

Supplemental Data 3

R scripts to perform the selection and validation of the calibration model

A video “user guide” containing instructions on how to setup and execute the R scripts is available at <https://youtu.be/azpD2GG0qNA>. Please watch it before using the R scripts.

The “.R” files ready for use are available at <https://bit.ly/2V1LVPx>.

```
1 #####  
2 ## File to be run ##  
3 #####  
4  
5 ## Parameters to be set  
6 directory <- "~/Desktop/Bridge/Calibration" # set work directory : folder where the  
    calibration data is  
                                # and the two R sheets Calibration2Ks and  
    Calibration2CVM  
7  
8 filename <- "Model.txt" # name of the file that contains the raw data (calibration data)  
9 result_filename <- "My_Results.txt" # name of the file that contains the results  
10 dec <- ".#" # , if the decimals are ex. 0,5 . if the decimals are ex 0.5  
11 nbgrille = 50 # global parameter for integral approximations  
12 b <- 1000 #global parameter for bootstrapping  
13 stat <- 2 # 1: KS, 2: CVM  
14 alpha <- 0.05 ## The alpha to use in hypothesis testing  
15 ## end of parameter setting  
16  
17  
18 #####  
19 ## Code ##  
20 #####  
21 source("CodeV5.R") ## Contains the main coded functions  
22 A <- read.table(filename,header=F,dec=dec) ## reading the data  
23 A <- as.matrix(A)  
24 stats = c("Kolmogorov Smirnov","Cramer von Mises")  
25  
26  
27 Decision <- rep(0,3)  
28 DCV <- rep(0,3)  
29  
30 # weight : 0  
31 weight <- 0;  
32 P0 = fitPlus(A,weight,b,nbgrille,stat)  
33 coP0 <- matrix(c(rev(P0$Linear$param),0,rev(P0$Quadratic$param)),ncol=3,byrow=T) ## Contains  
    the fitted coefficients  
34 colnames(coP0) <- c("b0","b1","b2")
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35 rownames(coP0) <- c("Linear", "Quadratic")
36
37
38 List0 <- list(Fitted.Param = coP0, Pval.Lin = P0$Linear$pvalN, Stat.Lin = P0$Linear$statN ,
39   Pval.Quad = P0$Quadratic$pvalN, Stat.Quad = P0$Quadratic$statN, Pval.PartialF = P0$ 
40   pvalFtest)
41 Decision [1] = 1 + (List0$Pval.PartialF < 0.05)
42
43 # weight : 1
44 weight <- 1;
45 P1 = fitPlus(A, weight, b, nbgrille, stat)
46 coP1 <- matrix(c(rev(P1$Linear$param), 0, rev(P1$Quadratic$param)), ncol=3, byrow=T) ## Contains
47   the fitted coefficients
48 colnames(coP1) <- c("b0", "b1", "b2")
49 rownames(coP1) <- c("Linear", "Quadratic")
50
51
52 List1 <- list(Fitted.Param = coP1, Pval.Lin = P1$Linear$pvalN, Stat.Lin = P1$Linear$statN ,
53   Pval.Quad = P1$Quadratic$pvalN, Stat.Quad = P1$Quadratic$statN, Pval.PartialF = P1$ 
54   pvalFtest)
55 Decision [2] = 1 + (List1$Pval.PartialF < 0.05)
56
57 # weight : 2
58 weight <- 2;
59 P2 = fitPlus(A, weight, b, nbgrille, stat)
