

Parameter Table

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1 Purpose

This script picks up after model.Rnw to process bootstrap results and make a parameter table. It assumes the current working directory is the script directory containing this file.

1.1 Package

Listing 1:

```
> library(metrumrg)
```

2 inputs

'wikitab' gives us a quick synthesis of 'rlog' and the 'lookup' of wiki notation in 1005.ctl. We do some science on the result first, and then some aesthetics for printing in a \LaTeX table. Table 1.

Listing 2:

```
> tab <- wikitab(1005, '../nonmem')
> tab$estimate <- signif(as.numeric(tab$estimate), 3)
> tab$tool <- NULL
> tab$run <- NULL
> tab$se <- NULL
> tab
```

| | parameter | description | | model estimate |
|----|--|---|--|----------------|
| 1 | THETA1 | apparent oral clearance | | |
| 2 | THETA2 | central volume of distribution | | |
| 3 | THETA3 | absorption rate constant | | |
| 4 | THETA4 | intercompartmental clearance | | |
| 5 | THETA5 | peripheral volume of distribution | | |
| 6 | THETA6 | male effect on clearance | | |
| 7 | THETA7 | weight effect on clearance | | |
| 8 | OMEGA1.1 | interindividual variability of clearance | | |
| 9 | OMEGA2.1 | interindividual clearance-volume covariance | | |
| 10 | OMEGA2.2 | interindividual variability of central volume | | |
| 11 | OMEGA3.1 | interindividual clearance-Ka covariance | | |
| 12 | OMEGA3.2 | interindividual volume-Ka covariance | | |
| 13 | OMEGA3.3 | interindividual variability of Ka | | |
| 14 | SIGMA1.1 | proportional error | | |
| 15 | SIGMA2.2 | additive error | | |
| 1 | CL/F (L/h) ~ theta_1 * theta_6 ^MALE * (WT/70)^theta_7 * e^eta_1 | | | 16 |
| 2 | V_c /F (L) ~ theta_2 * (WT/70)^1 * e^eta_2 | | | 14 |
| 3 | K_a (h^-1) ~ theta_3 * e^eta_3 | | | 6 |
| 4 | Q/F (L/h) ~ theta_4 | | | 15 |
| 5 | V_p /F (L) ~ theta_5 | | | 12 |
| 6 | MALE_CL/F ~ theta_6 | | | 11 |

```

7          WT_CL/F ~ theta_7          13
8          IIV_CL/F ~ Omega_1.1       10
9          cov_CL,V ~ Omega_2.1       8
10         IIV_V_c /F ~ Omega_2.2      7
11         cov_CL,Ka ~ Omega_3.1       2
12         cov_V,Ka ~ Omega_3.2       3
13         IIV_K_a ~ Omega_3.3       4
14         err_prop ~ Sigma_1.1       5
15         err_add ~ Sigma_2.2       9

prse
1  15
2  14
3  13
4   3
5   5
6   2
7   8
8   6
9   7
10  9
11  4
12 12
13 11
14  1
15 10

```

Now we can extract some information from the model statements.

Listing 3:

```

> tab$units <- justUnits(tab$model)
> tab$model <- noUnits(tab$model)
> tab$name <- with(tab, wiki2label(model))
> tab[c('model','units','name')]

          model units
1 CL/F ~ theta_1 * theta_6 ^MALE * (WT/70)^theta_7 * e^eta_1 L/h
2          V_c /F ~ theta_2 * (WT/70)^1 * e^eta_2      L
3          K_a ~ theta_3 * e^eta_3 h^-1
4          Q/F ~ theta_4      L/h
5          V_p /F ~ theta_5      L
6          MALE_CL/F ~ theta_6
7          WT_CL/F ~ theta_7
8          IIV_CL/F ~ Omega_1.1
9          cov_CL,V ~ Omega_2.1
10         IIV_V_c /F ~ Omega_2.2
11         cov_CL,Ka ~ Omega_3.1
12         cov_V,Ka ~ Omega_3.2
13         IIV_K_a ~ Omega_3.3
14         err_prop ~ Sigma_1.1
15         err_add ~ Sigma_2.2

