

Simulation

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Introduction

We would like to study if under the setting of outcome depending sampling the conditional likelihood approach is still a valid approach to inference. To study this, I began writing a simulation.

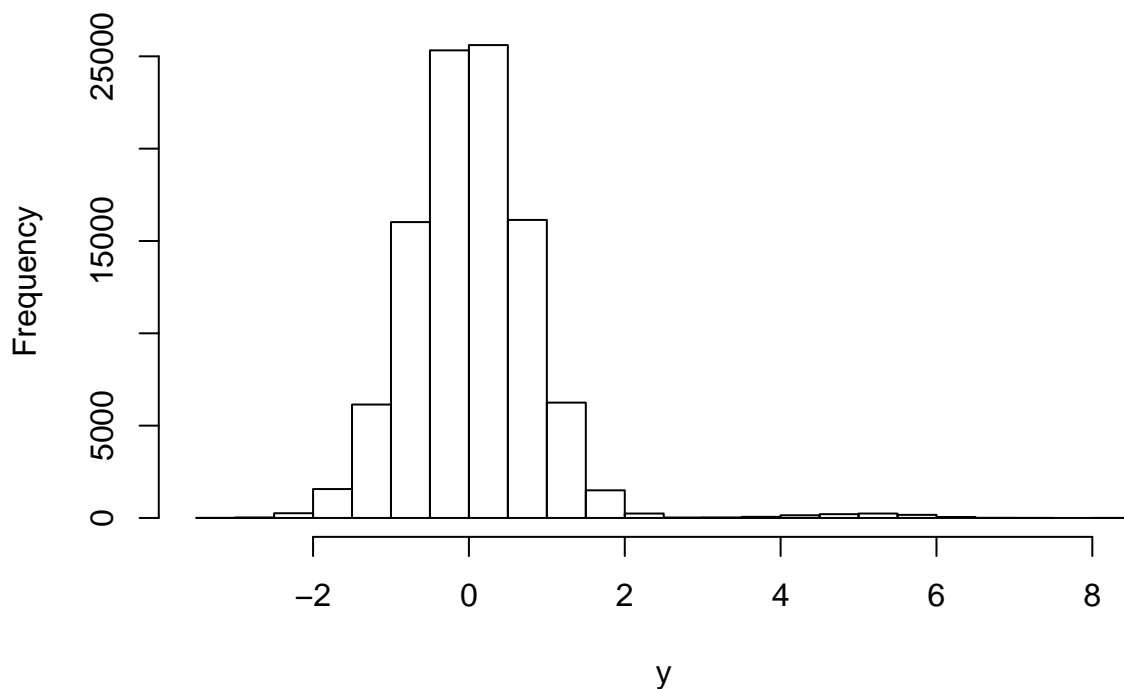
Model

The model I study in my simulation is,

$$Y_{ij} = \beta_0 + \beta X_i + U_i + \epsilon_{ij}.$$

Which is simply a linear model with subject-specific intercept. In my simulation I made X_i a bernoulli random variable with $P(X_i = 1) = 1 - P(X_i = 0) = 0.01$. Where $X_i = 1$ denotes a case and $X_i = 0$ denotes a control. Also, I made X_i fixed within-subject (which may be obvious from notation). However, the errors within-subject are positively correlated with each other. Code for the simulation is shown below, but here is a sample of the histogram of Y values for the population for one repetition

Histogram of y



Clearly, there appears to be 2 subpopulations here, where one has many more members than the other. In my simulated data, I make $\beta_0 = 0$ and $\beta = 5$.

Simulation

The goal here is to convince myself that if we sample based off the sum of outcomes for a given subject, (here I made the number of measurements per subject equal to 5) we still get a consistent estimate of for beta. What I did here, was generate a “population” of subjects (number of subjects equals 20,000, each subject has 5 measurements) from the model above. Then I aggregate the data by taking the sum of all measurements for each subject. Then I sample 50 subjects from each group based off that sum. With the sampled data, I fit the model given above and record the estimated value of beta. I repeat this process many times and create a histogram for estimated values for the fixed effect. The code and resulting histogram are shown below.

```
library(nlme)
set.seed(1104)

pop.m<-20000 # number of clusters
pop.n<- 5 # number within clusters
case.prob<- .01 #probability of case in underlying population
beta<-5 #slope for indicator
sigma<- .5 #overall standard deviation in the linear model
tau_e<-0.8 #error correlation
reps=2000

beta.est<-vector(length = reps)
for(i in 1:reps){
  x<-rbinom(pop.m,1,prob=case.prob)
  x<-rep(x,each=pop.n)

  u<-rnorm(pop.m,mean = 0, sd=sqrt(sigma*tau_e)) #cluster samples
  u1<-rep(u,each=pop.n) # repeat each cluster sample n times
  estar<-rnorm(pop.m*pop.n,mean = 0, sd=sqrt(sigma*(1-tau_e))) # samples within each cluster
  err<-u1+estar #total error

  y<-beta*x+err
  dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
  agg.dat<-aggregate(y~id, dat, sum) # sum y by id
  case.samp<-sample(agg.dat$id[agg.dat$y>15],50) #sample cases
  control.samp<-sample(agg.dat$id[agg.dat$y<15],50) # sample controls
  samp<-c(case.samp, control.samp)
  samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids

  fit<-lme(y~x, data = samp.dat, random = ~1|id)
  beta.est[i]<-fixed.effects(fit)[2]
}

hist(beta.est)### histogram looks good
```

Looking back at my notes, I’m not fully sure what to do from here. I have a note that says subtract off the group means of the X’s and Y’s. That would give me something like that within-subject part of the model. But what do I do with that? look at the estimate for the fixed effects?

Updated Simulation

Discrete X

I've updated the model I'm studying, now we have,

$$Y_{ij} = \beta_0 + \beta X_{ij} + U_i + \epsilon_{ij}.$$

Where I set $\beta_0 = 0$ and $\beta = 5$. The X_{ij} terms are generated as follows: 1.) We generate $X_{ij}^* \sim N(U_i, 1)$, 2.) We then make the terms into a binary random variable by coding $X_{ij}^* > 2.5$ as 1 and 0 otherwise. The rest of the simulation is the same as above except at the end we fit a model of the form,

$$Y_{ij} = \beta_0 + \beta_1 \bar{X}_i + \beta_2 (X_{ij} - \bar{X}_i) + U_i + \epsilon_{ij}$$

```
####ODS Conditional Likelihood Simulation: Jacob M. Maronge 01/09/17

#### Inspired by the paper: Separating between- and within-cluster covariate effects by using conditionalization
#### By John M. Neuhaus and Charles E. McCulloch
set.seed(1104)
library(foreach)
library(doParallel)
cores=detectCores()
cl <- makeCluster(3)
registerDoParallel(cl)
clusterEvalQ(cl,
  {library(nlme)
    pop.m<-20000 # number of clusters
    pop.n<- 5 # number within clusters
    beta<-5 #slope for indicator
    sigma<- 1 #overall standard deviation in the linear model
    tau_e<-0.8 #error correlation
    norm.sim<-function(pop.m, pop.n, beta,tau_e, sigma){
      u<-rnorm(pop.m,mean = 0, sd=sqrt(sigma*tau_e)) #cluster samples
      u1<-rep(u,each=pop.n) # repeat each cluster sample n times
      estar<-rnorm(pop.m*pop.n,mean = 0, sd=sqrt(sigma*(1-tau_e))) # samples within each cluster
      err<-u1+estar #total error

      x<-rnorm(u1,1)
      x<-as.numeric(x>2.5)
      y<-beta*x+err
      dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
      agg.dat<-aggregate(y~id, dat, sum) # sum y by id
      case.samp<-sample(agg.dat$id[agg.dat$y>12],50) #sample cases
      control.samp<-sample(agg.dat$id[agg.dat$y<12],50) # sample controls
      samp<-c(case.samp, control.samp)
      samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids
      samp.agg.dat<-aggregate(x~id, samp.dat, mean) #calculate means for x
      x_ibar<-rep(samp.agg.dat$x,each=pop.n) # match means dimensions with dat
      samp.dat$x_ibar<-x_ibar
      samp.dat$diff<-samp.dat$x-samp.dat$x_ibar # calculate x_ij-x_ibar

      fit<-lme(y~diff+x_ibar, data = samp.dat, random = ~1|id)
      beta.diff.est<-fixed.effects(fit)[2]
      beta.diff.cov.prob<-(intervals(fit)$fixed[2,1]<=beta&intervals(fit)$fixed[2,3]>=beta)
```

```

        beta.x_ibar.est<-fixed.effects(fit)[3]
        beta.x_ibar.cov.prob<-(intervals(fit)$fixed[3,1]<=beta&intervals(fit)$fixed[3,3]>=beta)
        out<-list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
        names(out)<-c("Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
        return(out)
    })

## [[1]]
## function (pop.m, pop.n, beta, tau_e, sigma)
## {
##     u <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_e))
##     u1 <- rep(u, each = pop.n)
##     estar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_e)))
##     err <- u1 + estar
##     x <- rnorm(u1, 1)
##     x <- as.numeric(x > 2.5)
##     y <- beta * x + err
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     case.samp <- sample(agg.dat$id[agg.dat$y > 12], 50)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 12], 50)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)
##     samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##     x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##     samp.dat$x_ibar <- x_ibar
##     samp.dat$diff <- samp.dat$x - samp.dat$x_ibar
##     fit <- lme(y ~ diff + x_ibar, data = samp.dat, random = ~1 |
##         id)
##     beta.diff.est <- fixed.effects(fit)[2]
##     beta.diff.cov.prob <- (intervals(fit)$fixed[2, 1] <= beta &
##         intervals(fit)$fixed[2, 3] >= beta)
##     beta.x_ibar.est <- fixed.effects(fit)[3]
##     beta.x_ibar.cov.prob <- (intervals(fit)$fixed[3, 1] <= beta &
##         intervals(fit)$fixed[3, 3] >= beta)
##     out <- list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est,
##         beta.x_ibar.cov.prob)
##     names(out) <- c("Diff Estimate", "Diff Covered", "x_ibar Estimate",
##         "x_ibar Covered")
##     return(out)
## }
##
## [[2]]
## function (pop.m, pop.n, beta, tau_e, sigma)
## {
##     u <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_e))
##     u1 <- rep(u, each = pop.n)
##     estar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_e)))
##     err <- u1 + estar
##     x <- rnorm(u1, 1)
##     x <- as.numeric(x > 2.5)
##     y <- beta * x + err

```

```

## dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
## agg.dat <- aggregate(y ~ id, dat, sum)
## case.samp <- sample(agg.dat$id[agg.dat$y > 12], 50)
## control.samp <- sample(agg.dat$id[agg.dat$y < 12], 50)
## samp <- c(case.samp, control.samp)
## samp.dat <- subset(dat, dat$id %in% samp)
## samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
## x_ibar <- rep(samp.agg.dat$x, each = pop.n)
## samp.dat$x_ibar <- x_ibar
## samp.dat$diff <- samp.dat$x - samp.dat$x_ibar
## fit <- lme(y ~ diff + x_ibar, data = samp.dat, random = ~1 |
## id)
## beta.diff.est <- fixed.effects(fit)[2]
## beta.diff.cov.prob <- (intervals(fit)$fixed[2, 1] <= beta &
## intervals(fit)$fixed[2, 3] >= beta)
## beta.x_ibar.est <- fixed.effects(fit)[3]
## beta.x_ibar.cov.prob <- (intervals(fit)$fixed[3, 1] <= beta &
## intervals(fit)$fixed[3, 3] >= beta)
## out <- list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est,
## beta.x_ibar.cov.prob)
## names(out) <- c("Diff Estimate", "Diff Covered", "x_ibar Estimate",
## "x_ibar Covered")
## return(out)
## }
##
## [[3]]
## function (pop.m, pop.n, beta, tau_e, sigma)
## {
## u <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_e))
## u1 <- rep(u, each = pop.n)
## estar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
## (1 - tau_e)))
## err <- u1 + estar
## x <- rnorm(u1, 1)
## x <- as.numeric(x > 2.5)
## y <- beta * x + err
## dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
## agg.dat <- aggregate(y ~ id, dat, sum)
## case.samp <- sample(agg.dat$id[agg.dat$y > 12], 50)
## control.samp <- sample(agg.dat$id[agg.dat$y < 12], 50)
## samp <- c(case.samp, control.samp)
## samp.dat <- subset(dat, dat$id %in% samp)
## samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
## x_ibar <- rep(samp.agg.dat$x, each = pop.n)
## samp.dat$x_ibar <- x_ibar
## samp.dat$diff <- samp.dat$x - samp.dat$x_ibar
## fit <- lme(y ~ diff + x_ibar, data = samp.dat, random = ~1 |
## id)
## beta.diff.est <- fixed.effects(fit)[2]
## beta.diff.cov.prob <- (intervals(fit)$fixed[2, 1] <= beta &
## intervals(fit)$fixed[2, 3] >= beta)
## beta.x_ibar.est <- fixed.effects(fit)[3]
## beta.x_ibar.cov.prob <- (intervals(fit)$fixed[3, 1] <= beta &
## intervals(fit)$fixed[3, 3] >= beta)