60 coP2 <- matrix(c(rev(P2$Linear$param), 0, rev(P2$Quadratic$param)), ncol=3, byrow=T) ## Contains
61   the fitted coefficients
62 colnames(coP2) <- c("b0", "b1", "b2")
63 rownames(coP2) <- c("Linear", "Quadratic")
64
65 List2 <- list(Fitted.Param = coP2, Pval.Lin = P2$Linear$pvalN, Stat.Lin = P2$Linear$statN ,
66   Pval.Quad = P2$Quadratic$pvalN, Stat.Quad = P2$Quadratic$statN, Pval.PartialF = P2$ 
67   pvalFtest)
68 Decision [3] = 1+ (List2$Pval.PartialF < 0.05)
69
70 ## Variance Test for weight selection
71 Ano = A[, -1]/sqrt(A[, 1])
72 spoids = sum(1/sqrt(A[, 1]))
73 var1 = apply(Ano/spoids, 1, var)
74 Ano2 = A[, -1]/A[, 1]
75 spoids2 = sum(1/(A[, 1]))
76 var2 = apply(Ano2/spoids2, 1, var)
77 var0 = apply(A[, -1]/length(A[, 1]), 1, var)
78 s1 = signif(sum((var0 - mean(var0))^2), 3)
79 s2 = signif(sum((var1 - mean(var1))^2), 3)
80 s3 = signif(sum((var2 - mean(var2))^2), 3)
81 sss = c(s1, s2, s3)
82 z1 = paste("Variance test for weight selection")
83 z2 = paste("Scores:", "No weight:", s1, "x^(-1):", s2, "x^(-2):", s3)
84 ccc = c("no weight", "x^(-1)", "x^(-2)")

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84 ccb = paste("Selected weight: ",ccc [which(sss==min(sss))],sep="")
85 zaa = paste(z1,z2,ccb,sep="\n")
86
87 best <- which(sss==min(sss))
88
89 awiner = eval(parse(text = paste("List",best-1,sep="")))
90
91 ## Ftest Hétéro
92 pop1 = A[1,-1]
93 pop2 = A[nrow(A),-1]
94 ttt = var.test(pop1,pop2,alternative = "less")
95 bb = paste("F-test for heteroscedasticity")
96 bb1 = paste("p-value: ",signif(ttt$p.value,4))
97 ddd = "No"
98 if(ttt$p.value<alpha){ddd="Yes"}
99 bb2 = paste("Weighting needed:",ddd,sep=" ")
100 pop1 = paste(bb,bb1,bb2,sep="\n")
101
102
103 v = paste("Partial F-test for model order selection")
104 v1 = paste("p-value",signif(awiner$Pval_PartialF,4),sep=":")
105 v2 = paste("Model selected: ",c("linear","quadratic")[Decision[best]],sep="")
106 vv = paste(v,v1,v2,sep="\n")
107
108
109 w = paste("Normality of the standerdized residuals")
110 w1 = paste("Test used: ",stats[stat],sep="")
111 aw= eval(parse(text = paste("awiner$Pval_",
112 ,c("Lin","Quad"))[Decision[best]],sep=""))
113 w2 = paste("p-value: ",signif(aw,4))
114 ww = "No"
115 if( alpha<aw)
116 {
117   ww = "Yes"
118 }
119 wwl = paste("Validation test passed:",ww,sep=" ")
120 vw =paste(w,w1,w2,wwl,sep="\n")
121
122 md = c("Linear","Quadratic")[Decision[best]]
123 c("1","1/x","1/x^2")[best]
124 if(ww=="Yes")
125 {
126 zw=paste("Model selected: ",md," , ",c("1","1/x","1/x^2")[best],sep=" ")
127 zw2 = paste("Calibration equation:")
128 if(md=="Linear")
129 {
130   co1 = awiner$Fitted.Param[1,1]
131   co2 = awiner$Fitted.Param[1,2]
