

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 LaTeXtable. Table 1.

Listing 2:

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

| | 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
     THETA7
                                weight effect on clearance
                  interindividual variability of clearance
   OMEGA1.1
              interindividual clearance-volume covariance
9
   OMEGA2.1
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
  CL/F (L/h) ~ theta_1 * theta_6 ^MALE * (WT/70) ^theta_7 * e^eta_1
                                                                        9.5100
2
                           V_c /F (L) \sim theta_2 * (WT/70)^1 * e^eta_2 22.8000
3
                                      K_a (h^-1) \sim theta_3 * e^-ta_3
                                                                        0.0714
4
                                                  Q/F (L/h) ~ theta_4
                                                                        3.4700
5
                                                 V_p /F (L) ~ theta_5 113.0000
6
                                                 MALE_CL/F ~ theta_6 1.0200
7
                                                    WT CL/F \sim theta 7
                                                                        1.1900
8
                                                 IIV_CL/F \sim Omega_1.1 \quad 0.2140
9
                                                 cov_CL, V ~ Omega_2.1
                                                                        0.1210
10
                                               IIV_V_c /F \sim Omega_2.2
                                                                        0.0945
11
                                               cov_CL, Ka ~ Omega_3.1 -0.0116
12
                                                cov_V, Ka \sim Omega_3.2 -0.0372
13
                                                IIV_K_a \sim Omega_3.3
                                                                         0.0466
14
                                                err_prop ~ Sigma_1.1
                                                                        0.0492
15
                                                 err_add ~ Sigma_2.2
                                                                        0.2020
   prse
1 9.84
2 9.56
3 7.35
4 15.4
5 21
6 11.2
7 28.4
```



```
8 22.8
9 26.4
10 33.2
11 173
12 36.1
13 34.7
14 10.9
15 33.5
```

Now we can extract some information from the model statements.

Listing 3:

```
> tab$units <- justUnits(tab$model)</pre>
> tab$model <- noUnits(tab$model)</pre>
> tab$name <- with(tab, wiki2label(model))</pre>
> 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 \sim theta_2 * (WT/70)^1 * e^eta_2
3
                                          K_a \sim theta_3 * e^eta_3 h^{-1}
4
                                                     Q/F \sim theta_4 L/h
5
                                                  V_p /F \sim theta_5
6
                                               MALE_CL/F \sim theta_6
7
                                                  WT_CL/F \sim theta_7
8
                                               IIV_CL/F ~ Omega_1.1
9
                                               cov_CL, V ~ Omega_2.1
10
                                            IIV_V_c /F \sim Omega_2.2
11
                                            cov_CL, Ka ~ Omega_3.1
12
                                             cov_V, Ka ~ Omega_3.2
13
                                              IIV_K_a \sim 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
    WT_CL/F
   IIV_CL/F
   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[,c('parameter','name','estimate','cv')]
  parameter
                name estimate
                                    CV
1
     THETA1
                CL/F
                      9.5100
                                    NA
2
     THETA2
               V_c/F 22.8000
                                    NA
3
               K_a
                      0.0714
     THETA3
                                    NA
                 Q/F
4
     THETA4
                      3.4700
                                    NA
5
     THETA5
               V_p/F 113.0000
                                    NA
6
     THETA6 MALE_CL/F
                      1.0200
                                    NA
7
     THETA7
              WT_CL/F
                      1.1900
                                    NA
   OMEGA1.1 IIV_CL/F
                      0.2140 0.4884902
   OMEGA2.1 cov_CL,V 0.1210
9
                                    NA
10 OMEGA2.2 IIV_V_c/F
                       0.0945 0.3148161
   OMEGA3.1 cov_CL, Ka -0.0116
                                    NA
12 OMEGA3.2 cov_V, Ka -0.0372
                                    NA
13 OMEGA3.3 IIV_K_a
                      0.0466 0.2184098
14 SIGMA1.1 err_prop
                      0.0492
                                    NA
15 SIGMA2.2 err_add
                      0.2020
                                    NA
```

> tab\$cv[expo] <- cvLognormal(tab\$estimate[expo])</pre>

3.2 proportional

Listing 6:

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

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



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

NA

NA

0.0466 0.2184098

Listing 7:

