

```
1 $PROBLEM PROJECT singledose i.v bolus and infusion ;DATE 6-2-04 PROGRAMMER:XXXX
2 ;UNITS: Time=min, Concentration=ng/ml
3 ;Clearance=L/min, Volume = L
4
5 $DATA ivlinfest_crso.csv IGNORE=C
6 $INPUT ID TIME CONC=DV AMT CMT RATE EVID OCC MDV
7 $SUBROUTINE ADVAN1 TRANS2
8 $PK
9 OC1=0
10 IF(OCC.EQ.1)OC1=1
11 OC2=0
12 IF(OCC.EQ.2)OC2=1
13
14 BOVCL = ETA(3)*OC1+ETA(4)*OC2
15 BOVV = ETA(5)*OC1+ETA(6)*OC2
16
17 POPCL = THETA(1)
18 CL = POPCL*EXP(ETA(1)+BOVCL) ;Clearance in L/min
19
20 POPV = THETA(2)
21 V = POPV*EXP(ETA(2)+BOVV) ;Volume of distribution in L
22 S1 = V/1000 ;To get concentration in ng/ml
23
24
25 $ERROR
26 IPRED=F
27 Y=F+F*ERRCV+ERRSD
28
29 $THETA (0.01,1) ;POPCL
30 $THETA (1,20) ;POPV
31
32 $OMEGA 0.09 ;BSVCL
33 $OMEGA 0.09 ;BSVV
34 $OMEGA BLOCK(1) 0.09 ;BOVCL occ=1
35 $OMEGA BLOCK(1) SAME ;BOVCL occ=2;SAME specifies random effects
36 ;sampled from the same distribution
37 $OMEGA BLOCK(1) 0.09 ;BOVV occ=1
38 $OMEGA BLOCK(1) SAME ;BOVV occ=2
39
40 $SIGMA 0.0025 ;ERRCV
41 $SIGMA 1 ;ERRSD
42
43 $ESTIMATION METHOD=0 MAXEVAL=9990 PRINT=10 POSTHOC
44 $COVARIANCE
45
46 $TABLE ID TIME DV IPRED
47 NOPRINT ONEHEADER FILE=ivlinfest_crso.fit
48
```