```

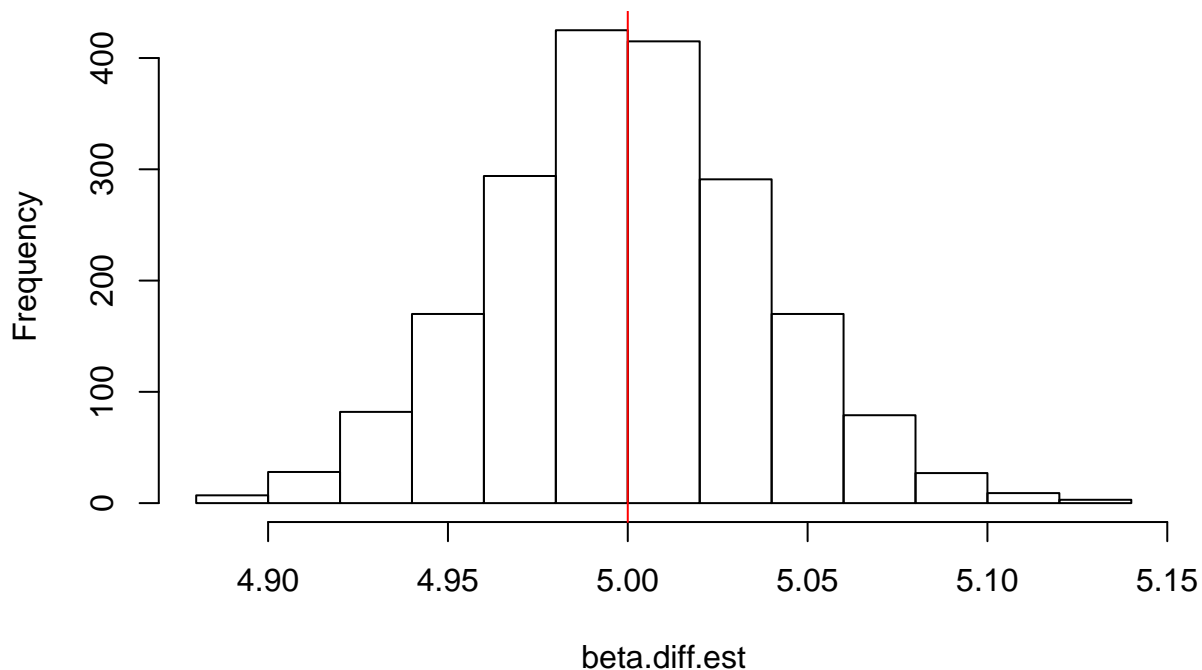
```
## out <- list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est,
##           beta.x_ibar.cov.prob)
## names(out) <- c("Diff Estimate", "Diff Covered", "x_ibar Estimate",
##               "x_ibar Covered")
## return(out)
## }
```

```
out<-foreach(i=1:2000, .combine=cbind) %dopar% {
  norm.sim(pop.m, pop.n, beta,tau_e, sigma)
```

```
}
beta.diff.est<-unlist(out[1,])
beta.diff.cov.prob<-unlist(out[2,])
beta.x_ibar.est <- unlist(out[3,])
beta.x_ibar.cov.prob<-unlist(out[4,])
```

```
hist(beta.diff.est)
abline(v = 5, col = "red")
```

Histogram of beta.diff.est

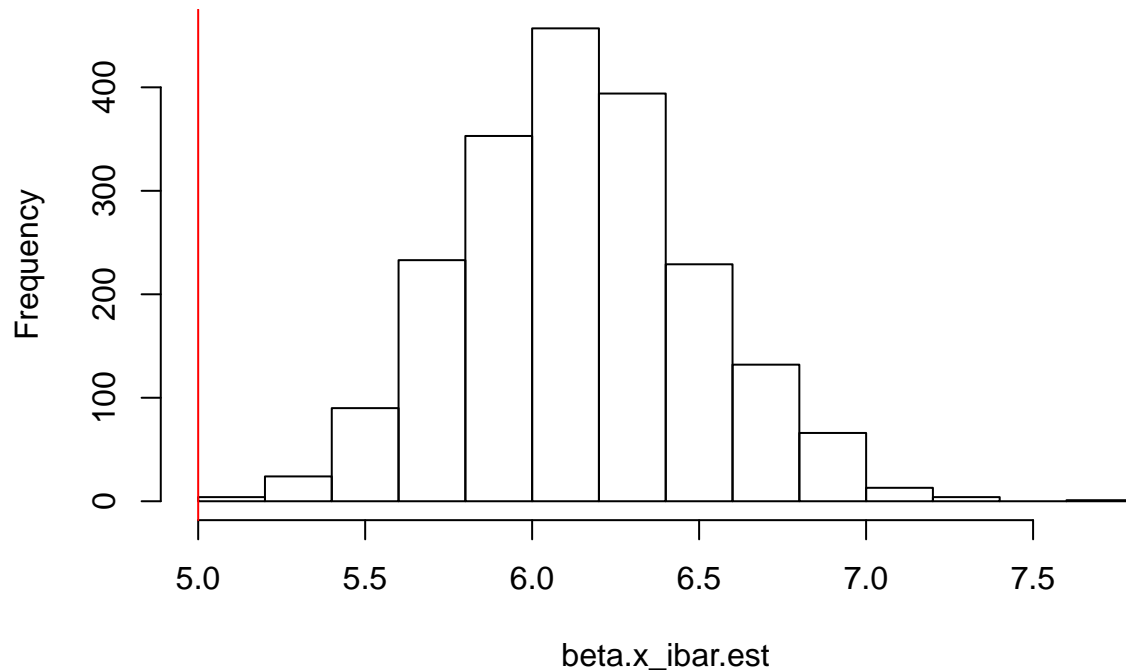


```
mean(beta.diff.cov.prob)
```

```
## [1] 0.954
```

```
hist(beta.x_ibar.est)
abline(v = 5, col = "red")
```

Histogram of beta.x_ibar.est



```
mean(beta.x_ibar.cov.prob)
```

```
## [1] 0.134
```

Continuous X

Here I generate the same model as above, except I don't convert the X's to discrete values.

```
set.seed(1104)
library(foreach)
library(doParallel)
cores=detectCores()
cl <- makeCluster(3)
registerDoParallel(cl)
clusterEvalQ(cl,
  {library(nlme)
    pop.m<-20000 # number of clusters
    pop.n<- 5 # number within clusters
    beta<-5 #slope for indicator
    sigma<- 1 #overall standard deviation in the linear model
    tau_e<-0.8 #error correlation
    norm.sim<-function(pop.m, pop.n, beta,tau_e, sigma){
      u<-rnorm(pop.m,mean = 0, sd=sqrt(sigma*tau_e)) #cluster samples
      u1<-rep(u,each=pop.n) # repeat each cluster sample n times
      estar<-rnorm(pop.m*pop.n,mean = 0, sd=sqrt(sigma*(1-tau_e))) # samples within each clu
      err<-u1+estar #total error

      x<-rnorm(u1,1)
      y<-beta*x+err
      dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
```

```

agg.dat<-aggregate(y~id, dat, sum) # sum y by id
case.samp<-sample(agg.dat$id[agg.dat$y>12],50) #sample cases
control.samp<-sample(agg.dat$id[agg.dat$y<12],50) # sample controls
samp<-c(case.samp, control.samp)
samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids
samp.agg.dat<-aggregate(x~id, samp.dat, mean) #calculate means for x
x_ibar<-rep(samp.agg.dat$x,each=pop.n) # mach means dimensions with dat
samp.dat$x_ibar<-x_ibar
samp.dat$diff<-samp.dat$x-samp.dat$x_ibar # calculate x_ij-x_ibar

fit<-lme(y~diff+x_ibar, data = samp.dat, random = ~1|id)
beta.diff.est<-fixed.effects(fit)[2]
beta.diff.cov.prob<-(intervals(fit)$fixed[2,1]<=beta&intervals(fit)$fixed[2,3]>=beta)
beta.x_ibar.est<-fixed.effects(fit)[3]
beta.x_ibar.cov.prob<-(intervals(fit)$fixed[3,1]<=beta&intervals(fit)$fixed[3,3]>=beta)
out<-list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
names(out)<-c("Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
return(out)
})

```

```

## [[1]]
## function (pop.m, pop.n, beta, tau_e, sigma)
## {
##   u <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_e))
##   u1 <- rep(u, each = pop.n)
##   estar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##     (1 - tau_e)))
##   err <- u1 + estar
##   x <- rnorm(u1, 1)
##   y <- beta * x + err
##   dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   case.samp <- sample(agg.dat$id[agg.dat$y > 12], 50)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 12], 50)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- x_ibar
##   samp.dat$diff <- samp.dat$x - samp.dat$x_ibar
##   fit <- lme(y ~ diff + x_ibar, data = samp.dat, random = ~1 |
##     id)
##   beta.diff.est <- fixed.effects(fit)[2]
##   beta.diff.cov.prob <- (intervals(fit)$fixed[2, 1] <= beta &
##     intervals(fit)$fixed[2, 3] >= beta)
##   beta.x_ibar.est <- fixed.effects(fit)[3]
##   beta.x_ibar.cov.prob <- (intervals(fit)$fixed[3, 1] <= beta &
##     intervals(fit)$fixed[3, 3] >= beta)
##   out <- list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est,
##     beta.x_ibar.cov.prob)
##   names(out) <- c("Diff Estimate", "Diff Covered", "x_ibar Estimate",
##     "x_ibar Covered")
##   return(out)
## }

```



```

##
## [[2]]
## function (pop.m, pop.n, beta, tau_e, sigma)
## {
##     u <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_e))
##     u1 <- rep(u, each = pop.n)
##     estar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_e)))
##     err <- u1 + estar
##     x <- rnorm(u1, 1)
##     y <- beta * x + err
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     case.samp <- sample(agg.dat$id[agg.dat$y > 12], 50)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 12], 50)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)
##     samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##     x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##     samp.dat$x_ibar <- x_ibar
##     samp.dat$diff <- samp.dat$x - samp.dat$x_ibar
##     fit <- lme(y ~ diff + x_ibar, data = samp.dat, random = ~1 |
##         id)
##     beta.diff.est <- fixed.effects(fit)[2]
##     beta.diff.cov.prob <- (intervals(fit)$fixed[2, 1] <= beta &
##         intervals(fit)$fixed[2, 3] >= beta)
##     beta.x_ibar.est <- fixed.effects(fit)[3]
##     beta.x_ibar.cov.prob <- (intervals(fit)$fixed[3, 1] <= beta &
##         intervals(fit)$fixed[3, 3] >= beta)
##     out <- list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est,
##         beta.x_ibar.cov.prob)
##     names(out) <- c("Diff Estimate", "Diff Covered", "x_ibar Estimate",
##         "x_ibar Covered")
##     return(out)
## }
##
## [[3]]
## function (pop.m, pop.n, beta, tau_e, sigma)
## {
##     u <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_e))
##     u1 <- rep(u, each = pop.n)
##     estar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_e)))
##     err <- u1 + estar
##     x <- rnorm(u1, 1)
##     y <- beta * x + err
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     case.samp <- sample(agg.dat$id[agg.dat$y > 12], 50)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 12], 50)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)
##     samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##     x_ibar <- rep(samp.agg.dat$x, each = pop.n)