132   zw1 = paste(signif(co2,4)," x + ",signif(co1,4),sep="")
133 }
134 if(md=="Quadratic")
135 {
136   co1 = awiner$Fitted.Param[2,1]
137   co2 = awiner$Fitted.Param[2,2]
138   co3 = awiner$Fitted.Param[2,3]

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138 zw1 = paste(signif(co3,4)," x^2 + ",signif(co2,4)," x + ",signif(co1,4),sep="")
139 }
140 vw2 = paste(zw,zw2,zw1,sep="\n")
141 }
142 if(ww == "No"){
143   vw2 = paste("No model selected , validation test failed")
144 }
145
146 ## Plots
147 namePlot = strsplit(result_filename,".",fixed=TRUE)[[1]][1]
148 # Plot of variance
149 var_level = apply(A[,-1],1,var)
150 pdf(file=paste(namePlot,"Variance.pdf",sep=""))
151 plot(A[,1],var_level,xlab="Concentration",ylab="Variance",main="Variance plot",mgp=c(2,
152 0.8, 0), axes=T)
152 dev.off()
153 ## Calibration curve
154 f <- function(x)
155 {
156   predic(rev(awiner$Fitted.Param[Decision[best],]),x)
157 }
158 Cal.dots = predic(rev(awiner$Fitted.Param[Decision[best],]),A[,1])
159 pdf(file=paste(namePlot,"Calibration curve.pdf",sep=""))
160 plot(rep(A[,1],each=(ncol(A[,-1]))),c(t(A[,-1])),xlab="Concentration",ylab="Signal",main=
161 "Calibration curve",mgp=c(2, 0.8, 0), axes=T)
162 curve(f,add=T)
163 dev.off()
164 mt = paste(popl,zaa,vv,vw,vw2,sep="\n \n")
165 write(mt,file=result_filename)
166 cat(mt)

```

```

1 # A : sample of data
2 # nbgrille : precision for the approximation of the grid
3 # variance to be used in the normal distribution function
4
5 ## Computes sqrt(n) * sup_x | F_n(x) - Phi(x) | where Phi is the normal distribution
6 ## function mean 0
7 ## variance sigma, F_n is the empirical distribution function computed from A
8
9 library("mvtnorm")
10
10 pval <- function(dis, cv){
11   mean(cv>dis)
12 }
13
14 OptParam <- function(X,Y,poids,indice){
15   ## trouve les param optimaux pour indice = 1: modèle linéaire , 2 : modèle quadratique
16   W<- diag(1/abs(X)^poids)
17   xf <- matrix(rep(X,each = indice +1),ncol=indice+1,byrow=T)
18   ex <- matrix(rep(0:(indice),length(X)),ncol=indice +1 ,byrow=T)
19   Xp = xf^ex
20   matInv <- solve(t(Xp)%*%W%*%Xp)
21   param.optimaux <- rev(matInv%*%t(Xp)%*%W%*%Y)
22   return(param.optimaux)
23 }
```

```

24  predic <- function(param,x)
25  {
26    xf <- matrix(rep(x,each = length(param)),ncol=length(param),byrow=T)
27    ex <- matrix(rep(0:(length(param)-1),length(x)),ncol=length(param),byrow=T)
28    par <- matrix(rep(rev(param),length(x)),ncol=length(param),byrow=T)
29    Temp <- par * xf^ex
30    return (apply(Temp,1,sum))
31  }
32
33 CV <- function(don,indice,poids)
34  {
35    X = rep(don[,1],ncol(don[,-1]))
36    Y = c(don[,-1])
37    CVc = 0
38    corr <- (1/X^poids)