```

```

      name
1      CL/F
2      V_c/F
3      K_a
4      Q/F
5      V_p/F
6 MALE_CL/F
7      WT_CL/F
8      IIV_CL/F
9      cov_CL,V
10 IIV_V_c/F
11 cov_CL,Ka
12 cov_V,Ka
13      IIV_K_a
14 err_prop
15      err_add

```

3 variance

The estimates for the matrix diagonals are variances, and their square roots have special meaning. In model 1005, interindividual variability was modelled exponentially, in which case square root of variance gives an approximate CV; alternatively, and exact CV can be calculated. For proportional error terms like ERR1, square root gives an exact CV. For additive error terms like ERR2, square root gives standard deviation.

We can use functions of 'parameter' to sort out the various error components, as they are used in this model.

3.1 exponential

Listing 4:

```

> expo <- is.iiv(tab$parameter) & is.diagonal(tab$parameter)
> tab$parameter[expo]

```

```

[1] "OMEGA1.1" "OMEGA2.2" "OMEGA3.3"

```

Listing 5:

```

> tab$cv[expo] <- cvLognormal(tab$estimate[expo])
> tab[,c('parameter','name','estimate','cv')]

```

| | parameter | name | estimate | cv |
|---|-----------|-------|----------|----|
| 1 | THETA1 | CL/F | 16 | NA |
| 2 | THETA2 | V_c/F | 14 | NA |
| 3 | THETA3 | K_a | 6 | NA |

| | | | | |
|----|----------|-----------|----|------------|
| 4 | THETA4 | Q/F | 15 | NA |
| 5 | THETA5 | V_p/F | 12 | NA |
| 6 | THETA6 | MALE_CL/F | 11 | NA |
| 7 | THETA7 | WT_CL/F | 13 | NA |
| 8 | OMEGA1.1 | IIV_CL/F | 10 | 148.409790 |
| 9 | OMEGA2.1 | cov_CL,V | 8 | NA |
| 10 | OMEGA2.2 | IIV_V_c/F | 7 | 33.100350 |
| 11 | OMEGA3.1 | cov_CL,Ka | 2 | NA |
| 12 | OMEGA3.2 | cov_V,Ka | 3 | NA |
| 13 | OMEGA3.3 | IIV_K_a | 4 | 7.321076 |
| 14 | SIGMA1.1 | err_prop | 5 | NA |
| 15 | SIGMA2.2 | err_add | 9 | NA |

3.2 proportional

Listing 6:

```
> writeLines(read.nmctl('../nonmem/ctl/1005.ctl')$err)

Y=F*(1+ERR(1)) + ERR(2)
IPRE=F
;<doc>
```

Listing 7:

```
> prop <- is.random(tab$parameter) & tab$name %contains% 'prop'
> tab$parameter[prop]

[1] "SIGMA1.1"
```

Listing 8:

```
> tab$cv[prop] <- sqrt(tab$estimate[prop])
> tab[,c('parameter','name','estimate','cv')]

  parameter      name estimate      cv
1  THETA1      CL/F        16      NA
2  THETA2     V_c/F        14      NA
3  THETA3      K_a         6      NA
4  THETA4      Q/F        15      NA
5  THETA5     V_p/F        12      NA
6  THETA6 MALE_CL/F        11      NA
7  THETA7     WT_CL/F        13      NA
8 OMEGA1.1 IIV_CL/F       10 148.409790
9 OMEGA2.1 cov_CL,V         8      NA
10 OMEGA2.2 IIV_V_c/F        7  33.100350
11 OMEGA3.1 cov_CL,Ka         2      NA
12 OMEGA3.2 cov_V,Ka         3      NA
13 OMEGA3.3 IIV_K_a         4   7.321076
14 SIGMA1.1 err_prop         5   2.236068
15 SIGMA2.2 err_add         9      NA
```