```

```
##      samp.dat$x_ibar <- x_ibar
##      samp.dat$diff <- samp.dat$x - samp.dat$x_ibar
##      fit <- lme(y ~ diff + x_ibar, data = samp.dat, random = ~1 |
##              id)
##      beta.diff.est <- fixed.effects(fit)[2]
##      beta.diff.cov.prob <- (intervals(fit)$fixed[2, 1] <= beta &
##              intervals(fit)$fixed[2, 3] >= beta)
##      beta.x_ibar.est <- fixed.effects(fit)[3]
##      beta.x_ibar.cov.prob <- (intervals(fit)$fixed[3, 1] <= beta &
##              intervals(fit)$fixed[3, 3] >= beta)
##      out <- list(beta.diff.est, beta.diff.cov.prob, beta.x_ibar.est,
##              beta.x_ibar.cov.prob)
##      names(out) <- c("Diff Estimate", "Diff Covered", "x_ibar Estimate",
##              "x_ibar Covered")
##      return(out)
## }

```

```
out<-foreach(i=1:2000, .combine=cbind) %dopar% {
  norm.sim(pop.m, pop.n, beta,tau_e, sigma)
}

```

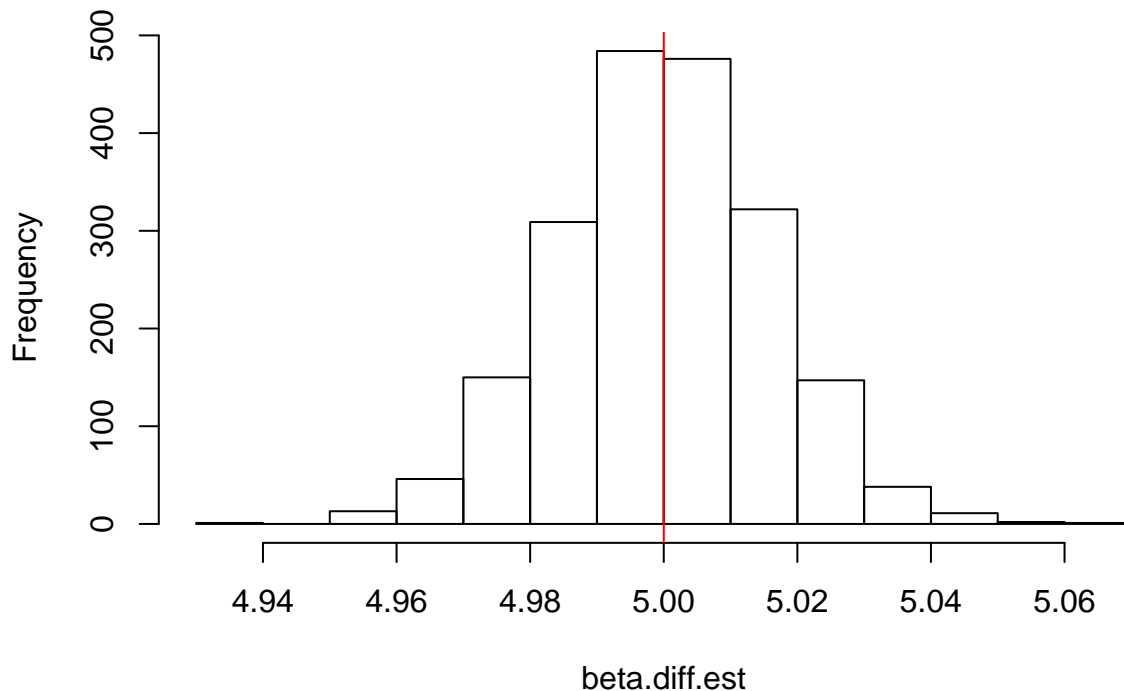
```
beta.diff.est<-unlist(out[1,])
beta.diff.cov.prob<-unlist(out[2,])
beta.x_ibar.est <- unlist(out[3,])
beta.x_ibar.cov.prob<-unlist(out[4,])

```

```
hist(beta.diff.est)
abline(v = 5, col = "red")

```

Histogram of beta.diff.est

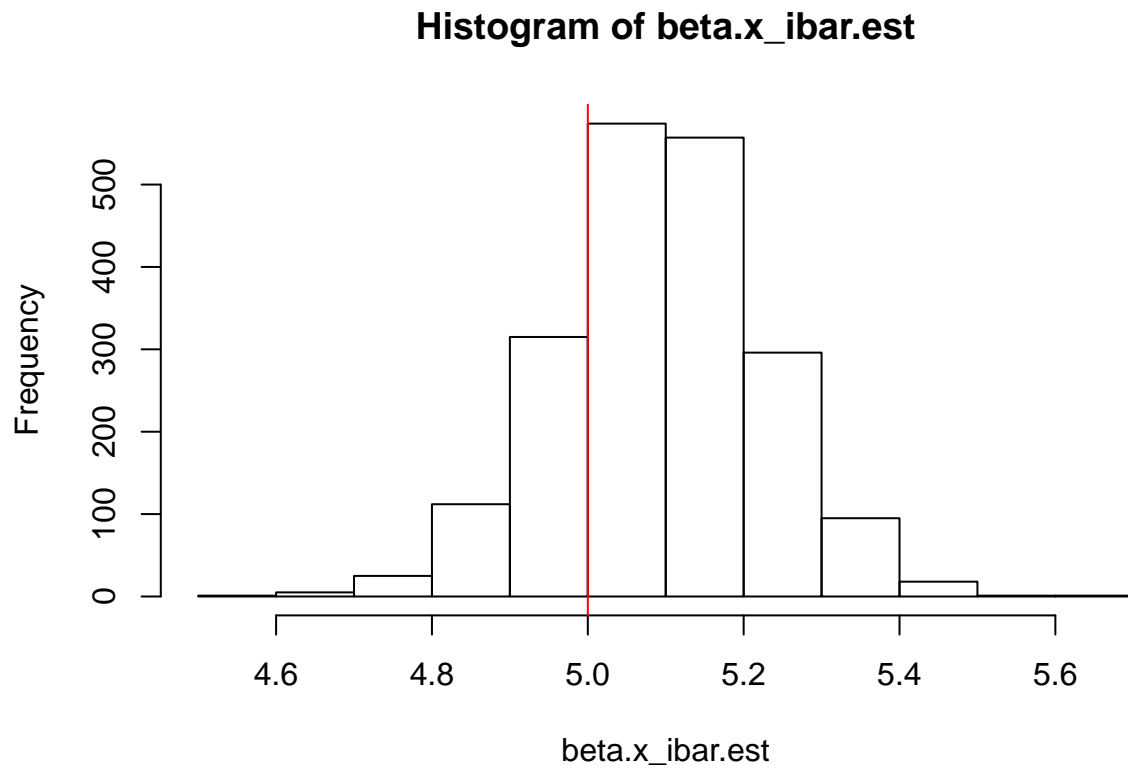


```
mean(beta.diff.cov.prob)
```

```
## [1] 0.948
```

```
hist(beta.x_ibar.est)
```

```
abline(v = 5, col = "red")
```



```
mean(beta.x_ibar.cov.prob)
```

```
## [1] 0.894
```

Logistic regression simulation 02/05/18

Discrete X

Since we've done the normal case, we decided to study a logistic model of the form,

$$\text{logit}(Y_{ij}) = \beta_0 + \beta X_{ij} + U_i.$$

Where I set $\beta_0 = -1.5$ and $\beta = 3$. The U_i terms are distributed as $U_i \sim N(0, 1/2)$. The X_{ij} terms are generated as follows: 1.) We generate $X_{ij}^* \sim N(U_i, 1)$, 2.) We then make the terms into a binary random variable by coding $X_{ij}^* > 2.5$ as 1 and 0 otherwise. Then we generate values Z_{ij} where,

$$Z_{ij} = \beta_0 + \beta X_{ij} + U_i.$$

Next, we take,

$$p_{ij} = \frac{1}{1 + e^{-Z_{ij}}}.$$

Finally, we generate our Y_{ij} by taking,

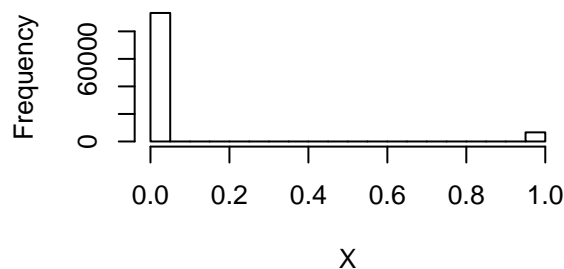
$$Y_{ij} = \text{Bern}(p_{ij}).$$

We then fit a model of the form,

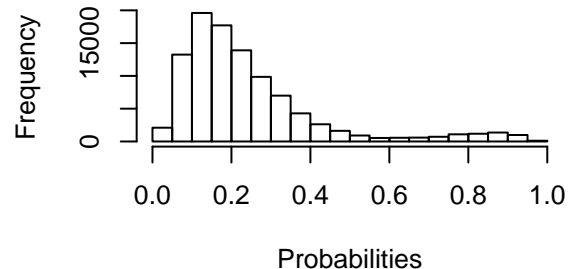
$$\text{logit}(Y_{ij}) = \beta_0 + \beta_1 \bar{X}_i + \beta_2 (X_{ij} - \bar{X}_i) + U_i.$$

Below, I show some plots from one repetition of my simulation.

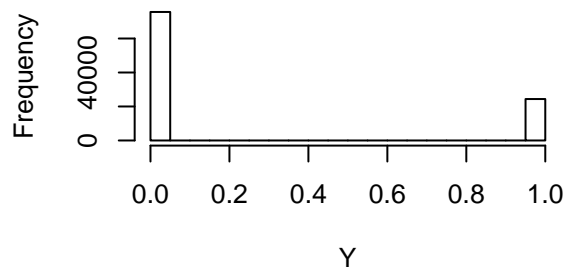
Histogram of X



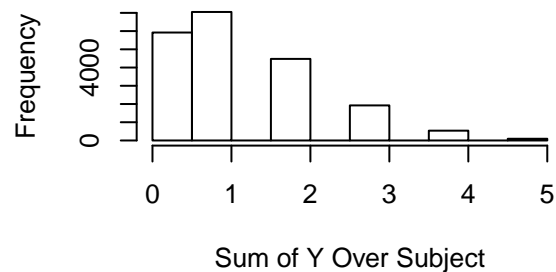
Histogram of Probabilities



Histogram of Y



Histogram of Aggregated Sums



```
set.seed(1104)
library(foreach)
library(doParallel)
```

```

cl <- makeCluster(3) #not to overload your computer
registerDoParallel(cl)
clusterEvalQ(cl,
  {library(lme4)
    pop.m<-80000 # number of clusters
    pop.n<- 5 # number within clusters
    beta1<-3 #slope for indicator
    beta0<--1.5 #intercept terms
    tau_sq<-.5 #variance of random intercept
    logit.sim<-function(pop.m, pop.n, beta1,beta0,tau_sq){
      library(lme4)
      u<-rnorm(pop.m,mean = 0, sd=sqrt(tau_sq)) #cluster samples
      u1<-rep(u,each=pop.n) # repeat each cluster sample n times
      x<-rnorm(u1,1)
      x<-as.numeric(x>2.5)
      z<-beta0+beta1*x+u1
      pr<-1/(1+exp(-z))
      y<-rbinom(n=pop.m*pop.n,size = 1,prob = pr )
      dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
      agg.dat<-aggregate(y~id, dat, sum) # sum y by id
      case.samp<-sample(agg.dat$id[agg.dat$y>=4],250) #sample cases
      control.samp<-sample(agg.dat$id[agg.dat$y<4],250)# sample controls
      samp<-c(case.samp, control.samp)
      samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids
      samp.agg.dat<-aggregate(x~id, samp.dat, mean) #calculate means for x
      x_ibar<-rep(samp.agg.dat$x,each=pop.n) # mach means dimensions with dat
      samp.dat$x_ibar<-(x_ibar)
      samp.dat$diff<-(samp.dat$x-samp.dat$x_ibar) # calculate x_ij-x_ibar

      fit.logit<-glmer(y~diff+x_ibar+(1|id),data = samp.dat,
        family = binomial(link = "logit"),
        glmerControl(optimizer = c("bobyqa","Nelder-Mead"))) )

      beta.int.est<-coef(summary(fit.logit))[1,1]
      beta.int.cov.prob<-((coef(summary(fit.logit))[1,1]-qt(.975, df=2000)*coef(summary(fit.
      beta.diff.est<-coef(summary(fit.logit))[2,1]
      beta.diff.cov.prob<-((coef(summary(fit.logit))[2,1]-qt(.975, df=2000)*coef(summary(fit
      beta.x_ibar.est<-coef(summary(fit.logit))[3,1]
      beta.x_ibar.cov.prob<-((coef(summary(fit.logit))[3,1]-qt(.975, df=2000)*coef(summary(f
      out<-list(beta.int.est, beta.int.cov.prob, beta.diff.est, beta.diff.cov.prob, beta.x_ib
      names(out)<-c("Intercept Estimate", "Intercept Covered", "Diff Estimate", "Diff Covered
      return(out)
    })
  })
)

## [[1]]
## function (pop.m, pop.n, beta1, beta0, tau_sq)
## {
##   library(lme4)
##   u <- rnorm(pop.m, mean = 0, sd = sqrt(tau_sq))
##   u1 <- rep(u, each = pop.n)
##   x <- rnorm(u1, 1)
##   x <- as.numeric(x > 2.5)
##   z <- beta0 + beta1 * x + u1