```
> tab$parameter[prop]
[1] "SIGMA1.1"
                                                     Listing 8:
> tab$cv[prop] <- sqrt(tab$estimate[prop])</pre>
> tab[,c('parameter','name','estimate','cv')]
  parameter
                 name estimate
                                       CV
      THETA1
                 CL/F
                        9.5100
                                      NA
2
                V_c/F 22.8000
      THETA2
                                      NA
3
     THETA3
                 K_a
                       0.0714
                                      NA
4
     THETA4
                  Q/F
                        3.4700
                                      NA
     THETA5
                V_p/F 113.0000
                                      NA
6
     THETA6 MALE_CL/F
                       1.0200
                                      NA
     THETA7
              WT_CL/F
                        1.1900
                                      NA
8
   OMEGA1.1 IIV_CL/F
                        0.2140 0.4884902
9
    OMEGA2.1 cov_CL, V 0.1210
                                      NA
   OMEGA2.2 IIV_V_c/F
                        0.0945 0.3148161
11 OMEGA3.1 cov_CL, Ka -0.0116
                                      NA
```

3.3 additive

12 OMEGA3.2 cov_V, Ka -0.0372

15 SIGMA2.2 err_add 0.2020

14 SIGMA1.1 err_prop 0.0492 0.2218107

13 OMEGA3.3 IIV_K_a

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])</pre>
> tab[,c('parameter','name','estimate','cv','sd')]
                 name estimate
                                               sd
  parameter
                                      CV
1
     THETA1
                 CL/F
                       9.5100
                                     NA
                                               NA
2
     THETA2
                V_c/F 22.8000
                                     NA
                                               NA
3
                       0.0714
     THETA3
                K_a
                                     NA
                                               NA
4
     THETA4
                  Q/F
                       3.4700
                                     NA
                                               NA
5
     THETA5
                V_p/F 113.0000
                                     NA
                                               NA
6
     THETA6 MALE_CL/F
                       1.0200
                                               NA
                                     NA
7
     THETA7
              WT_CL/F
                       1.1900
                                      NA
                                               NA
   OMEGA1.1 IIV_CL/F
                       0.2140 0.4884902
                                               NA
9
   OMEGA2.1 cov_CL, V
                       0.1210
                                               NA
                                      NA
10
   OMEGA2.2 IIV_V_c/F
                       0.0945 0.3148161
                                               NA
   OMEGA3.1 cov_CL, Ka -0.0116
                                               NA
                                      NA
12 OMEGA3.2 cov_V, Ka -0.0372
                                     NA
                                               NA
13 OMEGA3.3 IIV_K_a
                       0.0466 0.2184098
                                               NA
14 SIGMA1.1 err_prop 0.0492 0.2218107
                                               NA
15 SIGMA2.2 err_add
                      0.2020
                                     NA 0.4494441
```

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
```



```
[,2]
                                 [,3]
           [,1]
[1,] 1.0000000 0.8492811 -0.1163229
[2,] 0.8492811 1.0000000 -0.5607054
[3,] -0.1163229 -0.5607054 1.0000000
                                                        Listing 12:
> half(cor)
       1.1
                   2.1
                              2.2
                                          3.1
                                                      3.2
                                                                 3.3
 1.0000000 \quad 0.8492811 \quad 1.0000000 \quad -0.1163229 \quad -0.5607054 \quad 1.0000000
                                                        Listing 13:
> offdiag(half(cor))
       2.1
                  3.1
                              3.2
 0.8492811 -0.1163229 -0.5607054
                                                        Listing 14:
> off <- is.iiv(tab$parameter) & is.offdiagonal(tab$parameter)</pre>
> tab$parameter[off]
[1] "OMEGA2.1" "OMEGA3.1" "OMEGA3.2"
                                                        Listing 15:
> tab$cor[off] <- offdiag(half(cor))</pre>
> tab[,c('parameter','name','estimate','cv','sd','cor')]
   parameter
                  name estimate
                                         CV
                                                    sd
                                                              cor
1
      THETA1
                  CL/F 9.5100
                                         NA
                                                    NA
                                                               NA
2
                 V c/F 22.8000
      THETA2
                                         NA
                                                    NA
                                                               NA
      THETA3
                 K_a 0.0714
                                         NA
                                                    NA
                                                               NA
```