3.3 additive

Listing 9:

```
> add <- is.residual(tab$parameter) & tab$name %contains% 'add'
> tab$parameter[add]
```

```
[1] "SIGMA2.2"
```

Listing 10:

```
> tab$sd[add] <- sqrt(tab$estimate[add])
> tab[,c('parameter', 'name', 'estimate', 'cv', 'sd')]
```

| | parameter | name | estimate | cv | sd |
|----|-----------|-----------|----------|------------|----|
| 1 | THETA1 | CL/F | 16 | NA | NA |
| 2 | THETA2 | V_c/F | 14 | NA | NA |
| 3 | THETA3 | K_a | 6 | NA | NA |
| 4 | THETA4 | Q/F | 15 | NA | NA |
| 5 | THETA5 | V_p/F | 12 | NA | NA |
| 6 | THETA6 | MALE_CL/F | 11 | NA | NA |
| 7 | THETA7 | WT_CL/F | 13 | NA | NA |
| 8 | OMEGA1.1 | IIV_CL/F | 10 | 148.409790 | NA |
| 9 | OMEGA2.1 | cov_CL,V | 8 | NA | NA |
| 10 | OMEGA2.2 | IIV_V_c/F | 7 | 33.100350 | NA |
| 11 | OMEGA3.1 | cov_CL,Ka | 2 | NA | NA |
| 12 | OMEGA3.2 | cov_V,Ka | 3 | NA | NA |
| 13 | OMEGA3.3 | IIV_K_a | 4 | 7.321076 | NA |
| 14 | SIGMA1.1 | err_prop | 5 | 2.236068 | NA |
| 15 | SIGMA2.2 | err_add | 9 | NA | 3 |

4 covariance

The estimates of matrix off-diagonals are covariances, and are more useful if transformed to correlations. We could extract the matrices manually, or use package shortcuts.

Listing 11:

```
> cor <- omegacor(run=1005,project='../nonmem')
> cor
```

| | [,1] | [,2] | [,3] |
|------|------------|------------|------------|
| [1,] | 1.0000000 | 0.8494277 | -0.1162464 |
| [2,] | 0.8494277 | 1.0000000 | -0.5605290 |
| [3,] | -0.1162464 | -0.5605290 | 1.0000000 |

Listing 12:

```
> half(cor)
```

```

      1.1      2.1      2.2      3.1      3.2      3.3
1.0000000  0.8494277  1.0000000 -0.1162464 -0.5605290  1.0000000

```

Listing 13:

```
> offdiag(half(cor))
```

```

      2.1      3.1      3.2
0.8494277 -0.1162464 -0.5605290

```

Listing 14:

```
> off <- is.iiv(tab$parameter) & is.offdiagonal(tab$parameter)
> tab$parameter[off]
```

```
[1] "OMEGA2.1" "OMEGA3.1" "OMEGA3.2"
```

Listing 15:

```
> tab$cor[off] <- offdiag(half(cor))
> tab[,c('parameter','name','estimate','cv','sd','cor')]
```

| | parameter | name | estimate | cv | sd | cor |
|----|-----------|-----------------------|----------|------------|----|------------|
| 1 | THETA1 | CL/F | 16 | NA | NA | NA |
| 2 | THETA2 | V _c /F | 14 | NA | NA | NA |
| 3 | THETA3 | K _a | 6 | NA | NA | NA |
| 4 | THETA4 | Q/F | 15 | NA | NA | NA |
| 5 | THETA5 | V _p /F | 12 | NA | NA | NA |
| 6 | THETA6 | MALE_CL/F | 11 | NA | NA | NA |
| 7 | THETA7 | WT_CL/F | 13 | NA | NA | NA |
| 8 | OMEGA1.1 | IIV_CL/F | 10 | 148.409790 | NA | NA |
| 9 | OMEGA2.1 | cov _{CL,V} | 8 | NA | NA | 0.8494277 |
| 10 | OMEGA2.2 | IIV_V _c /F | 7 | 33.100350 | NA | NA |
| 11 | OMEGA3.1 | cov _{CL,Ka} | 2 | NA | NA | -0.1162464 |
| 12 | OMEGA3.2 | cov _{V,Ka} | 3 | NA | NA | -0.5605290 |
| 13 | OMEGA3.3 | IIV_K _a | 4 | 7.321076 | NA | NA |
| 14 | SIGMA1.1 | err _{prop} | 5 | 2.236068 | NA | NA |
| 15 | SIGMA2.2 | err _{add} | 9 | NA | 3 | NA |

