

# simulation

## Intro

This doc reproduces the simulation results of the article provided by Hanneh and Gerko.

## Full reproducible script

```
library(Hmisc)
```

```
Loading required package: lattice
```

```
Loading required package: survival
```

```
Warning: package 'survival' was built under R version 4.2.3
```

```
Loading required package: Formula
```

```
Loading required package: ggplot2
```

```
Attaching package: 'Hmisc'
```

```
The following objects are masked from 'package:base':
```

```
  format.pval, units
```

```
library(mice)
```

Warning: package 'mice' was built under R version 4.2.3

Attaching package: 'mice'

The following object is masked from 'package:stats':

filter

The following objects are masked from 'package:base':

cbind, rbind

```
library(tidyverse)
```

Warning: package 'tidyverse' was built under R version 4.2.3

Warning: package 'tibble' was built under R version 4.2.3

-- Attaching core tidyverse packages ----- tidyverse 2.0.0 --

v dplyr	1.1.0	v readr	2.1.4
v forcats	1.0.0	v stringr	1.5.0
v lubridate	1.9.2	v tibble	3.2.1
v purrr	1.0.1	v tidyr	1.3.0

-- Conflicts ----- tidyverse\_conflicts() --

x dplyr::filter() masks mice::filter(), stats::filter()

x dplyr::lag() masks stats::lag()

x dplyr::src() masks Hmisc::src()

x dplyr::summarize() masks Hmisc::summarize()

i Use the conflicted package (<http://conflicted.r-lib.org/>) to force all conflicts to become

```
d1 <- sasxport.get("../data/DEMO_I.xpt")
```

Processing SAS dataset DEMO\_I ..

```
d2 <- sasxport.get("../data/BPX_I.xpt")
```

Processing SAS dataset BPX\_I ..

```
d3 <- sasxport.get("../data/BMX_I.xpt")
```

Processing SAS dataset BMX\_I ..

```
d4 <- sasxport.get("../data/GHB_I.xpt")
```

Processing SAS dataset GHB\_I ..

```
d5 <- sasxport.get("../data/TCHOL_I.xpt")
```

Processing SAS dataset TCHOL\_I ..

```
d1.t <- subset(d1,select=c("seqn","riagendr","ridageyr"))
d2.t <- subset(d2,select=c("seqn","bpxsy1"))
d3.t <- subset(d3,select=c("seqn","bmxbmi"))
d4.t <- subset(d4,select=c("seqn","lbgxgh"))
d5.t <- subset(d5,select=c("seqn","lbdtcsl"))
d <- merge(d1.t,d2.t)
d <- merge(d,d3.t)
d <- merge(d,d4.t)
d <- merge(d,d5.t)
# =====
# rename variables:
# RIAGENDR - Gender
# RIDAGEYR - Age in years at screening
# BPSY1 - Systolic: Blood pres (1st rdg) mm Hg
# BMXBMI - Body Mass Index (kg/m**2)
# LBDTCSL - Total Cholesterol (mmol/L)
# LBGXGH - Glycohemoglobin (%)
d$age <- d$ridageyr
d$sex <- d$riagendr
d$bp <- d$bpxsy1
```

```

d$bmi <- d$bmxbmi
d$HbA1C <- d$lbxgh
d$chol <- d$lbdtcsi
d$age[d$age<18] <- NA
# =====
# select complete cases:
dc <- cc(subset(d,select=c("age","sex","bmi","HbA1C","bp")))
# analysis:
summary(lm(bp ~ HbA1C + age + as.factor(sex), data=dc))

```

Call:

```
lm(formula = bp ~ HbA1C + age + as.factor(sex), data = dc)
```

Residuals:

```

Systolic: Blood pres (1st rdg) mm Hg
      Min       1Q   Median       3Q      Max
-49.887 -10.509  -1.378   8.491 107.583

```

Coefficients:

	Estimate	Std. Error	t value	Pr(> t )
(Intercept)	98.75149	1.21418	81.332	< 2e-16 ***
HbA1C	1.12638	0.20291	5.551	2.98e-08 ***
age	0.44486	0.01284	34.648	< 2e-16 ***
as.factor(sex)2	-3.24792	0.45164	-7.191	7.34e-13 ***

---

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Residual standard error: 16.1 on 5088 degrees of freedom

