

# stat\_analysis

Ryan Wei

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```
source(knitr::purl("./data_manipulation.Rmd", quiet=TRUE))
source(knitr::purl("./EDA.Rmd", quiet=TRUE))
```

```
data.baseline <- data.comp %>% filter(day == 0)
```

## Complete case GEE

```
library(gee)
library(geepack)
library(glmtoolbox)
library(gtsummary)
library(tidyverse)
data.comp.lag <-
  data.comp %>%
  group_by(subject_id) %>%
  mutate(mem_comp_lag = lag(mem_comp), observed_lag = lag(observed)) %>%
  ungroup() %>%
  filter(observed_lag != 0) %>%
  mutate(drop = ifelse(observed == 0 & observed_lag == 1, 1, 0)) %>%
  dplyr::select(subject_id, day, day_fct, age, gender, treatment_group, mem_comp_lag, drop) %>%
  mutate(day_fct = factor(day, levels = c(5,19,90)))

model.complete.1 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct,
  model.complete.2 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct,
  model.complete.3 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct,

#tbl_regression(model.complete.1)
#tbl_regression(model.complete.2)
#tbl_regression(model.complete.3)

tbl_merge(
  list(tbl_regression(model.complete.1), tbl_regression(model.complete.2), tbl_regression(model.complete.3))
)
```

Characteristic

Beta

95% CI

**p-value**

**Beta**

**95% CI**

**p-value**

**Beta**

**95% CI**

**p-value**

day\_\_fct

0

—

—

—

—

—

—

5

-0.10

-0.34, 0.15

0.4

-0.10

-0.34, 0.15

0.4

-0.10

-0.34, 0.15

0.4

19

-0.28

-0.65, 0.08

0.13

-0.24

-0.58, 0.10

0.2

-0.22

-0.53, 0.10

0.2

90

-0.30  
-0.73, 0.13  
0.2  
-0.13  
-0.50, 0.25  
0.5  
-0.16  
-0.50, 0.19  
0.4  
treatment\_group

A

—  
—  
—  
—  
—  
—

B

-0.08  
-0.73, 0.57  
0.8  
-0.10  
-0.76, 0.56  
0.8  
-0.09  
-0.75, 0.57  
0.8

C

-0.19  
-0.72, 0.34  
0.5  
-0.18  
-0.71, 0.36  
0.5  
-0.19  
-0.72, 0.35

0.5  
 age  
 -0.02  
 -0.05, 0.02  
 0.4  
 -0.02  
 -0.05, 0.02  
 0.3  
 -0.02  
 -0.05, 0.01  
 0.2  
 gender  
 M  
 —  
 —  
 —  
 —  
 —  
 —  
 F  
 -0.06  
 -0.61, 0.50  
 0.8  
 0.02  
 -0.51, 0.55  
 >0.9  
 -0.03  
 -0.59, 0.53  
 >0.9  
 day\_fct \* treatment\_group  
 5 \* B  
 -0.06  
 -0.53, 0.42  
 0.8  
 -0.05  
 -0.51, 0.40

0.8  
-0.06  
-0.52, 0.40  
0.8  
19 \* B  
0.22  
-0.35, 0.79  
0.4  
0.24  
-0.22, 0.69  
0.3  
0.18  
-0.23, 0.60  
0.4  
90 \* B  
0.25  
-0.37, 0.88  
0.4  
0.29  
-0.18, 0.75  
0.2  
0.23  
-0.23, 0.69  
0.3  
5 \* C  
0.35  
0.00, 0.70  
0.053  
0.37  
0.03, 0.72  
0.032  
0.38  
0.04, 0.73  
0.031  
19 \* C  
0.71

0.21, 1.2  
 0.005  
 0.68  
 0.23, 1.1  
 0.003  
 0.66  
 0.22, 1.1  
 0.003  
 90 \* C  
 0.39  
 -0.17, 0.95  
 0.2  
 0.32  
 -0.20, 0.85  
 0.2  
 0.31  
 -0.17, 0.79  
 0.2

## Testing MCAR

