsheet[i,] = sheetline
  varnames = c(colnames(mustinclude), colnames(others)[possibilities[i,]==1])
  outputtext = c(outputtext, output_summaryreg(regression, varnames), separator)
}

outputfile = file("OtherRegressions.txt")
writeLines(c(legend_textfile, separator, outputtext), outputfile)
close(outputfile)

write.csv(x=data.frame(outputsheet), file="OtherRegressions.csv", na="", row.names=FALSE)
outputfile = file("OtherRegressions.csv", "a")
writeLines(c("", legend_sheet), outputfile)
close(outputfile)

# PowerCalculation.R
# Power calculations for sensitivity analyses regressions
# uses main regression as "pilot study" for obtaining estimates of true values of relevant parameters
# Output: PowerCalculation.txt; table with power calculations for detecting a nonzero coefficient for masks
# as a function of the number of independent variables included in the regression

powerlinreg = function(n,p,beta,Risq,Rsq,sig.level=0.05){
  # power calculation for linear regression Y=beta_0+beta_1X_1+...+beta_pX_p+epsilon
  # testing the null hypothesis beta_i=0 (for some fixed i in {1,...,p})
  # n = sample size, p = number of regressors
  # beta = beta_i sd(X_i)/sd(Y) = standardized regression coefficient for X_i
  # Risq = R-squared when X_i is regressed against all other X_j
  # Rsq = 1 - Var(epsilon)/Var(Y) = R-squared when Y is regressed against X_1, ..., X_p
  # sig.level = significance level (two-tailed p-value is considered)
  # power calculation is exact if X_1, ..., X_p are regarded as a fixed nonrandom sample of size n;
  # for exactness to hold, Risq should be the sample adjusted R-squared and sd(X_i) should be the sqrt of the unbiased sample variance of
  X_i
  df = n-p-1
  tcrit = qt(sig.level/2, df=df, lower.tail=FALSE)
  ncp = sqrt(1-Risq)*sqrt(n-p)*beta/sqrt(1-Rsq)
  return(pt(-tcrit, df=df, ncp=ncp)+pt(tcrit, df=df, ncp=ncp, lower.tail=FALSE))
}

```

```

sampleadjR2var = function(nrep,n,p,Rsq,varY){
# creates a sample of size nrep from the distribution of adjusted R squared and the unbiased sample variance of Y
# assuming Y is a sample of size n of  $Y = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p + \epsilon$ , with  $\epsilon$  normal with mean zero
#  $X_1, \dots, X_p$  multivariate normal;  $\epsilon$  independent of  $(X_1, \dots, X_p)$ 
# varY = true variance of Y
# Rsq = true R squared =  $\text{var}(\beta_1 X_1 + \dots + \beta_p X_p) / \text{var}(Y)$ 

  if(p == 0)
  {
    chi = rchisq(nrep,df=n-1)
    result = rbind(0,chi*varY*(1-Rsq)/(n-1))
  }else
  {
    chi1 = rchisq(nrep,df=n-p-1)
    chi2 = rchisq(nrep,df=p,ncp=Rsq/(1-Rsq)*rchisq(nrep,df=n-1))
    sumchi = chi1+chi2
    result = rbind(1-(chi1/sumchi)*((n-1)/(n-p-1)),sumchi*varY*(1-Rsq)/(n-1))
  }
  row.names(result) = c("adjRsq","hatvarY")
  return(result)
}

powerlinreg_randomX = function(n,p,beta,Risq,Rsq,nrep=10^5,sig.level=0.05){
# power calculation for linear regression  $Y = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p + \epsilon$  based on Monte Carlo simulation
# nrep = number of simulations
# testing the null hypothesis  $\beta_i = 0$  (for some fixed i in  $\{1, \dots, p\}$ )
# n = sample size, p = number of regressors
# beta =  $\beta_i$  sd( $X_i$ )/sd(Y) = standardized regression coefficient for  $X_i$ 
# Risq = true R-squared when  $X_i$  is regressed against all other  $X_j$ 
# Rsq =  $1 - \text{Var}(\epsilon) / \text{Var}(Y)$  = true R-squared when Y is regressed against  $X_1, \dots, X_p$ 
# sig.level = significance level (two-tailed p-value is considered)
# power calculation is exact if  $X_1, \dots, X_p$  is multivariate normal

  simulation = sampleadjR2var(nrep,n,p-1,Risq,1)
  # note that function sampleadjR2var is being used to simulate regressions of  $X_i$  against all other  $X_j$  (not regression of Y
  # against all  $X_i$ )
  # thus simulation["hatvarY",] is a simulation of sample variance of  $X_i$  and
  # simulation["adjRsq",] is a simulation of adjusted R squared of regression of  $X_i$  against all other  $X_j$ 

  return(mean(Vectorize(powerlinreg)(n,p,beta*sqrt(simulation["hatvarY",]),simulation["adjRsq",],Rsq,sig.level)))
}