Multiple R-squared: 0.2305, Adjusted R-squared: 0.23

F-statistic: 508 on 3 and 5088 DF, p-value: < 2.2e-16

```
confint(lm(bp ~ HbA1C + age + as.factor(sex), data=dc))
```

	2.5 %	97.5 %
(Intercept)	96.3711755	101.1317982
HbA1C	0.7285836	1.5241825
age	0.4196932	0.4700355
as.factor(sex)2	-4.1333281	-2.3625106

```
summary(lm(bp ~ HbA1C + bmi + age + as.factor(sex), data=dc))
```

Call:

```
lm(formula = bp ~ HbA1C + bmi + age + as.factor(sex), data = dc)
```

Residuals:

Systolic: Blood pres (1st rdg) mm Hg

Min	1Q	Median	3Q	Max
-51.068	-10.251	-1.504	8.264	107.410

Coefficients:

	Estimate	Std. Error	t value	Pr(> t )
(Intercept)	92.65583	1.39320	66.506	< 2e-16 ***
HbA1C	0.75177	0.20596	3.650	0.000265 ***
bmi	0.28632	0.03282	8.724	< 2e-16 ***
age	0.44586	0.01275	34.979	< 2e-16 ***
as.factor(sex)2	-3.63115	0.45049	-8.060	9.4e-16 ***

---

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Residual standard error: 15.98 on 5087 degrees of freedom

Multiple R-squared: 0.2418, Adjusted R-squared: 0.2412

F-statistic: 405.7 on 4 and 5087 DF, p-value: < 2.2e-16

```
confint(lm(bp ~ HbA1C + bmi + age + as.factor(sex), data=dc))
```

	2.5 %	97.5 %
(Intercept)	89.9245592	95.3871089
HbA1C	0.3479966	1.1555348
bmi	0.2219815	0.3506673
age	0.4208695	0.4708464
as.factor(sex)2	-4.5143014	-2.7479929

```
# =====
```

```
# simulation of measurement error:
```

```
#| output: FALSE
```

```
ref <- lm(bp ~ HbA1C + bmi + age + as.factor(sex), data=dc)$coef[2]
```

```

n.sim <- 1e3
perc.me.exp <- seq(0,.5,.1)
perc.me.conf<- seq(0,.5,.1)
scenarios <- expand.grid(perc.me.exp,perc.me.conf)
var.exp <- var(dc$HbA1C)
var.conf <- var(dc$bmi)
n <- dim(dc)[1]
beta.hat <- matrix(ncol=dim(scenarios)[1], nrow=n.sim)
for (k in 1:n.sim){
  print(k)
  set.seed(k)
  for (i in 1:dim(scenarios)[1]){
    var.me.exp <- var.exp*scenarios[i,1]/(1-scenarios[i,1])
    var.me.conf <- var.conf*scenarios[i,2]/(1-scenarios[i,2])
    dc$HbA1C.me <- dc$HbA1C + rnorm(dim(dc)[1], 0, sqrt(var.me.exp) )
    dc$bmi.me <- dc$bmi + rnorm(dim(dc)[1], 0, sqrt(var.me.conf) )
    beta.hat[k,i] <- lm(bp ~ HbA1C.me + age + bmi.me + as.factor(sex), data=dc)$coef[2]
  }}

```

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# =====
# create figure:
tot.mat <- cbind(100*scenarios,apply(beta.hat,2,mean))
colnames(tot.mat) <- c("me.exp","me.conf","estimate")
FIGURE <- ggplot(tot.mat, aes(me.exp, me.conf)) +
  geom_tile(color="white",aes(fill = estimate)) +
  geom_text(aes(label = round(estimate, 2))) +
  scale_fill_gradient2(low="#D55E00",mid="white",high = "#56B4E9", midpoint=ref) +
  labs(x=paste("% of total variance of HbA1c due to measurement error"),
       y=paste("% of total variance of BMI due to measurement error")) +
```

```

coord_equal()+
scale_y_continuous(breaks=unique(tot.mat[,1]))+
scale_x_continuous(breaks=unique(tot.mat[,1]))+
theme(panel.background = element_rect(fill='white', colour='grey'),
      plot.title=element_text(hjust=0),
      axis.ticks=element_blank(),
      axis.title=element_text(size=12),
      axis.text=element_text(size=10),
      legend.title=element_text(size=12),
      legend.text=element_text(size=10))

```

FIGURE