```
3.4700
                                               NA
     THETA4
                  Q/F
                                     NA
                                                          NA
5
     THETA5
                V_p/F 113.0000
                                     NA
                                               NA
                                                          NA
     THETA6 MALE_CL/F
                        1.0200
                                     NA
                                               NA
                                                          NA
7
     THETA7
              WT_CL/F
                        1.1900
                                     NA
                                               NA
                                                          NA
   OMEGA1.1 IIV_CL/F
                        0.2140 0.4884902
                                               NA
                                                          NA
9
   OMEGA2.1 cov_CL, V
                       0.1210
                                     NA
                                               NA 0.8492811
   OMEGA2.2 IIV_V_c/F
10
                       0.0945 0.3148161
                                               NA
                                                          NA
   OMEGA3.1 cov_CL, Ka -0.0116
                                     NA
                                               NA -0.1163229
12 OMEGA3.2 cov_V, Ka -0.0372
                                     NA
                                               NA -0.5607054
13 OMEGA3.3 IIV K a
                      0.0466 0.2184098
                                               NA
14 SIGMA1.1 err_prop
                       0.0492 0.2218107
                                                          NA
15 SIGMA2.2 err_add
                      0.2020
                                     NA 0.4494441
                                                          NA
```

5 confidence interval

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

> boot <- boot[boot\$moment=='estimate',]</pre>

Listing 16:

```
> boot <- read.csv('../nonmem/1005.bootlog.csv',as.is=TRUE)</pre>
> head(boot)
                                            value
  X tool run parameter
                          moment
1 1 nm7
                    ofv minimum 2353.21240698929
2 2
    nm7
                THETA1 estimate
                                          8.64914
3 3 nm7
          1
                THETA1
                                             <NA>
                            prse
4 4
     nm7
                THETA1
                                              <NA>
5 5
                                          21.5594
     nm7
                THETA2 estimate
6 6 nm7
          1
                                              <NA>
                THETA2
                            prse
                                                       Listing 17:
```



```
> boot <- data.frame(cast(boot,... ~ moment))</pre>
> head(boot)
   X tool run parameter estimate
1 2 nm7 1
                THETA1 8.64914
2 5 nm7 1
                THETA2 21.5594
3 8 nm7 1
              THETA3 0.0766718
4 11 nm7 1 THETA4 4.05759
5 14 nm7 1
                THETA5 106.257
6 17 nm7 1
                THETA6 1.07562
                                                    Listing 18:
> boot <- boot[,c('run','parameter','estimate')]</pre>
> sapply(boot,class)
                          estimate
       run parameter
  "integer" "character"
                          "factor"
                                                    Listing 19:
> boot$estimate <- as.numeric(as.character(boot$estimate))</pre>
> unique(boot$parameter)
[1] "THETA1" "THETA2"
                          "THETA3" "THETA4" "THETA5" "THETA6"
[7] "THETA7" "OMEGA1.1" "OMEGA2.1" "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))</pre>
> boot$1o <- with(boot, reapply(estimate,parameter,quan,probs=.05))</pre>
> boot$hi <- with(boot, reapply(estimate,parameter,quan,probs=.95))</pre>
> head(boot)
```



```
run parameter
                  estimate
                               10
                                      hi
         THETA1
                 8.6491400
                             6.19
                                      11
2
         THETA2 21.5594000
                             19.1
                                    26.4
                 0.0766718 0.0607 0.0801
3
   1
        THETA3
4
   1
         THETA4
                 4.0575900
                             2.82
                                    5.06
5
        THETA5 106.2570000
                             87.9
                                     930
6
   1
                1.0756200
                             0.86
                                    1.29
         THETA6
```

Listing 21:

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