# beginning of main code

set.seed(123) # for reproducibility

fulldataset = read.csv(file="fulldataset.csv")

# computes principal component of cardiovasc_death_rate and life_expectancy
prcomp_cardlife = prcomp(fulldataset[,c("cardiovasc_death_rate","life_expectancy")],scale=TRUE)
prcomp1_cardlife = prcomp_cardlife$sx[,1] # extracts the principal component
coef_cardlife = prcomp_cardlife$rot[,1] # coefficients used for calculating the principal component
prop1_cardlife = prcomp_cardlife$sdev[1]^2/length(prcomp_cardlife$sdev) # proportion of total variance contained in the principal component

# data for main regression
regdata = fulldataset[,c("people_fully_vaccinated_per_hundred","human_development_index","masks.avg","Per.Levitt.Age.Adjusted")]
regdata$cardlife = prcomp1_cardlife

mainreg = lm(Per.Levitt.Age.Adjusted~people_fully_vaccinated_per_hundred+human_development_index+cardlife+masks.avg,data=regdata)

# standardized coefficient for masks.avg in main regression
betamain = mainreg$coefficients["masks.avg"]*sd(regdata$masks.avg)/sd(regdata$Per.Levitt.Age.Adjusted)

# adjusted R squared for regression of masks.avg against other independent variables of main regression
Risqmain = summary(lm(masks.avg~people_fully_vaccinated_per_hundred+human_development_index+cardlife,data=regdata))$adj.r.squared

# adjusted R squared for main regression
Rsqmain = summary(mainreg)$adj.r.squared

separator = paste(rep("=",40),collapse="")

output = c("Parameter estimates from main regression:")
output = c(output,paste("Standardized coefficient for masks:",betamain))
output = c(output,paste("Adjusted R2 of regression of masks against other independent variables:",Risqmain))
output = c(output,paste("Adjusted R2 of main regression",Rsqmain))

n = nrow(regdata)
beta = 0.5
Risq = 0.4
Rsq = 0.75
output = c(output,separator)
output = c(output,"Parameters used for power calculation:")
output = c(output,paste("Sample size:",n),paste("True standardized coefficient for masks:",beta))
output = c(output,paste("true R2 of regression of masks against other independent variables:",Risq))
output = c(output,paste("true R2 of regression of PLAA against independent variables:",Rsq))
output = c(output,"Power estimates (0.05 significance level):")
output = c(output,capture.output(print(data.frame("number of variables"=4:10,power=Vectorize(powerlinreg_randomX)
(n=n,p=4:10,beta=beta,Risq=Risq,Rsq=Rsq)))))
outputfile = file("PowerCalculation.txt")
writeLines(output,outputfile)
close(outputfile)

# SimulationPlots.R
# Generates diagnostic plots for linear regressions based on simulated data satisfying linear model assumptions exactly
# used for comparisons with diagnostic plots from real data
# number of simulations = 10
# Outputs: simulated-fittedresidual.pdf; fitted values versus residuals
# simulated-fittedstdresidual.pdf; fitted values versus standardized residuals

```

```

# simulated-scalelocation.pdf; scale-location plots (fitted values versus squared roots of absolute values of standardized residuals)
# simulated-qqnormal.pdf; qq plots of standardized residuals against normal distribution

library(ggplot2)

stdresiduals = function(resultreg){
  # computes standardized regression residuals
  # resultreg = output of an lm command

  h = lm.influence(resultreg)$shat
  return(resultreg$residuals/(sqrt(1-h)*summary(resultreg)$sigma))
}

plot_fittedresidual = function(resultreg,abbreviation=NA){
  # generates plot of fitted values versus residuals
  # resultreg = output of an lm command; abbreviation = labels for points

