

ali_sim_params

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```
## Read in summary data (one row per patient)
summ_data = read.csv("~/Documents/Allostatic_load_audits/summary_data.csv") |>
  dplyr::mutate(ANY_ENCOUNTERS = as.numeric(NUM_ENCOUNTERS > 0)) ### Create indicator of >= 1 healthcare visit

## Subset to only the variables needed for the analysis
summ_data = summ_data |>
  select(PAT_MRN_ID, ANY_ENCOUNTERS, AGE_AT_ENCOUNTER, SEX,
         CREAT_C, ALB, BMI, BP_SYSTOLIC, BP_DIASTOLIC, A1C, CHOL, TRIG, CRP, HCST)

## Create a long version with one row per variable per patient
summ_data_long = summ_data |>
  gather(key = "COMP",
         value = "VAL",
         -c(1:4))

## Merge in the stratification values
strat_vals = read.csv("~/Documents/Allostatic_load_audits/Audit_Protocol/strat_vals.csv")
summ_data_long = summ_data_long |>
  left_join(strat_vals) |>
  mutate(POINT = case_when(
    INEQ == ">" ~ as.numeric(VAL > STRAT),
    INEQ == ">=" ~ as.numeric(VAL >= STRAT),
    INEQ == "<" ~ as.numeric(VAL < STRAT)
  ))
```

Joining with 'by = join_by(SEX, COMP)'

```
## Aggregate the stratified components by patient
summ_data = summ_data_long |>
  group_by(PAT_MRN_ID, ANY_ENCOUNTERS, AGE_AT_ENCOUNTER, SEX) |>
  summarize(ALI_STAR_NUM = sum(POINT, na.rm = TRUE), ### Number of components equal to 1
            ALI_STAR_DENOM = sum(!is.na(POINT)) ### Number of non-missing components
            ) |>
  mutate(ALI_STAR = ALI_STAR_NUM / ALI_STAR_DENOM,
         ALI_STAR_RESCALED = ALI_STAR / 0.1 ### Create ALI_STAR_RESCALED = ALI_STAR / 0.1
         )
```

'summarise()' has grouped output by 'PAT_MRN_ID', 'ANY_ENCOUNTERS',
'AGE_AT_ENCOUNTER'. You can override using the '.groups' argument.

Distribution of age at encounter

```
## Model age at first encounter (in 1-year increments) with a Poisson distribution
glm(formula = AGE_AT_ENCOUNTER ~ 1,
    family = poisson,
    data = summ_data) |>
coefficients() |>
exp()
```

```
## (Intercept)
##          45.662
```

```
### Rescale age at first encounter to be in 5-year increments
```

```
summ_data = summ_data |>
  ### Create AGE_AT_ENCOUNTER5 = AGE_AT_ENCOUNTER / 5
  dplyr::mutate(AGE_AT_ENCOUNTER5 = AGE_AT_ENCOUNTER / 5)
```

```
## Model age at first encounter (in 5-year increments) with a Poisson distribution
```

```
glm(formula = AGE_AT_ENCOUNTER5 ~ 1,
    family = poisson,
    data = summ_data) |>
coefficients() |>
exp()
```

```
## (Intercept)
##          9.1324
```

```
### Rescale age at first encounter to be in 10-year increments
```

```
summ_data = summ_data |>
  ### Create AGE_AT_ENCOUNTER10 = AGE_AT_ENCOUNTER / 10
  dplyr::mutate(AGE_AT_ENCOUNTER10 = AGE_AT_ENCOUNTER / 10)
```

```
## Model age at first encounter (in 10-year increments) with a Poisson distribution
```

```
glm(formula = AGE_AT_ENCOUNTER10 ~ 1,
    family = poisson,
    data = summ_data) |>
coefficients() |>
exp()
```

```
## (Intercept)
##          4.5662
```

Naive model of ALI* and healthcare utilization

```
## Naive model parameters (age in 10-year increments)
```

```
naive_mod = glm(formula = ANY_ENCOUNTERS ~ ALI_STAR + AGE_AT_ENCOUNTER10,
    family = "binomial",
    data = summ_data)
summary(naive_mod)
```

```
##
## Call:
## glm(formula = ANY_ENCOUNTERS ~ ALI_STAR + AGE_AT_ENCOUNTER10,
##      family = "binomial", data = summ_data)
##
## Coefficients:
##              Estimate Std. Error z value Pr(>|z|)
## (Intercept)   -1.56762    0.25149  -6.233 4.57e-10 ***
## ALI_STAR        0.94523    0.34183   2.765  0.00569 **
## AGE_AT_ENCOUNTER10 0.10266    0.05222   1.966  0.04928 *
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##    Null deviance: 1250.7  on 999  degrees of freedom
## Residual deviance: 1234.7  on 997  degrees of freedom
## AIC: 1240.7
##
## Number of Fisher Scoring iterations: 4
```

```
## Naive model parameters (age in 10-year increments)
naive_mod = glm(formula = ANY_ENCOUNTERS ~ ALI_STAR_RESCALED + AGE_AT_ENCOUNTER10,
                family = "binomial",
                data = summ_data)
summary(naive_mod)
```

