## **G** - Computation: Code Example

## **Estimating average response**

```
mean_response <- function(treat_sequence, coef_estimates, MC) {</pre>
    mu0 <- coef_estimates[1]; sigma0 <- coef_estimates[2]</pre>
    beta0 <- coef_estimates[3]; beta1 <- coef_estimates[4]; beta2 <- coef_estimates[5];</pre>
    beta3 <- coef_estimates[6]; beta4 <- coef_estimates[7];</pre>
    beta5 <- coef_estimates[8]; beta6 <- coef_estimates[9];</pre>
    beta7 <- coef_estimates[10];</pre>
    sigma <- coef_estimates[11];gamma0 <- coef_estimates[12];gamma1 <- coef_estimates[13]</pre>
    #### Time Period 0
        RO \leftarrow rep(0, MC)
        logCD40 <- rnorm(MC, mu0, sigma0)</pre>
        A0 <- rep(treat_sequence[1], MC)
        cumAO <- AO
    ### Time Period 1
        R1 <- rbinom(MC, 1, expit(gamma0 + gamma1*cumA0)
                      ) * (cumA0 > 0 & R0 == 0) + 0*(cumA0 == 0) + 1*(R0 == 1)
        logCD41 <-
          beta0 + beta1*logCD40 + beta2*A0 + beta3*R1 +
            beta4*logCD40*A0 + beta5*logCD40*R1 + beta6*A0*R1 + beta7*logCD40*A0*R1 +
            rnorm(MC, 0, sigma)
        A1 <- rep(treat_sequence[2], MC)
        cumA1 <- A0 + A1
    ### Time Period 2
```

```
R2 <- rbinom(MC, 1, expit(gamma0 + gamma1*cumA1)
                                                            *(cumA1 > 0 \& R1 == 0) + 0*(cumA1 == 0) + 1*(R1 == 1)
                       logCD42 <-
                            beta0 + beta1*logCD41 + beta2*A1 + beta3*R2 +
                                  beta4*logCD41*A1 + beta5*logCD41*R2 + beta6*A1*R2 + beta7*logCD41*A1*R2 +
                                  rnorm(MC, 0, sigma)
                       A2 <- rep(treat_sequence[3], MC)
                       cumA2 \leftarrow A0 + A1 + A2
           ### Time Period 3
                       R3 <- rbinom(MC, 1, expit(gamma0 + gamma1*cumA2)
                                                            *(cumA2 > 0 \& R2 == 0) + 0*(cumA2 == 0) + 1*(R2 == 1)
                       logCD43 <-
                            beta0 + beta1*logCD42 + beta2*A2 + beta3*R3 +
                                  beta4*logCD42*A2 + beta5*logCD42*R3 + beta6*A2*R3 + beta7*logCD42*A2*R3 +
                                  rnorm(MC, 0, sigma)
                       A3 <- rep(treat_sequence[4], MC)
                       cumA3 < - A0 + A1 + A2 + A3
           ### Time Period 4
                       R4 <- rbinom(MC, 1, expit(gamma0 + gamma1*cumA3)
                                                            *(cumA3 > 0 \& R3 == 0) + 0*(cumA3 == 0) + 1*(R3 == 1)
                       logCD44 <-
                            beta0 + beta1*logCD43 + beta2*A3 + beta3*R4 +
                                  beta4*logCD43*A3 + beta5*logCD43*R4 + beta6*A3*R4 + beta7*logCD43*A3*R4 + beta7*logCD43*A3*R4 + beta8*A3*R4 + be
                                  rnorm(MC, 0, sigma)
           mean_response <- c(mean(logCD44), mean(exp(logCD44)))</pre>
           names(mean_response) <- c("mean_logCD4", "mean_CD4")</pre>