5 confidence interval

We wish to include 95 percentiles in our table as confidence intervals.

Listing 16:

```
> boot <- read.csv('../nonmem/1005bootlog.csv',as.is=TRUE)
> head(boot)
```

```

X tool run parameter      moment      value
1 1  nm7    1      ofv minimum 2641.7825682304
2 2  nm7    1      THETA1 estimate    9.23638
3 3  nm7    1      THETA1      prse      <NA>
4 4  nm7    1      THETA1      se        <NA>
5 5  nm7    1      THETA2 estimate    23.3418
6 6  nm7    1      THETA2      prse      <NA>

```

Listing 17:

```

> boot <- boot[boot$moment=='estimate',]
> boot <- data.frame(cast(boot,... ~ moment))
> head(boot)

```

```

X tool run parameter estimate
1 2  nm7    1      THETA1    9.23638
2 5  nm7    1      THETA2   23.3418
3 8  nm7    1      THETA3  0.0677011
4 11 nm7    1      THETA4    3.82773
5 14 nm7    1      THETA5   114.89
6 17 nm7    1      THETA6    0.981208

```

Listing 18:

```

> boot <- boot[,c('run','parameter','estimate')]
> sapply(boot,class)

```

```

      run      parameter      estimate
"integer" "character"    "factor"

```

Listing 19:

```

> boot$estimate <- as.numeric(as.character(boot$estimate))
> unique(boot$parameter)

```

```

[1] "THETA1"  "THETA2"  "THETA3"  "THETA4"  "THETA5"  "THETA6"
[7] "THETA7"  "OMEGA1.1" "OMEGA2.1" "OMEGA2.2" "OMEGA3.1" "OMEGA3.2"
[13] "OMEGA3.3" "SIGMA1.1" "SIGMA2.1" "SIGMA2.2"

```

Listing 20:

```

> quan <- function(x,probs)as.character(signif(quantile(x,probs=probs,na.rm=TRUE)
,3))
> boot$lo <- with(boot, reapply(estimate,parameter,quan,probs=.05))
> boot$hi <- with(boot, reapply(estimate,parameter,quan,probs=.95))
> head(boot)

```

```

run parameter      estimate      lo      hi
1 1  THETA1    9.2363800    6.67    11.1
2 1  THETA2   23.3418000     19    27.6
3 1  THETA3    0.0677011  0.0636  0.0814
4 1  THETA4    3.8277300    2.77    4.97
5 1  THETA5  114.8900000    87.9    315
6 1  THETA6    0.9812080    0.845    1.27

```

Listing 21:

```
> boot <- unique(boot[,c('parameter','lo','hi')])
> boot
```

| | parameter | lo | hi |
|----|-----------|---------|----------|
| 1 | THETA1 | 6.67 | 11.1 |
| 2 | THETA2 | 19 | 27.6 |
| 3 | THETA3 | 0.0636 | 0.0814 |
| 4 | THETA4 | 2.77 | 4.97 |
| 5 | THETA5 | 87.9 | 315 |
| 6 | THETA6 | 0.845 | 1.27 |
| 7 | THETA7 | 0.685 | 1.96 |
| 8 | OMEGA1.1 | 0.127 | 0.325 |
| 9 | OMEGA2.1 | 0.0657 | 0.183 |
| 10 | OMEGA2.2 | 0.0457 | 0.159 |
| 11 | OMEGA3.1 | -0.0438 | 0.0248 |
| 12 | OMEGA3.2 | -0.0536 | -0.00759 |
| 13 | OMEGA3.3 | 0.0237 | 0.0789 |
| 14 | SIGMA1.1 | 0.0404 | 0.0594 |
| 15 | SIGMA2.1 | 0 | 0 |
| 16 | SIGMA2.2 | 0.073 | 0.323 |