```

```

## pr <- 1/(1 + exp(-z))
## y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
## dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
## agg.dat <- aggregate(y ~ id, dat, sum)
## case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
## control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
## samp <- c(case.samp, control.samp)
## samp.dat <- subset(dat, dat$id %in% samp)
## samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
## x_ibar <- rep(samp.agg.dat$x, each = pop.n)
## samp.dat$x_ibar <- (x_ibar)
## samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
## fit.logit <- glmer(y ~ diff + x_ibar + (1 | id), data = samp.dat,
##   family = binomial(link = "logit"), glmerControl(optimizer = c("bobyqa",
##     "Nelder_Mead")))
## beta.int.est <- coef(summary(fit.logit))[1, 1]
## beta.int.cov.prob <- ((coef(summary(fit.logit))[1, 1] - qt(0.975,
##   df = 2000) * coef(summary(fit.logit))[1, 2]) <= beta0 &
##   (coef(summary(fit.logit))[1, 1] + qt(0.975, df = 2000) *
##     coef(summary(fit.logit))[1, 2]) >= beta0)
## beta.diff.est <- coef(summary(fit.logit))[2, 1]
## beta.diff.cov.prob <- ((coef(summary(fit.logit))[2, 1] -
##   qt(0.975, df = 2000) * coef(summary(fit.logit))[2, 2]) <=
##   beta1 & (coef(summary(fit.logit))[2, 1] + qt(0.975, df = 2000) *
##     coef(summary(fit.logit))[2, 2]) >= beta1)
## beta.x_ibar.est <- coef(summary(fit.logit))[3, 1]
## beta.x_ibar.cov.prob <- ((coef(summary(fit.logit))[3, 1] -
##   qt(0.975, df = 2000) * coef(summary(fit.logit))[3, 2]) <=
##   beta1 & (coef(summary(fit.logit))[3, 1] + qt(0.975, df = 2000) *
##     coef(summary(fit.logit))[3, 2]) >= beta1)
## out <- list(beta.int.est, beta.int.cov.prob, beta.diff.est,
##   beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
## names(out) <- c("Intercept Estimate", "Intercept Covered",
##   "Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
## return(out)
## }
##
## [[2]]
## function (pop.m, pop.n, beta1, beta0, tau_sq)
## {
##   library(lme4)
##   u <- rnorm(pop.m, mean = 0, sd = sqrt(tau_sq))
##   u1 <- rep(u, each = pop.n)
##   x <- rnorm(u1, 1)
##   x <- as.numeric(x > 2.5)
##   z <- beta0 + beta1 * x + u1
##   pr <- 1/(1 + exp(-z))
##   y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##   dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)

```

```

##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   fit.logit <- glmer(y ~ diff + x_ibar + (1 | id), data = samp.dat,
##     family = binomial(link = "logit"), glmerControl(optimizer = c("bobyqa",
##       "Nelder_Mead"))))
##   beta.int.est <- coef(summary(fit.logit))[1, 1]
##   beta.int.cov.prob <- ((coef(summary(fit.logit))[1, 1] - qt(0.975,
##     df = 2000) * coef(summary(fit.logit))[1, 2]) <= beta0 &
##     (coef(summary(fit.logit))[1, 1] + qt(0.975, df = 2000) *
##       coef(summary(fit.logit))[1, 2]) >= beta0)
##   beta.diff.est <- coef(summary(fit.logit))[2, 1]
##   beta.diff.cov.prob <- ((coef(summary(fit.logit))[2, 1] -
##     qt(0.975, df = 2000) * coef(summary(fit.logit))[2, 2]) <=
##     beta1 & (coef(summary(fit.logit))[2, 1] + qt(0.975, df = 2000) *
##       coef(summary(fit.logit))[2, 2]) >= beta1)
##   beta.x_ibar.est <- coef(summary(fit.logit))[3, 1]
##   beta.x_ibar.cov.prob <- ((coef(summary(fit.logit))[3, 1] -
##     qt(0.975, df = 2000) * coef(summary(fit.logit))[3, 2]) <=
##     beta1 & (coef(summary(fit.logit))[3, 1] + qt(0.975, df = 2000) *
##       coef(summary(fit.logit))[3, 2]) >= beta1)
##   out <- list(beta.int.est, beta.int.cov.prob, beta.diff.est,
##     beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
##   names(out) <- c("Intercept Estimate", "Intercept Covered",
##     "Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
##   return(out)
## }
##
## [[3]]
## function (pop.m, pop.n, beta1, beta0, tau_sq)
## {
##   library(lme4)
##   u <- rnorm(pop.m, mean = 0, sd = sqrt(tau_sq))
##   u1 <- rep(u, each = pop.n)
##   x <- rnorm(u1, 1)
##   x <- as.numeric(x > 2.5)
##   z <- beta0 + beta1 * x + u1
##   pr <- 1/(1 + exp(-z))
##   y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##   dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   fit.logit <- glmer(y ~ diff + x_ibar + (1 | id), data = samp.dat,
##     family = binomial(link = "logit"), glmerControl(optimizer = c("bobyqa",
##       "Nelder_Mead"))))
##   beta.int.est <- coef(summary(fit.logit))[1, 1]

```

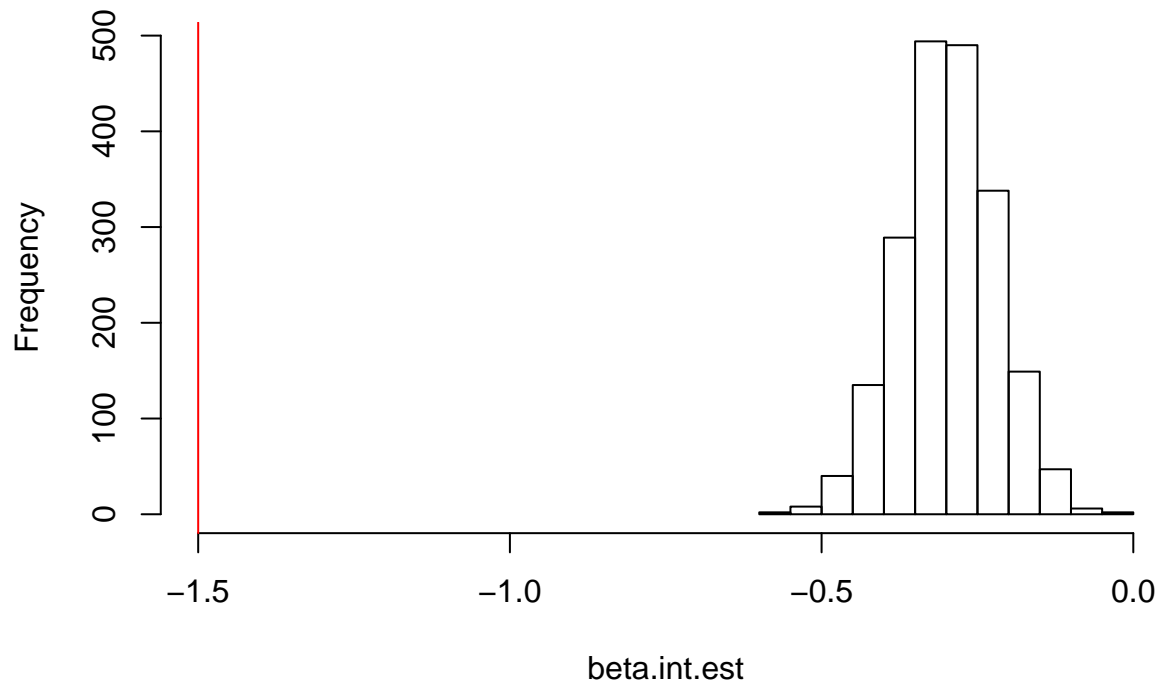
```
##      beta.int.cov.prob <- ((coef(summary(fit.logit))[1, 1] - qt(0.975,
##        df = 2000) * coef(summary(fit.logit))[1, 2]) <= beta0 &
##        (coef(summary(fit.logit))[1, 1] + qt(0.975, df = 2000) *
##          coef(summary(fit.logit))[1, 2]) >= beta0)
##      beta.diff.est <- coef(summary(fit.logit))[2, 1]
##      beta.diff.cov.prob <- ((coef(summary(fit.logit))[2, 1] -
##        qt(0.975, df = 2000) * coef(summary(fit.logit))[2, 2]) <=
##        beta1 & (coef(summary(fit.logit))[2, 1] + qt(0.975, df = 2000) *
##          coef(summary(fit.logit))[2, 2]) >= beta1)
##      beta.x_ibar.est <- coef(summary(fit.logit))[3, 1]
##      beta.x_ibar.cov.prob <- ((coef(summary(fit.logit))[3, 1] -
##        qt(0.975, df = 2000) * coef(summary(fit.logit))[3, 2]) <=
##        beta1 & (coef(summary(fit.logit))[3, 1] + qt(0.975, df = 2000) *
##          coef(summary(fit.logit))[3, 2]) >= beta1)
##      out <- list(beta.int.est, beta.int.cov.prob, beta.diff.est,
##        beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
##      names(out) <- c("Intercept Estimate", "Intercept Covered",
##        "Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
##      return(out)
##    }
}
```

```
out<-foreach(i=1:2000, .combine=cbind) %dopar% {
  logit.sim(pop.m, pop.n, beta1,beta0,tau_sq)
```

```
}
beta.int.est<-unlist(out[1,])
beta.int.cov.prob<-unlist(out[2,])
beta.diff.est<-unlist(out[3,])
beta.diff.cov.prob<-unlist(out[4,])
beta.x_ibar.est<-unlist(out[5,])
beta.x_ibar.cov.prob<-unlist(out[6,])
```

```
hist(beta.int.est,main = "Histogram of intercept estimates",xlim = c(-1.5,0))
abline(v = -1.5, col = "red")
```


Histogram of intercept estimates

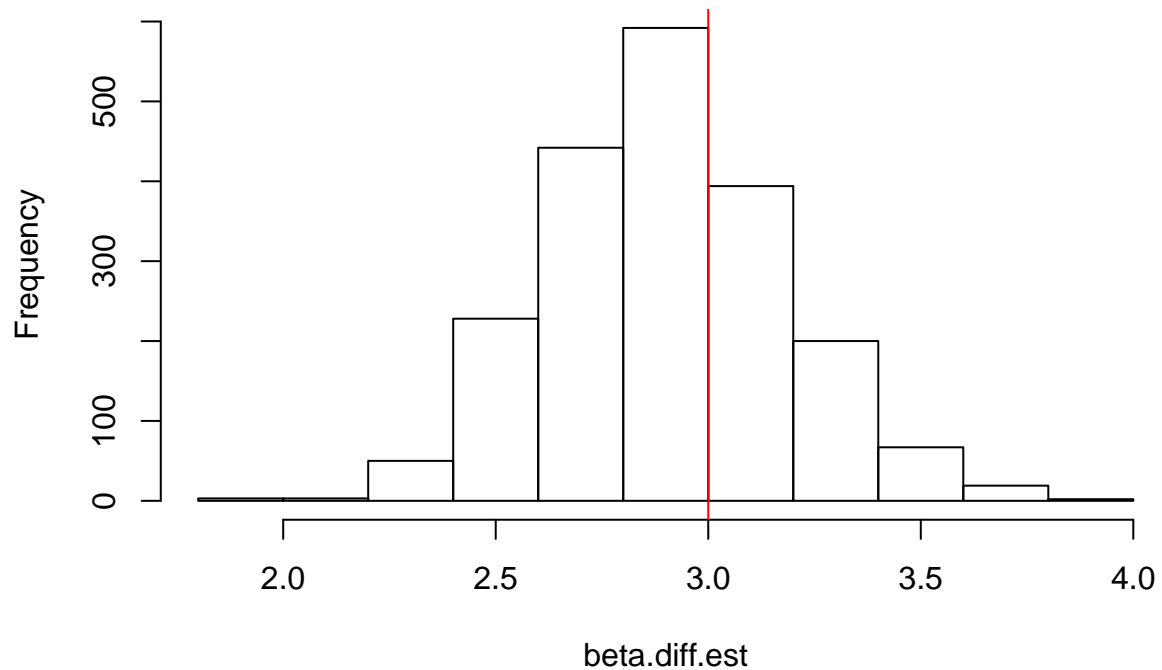


```
mean(beta.int.cov.prob)
```

```
## [1] 0
```

```
hist(beta.diff.est, main = "Coefficient of (x_ij-x_ibar) estimates")  
abline(v = 3, col = "red")
```