```
parameter
                 10
                          hi
     THETA1
               6.19
                          11
1
2
     THETA2
               19.1
                        26.4
3
                      0.0801
     THETA3 0.0607
4
               2.82
                        5.06
     THETA4
5
     THETA5
               87.9
                         930
6
     THETA6
               0.86
                        1.29
     THETA7
              0.571
                        1.87
8
   OMEGA1.1
              0.132
                       0.431
9
   OMEGA2.1 0.0663
                       0.222
   OMEGA2.2 0.0511
                       0.162
10
   OMEGA3.1 -0.046
                      0.0242
11
12 OMEGA3.2 -0.0565 -0.00863
13 OMEGA3.3 0.0231
                      0.0738
14 SIGMA1.1 0.0403
                      0.0592
15 SIGMA2.1
                           0
16 SIGMA2.2 0.0645
                       0.304
```

Listing 22:

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



```
10
                            hi
   parameter
                                                 ci
      THETA1
                 6.19
                            11
                                          (6.19, 11)
2
      THETA2
                19.1
                          26.4
                                       (19.1, 26.4)
3
      THETA3 0.0607
                        0.0801
                                   (0.0607, 0.0801)
4
                          5.06
                                       (2.82, 5.06)
     THETA4
                 2.82
5
      THETA5
                87.9
                           930
                                        (87.9,930)
6
     THETA6
                0.86
                          1.29
                                       (0.86, 1.29)
     THETA7
                0.571
                          1.87
                                      (0.571, 1.87)
8
    OMEGA1.1
               0.132
                         0.431
                                     (0.132, 0.431)
9
    OMEGA2.1 0.0663
                         0.222
                                    (0.0663, 0.222)
    OMEGA2.2 0.0511
                         0.162
                                    (0.0511, 0.162)
    OMEGA3.1 -0.046
                        0.0242
                                   (-0.046, 0.0242)
12
    OMEGA3.2 -0.0565 -0.00863 (-0.0565, -0.00863)
    OMEGA3.3 0.0231
                        0.0738
                                   (0.0231, 0.0738)
   SIGMA1.1 0.0403
                        0.0592
                                   (0.0403, 0.0592)
                             0
15 SIGMA2.1
                    0
                                              (0,0)
16 SIGMA2.2 0.0645
                         0.304
                                    (0.0645, 0.304)
```

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
     THETA7
                                weight effect on clearance
8
    OMEGA1.1
                  interindividual variability of clearance
9
    OMEGA2.1
               interindividual clearance-volume covariance
    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
   CL/F \sim theta_1 * theta_6 ^MALE * (WT/70)^theta_7 * e^eta_1
                                                                      9.5100 9.84
2
                          V_c / F \sim theta_2 * (WT/70)^1 * e^eta_2
                                                                    22.8000 9.56
3
                                          K_a \sim theta_3 * e^eta_3
                                                                      0.0714 7.35
4
                                                    Q/F \sim theta_4
                                                                      3.4700 15.4
5
                                                 V_p /F ~ theta_5 113.0000 21
6
                                               MALE_CL/F \sim theta_6
                                                                     1.0200 11.2
7
                                                 WT_CL/F \sim theta_7
                                                                     1.1900 28.4
8
                                              IIV_CL/F ~ Omega_1.1
                                                                     0.2140 22.8
9
                                              cov_CL, V ~ Omega_2.1
                                                                      0.1210 26.4
10
                                            IIV_V_c /F \sim Omega_2.2
                                                                      0.0945 33.2
11
                                            cov_CL, Ka ~ Omega_3.1
                                                                     -0.0116 173
12
                                             cov_V, Ka ~ Omega_3.2 -0.0372 36.1
13
                                              IIV K a ~ Omega 3.3
                                                                      0.0466 34.7
14
                                              err_prop ~ Sigma_1.1
                                                                      0.0492 10.9
15
                                                                      0.2020 33.5
                                              err_add ~ Sigma_2.2
   units
                                                cor
                                                                     ci
              name
                                     sd
                           CV
1
     L/h
              CL/F
                           NA
                                     NA
                                                 NA
                                                              (6.19, 11)
2
       L
             V_c/F
                           NA
                                     NA
                                                 NA
                                                            (19.1, 26.4)
3
  h^-1
                                                       (0.0607, 0.0801)
               K_a
                           NA
                                     NA
                                                 NA
4
     L/h
               Q/F
                           NA
                                     NA
                                                 NA
                                                            (2.82, 5.06)
5
             V_p/F
                           NA
                                     NA
                                                 NA
                                                            (87.9,930)
6
         MALE_CL/F
                                     NA
                                                            (0.86, 1.29)
                           NA
                                                 NA
7
           WT_CL/F
                           NA
                                     NA
                                                 NA
                                                          (0.571, 1.87)
8
          IIV_CL/F 0.4884902
                                     NA
                                                 NA
                                                          (0.132, 0.431)
9
          cov_CL, V
                           NA
                                     NA
                                         0.8492811
                                                        (0.0663, 0.222)
10
         IIV_V_c/F 0.3148161
                                     NA
                                                 NA
                                                        (0.0511, 0.162)
11
         cov_CL, Ka
                           NA
                                     NA -0.1163229
                                                        (-0.046, 0.0242)
12
          cov_V, Ka
                                     NA -0.5607054 (-0.0565, -0.00863)
13
          IIV_K_a 0.2184098
                                     NA
                                                 NA
                                                        (0.0231, 0.0738)
```



```
14 err_prop 0.2218107 NA NA (0.0403,0.0592)
15 err add NA 0.4494441 NA (0.0645,0.304)
```