Listing 22:

```
> boot$ci <- with(boot, parens(glue(lo,',',hi)))
> boot
```

| | parameter | lo | hi | ci |
|----|-----------|---------|----------|--------------------|
| 1 | THETA1 | 6.67 | 11.1 | (6.67,11.1) |
| 2 | THETA2 | 19 | 27.6 | (19,27.6) |
| 3 | THETA3 | 0.0636 | 0.0814 | (0.0636,0.0814) |
| 4 | THETA4 | 2.77 | 4.97 | (2.77,4.97) |
| 5 | THETA5 | 87.9 | 315 | (87.9,315) |
| 6 | THETA6 | 0.845 | 1.27 | (0.845,1.27) |
| 7 | THETA7 | 0.685 | 1.96 | (0.685,1.96) |
| 8 | OMEGA1.1 | 0.127 | 0.325 | (0.127,0.325) |
| 9 | OMEGA2.1 | 0.0657 | 0.183 | (0.0657,0.183) |
| 10 | OMEGA2.2 | 0.0457 | 0.159 | (0.0457,0.159) |
| 11 | OMEGA3.1 | -0.0438 | 0.0248 | (-0.0438,0.0248) |
| 12 | OMEGA3.2 | -0.0536 | -0.00759 | (-0.0536,-0.00759) |
| 13 | OMEGA3.3 | 0.0237 | 0.0789 | (0.0237,0.0789) |
| 14 | SIGMA1.1 | 0.0404 | 0.0594 | (0.0404,0.0594) |
| 15 | SIGMA2.1 | 0 | 0 | (0,0) |
| 16 | SIGMA2.2 | 0.073 | 0.323 | (0.073,0.323) |