39    for(i in 1:length(Y))
40    {
41      p <- OptParam(X[-i],Y[-i],poids,indice)
42      pr <- (predic(p,X[i]) - Y[i])
43      CVc = CVc + (pr)^2 * corr[i]
44    }
45    return(CVc/sum(corr))
46  }
47
48 fit <- function(A,poids,b,nbgrille,stat,indice){
49  ## A : matrix of data, column 1 contains the covariates
50  ## poids : 1/X^poids i.e poids = 0 : no additional weight, poids =2 : inverted quadratique
51  ## weight
52  ## b : number of bootstrap replicates to use, suggestion is 1000
53  ## nbgrille : number of points used to approximate the statistic, suggestion is 50
54  ## stat : 1 is Kolmogorov smirnoff statistic, 2 is cramer von mises
55  ## indice : 1 is linear, 2 is quadratic 3 is cubic
56
57  don <- as.matrix(A)
58  nbr <- ncol(A)-1
59
60  W <- diag(rep(1/abs(don[,1])^poids,nbr)) # Weights
61
62  # Optimal parameters
63  X = rep(don[,1],ncol(don[,-1]))
64  Y = c(don[,-1])
65  W <- diag(1/abs(X)^poids)
66  xf <- matrix(rep(X,each = indice +1),ncol=indice+1,byrow=T)
67  ex <- matrix(rep(0:(indice),length(X)),ncol=indice +1 ,byrow=T)
68  Xp = xf^ex
69  meanXp <- apply(Xp,2,mean)
70  matInv <- solve(t(Xp)%*%W%*%Xp)
71  matt <- matInv%*%t(Xp)%*%W
72  param.optimaux <- rev(matt%*%Y)
73
74  #Residuals
75  poidsob <- 1/abs(X)^poids
76  poidsob <- poidsob/sum(poidsob)
    epsilonp <- (predic(param.optimaux,X) - Y) * sqrt(poidsob)

```

```

77 var.param <- matInv * var(epsilonp) ## variance of estimated residuals
78
79 ## Bootstrap
80 n <- length(epsilonp)
81 sigma <- sqrt(var(epsilonp))
82
83 ## Computation of the data process
84 grille = (1:(nbgrille))/(nbgrille+1) ## Approximation over the grid
85 grrep <- t(matrix(rep(grille,each=n),byrow=T,ncol=n))
86 epsilonrep <- matrix(rep(epsilonp,each=nbgrille),ncol=nbgrille,byrow=T)/sigma
87 Fn <- apply(pnorm(epsilonrep)<=grrep,2,mean)
88 monproc <- sqrt(n)*(Fn - grille)
89
90 proc=matrix(rep(0,b*nbgrille),ncol=b)
91 Xb = predic(param.optimaux,X);
92 for(i in 1:b)
93 {
94   ech_boot = sample(epsilonp,n,replace=T)
95   Yb = Xb + ech_boot / sqrt(poidsob)
96   param.optimauxb <- rev(matt%*%Yb)
97   epsilonpb <- (predic(param.optimauxb,X) - Yb) * sqrt(poidsob)
98   epsilonrepb <- matrix(rep(epsilonpb,each=nbgrille),ncol=nbgrille,byrow=T)/sqrt(var(
99     epsilonpb))
100  Fnb <- apply(pnorm(epsilonrepb)<=grrep,2,mean)
101  monprocb <- sqrt(n)*(Fnb - Fn)
102  proc[,i] = monprocb
103 }
104
105 ## Computation of the pvalue
106 distr <- rep(0,b)
107 mastat <- 0
108 if(stat==1){ distr <- apply(abs(proc),2,max)
109   mastat <- max(abs(monproc)) }
110 if(stat==2){ distr <- apply(proc^2,2,mean)
111   mastat<- mean(monproc^2) }
112 quantileb = quantile(distr,0.95)
113 pvalN = mean(distr>mastat)
114