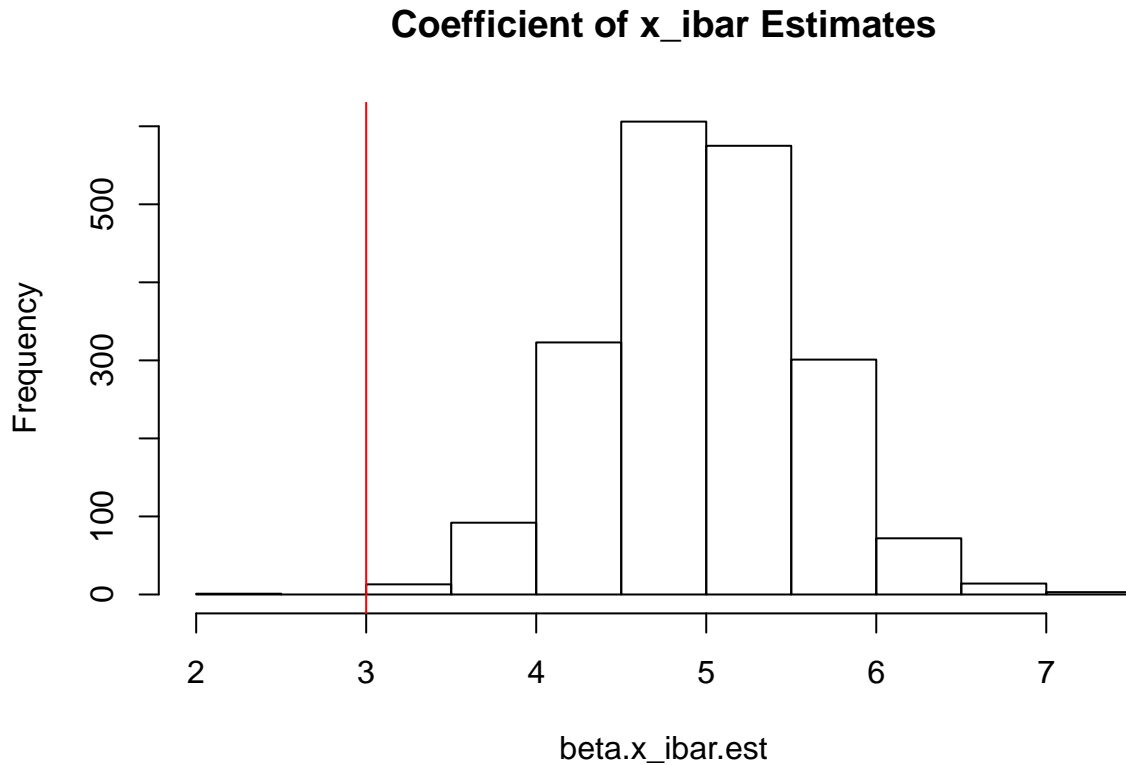
Coefficient of (x_{ij}-x_{ibar}) estimates



```
mean(beta.diff.cov.prob)
```

```
## [1] 0.934
```

```
hist(beta.x_ibar.est, main = "Coefficient of x_ibar Estimates")  
abline(v = 3, col = "red")
```



```
mean(beta.int.cov.prob)
```

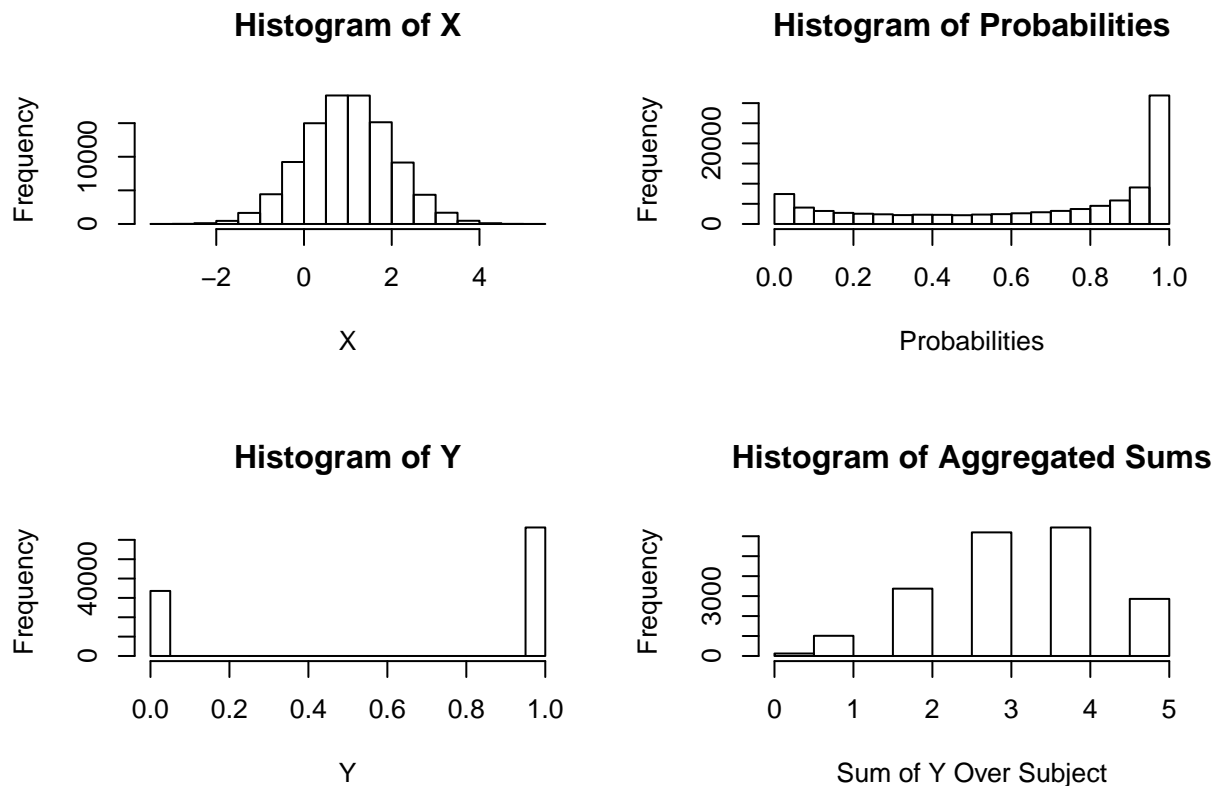
```
## [1] 0
```

The vertical red lines are drawn at the true value for each parameter. The intercept term appears to be biased, as well as the between-subject effect, and possibly also the within-subject effect.

Continuous X

Here I generate the same model as above, except I don't convert the X's to discrete values.

Below, I show some plots from one repetition of my simulation.



```
set.seed(1104)
library(foreach)
library(doParallel)
cores=detectCores()
cl <- makeCluster(3)
registerDoParallel(cl)
clusterEvalQ(cl,
  {library(lme4)
    pop.m<-80000 # number of clusters
    pop.n<- 5 # number within clusters
    beta1<-3 #slope for indicator
    beta0<--1.5 #intercept terms
    tau_sq<-.5 #variance of random intercept
    logit.sim<-function(pop.m, pop.n, beta1,beta0,tau_sq){
      library(lme4)
      u<-rnorm(pop.m,mean = 0, sd=sqrt(tau_sq)) #cluster samples
      u1<-rep(u,each=pop.n) # repeat each cluster sample n times
      x<-rnorm(u1,1)
      z<-beta0+beta1*x+u1
      pr<-1/(1+exp(-z))
      y<-rbinom(n=pop.m*pop.n,size = 1,prob = pr )
      dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
      agg.dat<-aggregate(y~id, dat, sum) # sum y by id
      case.samp<-sample(agg.dat$id[agg.dat$y>=4],250) #sample cases
      control.samp<-sample(agg.dat$id[agg.dat$y<4],250)# sample controls
      samp<-c(case.samp, control.samp)
      samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids
      samp.agg.dat<-aggregate(x~id, samp.dat, mean) #calculate means for x
      x_ibar<-rep(samp.agg.dat$x,each=pop.n) # mach means dimensions with dat
```

```

samp.dat$x_ibar<-(x_ibar)
samp.dat$diff<-(samp.dat$x-samp.dat$x_ibar) # calculate  $x_{ij}-x_{ib\bar{a}r}$ 

fit.logit<-glmer(y~diff+x_ibar+(1|id),data = samp.dat,
  family = binomial(link = "logit"),
  glmerControl(optimizer = c("bobyqa","Nelder_Mead")) )

beta.int.est<-coef(summary(fit.logit))[1,1]
beta.int.cov.prob<-((coef(summary(fit.logit))[1,1]-qt(.975, df=2000)*coef(summary(fit.
beta.diff.est<-coef(summary(fit.logit))[2,1]
beta.diff.cov.prob<-((coef(summary(fit.logit))[2,1]-qt(.975, df=2000)*coef(summary(fit
beta.x_ibar.est<-coef(summary(fit.logit))[3,1]
beta.x_ibar.cov.prob<-((coef(summary(fit.logit))[3,1]-qt(.975, df=2000)*coef(summary(f
out<-list(beta.int.est, beta.int.cov.prob, beta.diff.est, beta.diff.cov.prob, beta.x_ib
names(out)<-c("Intercept Estimate", "Intercept Covered", "Diff Estimate", "Diff Covered
return(out)
}}
)

```

```

## [[1]]
## function (pop.m, pop.n, beta1, beta0, tau_sq)
## {
##   library(lme4)
##   u <- rnorm(pop.m, mean = 0, sd = sqrt(tau_sq))
##   u1 <- rep(u, each = pop.n)
##   x <- rnorm(u1, 1)
##   z <- beta0 + beta1 * x + u1
##   pr <- 1/(1 + exp(-z))
##   y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##   dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   fit.logit <- glmer(y ~ diff + x_ibar + (1 | id), data = samp.dat,
##     family = binomial(link = "logit"), glmerControl(optimizer = c("bobyqa",
##       "Nelder_Mead")))
##   beta.int.est <- coef(summary(fit.logit))[1, 1]
##   beta.int.cov.prob <- ((coef(summary(fit.logit))[1, 1] - qt(0.975,
##     df = 2000) * coef(summary(fit.logit))[1, 2]) <= beta0 &
##     (coef(summary(fit.logit))[1, 1] + qt(0.975, df = 2000) *
##     coef(summary(fit.logit))[1, 2]) >= beta0)
##   beta.diff.est <- coef(summary(fit.logit))[2, 1]
##   beta.diff.cov.prob <- ((coef(summary(fit.logit))[2, 1] -
##     qt(0.975, df = 2000) * coef(summary(fit.logit))[2, 2]) <=
##     beta1 & (coef(summary(fit.logit))[2, 1] + qt(0.975, df = 2000) *
##     coef(summary(fit.logit))[2, 2]) >= beta1)
##   beta.x_ibar.est <- coef(summary(fit.logit))[3, 1]
##   beta.x_ibar.cov.prob <- ((coef(summary(fit.logit))[3, 1] -

