

```

1  $PROBLEM ONE COMPARTMENT ORAL                ;DATE = 6/2/04 PROGRAMMER=XXXX
2                                                ;Units: Time=hr, Concentration=ug/ml,
3                                                ;Dose = 100mg and 250mg
4
5  $DATA ORAL1EST_CRSO.CSV IGNORE=C
6
7  $INPUT ID TIME CONC=DV AMT DOSE EVID OCC MDV
8
9  $SUBROUTINE ADVAN2 TRANS2
10
11 $PK
12   OC1=0
13   IF(OCC.EQ.1)OC1=1
14   OC2=0
15   IF(OCC.EQ.2)OC2=1
16
17   BOVCL = ETA(4)*OC1+ETA(5)*OC2
18   BOVV  = ETA(6)*OC1+ETA(7)*OC2
19
20   POPCL = THETA(1)
21   POPV  = THETA(2)
22   POPKA = THETA(3)
23
24   CL = POPCL*EXP(ETA(1)+BOVCL)      ;CL/F
25   V  = POPV*EXP(ETA(2)+BOVV)       ;V/F
26   KA = POPKA*EXP(ETA(3))           ;Absorption rate constant
27
28   S2 = V                            ;Scaling factor
29
30
31 $ERROR
32   IPRED=F
33   Y=F+F*ERRCV+ERRSD
34                                     ;Additive and proportional residual error model
35
36 $THETA (1,10)          ;POPCL/F
37 $THETA (1,30)         ;POPV/F
38 $THETA (0.1,3)        ;POPKA
39
40 $OMEGA 0.09            ;BSVCL
41 $OMEGA 0.09            ;BSVV
42 $OMEGA 0.09            ;BSVKA
43 $OMEGA BLOCK(1) 0.09  ;BOVCL occ=1
44 $OMEGA BLOCK(1) SAME  ;BOVCL occ=2;SAME specifies random effects
45                       ;sampled from the same distribution
46 $OMEGA BLOCK(1) 0.09  ;BOVV occ=1
47 $OMEGA BLOCK(1) SAME  ;BOVV occ=2
48
49
50 $SIGMA 0.01           ;ERRCV
51 $SIGMA 1              ;ERRSD
52
53 $ESTIMATION METHOD=0 MAXEVAL=9999 PRINT=5 POSTHOC
54 $COVARIANCE
55
56 $TABLE ID TIME DV IPRED DOSE
57   NOPRINT ONEHEADER FILE=oralest_crso.fit
58
59

```