

Additional file 1 - metafor syntax for GLMM in R

In *metafor*, the NCHGN model is specified as

```
rma.glmm(measure = "OR", ai =, bi =, ci =, di =, data =, model = "CM.EL").
```

To use the “dnoncenhypergeom” function for noncentral-hypergeometric distribution or a specific optimizer such as “optim”, “bobyqa”, “newuoa”, or “uobyqa”, the model should be specified as

```
rma.glmm(measure = "OR", ai =, bi =, ci =, di =,
          model = "CM.EL", control = list(dnchgc = "dnoncenhypergeom"))
```

and

```
rma.glmm(measure = "OR", ai =, bi =, ci =, di =,
          model = "CM.EL", control = list(optimizer = "optim"))
```

where ai , bi , ci , di are the binomial data from the table

	Event	No event	Total
Treatment	ai	bi	$ai + bi$
Control	ci	di	$ci + di$
Total	$ai + ci$	$bi + di$	$ai + bi + ci + di$

The two methods (“dFNCHypergeo” and “dnoncenhypergeom”) should perform similarly, when fitting the conditional GLMM (exact likelihood). However, convergence problems might occur when trying to fit a saturated model. Switching to an alternative method can help to solve the problem.

With small numbers of events, the binomial-normal approximation may be used. Then the model is a conditional GLMM (approximate likelihood). This model is specified as

```
rma.glmm(measure = "OR", ai =, bi =, ci =, di =, data =, model = "CM.AL").
```