```

```

##      qt(0.975, df = 2000) * coef(summary(fit.logit))[3, 2]) <=
##      beta1 & (coef(summary(fit.logit))[3, 1] + qt(0.975, df = 2000) *
##      coef(summary(fit.logit))[3, 2]) >= beta1)
##    out <- list(beta.int.est, beta.int.cov.prob, beta.diff.est,
##      beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
##    names(out) <- c("Intercept Estimate", "Intercept Covered",
##      "Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
##    return(out)
##  }
##
##  [[2]]
##  function (pop.m, pop.n, beta1, beta0, tau_sq)
##  {
##    library(lme4)
##    u <- rnorm(pop.m, mean = 0, sd = sqrt(tau_sq))
##    u1 <- rep(u, each = pop.n)
##    x <- rnorm(u1, 1)
##    z <- beta0 + beta1 * x + u1
##    pr <- 1/(1 + exp(-z))
##    y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##    dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##    agg.dat <- aggregate(y ~ id, dat, sum)
##    case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##    control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##    samp <- c(case.samp, control.samp)
##    samp.dat <- subset(dat, dat$id %in% samp)
##    samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##    x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##    samp.dat$x_ibar <- (x_ibar)
##    samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##    fit.logit <- glmer(y ~ diff + x_ibar + (1 | id), data = samp.dat,
##      family = binomial(link = "logit"), glmerControl(optimizer = c("bobyqa",
##      "Nelder_Mead")))
##    beta.int.est <- coef(summary(fit.logit))[1, 1]
##    beta.int.cov.prob <- ((coef(summary(fit.logit))[1, 1] - qt(0.975,
##      df = 2000) * coef(summary(fit.logit))[1, 2]) <= beta0 &
##      (coef(summary(fit.logit))[1, 1] + qt(0.975, df = 2000) *
##      coef(summary(fit.logit))[1, 2]) >= beta0)
##    beta.diff.est <- coef(summary(fit.logit))[2, 1]
##    beta.diff.cov.prob <- ((coef(summary(fit.logit))[2, 1] -
##      qt(0.975, df = 2000) * coef(summary(fit.logit))[2, 2]) <=
##      beta1 & (coef(summary(fit.logit))[2, 1] + qt(0.975, df = 2000) *
##      coef(summary(fit.logit))[2, 2]) >= beta1)
##    beta.x_ibar.est <- coef(summary(fit.logit))[3, 1]
##    beta.x_ibar.cov.prob <- ((coef(summary(fit.logit))[3, 1] -
##      qt(0.975, df = 2000) * coef(summary(fit.logit))[3, 2]) <=
##      beta1 & (coef(summary(fit.logit))[3, 1] + qt(0.975, df = 2000) *
##      coef(summary(fit.logit))[3, 2]) >= beta1)
##    out <- list(beta.int.est, beta.int.cov.prob, beta.diff.est,
##      beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
##    names(out) <- c("Intercept Estimate", "Intercept Covered",
##      "Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
##    return(out)
##  }

```

```

##
## [[3]]
## function (pop.m, pop.n, beta1, beta0, tau_sq)
## {
##   library(lme4)
##   u <- rnorm(pop.m, mean = 0, sd = sqrt(tau_sq))
##   u1 <- rep(u, each = pop.n)
##   x <- rnorm(u1, 1)
##   z <- beta0 + beta1 * x + u1
##   pr <- 1/(1 + exp(-z))
##   y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##   dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   fit.logit <- glmer(y ~ diff + x_ibar + (1 | id), data = samp.dat,
##     family = binomial(link = "logit"), glmerControl(optimizer = c("bobyqa",
##       "Nelder_Mead")))
##   beta.int.est <- coef(summary(fit.logit))[1, 1]
##   beta.int.cov.prob <- ((coef(summary(fit.logit))[1, 1] - qt(0.975,
##     df = 2000) * coef(summary(fit.logit))[1, 2]) <= beta0 &
##     (coef(summary(fit.logit))[1, 1] + qt(0.975, df = 2000) *
##       coef(summary(fit.logit))[1, 2]) >= beta0)
##   beta.diff.est <- coef(summary(fit.logit))[2, 1]
##   beta.diff.cov.prob <- ((coef(summary(fit.logit))[2, 1] -
##     qt(0.975, df = 2000) * coef(summary(fit.logit))[2, 2]) <=
##     beta1 & (coef(summary(fit.logit))[2, 1] + qt(0.975, df = 2000) *
##       coef(summary(fit.logit))[2, 2]) >= beta1)
##   beta.x_ibar.est <- coef(summary(fit.logit))[3, 1]
##   beta.x_ibar.cov.prob <- ((coef(summary(fit.logit))[3, 1] -
##     qt(0.975, df = 2000) * coef(summary(fit.logit))[3, 2]) <=
##     beta1 & (coef(summary(fit.logit))[3, 1] + qt(0.975, df = 2000) *
##       coef(summary(fit.logit))[3, 2]) >= beta1)
##   out <- list(beta.int.est, beta.int.cov.prob, beta.diff.est,
##     beta.diff.cov.prob, beta.x_ibar.est, beta.x_ibar.cov.prob)
##   names(out) <- c("Intercept Estimate", "Intercept Covered",
##     "Diff Estimate", "Diff Covered", "x_ibar Estimate", "x_ibar Covered")
##   return(out)
## }

```

```

out<-foreach(i=1:2000, .combine=cbind) %dopar% {
  logit.sim(pop.m, pop.n, beta1,beta0,tau_sq)
}
beta.int.est<-unlist(out[1,])
beta.int.cov.prob<-unlist(out[2,])
beta.diff.est<-unlist(out[3,])
beta.diff.cov.prob<-unlist(out[4,])

```

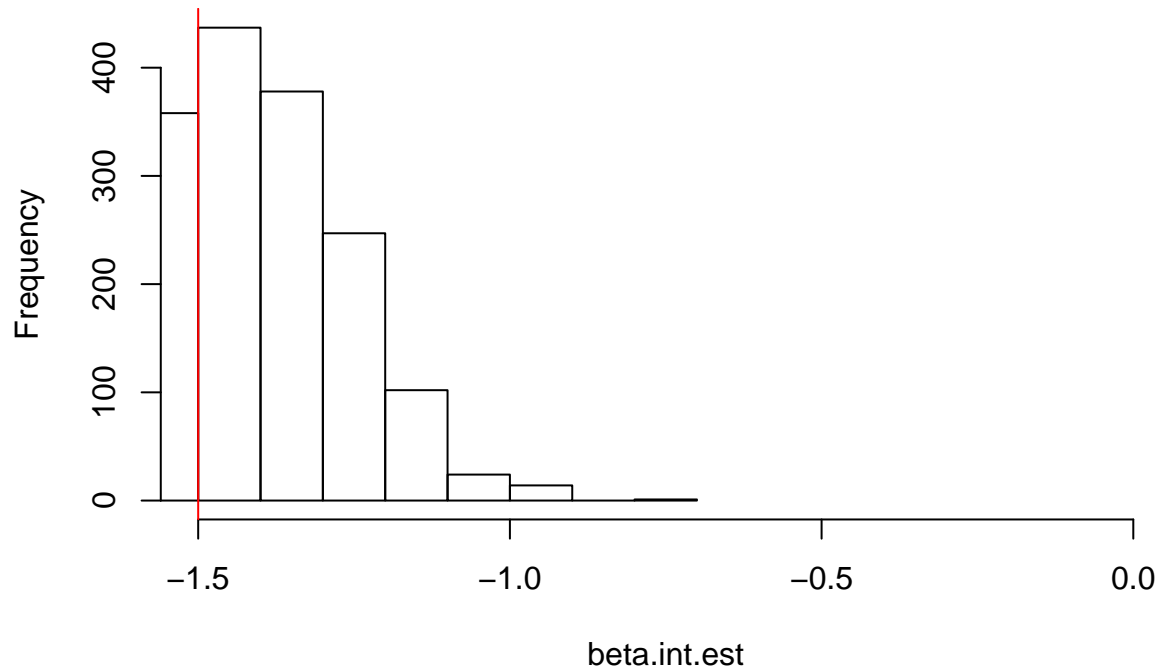
```

beta.x_ibar.est<-unlist(out[5,])
beta.x_ibar.cov.prob<-unlist(out[6,])

hist(beta.int.est,main = "Histogram of intercept estimates",xlim = c(-1.5,0))
abline(v = -1.5, col = "red")

```

Histogram of intercept estimates



```

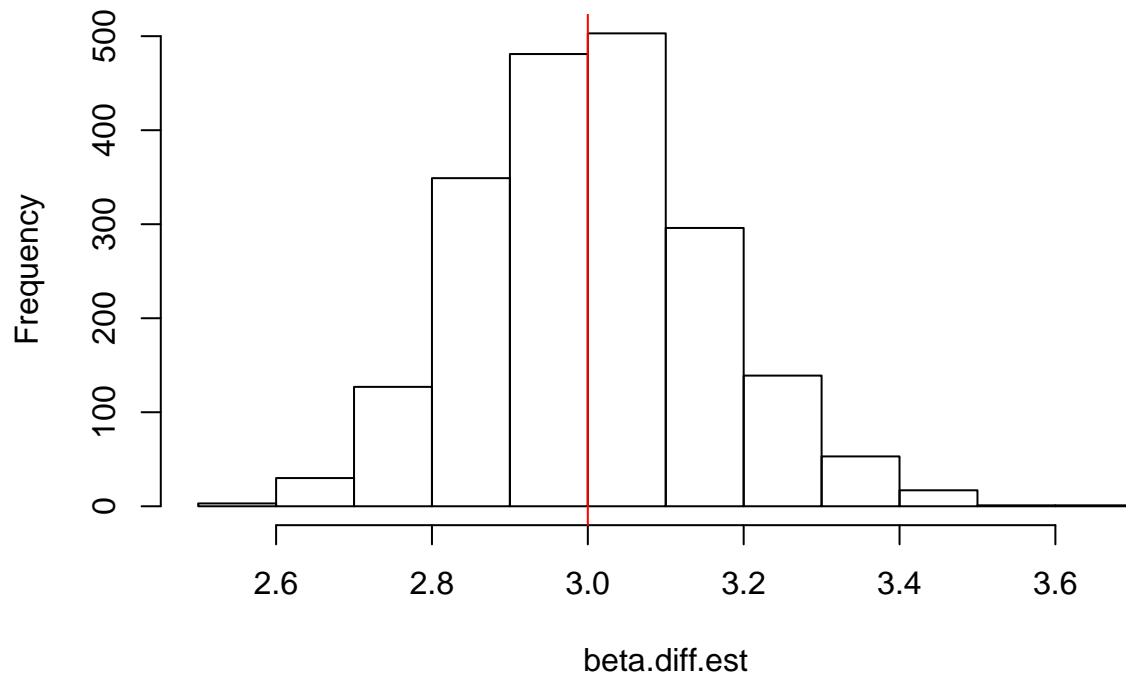
mean(beta.int.cov.prob)

## [1] 0.9195

hist(beta.diff.est, main = "Coefficient of (x_ij-x_ibar) estimates")
abline(v = 3, col = "red")