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</pre>
```

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$variability)] <- NA
> i <- !is.na(m)
> i[!i] <- NA
> tab$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'))</pre>
```



```
> tab$variability <- paste(tab$statistic,tab$variability,sep=' = ')
> tab$variability[is.na(tab$statistic)] <- NA
> tab$statistic <- NULL
> tab$cv <- NULL
> tab$cor <- NULL
> tab$scd <- NULL
> tab$scd <- NULL
```

Table 1: Parameter Estimates from Population Pharmacokinetic Model Run 1005

| description | model | estimate | prse | ci | variability |
|---|--|---------------|------|--------------------|---------------|
| apparent oral clearance | $\mathrm{CL/F} \sim \theta_1 \cdot \theta_6^{\mathrm{MALE}} \cdot (\mathrm{WT/70})^{\theta_7} \cdot \mathrm{e}^{\eta_1}$ | 9.51 L/h | 9.84 | (6.19,11) | |
| central volume of distribution | $V_c/F \sim \theta_2 \cdot (WT/70)^1 \cdot e^{\eta_2}$ | 22.8 L | 9.56 | (19.1,26.4) | |
| absorption rate constant | $ m K_a \sim 	heta_3 \cdot e^{\eta_3}$ | 0.0714 h^-1 | 7.35 | (0.0607, 0.0801) | |
| intercompartmental clearance | $\mathrm{Q/F}\sim	heta_4$ | 3.47 L/h | 15.4 | (2.82,5.06) | |
| peripheral volume of distribution | $ m V_p/F \sim 	heta_5$ | 113 L | 21 | (87.9,930) | |
| male effect on clearance | $\mathrm{MALE_{CL/F}} \sim 	heta_6$ | 1.02 | 11.2 | (0.86, 1.29) | |
| weight effect on clearance | ${ m WT}_{ m CL/F} \sim 	heta_7$ | 1.19 | 28.4 | (0.571, 1.87) | |
| interindividual variability of clearance | $\mathrm{IIV}_{\mathrm{CL/F}} \sim \Omega_{1.1}$ | 0.214 | 22.8 | (0.132, 0.431) | %CV = 48.8 |
| interindividual clearance-volume covariance | ${\rm cov}_{{ m CL,V}} \sim \Omega_{2.1}$ | 0.121 | 26.4 | (0.0663, 0.222) | CORR = 0.849 |
| interindividual variability of central volume | $\mathrm{IIV}_{\mathrm{V_c}/\mathrm{F}} \sim \Omega_{2.2}$ | 0.0945 | 33.2 | (0.0511,0.162) | %CV = 31.5 |
| interindividual clearance-Ka covariance | $\mathrm{cov}_{\mathrm{CL,Ka}} \sim \Omega_{3.1}$ | -0.0116 | 173 | (-0.046, 0.0242) | CORR = -0.116 |
| interindividual volume-Ka covariance | ${ m cov_{V,Ka}} \sim \Omega_{3.2}$ | -0.0372 | 36.1 | (-0.0565,-0.00863) | CORR = -0.561 |
| interindividual variability of Ka | $\mathrm{IIV}_{\mathrm{K_a}} \sim \Omega_{3.3}$ | 0.0466 | 34.7 | (0.0231, 0.0738) | %CV = 21.8 |
| proportional error | $\mathrm{err}_{\mathrm{prop}} \sim \Sigma_{1.1}$ | 0.0492 | 10.9 | (0.0403, 0.0592) | %CV = 22.2 |
| additive error | $\mathrm{err}_{\mathrm{add}} \sim \Sigma_{2.2}$ | 0.202 | 33.5 | (0.0645, 0.304) | SD = 0.449 |