```
model.drop <-  
  geeglm(drop ~ day_fct + treatment_group + age + gender + mem_comp_lag, data = data.comp.lag, family =  
  tbl_regression(model.drop, exponentiate = F)
```

**Characteristic**

**log(OR)**

**95% CI**

**p-value**

day\_\_fct

5

—

—

19

2.0

0.41, 3.7

0.014  
 90  
 2.3  
 0.61, 4.1  
 0.008  
 treatment\_group  
 A  
 —  
 —  
 B  
 0.06  
 -1.2, 1.3  
 >0.9  
 C  
 -0.25  
 -1.3, 0.84  
 0.6  
 age  
 -0.02  
 -0.06, 0.03  
 0.4  
 gender  
 M  
 —  
 —  
 F  
 0.70  
 -0.25, 1.7  
 0.2  
 mem\_comp\_lag  
 0.24  
 -0.35, 0.83  
 0.4  
 No significant evidence of violating MCAR. . .

## Linear mixed effect models

```
library(lme4)
library(nlme)
library(lattice)
library(knitr)
library(kableExtra)

model.lmer = lmer(
  mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct + (1 | subject_id), data = data
)

tbl_regression(model.lmer)
```

### Characteristic

#### Beta

#### 95% CI

day\_fct

0

—

—

5

-0.10

-0.37, 0.18

19

-0.21

-0.51, 0.08

90

-0.16

-0.47, 0.16

treatment\_group

A

—

—

B

-0.09

-0.77, 0.59

C

-0.19

-0.84, 0.46



```

age
-0.02
-0.05, 0.01
gender
M
—
—
F
-0.03
-0.56, 0.50
day_fct * treatment_group
5 * B
-0.06
-0.47, 0.35
19 * B
0.18
-0.28, 0.64
90 * B
0.23
-0.23, 0.69
5 * C
0.38
-0.01, 0.78
19 * C
0.66
0.23, 1.1
90 * C
0.31
-0.15, 0.77

```

## IPW GEE

```

# propensity score models, using logistic regression and coxph model
model.prob.drop <-
  geeglm(dropped ~ day_fct + treatment_group + age + gender, data = data.surv %>% mutate(day_fct = factor(
    day_fct)))
model.prob.drop.coxph <- coxph(Surv(start, stop, dropped) ~ treatment_group + age + gender, data = data.surv)
tbl_regression(model.prob.drop, exponentiate = F)

```

**Characteristic**

**log(OR)**

**95% CI**

**p-value**

day\_fct

5

—

—

19

2.1

0.56, 3.6

0.007

90

2.5

0.84, 4.2

0.003

treatment\_group

A

—

—

B

0.27

-0.86, 1.4

0.6

C

0.10

-0.98, 1.2

0.9

age

-0.03

-0.07, 0.01

0.2

gender

M

—

—

F  
0.48  
-0.45, 1.4  
0.3

```
tbl_regression(model.prob.drop.coxph, exponentiate = F)
```

**Characteristic**

**log(HR)**

**95% CI**

**p-value**

treatment\_group

A

—

—

B

0.20

-0.69, 1.1

0.7

C

0.09

-0.79, 0.97

0.8

age

-0.03

-0.07, 0.02

0.2

gender

M

—

—

F

0.38

-0.36, 1.1

0.3