  if(is.na(abbreviation[1]))
  {
    abbreviation = rep("",nrow(resultreg$model))
  }
  dataplot = data.frame(labels=abbreviation,x=resultreg$fitted.values,y=resultreg$residuals)
  return(ggplot(dataplot,aes(x=x,y=y,label=labels))+geom_point()+geom_text(vjust=1.5)
    +labs(x="fitted values",y="residuals")+stat_smooth(formula=y~x,method="loess"))
}

plot_fittedstdresidual = function(resultreg,abbreviation=NA){
  # generates plot of fitted values versus standardized residuals
  # resultreg = output of an lm command; abbreviation = labels for points

  if(is.na(abbreviation[1]))
  {
    abbreviation = rep("",nrow(resultreg$model))
  }
  dataplot = data.frame(labels=abbreviation,x=resultreg$fitted.values,y=stdresiduals(resultreg))
  return(ggplot(dataplot,aes(x=x,y=y,label=labels))+geom_point()+geom_text(vjust=1.5)
    +labs(x="fitted values",y="standardized residuals")+stat_smooth(formula=y~x,method="loess"))
}

plot_scalelocation = function(resultreg,abbreviation=NA){
  # generates scale-location plot (fitted values versus square root of absolute values of standardized residuals)
  # resultreg = output of an lm command; abbreviation = labels for points

  if(is.na(abbreviation[1]))
  {
    abbreviation = rep("",nrow(resultreg$model))
  }
  dataplot = data.frame(labels=abbreviation,x=resultreg$fitted.values,y=sqrt(abs(stdresiduals(resultreg))))
  return(ggplot(dataplot,aes(x=x,y=y,label=labels))+geom_point()+geom_text(vjust=1.5)
    +labs(x="fitted values",y="sqrt of abs value of standardized residuals")+stat_smooth(formula=y~x,method="loess"))
}

plot_qqnormal = function(resultreg){
  # generates qq plot of standardized residuals against a normal distribution
  # resultreg = output of an lm command

  dataplot = data.frame(y=stdresiduals(resultreg))
  return(ggplot(dataplot,aes(sample=y))+stat_qq()+stat_qq_line()+labs(x="theoretical quantile",y="standardized residuals"))
}

# beginning of main code

fulldataset = read.csv(file="fulldataset.csv")

# computes principal component of cardiovasc_death_rate and life_expectancy
prcomp_cardlife = prcomp(fulldataset[,c("cardiovasc_death_rate","life_expectancy")],scale=TRUE)
prcomp1_cardlife = prcomp_cardlife$x[,1] # extracts the principal component
coef_cardlife = prcomp_cardlife$rot[,1] # coefficients used for calculating the principal component
prop1_cardlife = prcomp_cardlife$sdev[1]^2/length(prcomp_cardlife$sdev) # proportion of total variance contained in the principal component

# data for main regression
regdata = fulldataset[,c("people_fully_vaccinated_per_hundred","human_development_index","masks.avg","Per.Levitt.Age.Adjusted")]
regdata$cardlife = prcomp1_cardlife

mainreg = lm(Per.Levitt.Age.Adjusted~people_fully_vaccinated_per_hundred+human_development_index+cardlife+masks.avg,data=regdata)
sigma = summary(mainreg)$sigma # residual variance estimate from main regression
X = as.matrix(mainreg$model[, -1]) # matrix with independent variables from main regression

nsimul = 10 # number of simulations

set.seed(123) # for reproducibility

# generates simulated regressions
simulatedregs = list()
for(i in 1:nsimul)
{
  simulatedy = cbind(1,X)%*%mainreg$coefficients+rnorm(nrow(regdata),0,sigma)
  simulatedregs[[i]] = lm(simulatedy~X)
}

pdf("simulated-fittedresidual.pdf",width=14.23,height=6.77)
for(i in 1:nsimul)
{
  print(plot_fittedresidual(simulatedregs[[i]],NA))
}
dev.off()

pdf("simulated-fittedstdresidual.pdf",width=14.23,height=6.77)

```



```
for(i in 1:nsimul)
{
    print(plot_fittedstdresidual(simulatedregs[[i]],NA))
}
dev.off()

pdf("simulated-scalelocation.pdf",width=14.23,height=6.77)
for(i in 1:nsimul)
{
    print(plot_scalelocation(simulatedregs[[i]],NA))
}
dev.off()

pdf("simulated-qqnormal.pdf",width=14.23,height=6.77)
for(i in 1:nsimul)
{
    print(plot_qqnormal(simulatedregs[[i]]))
}
dev.off()
```