```
##
## Call:
## glm(formula = ANY_ENCOUNTERS ~ ALI_STAR_RESCALED + AGE_AT_ENCOUNTER10,
##      family = "binomial", data = summ_data)
##
## Coefficients:
##              Estimate Std. Error z value Pr(>|z|)
## (Intercept)   -1.56762    0.25149  -6.233 4.57e-10 ***
## ALI_STAR_RESCALED 0.09452    0.03418   2.765  0.00569 **
## AGE_AT_ENCOUNTER10 0.10266    0.05222   1.966  0.04928 *
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##    Null deviance: 1250.7  on 999  degrees of freedom
## Residual deviance: 1234.7  on 997  degrees of freedom
## AIC: 1240.7
##
## Number of Fisher Scoring iterations: 4
```

Naive proportions “yes” and missing data per ALI component

```
## probability of stressor = YES
summ_data_long |>
  group_by(COMP) |>
  summarize(PROP_YES = mean(POINT, na.rm = TRUE),
            PROP_MISS = mean(is.na(POINT)))
```

```
## # A tibble: 10 x 3
##   COMP          PROP_YES PROP_MISS
##   <chr>          <dbl>     <dbl>
## 1 A1C            0.249      0.494
## 2 ALB            0.987      0.153
## 3 BMI            0.455      0.002
## 4 BP_DIASTOLIC   0.058      0
## 5 BP_SYSTOLIC    0.145      0
## 6 CHOL           0.314      0.213
## 7 CREAT_C        0.25       0.996
## 8 CRP            0.311      0.955
## 9 HCST           0         0.983
## 10 TRIG          0.332      0.213
```

SMLE model of ALI and healthcare utilization (pilot + wave 1)

```
## Read in long audit data (one row per patient per audited variable)
audit_dat = read.csv("~/Documents/ali-data/summary_data_audit1.csv")

## Create a long version with one row per variable per patient
audit_dat_long = audit_dat |>
  gather(key = "COMP",
         value = "VAL",
         -c(1))

## Merge in the stratification values
strat_vals = strat_vals |>
  filter(SEX == "Female") |>
  select(-SEX)
audit_dat_long = audit_dat_long |>
  left_join(strat_vals) |>
  mutate(POINT = case_when(
    INEQ == ">" ~ as.numeric(VAL > STRAT),
    INEQ == ">=" ~ as.numeric(VAL >= STRAT),
    INEQ == "<" ~ as.numeric(VAL < STRAT)
  ))

## Joining with 'by = join_by(COMP)'

## Aggregate the stratified components by patient
audit_dat = audit_dat_long |>
  group_by(PAT_MR_N_ID) |>
  summarize(ALI_NUM = sum(POINT, na.rm = TRUE), ### Number of components equal to 1
            ALI_DENOM = sum(!is.na(POINT)) ### Number of non-missing components
            ) |>
  mutate(ALI = ALI_NUM / ALI_DENOM,
         ALI_RESCALED = ALI / 0.1 ### Create ALI_RESCALED = ALI / 0.1
         )

## Merge with summ_data
summ_data = summ_data |>
  left_join(audit_dat)

## Joining with 'by = join_by(PAT_MR_N_ID)'

# Estimate parameters using Phase IIa audits + the rest of Phase I -----
## Setup B-splines
B = splines::bs(x = summ_data$ALI_STAR,
               df = 8,
               Boundary.knots = range(summ_data$ALI_STAR),
               intercept = TRUE,
               degree = 3)
colnames(B) = paste0("bs", seq(1, 8))
summ_data = summ_data |>
  bind_cols(B) |>
  data.frame()
```

```

### Fit SMLE -----
library(sleev)
suppressMessages(fit <- logistic2ph(
  Y_unval = NULL,
  Y = "ANY_ENCOUNTERS",
  X_unval = "ALI_STAR",
  X = "ALI",
  Z = "AGE_AT_ENCOUNTER10",
  Bspline = colnames(B),
  data = summ_data,
  hn_scale = 1,
  noSE = FALSE,
  TOL = 1e-04,
  MAX_ITER = 1000
))
fit$coefficients

```

##		Estimate	SE	Statistic	p-value
##	Intercept	-1.9309188	0.32092231	-6.016779	1.779213e-09
##	ALI	1.8779397	0.27712564	6.776492	1.231282e-11
##	AGE_AT_ENCOUNTER10	0.0991235	0.04488015	2.208627	2.720060e-02

```

# Estimate parameters using Phase IIa audits + the rest of Phase I -----
## Setup B-splines
summ_data = summ_data[, -grep(pattern = "bs", x = colnames(summ_data))]
B = splines::bs(x = summ_data$ALI_STAR_RESCALED,
  df = 8,
  Boundary.knots = range(summ_data$ALI_STAR_RESCALED),
  intercept = TRUE,
  degree = 3)
colnames(B) = paste0("bs", seq(1, 8))
summ_data = summ_data |>
  bind_cols(B)

summ_data = data.frame(summ_data)

```

```

### Fit SMLE -----
suppressMessages(fit <- logistic2ph(
  Y_unval = NULL,
  Y = "ANY_ENCOUNTERS",
  X_unval = "ALI_STAR_RESCALED",
  X = "ALI_RESCALED",
  Z = "AGE_AT_ENCOUNTER10",
  Bspline = colnames(B),
  data = summ_data,
  hn_scale = 1 / 2, ##### since the parameter estimates are smaller, need smaller perturbation for SEs
  noSE = FALSE,
  TOL = 1e-04,
  MAX_ITER = 1000
))
fit$coefficients

```

##		Estimate	SE	Statistic	p-value
----	--	----------	----	-----------	---------

## Intercept	-1.9309188	0.02259760	-85.447951	0.000000000
## ALI_RESCALED	0.1877940	0.05999031	3.130405	0.001745653
## AGE_AT_ENCOUNTER10	0.0991235	0.04150285	2.388354	0.016924040