```
tbl_merge(  
  list(tbl_regression(model.prob.drop, exponentiate = F),tbl_regression(model.prob.drop.coxph, exponentiate = F))  
)
```

**Characteristic**

**log(OR)**

**95% CI**

**p-value**

**log(HR)**

**95% CI**

**p-value**

day\_\_fct

5

—

—

19

2.1

0.56, 3.6

0.007

90

2.5

0.84, 4.2

0.003

treatment\_group

A

—

—

—

—

B

0.27

-0.86, 1.4

0.6

0.20

-0.69, 1.1

0.7

C

0.10

-0.98, 1.2

0.9

0.09  
-0.79, 0.97

0.8

age

-0.03

-0.07, 0.01

0.2

-0.03

-0.07, 0.02

0.2

gender

M

—

—

—

—

F

0.48

-0.45, 1.4

0.3

0.38

-0.36, 1.1

0.3

```
predict.prob.drop <- predict(model.prob.drop, newdata = data.surv, type = "response")
predict.prob.drop.coxph <- predict(model.prob.drop.coxph, newdata = data.surv, type = "survival")

data.surv$prob.stay <- 1 - predict.prob.drop
data.surv$prob.stay.coxph <- predict.prob.drop.coxph

data.comp <- left_join(data.comp, data.surv %>% dplyr::select(subject_id, day_fct, treatment_group, prob.stay, prob.stay.coxph))
data.comp <-
  data.comp %>%
  mutate(
    prob.stay = ifelse(day_fct==0, 1, prob.stay),
    prob.stay.coxph = ifelse(day_fct==0, 1, prob.stay.coxph)
  )
```

```
model.ipw.1 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct, data = data.comp, weights = prob.stay, family = geeglm.family)
model.ipw.2 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct, data = data.comp, weights = prob.stay.coxph, family = geeglm.family)
model.ipw.3 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct, data = data.comp, weights = 1, family = geeglm.family)
```

```
tbl_merge(
  list(tbl_regression(model.ipw.1), tbl_regression(model.ipw.2), tbl_regression(model.ipw.3))
)
```

**Characteristic**

**Beta**

**95% CI**

**p-value**

**Beta**

**95% CI**

**p-value**

**Beta**

**95% CI**

**p-value**

day\_\_fct

0

—

—

—

—

—

—

5

-0.10

-0.35, 0.15

0.4

-0.10

-0.35, 0.15

0.4

-0.10

-0.34, 0.15

0.4

19

-0.29

-0.66, 0.07

0.12

-0.24

-0.59, 0.10  
 0.2  
 -0.22  
 -0.54, 0.09  
 0.2  
 90  
 -0.32  
 -0.75, 0.11  
 0.14  
 -0.14  
 -0.51, 0.23  
 0.5  
 -0.19  
 -0.53, 0.15  
 0.3  
 treatment\_group  
 A  
 —  
 —  
 —  
 —  
 —  
 —  
 B  
 -0.07  
 -0.72, 0.58  
 0.8  
 -0.10  
 -0.75, 0.55  
 0.8  
 -0.11  
 -0.76, 0.54  
 0.7  
 C  
 -0.18  
 -0.71, 0.35

0.5  
 -0.17  
 -0.70, 0.35  
 0.5  
 -0.22  
 -0.74, 0.31  
 0.4  
 age  
 -0.01  
 -0.05, 0.02  
 0.5  
 -0.01  
 -0.05, 0.02  
 0.4  
 -0.02  
 -0.06, 0.02  
 0.4  
 gender  
 M  
 —  
 —  
 —  
 —  
 —  
 —  
 F  
 -0.05  
 -0.61, 0.52  
 0.9  
 0.06  
 -0.47, 0.60  
 0.8  
 -0.01  
 -0.61, 0.59  
 >0.9  
 day\_fct \* treatment\_group



$5 * B$   
 -0.05  
 -0.53, 0.43  
 0.8  
 -0.05  
 -0.50, 0.41  
 0.8  
 -0.05  
 -0.51, 0.41  
 0.8  
 $19 * B$   
 0.22  
 -0.36, 0.80  
 0.4  
 0.25  
 -0.21, 0.70  
 0.3  
 0.19  
 -0.23, 0.60  
 0.4  
 $90 * B$   
 0.25  
 -0.39, 0.89  
 0.4  
 0.26  
 -0.21, 0.73  
 0.3  
 0.23  
 -0.24, 0.70  
 0.3  
 $5 * C$   
 0.36  
 0.00, 0.71  
 0.050  
 0.38  
 0.03, 0.72

0.031

0.39

0.04, 0.74

0.027

19 \* C

0.74

0.25, 1.2

0.003

0.70

0.25, 1.1

0.002

0.68

0.25, 1.1

0.002

90 \* C

0.40

-0.19, 0.99

0.2

0.31

-0.24, 0.85

0.3

0.32

-0.18, 0.81

0.2