           return(mean_response)
}
trt_sequences <- list(</pre>
           c(0, 0, 0, 0),
           c(0, 0, 0, 1),
           c(0, 0, 1, 1),
```

```
c(0, 1, 1, 1),
    c(1, 1, 1, 1),
    c(1, 1, 1, 0),
    c(1, 1, 0, 0),
    c(1, 0, 0, 0)
)
set.seed(8172013)
mean_trt_response <-</pre>
  sapply(trt_sequences,
         mean_response,
         coef_estimates = as.vector(coef_estimates[, 1]), MC = 100000)
mean_trt_response <- t(mean_trt_response)</pre>
rownames(mean_trt_response) <-</pre>
  c("(0, 0, 0, 0)",
      "(0, 0, 0, 1)",
      "(0, 0, 1, 1)",
      "(0, 1, 1, 1)",
      "(1, 1, 1, 1)",
      "(1, 1, 1, 0)",
      "(1, 1, 0, 0)",
      "(1, 0, 0, 0)")
#print(mean_trt_response, digits = 3)
```

## **Bootstrap to Estimate SE**

```
dataHW3_orig <- dataHW3</pre>
n <- nrow(dataHW3_orig)</pre>
boot <- 50
mean_trt_response_boot <- array(0, dim = c(8, 2, boot))
for(j in 1:boot) {
    dataHW3 <- dataHW3_orig[sample(1:n, n, replace = TRUE), ]</pre>
    # get estimates of mu0 and sigma0
    mu0 <- mean(dataHW3$logCD40)</pre>
    sigma0 <- sd(dataHW3$logCD40)</pre>
    dataHW3$ID <- 1:n</pre>
    dataHW3 long <-
      melt(setDT(dataHW3),
            id = 1L,
          measure = patterns("^logCD4", "^CD4", "^R", "^A"),
          value.name = c("logCD4", "CD4", "R", "A")
    dataHW3_long <- data.frame(dataHW3_long)</pre>
    dataHW3_long <- dataHW3_long[order(dataHW3_long$ID), ]</pre>
    dataHW3_long <-
        dataHW3_long %>%
        group_by(ID) %>%
        mutate(
          lag_logCD4 = lag(logCD4, n = 1, default = NA),
            lag_A = lag(A, n = 1, default = NA),
            cum_A = cumsum(A),
            lag_cum_A = lag(cum_A, n = 1, default = NA),
            lag_R = lag(R, n = 1, default = NA)
```

```
)
    dataHW3_long <-
      mutate(
        dataHW3_long,
      time = as.numeric(variable) - 1,
      logCD4_diff = logCD4 - lag_logCD4)
    dataHW3_long_t0 <- filter(dataHW3_long, time > 0)
    m1 \leftarrow lm(logCD4 \sim lag_logCD4 + lag_A + R +
        lag_logCD4*lag_A + lag_logCD4*R +
        lag_A*R + lag_logCD4*lag_A*R, data = dataHW3_long_t0)
    dataHW3_long_resist_model <- filter(dataHW3_long_t0, lag_R == 0 & lag_cum_A > 0)
    m2 <- glm(R ~ lag_cum_A, family = "binomial",</pre>
        data = dataHW3_long_resist_model)
    coef_estimates <- c(mu0, sigma0, coef(m1), summary(m1)$sigma, coef(m2))</pre>
    mean_trt_response <-</pre>
      sapply(trt_sequences,
             mean_response,
           coef_estimates = coef_estimates,
             MC = 100000)
    mean_trt_response <- t(mean_trt_response)</pre>
    mean_trt_response_boot[, , j] <- mean_trt_response</pre>
}
se <- apply(mean_trt_response_boot, c(1, 2), sd)</pre>
rownames(se) <- c("(0, 0, 0, 0)",
    "(0, 0, 0, 1)",
    "(0, 0, 1, 1)",
    "(0, 1, 1, 1)",
    "(1, 1, 1, 1)",
    "(1, 1, 1, 0)",
    "(1, 1, 0, 0)",
    "(1, 0, 0, 0)")
```

```
colnames(se) <- c("SE_logCD4", "SE_CD4")
print(se, digits = 2)</pre>
```