115 liste <- list(param=param.optimaux,varparam=var.param,pvalN=pvalN,statN=mastat,qBoot=
116   quantileb,esp=epsilonp)
117 return(liste)
118 }
119
120 fitPlus <- function(A,poids,b,nbgrille,stat)
121 {
122   ## Performs the fit plus the partial F test
123   fitLin <- fit(A,poids,b,nbgrille,stat,1)
124   fitQ <- fit(A,poids,b,nbgrille,stat,2)
125
126   SSREGL <- sum(fitLin$esp^2)
127   SSREGQ <- sum(fitQ$esp^2)
128   degf <- length(fitQ$esp)-3
129   stat <- (SSREGL-SSREGQ)/(SSREGQ/(degf))
130   pvalF <- 1-pf(stat,1,degf)

```

```

129
130 L2 <- list(Linear= fitLin ,Quadratic = fitQ ,Ftest=stat ,pvalFtest=pvalF)
131 return(L2)
132 }

1 ## Code KS
2
3 KS <- function(A,nbgrille ,sigma){
4 # A : sample of data
5 # nbgrille : preision for the approximation of the grid
6 # variance to be used in the normal distribution function
7
8 ## Computes sqrt(n) * sup_x | F_n(x) - Phi(x) | where Phi is the normal distribution
9 # function mean 0
10 ## variance sigma , F_n is the empirical distribution function computed from A
11
12 n<- length(A)
13 fn <- function(t){sqrt(n) * abs(mean(A <= qnorm(t,0, sd=sqrt(sigma))) - t) }
14 max(sapply((1:nbgrille)/nbgrille ,fn))
15 }
16 pval <- function(dis ,cv){
17 mean(cv>dis)
18 }
19
20 # Linear fit using KS statistic
21 fit1KS <- function(A,poids ,b, nbgrille){
22 don <- as.matrix(A)
23 nbr <- ncol(A)-1
24
25 W <- diag(rep(1/abs(don[,1])^poids ,nbr)) # matrix of weights
26
27 # Finding the optimal parameters for the regression
28 x1 <- don[,1]
29 Xp <- matrix(c(rep(1,length(don[,1])*nbr) ,rep(x1,nbr)) ,ncol=2)
30 matInv <- solve(t(Xp)%*%W%*%Xp)
31 param.optimaux <- rev(matInv%*%t(Xp)%*%W%*%c(don[,-1]))
32
33 # Computing the residuals
34 temp <- param.optimaux[1]*don[,1]+ param.optimaux[2]
35 temp2 <- matrix(rep(temp ,nbr) ,nrow=length(don[,1]))
36 temp3 <- (don[,-1] - temp2)
37
38 poidsob <- 1/abs(don[,1])^poids
39 poidsob <- poidsob/sum(poidsob)
40 epsilonp <- c(temp3)*(sqrt(rep(poidsob ,nbr))) # residuals
41
42 var.param <- matInv * var(epsilonp) # the variance of the estimated paramters
43
44 ## Bootstrap
45 n <- length(epsilonp)
46 sigma <- sqrt(var(epsilonp))
47 distrCv <- rep(0,b)
48
49 for(j in 1:b){

```

```

50
51 B <- rnorm(n,0,sqrt(sigma))
52 C <- rmvnorm(1,mean=c(0,0),var.param)
53 drift <- (C[1] + C[2]*mean(don[,1])) # Drift function to used when esimated residuals are
54     involved
55 n<- length(A)
56 fn <- function(t){
57     sqrt(n) * abs(mean(A <= qnorm(t,0, sd=sqrt(sigma))) -
58         t + drift*dnorm(qnorm(t,0, sqrt(sigma)),0, sd=sqrt(sigma))/sqrt(n)) }
59 distrCv[j] <- max(sapply((1:nbgrille)/nbgrille,fn))
60
61 cv0 <- KS(epsilon,np,nbgrille,sigma)
62 pvalN <- pval(cv0,distrCv)
63