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)</pre>
> head(tab)
 tool run parameter
                                         value
                       moment
1 nm7 1005
                 ofv minimum 2405.91625845151
2 nm7 1005
              THETA1 estimate
                                      9.50754
3 nm7 1005 THETA1
                      prse
                                          9.84
4 nm7 1005
              THETA1
                                      0.935942
                                       22.7907
5 nm7 1005
              THETA2 estimate
6 nm7 1005
                                          9.56
              THETA2
                         prse
                                                    Listing 27:
> tab$tool <- NULL
> tab$run <- NULL
> tab <- tab[tab$moment %in% c('estimate','prse'),]</pre>
> unique(tab$parameter)
[1] "THETA1" "THETA2"
                          "THETA3" "THETA4" "THETA5" "THETA6"
[7] "THETA7" "OMEGA1.1" "OMEGA2.1" "OMEGA3.1" "OMEGA3.2"
[13] "OMEGA3.3" "SIGMA1.1" "SIGMA2.1" "SIGMA2.2"
                                                    Listing 28:
> tab$value <- signif(as.numeric(tab$value),3)</pre>
> tab$parameter <- factor(tab$parameter,levels=unique(tab$parameter))#to preserve row order during cast
> tab <- cast(tab,parameter ~ moment)</pre>
> tab
```



```
parameter estimate
                     prse
     THETA1 9.5100
                     9.84
2
     THETA2 22.8000
                     9.56
3
    THETA3 0.0714
                     7.35
4
    THETA4 3.4700 15.40
5
    THETA5 113.0000 21.00
6
    THETA6 1.0200 11.20
    THETA7 1.1900 28.40
   OMEGA1.1 0.2140 22.80
8
9
   OMEGA2.1 0.1210 26.40
   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 34.70
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
    theta_1
              9.5100
                      9.84
    theta_2 22.8000
                     9.56
3
    theta_3 0.0714
                     7.35
    theta_4
              3.4700 15.40
    theta_5 113.0000 21.00
6
              1.0200 11.20
    theta_6
7
    theta_7
              1.1900 28.40
              0.2140 22.80
8 Omega_1.1
9 Omega_2.1
              0.1210 26.40
10 Omega_2.2
              0.0945 33.20
11 Omega_3.1
             -0.0116 173.00
```



Listing 30:

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

```
parameter estimate
                                      prse
    $\\mathrm{\\theta_{1}}$
1
                             9.5100
                                      9.84
2
    $\\mathrm{\\theta_{2}}$ 22.8000
                                      9.56
3
    $\\mathrm{\\theta_{3}}$
0.0714
                                      7.35
4
    $\\mathrm{\\theta_{4}}$
                                     15.40
                             3.4700
    $\\mathrm{\\theta_{5}}$ 113.0000 21.00
    $\\mathrm{\\theta_{6}}$
1.0200
                                     11.20
    $\\mathrm{\\theta_{7}}$
1.1900
                                     28.40
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 34.70
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 | 9.84 |
| $	heta_2$ | 22.8000 | 9.56 |
| θ_3 | 0.0714 | 7.35 |
| $	heta_4$ | 3.4700 | 15.40 |
| $	heta_5$ | 113.0000 | 21.00 |
| θ_6 | 1.0200 | 11.20 |
| θ_7 | 1.1900 | 28.40 |
| $\Omega_{1.1}$ | 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 | 34.70 |
| $\Sigma_{1.1}$ | 0.0492 | 10.90 |
| $\Sigma_{2.1}$ | 0.0000 | Inf |
| $\Sigma_{2.2}$ | 0.2020 | 33.50 |