```
model.ipw.coxph.1 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct, data = data, weights = ipw, family = poisson)
model.ipw.coxph.2 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct, data = data, weights = ipw, family = poisson)
model.ipw.coxph.3 <- geeglm(mem_comp ~ day_fct + treatment_group + age + gender + treatment_group * day_fct, data = data, weights = ipw, family = poisson)
```

`tbl_merge(`

`list(tbl_regression(model.ipw.coxph.1), tbl_regression(model.ipw.coxph.2), tbl_regression(model.ipw.coxph.3))`

**Characteristic**

**Beta**

**95% CI**

**p-value**

**Beta**

**95% CI**

**p-value**

**Beta**

**95% CI**

**p-value**

day\_\_fct

0

—

—

—

—

—

—

5

-0.10

-0.35, 0.15

0.4

-0.10

-0.34, 0.15

0.4

-0.10

-0.34, 0.15

0.4

19

-0.29

-0.66, 0.07

0.12

-0.24

-0.59, 0.10

0.2

-0.22

-0.54, 0.09

0.2

90

-0.32

-0.75, 0.11

0.14

-0.14  
 -0.51, 0.23  
 0.5  
 -0.19  
 -0.53, 0.15  
 0.3  
 treatment\_group  
 A  
 —  
 —  
 —  
 —  
 —  
 —  
 B  
 -0.07  
 -0.72, 0.58  
 0.8  
 -0.11  
 -0.76, 0.54  
 0.7  
 -0.12  
 -0.76, 0.53  
 0.7  
 C  
 -0.18  
 -0.71, 0.35  
 0.5  
 -0.18  
 -0.70, 0.35  
 0.5  
 -0.22  
 -0.75, 0.31  
 0.4  
 age  
 -0.01

-0.05, 0.02  
 0.5  
 -0.01  
 -0.05, 0.02  
 0.4  
 -0.02  
 -0.06, 0.02  
 0.4  
 gender  
 M  
 —  
 —  
 —  
 —  
 —  
 —  
 F  
 -0.05  
 -0.61, 0.52  
 0.9  
 0.06  
 -0.48, 0.60  
 0.8  
 -0.01  
 -0.61, 0.59  
 >0.9  
 day\_fct \* treatment\_group  
 5 \* B  
 -0.05  
 -0.53, 0.43  
 0.8  
 -0.05  
 -0.50, 0.41  
 0.8  
 -0.05  
 -0.51, 0.41

0.8  
19 \* B  
0.22  
-0.36, 0.80  
0.5  
0.24  
-0.21, 0.70  
0.3  
0.19  
-0.22, 0.60  
0.4  
90 \* B  
0.25  
-0.40, 0.89  
0.5  
0.26  
-0.21, 0.73  
0.3  
0.23  
-0.24, 0.69  
0.3  
5 \* C  
0.35  
0.00, 0.71  
0.050  
0.38  
0.03, 0.72  
0.031  
0.39  
0.04, 0.74  
0.027  
19 \* C  
0.74  
0.25, 1.2  
0.003  
0.69

0.25, 1.1

0.002

0.68

0.25, 1.1

0.002

90 \* C

0.40

-0.19, 0.99

0.2

0.30

-0.24, 0.85

0.3

0.32

-0.18, 0.81

0.2