64 liste <- list(param=param.optimaux,res1=cbind(rep(poidsob,nbr),c(temp3)),epsp=epsilon,np,
65 varparam=var.param,pvalN=pvalN)
66 return(liste)
67
68 #quadratic fit using KS statistic
69 fit2KS <- function(A,poids,b,nbgrille){
70 don <- as.matrix(A)
71 nbr <- ncol(A)-1
72
73 W <- diag(rep(1/abs(don[,1])^poids,nbr)) # Weights
74
75 # Optimal parameters
76 x1 <- don[,1]
77 Xp <- matrix(c(rep(1,length(don[,1])*nbr),rep(x1,nbr),rep(x1^2,nbr)),ncol=3)
78 matInv <- solve(t(Xp)%*%W%*%Xp)
79 param.optimaux <- rev(matInv%*%t(Xp)%*%W%*%c(don,,-1)))
80
81 #Residuals
82 temp <- param.optimaux[1]*don[,1]^2 + param.optimaux[2]*don[,1] + param.optimaux[3]
83 temp2 <- matrix(rep(temp,nbr),nrow=nrow(don))
84 temp3 <- (don[,,-1] - temp2)
85
86 poidsob <- 1/abs(don[,1])^poids
87 poidsob <- poidsob/sum(poidsob)
88 epsilonnp <- c(temp3)*(sqrt(rep(poidsob,nbr)))
89 var.param <- matInv * var(epsilonnp)
90
91 ## Bootstrap
92 n <- length(epsilonnp)
93 sigma <- sqrt(var(epsilonnp))
94 distrCv <- rep(0,b)
95
96 for(j in 1:b){
97     B <- rnorm(n,0,sqrt(sigma))
98     C <- rmvnorm(1,mean=c(0,0,0),var.param)
99     drift <- (C[1] + C[2]*mean(don[,1]) + C[3]*mean(don[,1]^2)) # Drift function to used when
100      esimated residuals are involved
101     n<- length(A)
102     fn <- function(t){

```

```

101     sqrt(n) * abs(mean(A <= qnorm(t, 0, sd=sqrt(sigma))) -
102         t + drift*dnorm(qnorm(t, 0, sqrt(sigma)), 0, sd=sqrt(sigma))/sqrt(n)) }
103 distrCv[j] <- max(sapply((1:nbgrille)/nbgrille, fn))
104 }
105 cv0 <- KS(epsilon, nbgrille, sigma)
106 pvalN <- pval(cv0, distrCv)
107 pvalN <- pval(cv0, distrCv)
108
109 liste <- list(param=param.optimaux, varparam=var.param, pvalN=pvalN)
110 return(liste)
111 }
```

```

1 ## Code CVM
2 CVM <- function(A, nbgrille, sigma){
3 {
4 # A : sample of data
5 # nbgrille : precision for the approximation of the grid
6 # variance to be used in the normal distribution function
7
8 ## Computes sqrt(n) * \int (F_n(x) - Phi(x))^2 dx where Phi is the normal
9 ## distribution function mean 0
10 ## variance sigma, F_n is the empirical distribution function computed from A
11 cv =0
12 n <- length(A)
13 for(i in 1:nbgrille){
14   cv <- cv + n/nbgrille * (mean(A <= qnorm(i/nbgrille, mean=0, sd=sqrt(sigma))) - i/
15     nbgrille)^2
16 }
17 cv
18 }
19 pval <- function(dis, cv){
20   mean(cv>dis)
21 }
22
23 #Linear fit with CVM statistic
24 fit1CVM <- function(A, poids, b, nbgrille){
25 don <- as.matrix(A)
26 nbr <- ncol(A)-1
27 nbniv <- nrow(don)
28
29 # Weights
30 W <- diag(rep(1/abs(don[,1])^poids, nbr))
31 # Optimal parameters
32 x1 <- don[,1]
33 Xp <- matrix(c(rep(1, length(don[,1])*nbr), rep(x1, nbr)), ncol=2)
34 matInv <- solve(t(Xp)%*%W%*%Xp)