```

Coefficient of ($x_{ij}-x_{\text{ibar}}$) estimates

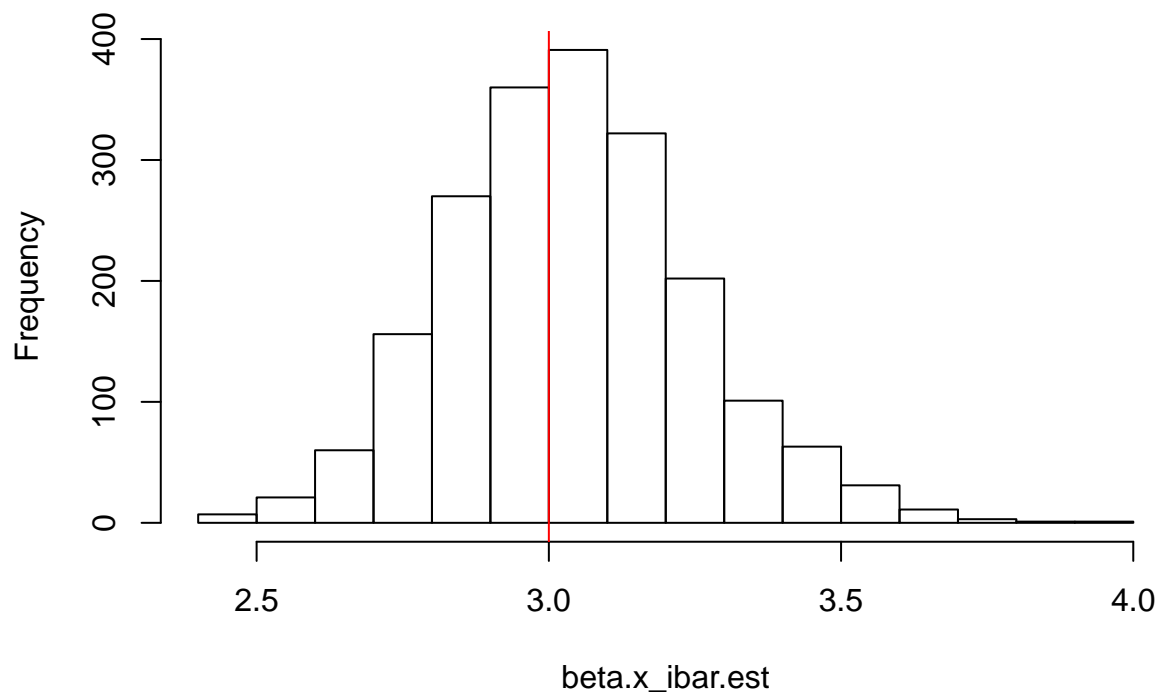


```
mean(beta.diff.cov.prob)
```

```
## [1] 0.9195
```

```
hist(beta.x_ibar.est, main = "Coefficient of x_ibar Estimates")  
abline(v = 3, col = "red")
```

Coefficient of x_{ibar} Estimates




```
mean(beta.int.cov.prob)
```

```
## [1] 0.9195
```

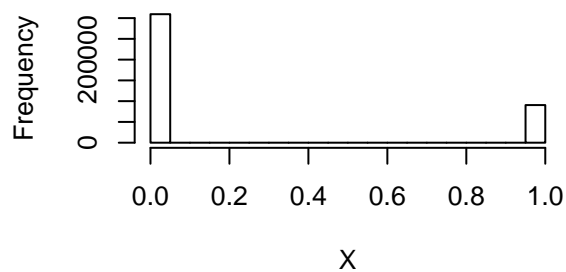
Conditional logistic regression

Discrete X

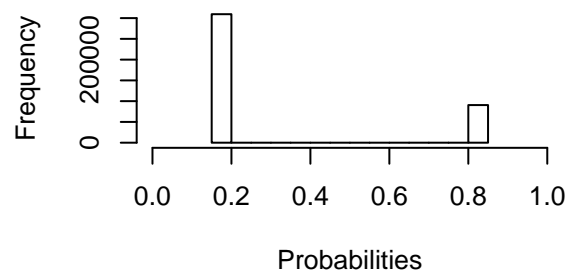
Below, I show some plots from one repetition of my simulation.

```
##  
##      0      1      2      3      4      5  
## 15329 25583 20653 11903  5101  1431
```

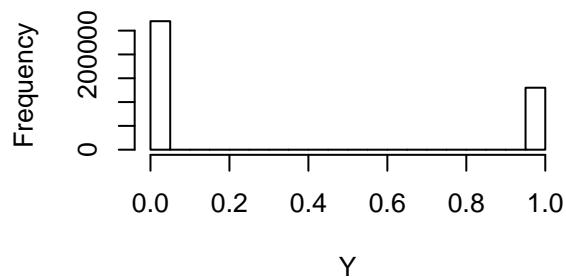
Histogram of X



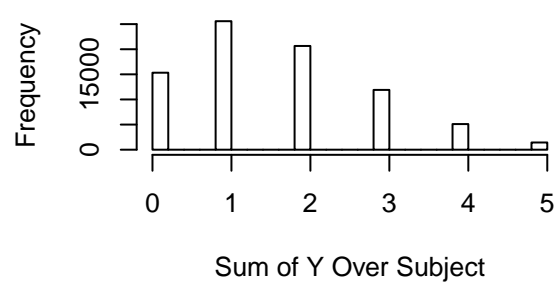
Histogram of Probabilities



Histogram of Y



Histogram of Aggregated Sums



```
set.seed(1104)  
library(foreach)  
library(doParallel)  
cores=detectCores()  
cl <- makeCluster(3)  
registerDoParallel(cl)  
clusterEvalQ(cl,  
  {library(survival)  
    pop.m<-80000 # number of clusters  
    pop.n<- 5 # number within clusters  
    beta1<-3 #slope for indicator  
    beta0<-1.5 #intercept terms  
    tau_x<-0.5  
    sigma<-1  
    clogit.sim<-function(pop.m, pop.n, beta1,beta0,tau_x, sigma){
```

```

v<-rnorm(pop.m,mean = 0, sd=sqrt(sigma*tau_x)) #cluster samples
v1<-rep(v,each=pop.n) # repeat each cluster sample n times
xstar<-rnorm(pop.m*pop.n,mean = 0, sd=sqrt(sigma*(1-tau_x))) # samples within each clu
x<-v1+xstar #total x
x<-as.numeric(x>.75)
z<-beta0+beta1*x
pr<-1/(1+exp(-z))
y<-rbinom(n=pop.m*pop.n,size = 1,prob = pr )
dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
agg.dat<-aggregate(y~id, dat, sum) # sum y by id
table(agg.dat$y)
case.samp<-sample(agg.dat$id[agg.dat$y>=4],250) #sample cases
control.samp<-sample(agg.dat$id[agg.dat$y<4],250)# sample controls
samp<-c(case.samp, control.samp)
samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids
samp.agg.dat<-aggregate(x~id, samp.dat, mean) #calculate means for x
x_ibar<-rep(samp.agg.dat$x,each=pop.n) # mach means dimensions with dat
samp.dat$x_ibar<-(x_ibar)
samp.dat$diff<-(samp.dat$x-samp.dat$x_ibar) # calculate x_ij-x_ibar
table(samp.dat$diff,samp.dat$y)
fit.clogit<-clogit(y ~ diff + strata(id), samp.dat)
beta.diff.est<-coef(fit.clogit)[1]
ci<-confint(fit.clogit)
beta.diff.cov.prob<-(ci[1,1]<=beta1&ci[1,2]>=beta1)
out<-list(beta.diff.est, beta.diff.cov.prob)
names(out)<-c("Diff Estimate", "Diff Covered")
return(out)
})

```

```

## [[1]]
## function (pop.m, pop.n, beta1, beta0, tau_x, sigma)
## {
##   v <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_x))
##   v1 <- rep(v, each = pop.n)
##   xstar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##     (1 - tau_x)))
##   x <- v1 + xstar
##   x <- as.numeric(x > 0.75)
##   z <- beta0 + beta1 * x
##   pr <- 1/(1 + exp(-z))
##   y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##   dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   table(agg.dat$y)
##   case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   table(samp.dat$diff, samp.dat$y)
##   fit.clogit <- clogit(y ~ diff + strata(id), samp.dat)

```

```

##      beta.diff.est <- coef(fit.clogit)[1]
##      ci <- confint(fit.clogit)
##      beta.diff.cov.prob <- (ci[1, 1] <= beta1 & ci[1, 2] >= beta1)
##      out <- list(beta.diff.est, beta.diff.cov.prob)
##      names(out) <- c("Diff Estimate", "Diff Covered")
##      return(out)
## }
##
## [[2]]
## function (pop.m, pop.n, beta1, beta0, tau_x, sigma)
## {
##     v <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_x))
##     v1 <- rep(v, each = pop.n)
##     xstar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_x)))
##     x <- v1 + xstar
##     x <- as.numeric(x > 0.75)
##     z <- beta0 + beta1 * x
##     pr <- 1/(1 + exp(-z))
##     y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     table(agg.dat$y)
##     case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)
##     samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##     x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##     samp.dat$x_ibar <- (x_ibar)
##     samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##     table(samp.dat$diff, samp.dat$y)
##     fit.clogit <- clogit(y ~ diff + strata(id), samp.dat)
##     beta.diff.est <- coef(fit.clogit)[1]
##     ci <- confint(fit.clogit)
##     beta.diff.cov.prob <- (ci[1, 1] <= beta1 & ci[1, 2] >= beta1)
##     out <- list(beta.diff.est, beta.diff.cov.prob)
##     names(out) <- c("Diff Estimate", "Diff Covered")
##     return(out)
## }
##
## [[3]]
## function (pop.m, pop.n, beta1, beta0, tau_x, sigma)
## {
##     v <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_x))
##     v1 <- rep(v, each = pop.n)
##     xstar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_x)))
##     x <- v1 + xstar
##     x <- as.numeric(x > 0.75)
##     z <- beta0 + beta1 * x
##     pr <- 1/(1 + exp(-z))
##     y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))

```

```
##   agg.dat <- aggregate(y ~ id, dat, sum)
##   table(agg.dat$y)
##   case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##   control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##   samp <- c(case.samp, control.samp)
##   samp.dat <- subset(dat, dat$id %in% samp)
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   table(samp.dat$diff, samp.dat$y)
##   fit.clogit <- clogit(y ~ diff + strata(id), samp.dat)
##   beta.diff.est <- coef(fit.clogit)[1]
##   ci <- confint(fit.clogit)
##   beta.diff.cov.prob <- (ci[1, 1] <= beta1 & ci[1, 2] >= beta1)
##   out <- list(beta.diff.est, beta.diff.cov.prob)
##   names(out) <- c("Diff Estimate", "Diff Covered")
##   return(out)
## }
```

```
out<-foreach(i=1:2000, .combine=cbind) %dopar% {
  clogit.sim(pop.m, pop.n, beta1,beta0,tau_x, sigma)
```

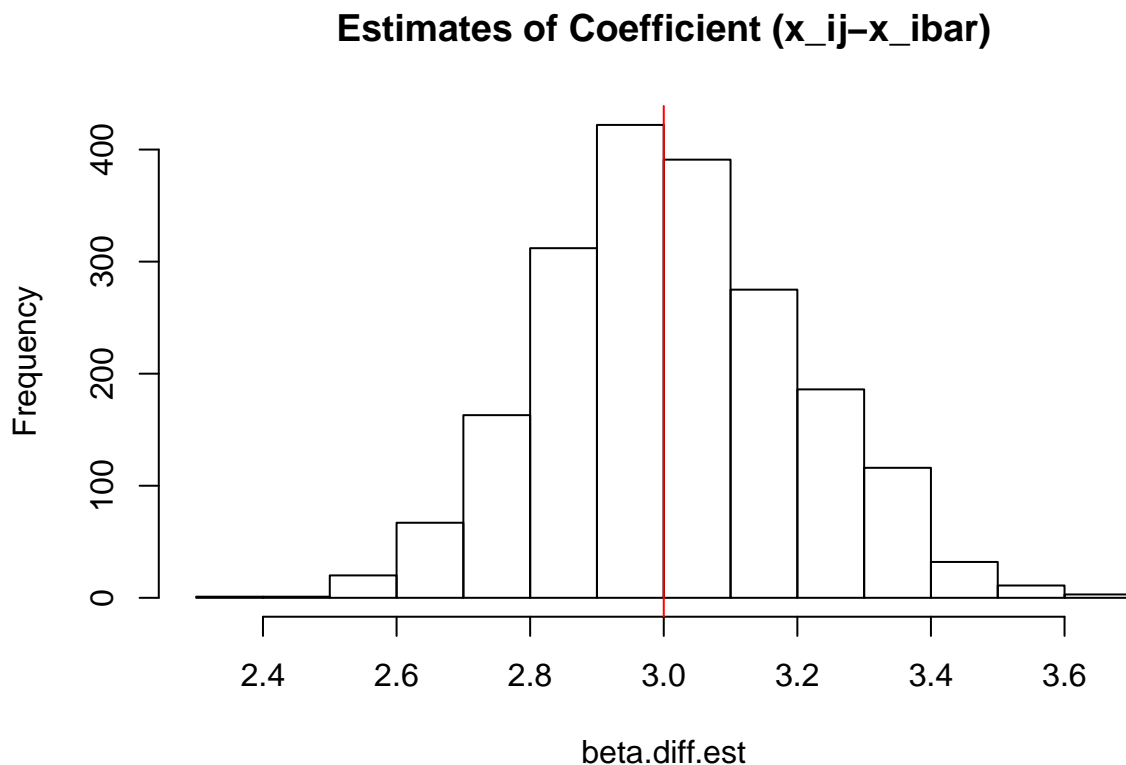
```
}
```

```
beta.diff.est<-unlist(out[1,])
```

```
beta.diff.cov.prob<-unlist(out[2,])
```

```
hist(beta.diff.est, main = "Estimates of Coefficient (xij-xibar)")
```

```
abline(v = 3, col = "red")
```



```
mean(beta.diff.cov.prob)
```