Listing 23:

```
> tab <- stableMerge(tab,boot[,c('parameter','ci')])
> tab
```

| parameter | description | | | | | |
|-----------|-------------|---|------------|-----------|--------------------|--|
| 1 | THETA1 | apparent oral clearance | | | | |
| 2 | THETA2 | central volume of distribution | | | | |
| 3 | THETA3 | absorption rate constant | | | | |
| 4 | THETA4 | intercompartmental clearance | | | | |
| 5 | THETA5 | peripheral volume of distribution | | | | |
| 6 | THETA6 | male effect on clearance | | | | |
| 7 | THETA7 | weight effect on clearance | | | | |
| 8 | OMEGA1.1 | interindividual variability of clearance | | | | |
| 9 | OMEGA2.1 | interindividual clearance-volume covariance | | | | |
| 10 | OMEGA2.2 | interindividual variability of central volume | | | | |
| 11 | OMEGA3.1 | interindividual clearance-Ka covariance | | | | |
| 12 | OMEGA3.2 | interindividual volume-Ka covariance | | | | |
| 13 | OMEGA3.3 | interindividual variability of Ka | | | | |
| 14 | SIGMA1.1 | proportional error | | | | |
| 15 | SIGMA2.2 | additive error | | | | |
| | | model | estimate | prse | | |
| 1 | CL/F | ~ theta_1 * theta_6 ^MALE * (WT/70)^theta_7 * e^eta_1 | 16 | 15 | | |
| 2 | V_c /F | ~ theta_2 * (WT/70)^1 * e^eta_2 | 14 | 14 | | |
| 3 | K_a | ~ theta_3 * e^eta_3 | 6 | 13 | | |
| 4 | Q/F | ~ theta_4 | 15 | 3 | | |
| 5 | V_p /F | ~ theta_5 | 12 | 5 | | |
| 6 | MALE_CL/F | ~ theta_6 | 11 | 2 | | |
| 7 | WT_CL/F | ~ theta_7 | 13 | 8 | | |
| 8 | IIV_CL/F | ~ Omega_1.1 | 10 | 6 | | |
| 9 | cov_CL,V | ~ Omega_2.1 | 8 | 7 | | |
| 10 | IIV_V_c /F | ~ Omega_2.2 | 7 | 9 | | |
| 11 | cov_CL,Ka | ~ Omega_3.1 | 2 | 4 | | |
| 12 | cov_V,Ka | ~ Omega_3.2 | 3 | 12 | | |
| 13 | IIV_K_a | ~ Omega_3.3 | 4 | 11 | | |
| 14 | err_prop | ~ Sigma_1.1 | 5 | 1 | | |
| 15 | err_add | ~ Sigma_2.2 | 9 | 10 | | |
| units | name | cv | sd | cor | ci | |
| 1 | L/h | CL/F | NA | NA | (6.67,11.1) | |
| 2 | L | V_c/F | NA | NA | (19,27.6) | |
| 3 | h^-1 | K_a | NA | NA | (0.0636,0.0814) | |
| 4 | L/h | Q/F | NA | NA | (2.77,4.97) | |
| 5 | L | V_p/F | NA | NA | (87.9,315) | |
| 6 | | MALE_CL/F | NA | NA | (0.845,1.27) | |
| 7 | | WT_CL/F | NA | NA | (0.685,1.96) | |
| 8 | | IIV_CL/F | 148.409790 | NA | (0.127,0.325) | |
| 9 | | cov_CL,V | NA | 0.8494277 | (0.0657,0.183) | |
| 10 | | IIV_V_c/F | 33.100350 | NA | (0.0457,0.159) | |
| 11 | | cov_CL,Ka | NA | NA | (-0.0438,0.0248) | |
| 12 | | cov_V,Ka | NA | NA | (-0.0536,-0.00759) | |
| 13 | | IIV_K_a | 7.321076 | NA | (0.0237,0.0789) | |
| 14 | | err_prop | 2.236068 | NA | (0.0404,0.0594) | |
| 15 | | err_add | NA | 3 | (0.073,0.323) | |

6 aesthetics

Here we format the table for printing.

Listing 24:

```
> tab$name <- NULL
> tab$parameter <- NULL
> tab$model <- wiki2latex(tab$model)
> tab$estimate <- as.character(tab$estimate)
> tab$estimate <- paste(tab$estimate, '$', tab$units, '$')
> tab$units <- NULL
```

Note that no parameter defines more than one of CV, SD, and COR. We could collapse these into a single column, and add a descriptive flag.

Listing 25:

```
> m <- as.matrix(tab[, c('cv', 'sd', 'cor')])
> tab$variability <- suppressWarnings(apply(m, 1, max, na.rm=TRUE))
> tab$variability[is.infinite(tab$variability)] <- NA
> i <- is.defined(m)
> i[!i] <- NA
> tab$statistic <- apply(i, 1, function(x) {
+   p <- colnames(i)[x]
+   ifelse(all(is.na(p)), NA, p[!is.na(p)])
+ })
> toPercent <- with(tab, !is.na(statistic) & statistic=='cv')
> tab$variability[toPercent] <- percent(tab$variability[toPercent])
> tab$variability <- as.character(signif(tab$variability, 3))
> tab$statistic <- map(tab$statistic, from=c(NA, 'cv', 'cor', 'sd'), to=c(NA, '\\\\%CV', 'CORR', 'SD'))
> tab$variability <- paste(tab$statistic, tab$variability, sep=' = ')
> tab$variability[is.na(tab$statistic)] <- NA
> tab$statistic <- NULL
> tab$cv <- NULL
> tab$sd <- NULL
> tab$cor <- NULL
```

7 simple parameter table

We can make a quick parameter table that does not use wikitab markup. Table 2.

