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#### Supporting Information: WinBUGS code for label-invariant models

#### Univariable model for the influence of a single study-level characteristic

#### Data required:

```
N study=no. of studies
        N=no. of studies x 2 arms
        N_ma=no. of meta-analyses
Study-level:
        treat=1 if treatment arm, 0 if control
        r=no. of events
        n= no. of participants
        C1[i] = 1 if the ith study has the characteristic of interest, 0 otherwise
        ma[i]<- meta-analysis index
        s[i]<-study index
Meta-analysis level:
        kappa_ok[m] = 1 if there are at least two studies with, and two studies without, the
        characteristic in the mth meta-analysis, 0 otherwise
The WinBUGS model
model{
for (i in 1:N) {
        r[i] ~ dbin(p[i],n[i]) # likelihood (binary data r/n=no.of events/no. of participants)
        logit(p[i]) <- alpha[s[i]] + treat[i]*(theta.minus[i]*+ beta[i] *C1[i])</pre>
                # model (treat indicator of treatment group; C1 indicator of study characteristic)
        beta[i]~dnorm(mean[ma[i]],p.k2[ma[i]])I(-10,10)
                # between study, within meta-analysis, variation in difference in effect associated
with characteristic
theta.minus[i]~dnorm(d[ma[i]],p.d[ma[i]])I(-10,10) #RE for intervention effect within meta-analysis
        rhat[i] <- p[i] * n[i]
                                 #calculate residual deviance
```

```
dev[i] < 2 * (r[i] * (log(r[i])-log(rhat[i])) + (n[i]-r[i]) * (log(n[i]-r[i]) - log(n[i]-rhat[i])))
}
```

```
resdev <- sum(dev[])
```

```
for (j in 1:N_study) {alpha[j] ~ dnorm(0,.001)}
```

# priors for study baseline effects - unrelated

```
for (m in 1:N_ma) {
       mean[m] <- d[m] + b[m]
       d[m] ~ dnorm(0,.001)
                                       # priors for true fixed (unrelated) intervention effects
       b[m] \sim dnorm(b0, p.phi)
                                       # between meta-analysis variation in average difference in
effect associated with characteristic
       var d[m]~dlnorm(mu,p.tau)
                                       # log-normal distribution for between-study variances
       p.d[m] <- 1/var d[m]
       p.k2[m] <- equals(kappa ok[m],1)/(var d[m]*lambda)
               +equals(kappa_ok[m],0)/(var_d[m]*cut(lambda))
       }
#Prior for unknown parameters
b0 \sim dnorm(0,.001)
                       # vague prior for overall average difference in effect associated with
characteristic
```

lambda ~dlnorm(0,1) # vague prior for change in between-study variation associated with characteristic

p.phi1 ~ dgamma(.001,.001) # vague prior for between meta-analysis variation in average difference in effect associated with characteristic phi <- pow(p.phi,-0.5) p.phi <- p.phi1/(1-patom.phi) patom.phi ~ dbeta(1,1) mu~dnorm(0,0.001) p.tau<-1/(sd.tau\*sd.tau) sd.tau~dunif(0,2)

log.tau2.new~dlnorm(mu,p.tau) # predictive distn for heterogeneity among studies without the characteristic tau2.new<-exp(log.tau2.new)

# Parameters to monitor q[1] <- b0 q[2] <- exp(b0) q[3] <- lambda q[4] <- phi q[5]<- log.tau2.new q[6]<-tau2.new</pre>

}

Multivariable model for the influence of three study-level characteristics

#### Data required:

N\_study=no. of studies N=no. of studies x 2 arms N\_ma=no. of meta-analyses

Study-level:

treat=1 if treatment arm, 0 if control
r=no. of events
n= no. of participants
C1[i,j] = 1 if the *i*th study has the *j*th characteristic, 0 otherwise
ma[i]<- meta-analysis index
s[i]<-study index</pre>

Meta-analysis level:

kappa\_ok[m,j] = 1 if there are at least two studies with, and two studies without, the *j*th characteristic in the *m*th meta-analysis, 0 otherwise

clambda[m,j] = 1 if there are 1, K-1 or K studies with or without the *j*th characteristic in the *m*th meta-analysis, 0 otherwise, where K is the no. of studies in the meta-analysis. C0[m,j]=1 if there are no studies in the *m*th meta-analysis with the *j*th characteristic