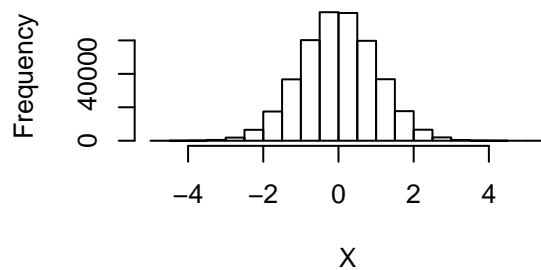
```
## [1] 0.959
```

Continuous X

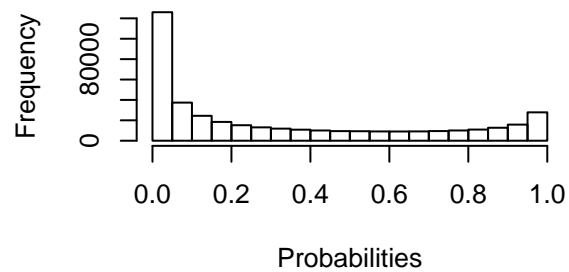
Below, I show some plots from one repetition of my simulation.

```
##
##      0      1      2      3      4      5
## 21912 20016 15675 11503  7344  3550
```

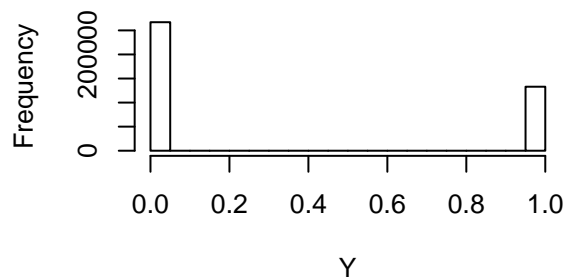
Histogram of X



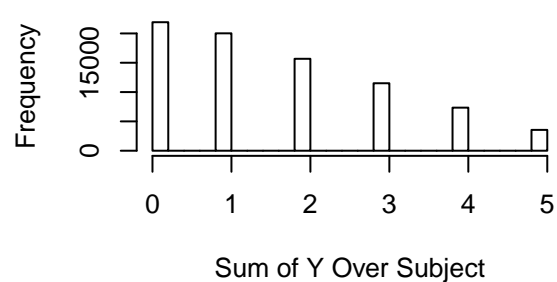
Histogram of Probabilities



Histogram of Y



Histogram of Aggregated Sums



```
set.seed(1104)
library(foreach)
library(doParallel)
cores=detectCores()
cl <- makeCluster(3)
registerDoParallel(cl)
clusterEvalQ(cl,
  {library(survival)
    pop.m<-80000 # number of clusters
    pop.n<- 5 # number within clusters
    beta1<-3 #slope for indicator
    beta0<- -1.5 #intercept terms
    tau_x<-0.5
    sigma<-1
    clogit.sim<-function(pop.m, pop.n, beta1,beta0,tau_x, sigma){
      v<-rnorm(pop.m,mean = 0, sd=sqrt(sigma*tau_x)) #cluster samples
      v1<-rep(v,each=pop.n) # repeat each cluster sample n times
```

```

xstar<-rnorm(pop.m*pop.n,mean = 0, sd=sqrt(sigma*(1-tau_x))) # samples within each clu
x<-v1+xstar #total x
z<-beta0+beta1*x
pr<-1/(1+exp(-z))
y<-rbinom(n=pop.m*pop.n,size = 1,prob = pr )
dat<-data.frame(y=y,x=x,id=rep(c(1:pop.m),each=pop.n)) #make data
agg.dat<-aggregate(y~id, dat, sum) # sum y by id
table(agg.dat$y)
case.samp<-sample(agg.dat$id[agg.dat$y>=4],250) #sample cases
control.samp<-sample(agg.dat$id[agg.dat$y<4],250)# sample controls
samp<-c(case.samp, control.samp)
samp.dat<-subset(dat,dat$id%in%samp) # get dataframe for sampled ids
samp.agg.dat<-aggregate(x~id, samp.dat, mean) #calculate means for x
x_ibar<-rep(samp.agg.dat$x,each=pop.n) # mach means dimensions with dat
samp.dat$x_ibar<-(x_ibar)
samp.dat$diff<-(samp.dat$x-samp.dat$x_ibar) # calculate x_ij-x_ibar
table(samp.dat$diff,samp.dat$y)
fit.clogit<-clogit(y ~ diff + strata(id), samp.dat)
beta.diff.est<-coef(fit.clogit)[1]
ci<-confint(fit.clogit)
beta.diff.cov.prob<-(ci[1,1]<=beta1&ci[1,2]>=beta1)
out<-list(beta.diff.est, beta.diff.cov.prob)
names(out)<-c("Diff Estimate", "Diff Covered")
return(out)
})

```

```

## [[1]]
## function (pop.m, pop.n, beta1, beta0, tau_x, sigma)
## {
##     v <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_x))
##     v1 <- rep(v, each = pop.n)
##     xstar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_x)))
##     x <- v1 + xstar
##     z <- beta0 + beta1 * x
##     pr <- 1/(1 + exp(-z))
##     y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     table(agg.dat$y)
##     case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)
##     samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##     x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##     samp.dat$x_ibar <- (x_ibar)
##     samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##     table(samp.dat$diff, samp.dat$y)
##     fit.clogit <- clogit(y ~ diff + strata(id), samp.dat)
##     beta.diff.est <- coef(fit.clogit)[1]
##     ci <- confint(fit.clogit)
##     beta.diff.cov.prob <- (ci[1, 1] <= beta1 & ci[1, 2] >= beta1)
##     out <- list(beta.diff.est, beta.diff.cov.prob)
##

```

```

##     names(out) <- c("Diff Estimate", "Diff Covered")
##     return(out)
## }
##
## [[2]]
## function (pop.m, pop.n, beta1, beta0, tau_x, sigma)
## {
##     v <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_x))
##     v1 <- rep(v, each = pop.n)
##     xstar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_x)))
##     x <- v1 + xstar
##     z <- beta0 + beta1 * x
##     pr <- 1/(1 + exp(-z))
##     y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     table(agg.dat$y)
##     case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)
##     samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##     x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##     samp.dat$x_ibar <- (x_ibar)
##     samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##     table(samp.dat$diff, samp.dat$y)
##     fit.clogit <- clogit(y ~ diff + strata(id), samp.dat)
##     beta.diff.est <- coef(fit.clogit)[1]
##     ci <- confint(fit.clogit)
##     beta.diff.cov.prob <- (ci[1, 1] <= beta1 & ci[1, 2] >= beta1)
##     out <- list(beta.diff.est, beta.diff.cov.prob)
##     names(out) <- c("Diff Estimate", "Diff Covered")
##     return(out)
## }
##
## [[3]]
## function (pop.m, pop.n, beta1, beta0, tau_x, sigma)
## {
##     v <- rnorm(pop.m, mean = 0, sd = sqrt(sigma * tau_x))
##     v1 <- rep(v, each = pop.n)
##     xstar <- rnorm(pop.m * pop.n, mean = 0, sd = sqrt(sigma *
##         (1 - tau_x)))
##     x <- v1 + xstar
##     z <- beta0 + beta1 * x
##     pr <- 1/(1 + exp(-z))
##     y <- rbinom(n = pop.m * pop.n, size = 1, prob = pr)
##     dat <- data.frame(y = y, x = x, id = rep(c(1:pop.m), each = pop.n))
##     agg.dat <- aggregate(y ~ id, dat, sum)
##     table(agg.dat$y)
##     case.samp <- sample(agg.dat$id[agg.dat$y >= 4], 250)
##     control.samp <- sample(agg.dat$id[agg.dat$y < 4], 250)
##     samp <- c(case.samp, control.samp)
##     samp.dat <- subset(dat, dat$id %in% samp)

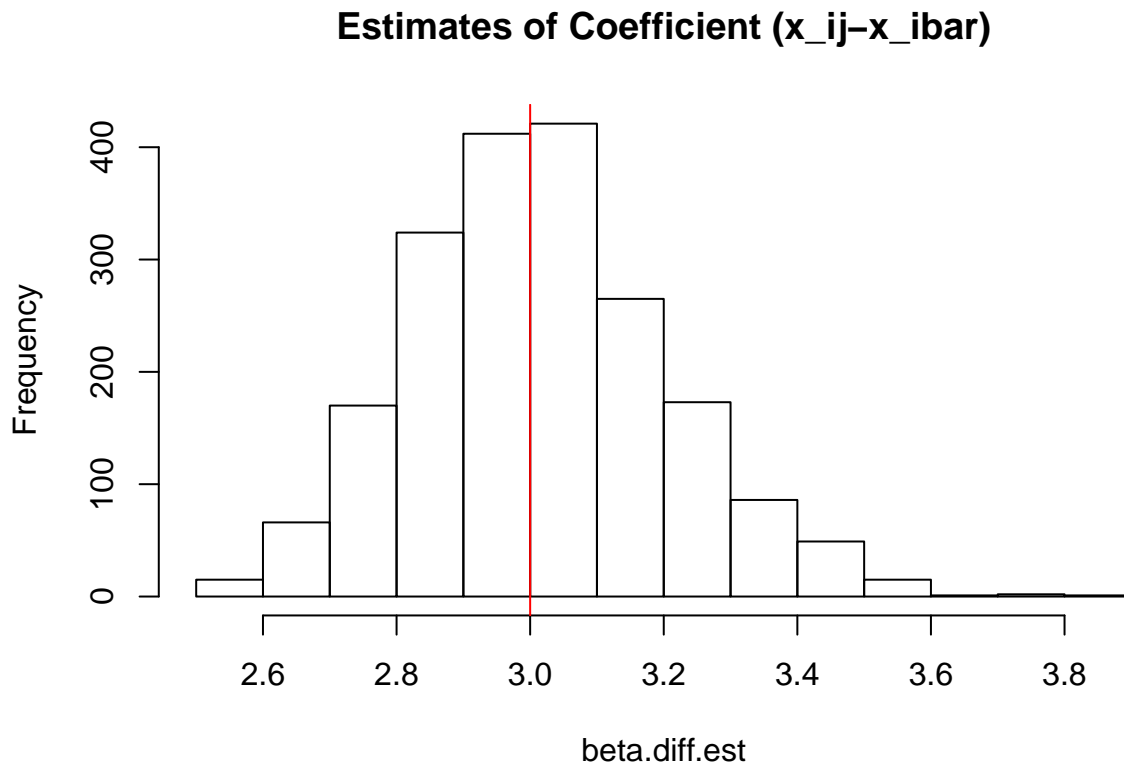
```

```
##   samp.agg.dat <- aggregate(x ~ id, samp.dat, mean)
##   x_ibar <- rep(samp.agg.dat$x, each = pop.n)
##   samp.dat$x_ibar <- (x_ibar)
##   samp.dat$diff <- (samp.dat$x - samp.dat$x_ibar)
##   table(samp.dat$diff, samp.dat$y)
##   fit.clogit <- clogit(y ~ diff + strata(id), samp.dat)
##   beta.diff.est <- coef(fit.clogit)[1]
##   ci <- confint(fit.clogit)
##   beta.diff.cov.prob <- (ci[1, 1] <= beta1 & ci[1, 2] >= beta1)
##   out <- list(beta.diff.est, beta.diff.cov.prob)
##   names(out) <- c("Diff Estimate", "Diff Covered")
##   return(out)
## }
```

```
out<-foreach(i=1:2000, .combine=cbind) %dopar% {
  clogit.sim(pop.m, pop.n, beta1,beta0,tau_x, sigma)
}
```

```
beta.diff.est<-unlist(out[1,])
beta.diff.cov.prob<-unlist(out[2,])
```

```
hist(beta.diff.est, main = "Estimates of Coefficient (x_ij-x_ibar)")
abline(v = 3, col = "red")
```



```
mean(beta.diff.cov.prob)
```

```
## [1] 0.954
```