

A Appendix: simulation code and model files

A.1 R code to simulate data

```
1 library(LaplacesDemon)
2 library(MASS)
3
4 #####
5 ##### function to simulate survival time #####
6 #####
7 # survival function is given by:  $S(t) = \exp(-\exp(B) * (\exp(A*t) - 1)) / A$ 
8 sim_Ti = function(n=500, alpha, delta=c(1,1), gamma=c(1,1)){
9   Time = numeric(n)
10  S = runif(n) # survival probability
11  H = matrix(rnorm(2*n), ncol=2)
12  W = matrix(rnorm(2*n), ncol=2)
13  # random effects
14  U = mvrnorm(n, mu=c(0,0), Sigma=matrix(c(0.09, 0.09*0.16, 0.09*0.16, 0.09),
15    nrow=2, byrow=T))
16  attributes(U)[[2]]=NULL # remove 'dimnames' attribute
17
18  # calculate survival time
19  if(alpha[1]==0 & alpha[2]==0) Time = - log(S) / exp(gamma %*% t(W))
20
21  else{
22    B = alpha[1] * delta[1] * H[,1] + alpha[2] * U[,1] + gamma %*% t(W)
23    A = alpha[2] * U[,2] + alpha[1] * delta[2] * H[,2]
24    Time = log(1-log(S)*A/exp(B))/A
25  }
26
27  Ti_id = which(!is.na(Time))
28  Time = Time[Ti_id][1:250] # true survival time: take the first 250 that are
29    not NA
30  Ci = rbeta(250, 4, 1)*2 # censoring time
31  Ti = pmin(Time, Ci) # observed survival time: choose the smaller one
32  event = as.numeric(Time == Ti) # 1 for event, 0 for censor
33  U = U[Ti_id, ][1:250, ]
34  H = H[Ti_id, ][1:250, ]
35  W = W[Ti_id, ][1:250, ]
36
37  list(Ti=Ti, event=event, H=H, U=U, W=W)
38 }
39
40 #####
41 ##### function to simulate longitudinal data #####
42 #####
43 sim_longitudinal_data = function(survival_data=surdata, n=250, time=c(0, 0.25,
44   0.5, 0.75, 1, 3), tau, sigma=1, beta=c(1,1), delta=c(1,1)){
45   # survival_data - data simulated from survival model
46   # n - # of subjects
47   # time - time points of observations
```

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45 # tau - quantile
46 # sigma - scale parameter for ALD
47 time = time # at most # = length(time) observations per patient
48 y = matrix(NA, nrow=n, ncol=length(time)) # wide format
49 Ti = survival_data$Ti
50 U = survival_data$U # random effects
51 H = survival_data$H
52 X = cbind(1, rnorm(n))
53 count = sapply(Ti, function(x) sum(x > time)) # number of observations after
    drop-outs
54
55 for (i in 1:n){
56   for (j in 1:count[i]){
57     location = beta %*% X[i, ] + delta %*% c(H[i,1], H[i,2]*time[j]) + U[i,]
58     %*% c(1, time[j])
59     y[i,j] = rlaplace(1, location, scale=sigma, kappa=tau)
60   }
61 }
62 list(y = y, X = X, J=count)
63 }
64
65 #####
66 ##### function to simulate multiple joint data sets #####
67 #####
68 sim_multiple_data = function(N, sur_fun=sim_Ti, longi_fun=sim_longitudinal_
    data, alpha, tau){
69   # N - number of data sets to generate
70   # sur_fun - function to simulate survival data
71   # longi_fun - function to simulate longitudinal data
72   # alpha - association parameters for JM
73   # tau - quantile
74
75   outdata = vector(mode='list', N)
76   for (i in 1:N){
77     sur_data = sur_fun(alpha=alpha)
78     longi_data = longi_fun(sur_data, tau=tau)
79     outdata[[i]] = list(survival_data=sur_data, longitudinal_data=longi_data)
80   }
81   outdata
82 }

```

A.2 JAGS model file

```

1 model{
2   zero[1] <- 0
3   zero[2] <- 0
4   k1 <- (1-2*qt)/(qt*(1-qt))
5   k2 <- 2/(qt*(1-qt))
6
7   for (i in 1:I){
8     # prior for random effects

```

```

9   u[i, 1:2] ~ dmnorm(zero[], precision[,])
10
11  # longitudinal process, BQR mixed model using ALD representation
12  for (j in 1:J[i]){
13    er[i,j] ~ dexp(sigma)
14    mu[i,j] <- u[i,1] + u[i,2]*t[j] + inprod(X[i,], beta[]) + delta[1]*H[i
,1] + delta[2]*H[i,2]*t[j] + k1*er[i,j]
15    prec[i,j] <- sigma/(k2*er[i,j])
16    y[i,j] ~ dnorm(mu[i,j], prec[i,j])
17  } #end of j loop
18
19  # survival process, baseline hazard is set to 1
20  A[i] <- alpha2*u[i,2] + alpha1*delta[2]*H[i,2]
21  B[i] <- alpha1*delta[1]*H[i,1] + alpha2*u[i,1] + inprod(gamma, W[i,])
22  S[i] <- exp(- exp(B[i])*(pow(exp(A[i]), Ti[i])-1)/A[i])
23  h[i] <- exp(inprod(gamma, W[i,]) + alpha1*(delta[1]*H[i,1] + delta[2]*H[i
,2]*Ti[i]) + alpha2*(u[i,1] + u[i,2]*Ti[i]))
24  L[i] <- pow(h[i], event[i])*S[i]/1.0E+08
25
26  # zero trick
27  phi[i] <- -log(L[i])
28  zeros[i] ~ dpois(phi[i])
29
30  }#end of i loop
31
32  precision[1:2,1:2] <- inverse(Sigma[,])
33  Sigma[1,1] <- 1
34  Sigma[1,2] <- rho*sig1
35  Sigma[2,1] <- Sigma[1,2]
36  Sigma[2,2] <- sig1*sig1
37
38  # priors for other parameters
39  alpha1 ~ dnorm(0, 0.001)
40  alpha2 ~ dnorm(0, 0.001)
41  beta[1] ~ dnorm(0, 0.001)
42  beta[2] ~ dnorm(0, 0.001)
43  delta[1] ~ dnorm(0, 0.001)
44  delta[2] ~ dnorm(0, 0.001)
45  gamma[1] ~ dnorm(0, 0.001)
46  gamma[2] ~ dnorm(0, 0.001)
47  sigma ~ dgamma(0.001, 0.001)
48  rho ~ dunif(-1, 1)
49  sig1 ~ dgamma(0.01, 0.01)
50 }

```