Listing 26:

```
> tab <- rlog(1005, '../nonmem', tool='nm7', file=NULL)
> head(tab)
```

Table 1: Parameter Estimates from Population Pharmacokinetic Model Run 1005

| description | model | estimate | prse | ci |
|---|--|-------------------|------|--------------------|
| apparent oral clearance | $CL/F \sim \theta_1 \cdot \theta_6^{MALE} \cdot (WT/70)^{\theta_7} \cdot e^{\eta_1}$ | 16 L/h | 15 | (6.67,11.1) |
| central volume of distribution | $V_c/F \sim \theta_2 \cdot (WT/70)^1 \cdot e^{\eta_2}$ | 14 L | 14 | (19,27.6) |
| absorption rate constant | $K_a \sim \theta_3 \cdot e^{\eta_3}$ | 6 h ⁻¹ | 13 | (0.0636,0.0814) |
| intercompartmental clearance | $Q/F \sim \theta_4$ | 15 L/h | 3 | (2.77,4.97) |
| peripheral volume of distribution | $V_p/F \sim \theta_5$ | 12 L | 5 | (87.9,315) |
| male effect on clearance | $MALE_{CL/F} \sim \theta_6$ | 11 | 2 | (0.845,1.27) |
| weight effect on clearance | $WT_{CL/F} \sim \theta_7$ | 13 | 8 | (0.685,1.96) |
| interindividual variability of clearance | $IIV_{CL/F} \sim \Omega_{1.1}$ | 10 | 6 | (0.127,0.325) |
| interindividual clearance-volume covariance | $cov_{CL,V} \sim \Omega_{2.1}$ | 8 | 7 | (0.0657,0.183) |
| interindividual variability of central volume | $IIV_{V_c/F} \sim \Omega_{2.2}$ | 7 | 9 | (0.0457,0.159) |
| interindividual clearance-Ka covariance | $cov_{CL,Ka} \sim \Omega_{3.1}$ | 2 | 4 | (-0.0438,0.0248) |
| interindividual volume-Ka covariance | $cov_{V,Ka} \sim \Omega_{3.2}$ | 3 | 12 | (-0.0536,-0.00759) |
| interindividual variability of Ka | $IIV_{K_a} \sim \Omega_{3.3}$ | 4 | 11 | (0.0237,0.0789) |
| proportional error | $err_{prop} \sim \Sigma_{1.1}$ | 5 | 1 | (0.0404,0.0594) |
| additive error | $err_{add} \sim \Sigma_{2.2}$ | 9 | 10 | (0.073,0.323) |

```

tool  run parameter  moment      value
1  nm7 1005      ofv  minimum 2405.91626347113
2  nm7 1005      THETA1 estimate    9.5079
3  nm7 1005      THETA1 prse         9.72
4  nm7 1005      THETA1 se          0.923845
5  nm7 1005      THETA2 estimate    22.7899
6  nm7 1005      THETA2 prse         9.56

```

Listing 27:

```

> tab$tool <- NULL
> tab$run <- NULL
> tab <- tab[tab$moment %in% c('estimate','prse'),]
> unique(tab$parameter)

[1] "THETA1" "THETA2" "THETA3" "THETA4" "THETA5" "THETA6"
[7] "THETA7" "OMEGA1.1" "OMEGA2.1" "OMEGA2.2" "OMEGA3.1" "OMEGA3.2"
[13] "OMEGA3.3" "SIGMA1.1" "SIGMA2.1" "SIGMA2.2"

```

Listing 28:

```

> tab$value <- signif(as.numeric(tab$value),3)
> tab$parameter <- factor(tab$parameter,levels=unique(tab$parameter))#to preserve
  row order during cast
> tab <- cast(tab,parameter ~ moment)
> tab