35 param.optimaux <- rev(matInv%*%t(Xp)%*%W%*%c(don, -1)))
36
37 #Residuals
38 temp <- param.optimaux[1]*don[,1]+ param.optimaux[2]
39 temp2 <- matrix(rep(temp, nbr), nrow=length(don[,1]))
40 temp3 <- (don[, -1] - temp2)
41 poidsob <- 1/abs(don[,1])^poids
```

```

42 poid sob <- poid sob /sum(poid sob)
43 epsilon p <- c(temp3)*(sqrt(rep(poid sob ,nbr)))
44
45 var . param <- matInv * var(epsilon p)
46
47 ## Bootstrap
48 n <- length(epsilon p)
49 sigma <- sqrt(var(epsilon p))
50 distrCv <- rep(0,b)
51
52 for(j in 1:b){
53
54 B <- rnorm(n,0,sqrt(sigma))
55 C <- rmvnorm(1,mean=c(0,0),var.param)
56 drift <- (C[1] + C[2]*mean(don[,1])) # Drift function to used when esimated residuals are
      involved
57 cv <- 0
58 for(i in 1:nbgrille){
59   cv <- cv + n/nbgrille * (mean(B <= qnorm(i/nbgrille,mean=0,sd=sqrt(sigma))) - i/
      nbgrille + drift*dnorm(qnorm(i/nbgrille,0,sqrt(sigma)),0,sd=sqrt(sigma))/sqrt(n))^2
60 }
61 distrCv[j] <- cv
62 }
63 cv0 <- CVM(epsilon p,nbgrille,sigma)
64 pvalN <- pval(cv0,distrCv)
65
66 liste <- list(param=param.optimaux,varparam=var.param,pvalN=pvalN)
67 return(liste)
68 }
69
70 # Quadratic fit statistic CVM
71 fit2CVM <- function(A,poids,b,nbgrille){
72 don <- as.matrix(A)
73 nbr <- ncol(A)-1
74 nbniv <- nrow(A)
75
76
77 #Weights
78 W <- diag(rep(1/abs(don[,1])^poids,nbr))
79 #Optimal paramters
80 x1 <- don[,1]
81 Xp <- matrix(c(rep(1,length(don[,1])*nbr),rep(x1,nbr),rep(x1^2,nbr)),ncol=3)
82 matInv <- solve(t(Xp)%*%W%*%Xp)
83 param.optimaux <- rev(matInv%*%t(Xp)%*%W%*%c(don,,-1)))
84
85
86 #Residuals
87 temp <- param.optimaux[1]*don[,1]^2 + param.optimaux[2]*don[,1] + param.optimaux[3]
88 temp2 <- matrix(rep(temp,nbr),nrow=nbniv)
89 temp3 <- (don,,-1] - temp2)
90
91 poid sob <- 1/abs(don[,1])^poids
92 poid sob <- poid sob /sum(poid sob)
93 epsilon p <- c(temp3)*(sqrt(rep(poid sob ,nbr)))

```

```

94 var.param <- matInv * var(epsilonp)
95
96 ## Bootstrap
97 n <- length(epsilonp)
98 sigma <- sqrt(var(epsilonp))
99 distrCv <- rep(0,b)
100
101 for(j in 1:b){
102
103   B <- rnorm(n,0,sqrt(sigma))
104   C <- rmvnorm(1,mean=c(0,0,0),var=var.param)
105   drift <- (C[1] + C[2]*mean(don[,1]) + C[3]*mean(don[,1]^2)) # Drift function to used when
106   # esimated residuals are involved
107   cv <- 0
108   for(i in 1:nbgrille){
109     cv <- cv + n/nbgrille * (mean(B <= qnorm(i/nbgrille,mean=0,sd=sqrt(sigma))) - i/
110     nbgrille + drift*dnorm(qnorm(i/nbgrille,0,sqrt(sigma)),0,sd=sqrt(sigma))/sqrt(n))^2
111   }
112   distrCv[j] <- cv
113 }
114
115 cv0 <- CVM(epsilonp,nbgrille,sigma)
116 pvalN <- pval(cv0,distrCv)
117
118 liste <- list(param=param.optimaux,varparam=var.param,pvalN=pvalN)
119 return(liste)
120
121 }
```