#### The WinBUGS model

```
model {
for (i in 1:N) {
    r[i] ~ dbin(p[i],n[i]) # likelihood (binary data r/n=no.of events/no. of participants)
    logit(p[i]) <- alpha[s[i]] + theta[i]*treat[i]
```

theta[i] < theta.minus[i]\* (1-C1[i,1])\*(1-C1[i,2])\*(1-C1[i,3]) #effect in study without any of the

characteristics
 +theta.plus[i]\* (1-(1-C1[i,1]) \*(1-C1[i,2]) \*(1-C1[i,3])) effect in studies with
one or more characteristics

theta.plus[i] ~ dnorm(mean[i], p.k2[i])I(-10,10)

theta.minus[i]~dnorm(d[ma[i]],p.d[ma[i]])I(-10,10) #RE for treatment effects within meta-analysis

```
mean[i]<-d[ma[i]]+b[ma[i],1]*C1[i,1]+b[ma[i],2]*C1[i,2]+b[ma[i],3]*C1[i,3]
```

k2[i]<- ((1-C1[i,1]) + # without characteristic 1

C1[i,1] \* kappa\_ok[ma[i],1] \* lambda[1] + # with characteristic 1 and inform C1[i,1] \* clambda[ma[i],1] \* cut(lambda[1]) + # with characteristic 1 but don't inform C1[i,1] \* CO[ma[i],1] \* 1) \* # no studies have characteristic 1 in the MA ( (1-C1[i,2]) + # without characteristic 2 C1[i,2] \* kappa\_ok[ma[i],2] \* lambda[2] + # with characteristic 2 and inform C1[i,2] \* clambda[ma[i],2] \* cut(lambda[2]) + # with characteristic 2 but don't inform C1[i,2] \* CO[ma[i],2] \* 1) \*# no studies have characteristic 2 in the MA ( (1-C1[i,3]) + # without characteristic 3 C1[i,3] \* kappa\_ok[ma[i],3] \* lambda[3] + # with characteristic 3 and inform C1[i,3] \* clambda[ma[i],3] \* cut(lambda[3]) + # with characteristic 3 but don't inform C1[i,3] \* C0[ma[i],3] \* 1)\* # no studies have characteristic 1 in the MA var d[ma[i]]

p.k2[i]<-1/k2[i]

rhat[i] <- p[i] \* n[i] #calculate residual deviance dev[i] <- 2 \* (r[i] \* (log(r[i])-log(rhat[i])) + (n[i]-r[i]) \* (log(n[i]-r[i]) - log(n[i]-rhat[i])))

}

```
resdev <- sum(dev[])
```

for (j in 1:N\_study) {alpha[j] ~ dnorm(0,.01)} # priors for study baseline effects - unrelated

```
for(m in 1:N_ma){
```

```
d[m] ~ dnorm(0,0.01) # priors for true fixed (unrelated) intervention effects
    for(j in 1:3){b[m,j] ~ dnorm(b0[j], p.phi[j])} # between meta-analysis variation in average
difference in effect associated with characteristic
```

```
var_d[m]~dlnorm(mu,p.tau) # log-normal distribution for between-study variances
p.d[m] <- 1/var_d[m]
```

```
}
```

#Prior for unknown parameters

```
for(j in 1:3){
```

b0[j] ~dnorm(0,0.001) # vague prior for overall average difference in effect associated with characteristic

 $lambda[j] \sim dlnorm(0,0.1) \text{ # vague prior for change in between-study variation associated with characteristic j} p.phi1[j] \sim dgamma(0.001, 0.001)$ 

```
phi[j] <- pow(p.phi[j],-0.5)
```

p.phi[j] <-p.phi1[j]/(1-patom.phi[j]) # vague prior for between meta-analysis variation in average difference in effect associated with characteristic

```
patom.phi[j] ~ dbeta(1,1)
```

# exp.b0[j]<-exp(b0[j]) }

mu~dnorm(0,0.01) p.tau<-1/(sd\_tau\*sd\_tau) sd\_tau~dunif(0,2)

log.tau2.new~dlnorm(mu,p.tau) tau2.new<-exp(log.tau2.new)

# predictive distn for heterogeneity among studies without the characteristics

}