```

```

      parameter estimate    prse
1      THETA1      9.5100    9.72
2      THETA2     22.8000    9.56
3      THETA3      0.0714    7.34
4      THETA4      3.4700   15.40
5      THETA5    113.0000   20.90
6      THETA6      1.0200   11.00
7      THETA7      1.1900   28.30
8      OMEGA1.1     0.2140   22.80
9      OMEGA2.1     0.1210   26.40
10     OMEGA2.2     0.0945   33.20
11     OMEGA3.1    -0.0116  173.00
12     OMEGA3.2    -0.0372   36.10
13     OMEGA3.3     0.0466   34.80
14     SIGMA1.1     0.0492   10.90
15     SIGMA2.1     0.0000    Inf
16     SIGMA2.2     0.2020   33.50

```

Listing 29:

```

> tab$parameter <- parameter2wiki(tab$parameter)
> tab

```

```

      parameter estimate    prse
1      theta_1      9.5100    9.72
2      theta_2     22.8000    9.56
3      theta_3      0.0714    7.34
4      theta_4      3.4700   15.40
5      theta_5    113.0000   20.90
6      theta_6      1.0200   11.00
7      theta_7      1.1900   28.30
8      Omega_1.1     0.2140   22.80
9      Omega_2.1     0.1210   26.40
10     Omega_2.2     0.0945   33.20
11     Omega_3.1    -0.0116  173.00
12     Omega_3.2    -0.0372   36.10
13     Omega_3.3     0.0466   34.80
14     Sigma_1.1     0.0492   10.90
15     Sigma_2.1     0.0000    Inf
16     Sigma_2.2     0.2020   33.50

```

Listing 30:

```

> tab$parameter <- wiki2latex(tab$parameter)
> tab

```

```

      parameter estimate    prse
1   $\theta_1$       9.5100    9.72
2   $\theta_2$      22.8000    9.56
3   $\theta_3$       0.0714    7.34
4   $\theta_4$       3.4700   15.40

```

```

5    $\mathrm{\theta_5}$ 113.0000 20.90
6    $\mathrm{\theta_6}$ 1.0200 11.00
7    $\mathrm{\theta_7}$ 1.1900 28.30
8    $\mathrm{\Omega_{1.1}}$ 0.2140 22.80
9    $\mathrm{\Omega_{2.1}}$ 0.1210 26.40
10   $\mathrm{\Omega_{2.2}}$ 0.0945 33.20
11   $\mathrm{\Omega_{3.1}}$ -0.0116 173.00
12   $\mathrm{\Omega_{3.2}}$ -0.0372 36.10
13   $\mathrm{\Omega_{3.3}}$ 0.0466 34.80
14   $\mathrm{\Sigma_{1.1}}$ 0.0492 10.90
15   $\mathrm{\Sigma_{2.1}}$ 0.0000 Inf
16   $\mathrm{\Sigma_{2.2}}$ 0.2020 33.50

```

Table 2: Simple Parameter Table

| parameter | estimate | prse |
|----------------|----------|--------|
| θ_1 | 9.5100 | 9.72 |
| θ_2 | 22.8000 | 9.56 |
| θ_3 | 0.0714 | 7.34 |
| θ_4 | 3.4700 | 15.40 |
| θ_5 | 113.0000 | 20.90 |
| θ_6 | 1.0200 | 11.00 |
| θ_7 | 1.1900 | 28.30 |
| $\Omega_{1.1}$ | 0.2140 | 22.80 |
| $\Omega_{2.1}$ | 0.1210 | 26.40 |
| $\Omega_{2.2}$ | 0.0945 | 33.20 |
| $\Omega_{3.1}$ | -0.0116 | 173.00 |
| $\Omega_{3.2}$ | -0.0372 | 36.10 |
| $\Omega_{3.3}$ | 0.0466 | 34.80 |
| $\Sigma_{1.1}$ | 0.0492 | 10.90 |
| $\Sigma_{2.1}$ | 0.0000 | Inf |
| $\Sigma_{2.2}$ | 0.2020 | 33.50 |