

# The mcemGLM package

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## 1 A Generalized Linear Mixed Model

We start by assuming that we observe a vector of data  $Y = (Y_1, \dots, Y_n)$  corresponding to a probability model that depends on a  $(p + l)$ -dimensional parameter vector  $\theta$ , a known  $n \times p$  fixed effects design matrix  $X$ , a known  $n \times k$  known random effects design matrix  $Z$ , and a  $k$ -dimensional vector of unobservable random effects  $U$ .

Let  $\theta$  consist of  $p$  fixed effects coefficients  $\beta = (\beta_1, \dots, \beta_p)^T$  and  $l$  variance parameters,  $\sigma^2 = (\sigma_1^2, \dots, \sigma_l^2)^T$ , associated to the random effects  $U$ . Our goal is to find maximum likelihood estimates (MLEs) for the  $(p + l)$ -dimensional parameter  $\theta$  in a space  $\Theta$  for a generalized linear mixed model.

We assume that the expected variable of  $Y_i$ , can be written as a linear combination of the observable and unobservable variables through a bijective “link” function  $g$ . Let  $X_i$  and  $Z_i$  be the  $i$ th rows of the matrices  $X$  and  $Z$ , and let  $E(Y_i|U = u) = \mu_i$ . Then

$$g(\mu_i) = X_i \beta + Z_i u, \text{ for } i = 1, \dots, n.$$

Let  $U = (U_1^T, \dots, U_l^T)^T$ , and  $Z = (Z_1 \cdots Z_l)$  a decomposition for the vector  $U$  and the matrix  $Z$ . We assume that  $U_i$  is a  $k_i$ -dimensional vector with  $\sum_i^l k_i = k$ . Furthermore we assume that  $U_i$  has a known distribution with variance that depends on the parameter  $\sigma_i^2$ . In general let  $\mu = (\mu_1, \dots, \mu_n)$  and let  $g(\mu)$  denote the element-wise evaluation of  $g$  on the vector  $\mu$ , then we can write mean our

model as

$$g(\mu) = X\beta + \sum_{i=1}^l Z_i u_i. \quad (1)$$

Let  $h_U(u)$  be the probability density function of  $U$ . We assume that conditional on  $U$ , the data is generated from a probability model with probability mass function  $f(Y|\theta, X, Z, U)$  and that we can write its likelihood function in terms of  $\mu = g^{-1}(X\beta + \sum_{i=1}^l Z_i u_i)$ , and  $\sigma^2$ . Defining the model this way yields to the following likelihood functions:

1. A complete likelihood function:

$$L(\theta|X, Z, U) = f(y, u|\theta, X, Z) = f_Y(y|\theta, X, Z, u) h_U(u|\theta). \quad (2)$$

2. And a marginal likelihood function:

$$L(\theta|X, Z) = \int_{\mathbb{R}^l} f(Y|\theta, X, Z, U) h_U(u|\theta) du. \quad (3)$$

Since the vector  $U$  is not observable we need to obtain the MLEs from 3. This means that before maximizing the likelihood function we need to integrate out the vector of random effects.

The `mcemGLM` package focuses on three types of models for the marginal data:

1. Bernoulli data. We say that  $Y_i \stackrel{iid}{\sim} \text{Bernoulli}(p_i)$ , for  $i = 1, \dots, n$ , with  $0 < p_i < 1$ , if  $Y_i$  has probability mass function

$$f(y_i) = p_i^{y_i} (1 - p_i)^{1-y_i}, \text{ for } y_i = 0, 1.$$

With  $E(Y_i) = p_i$ ,  $\text{Var}(Y_i) = p_i(1 - p_i)$ , and  $g(p_i) = \log(p_i/(1 - p_i))$ .

2. Poisson data. We say that  $Y_i \stackrel{iid}{\sim} \text{Poisson}(\mu_i)$  for  $i = 1, \dots, n$ , if  $Y_i$  has probability mass function

$$f(y_i) = e^{-\mu_i} \frac{\mu_i^{y_i}}{y_i!}, \text{ for } y_i = 0, 1, 2, \dots$$

With  $E(Y_i) = \mu_i$ ,  $\text{Var}(Y_i) = \mu_i$ , and  $g(\mu_i) = \log(\mu_i)$ .

3. Negative binomial data. We say that  $Y_i \stackrel{iid}{\sim} \text{neg-binom}(\mu_i, \alpha)$ , for  $i = 1, \dots, n$ , with  $\mu_i > 0$ , and  $\alpha > 0$ , if  $Y_i$  has probability mass function

$$f(y_i) = \frac{\Gamma(y_i + \alpha)}{\Gamma(\alpha) y_i!} \left( \frac{\alpha}{\mu_i + \alpha} \right)^\alpha \left( \frac{\mu_i}{\mu_i + \alpha} \right)^{y_i}, \text{ for } y_i = 0, 1, 2, \dots$$

With  $E(Y_i) = \mu_i$ ,  $\text{Var}(Y_i) = \mu_i + \mu_i^2/\alpha$ , and  $g(\mu_i) = \log(\mu_i)$ .

The expectation and variance of  $Y_i$  can be found easily by using iterated expectation with respect to a random variable  $M$  distributed gamma with shape parameter  $\alpha$ , and rate parameter  $\alpha/\mu$  and setting  $Y_i|M = m \sim \text{Poisson}(m)$ .

By using this definition of the distribution of  $Y_i$  we can treat the parameter  $\alpha$  as the amount of over-dispersion with respect to the Poisson distribution. The value  $\alpha = \infty$  corresponds to no over-dispersion.

By introducing  $\alpha$  to the model notice that we need to estimate this extra parameter in addition to  $\beta$  and  $\sigma^2$ .

In addition to the model selection the `mcemGLM` package allows to specify two types of random effects. Let  $I_k$  be an  $n \times n$  identity matrix,  $N_n(\mu, \Sigma)$  an  $n$ -dimensional multivariate normal distribution with mean vector  $\mu$  and covariance matrix  $\Sigma$ , and  $t_n(\nu, \mu, \Sigma)$  an  $n$ -dimensional multivariate  $t$  distribution with  $\nu$  degrees of freedom, location vector  $\mu$  and scale matrix  $\Sigma$ .

1. Normal distribution. We set  $U_i \sim N_{k_i}(0, \sigma_i^2 I_{k_i})$  for  $i = 1, \dots, l$ , and set the  $U_i$ s to be jointly independent.
2.  $t$  distribution with known degrees of freedom  $\nu$ . We set  $U_i \sim t_{k_i}(\nu, 0, \sigma_i^2 I_{k_i})$  for  $i = 1, \dots, l$ , and set the  $U_i$ s to be jointly independent.

## 2 The MCEM algorithm

The MCEM algorithm is a modification of the EM algorithm. The later assumes two sets of data an observed data set  $Y$  and a set of missing data  $U$ .

The EM algorithm estimates the MLEs by an iterative algorithm. Let  $\theta^{(t)}$  denote the current

estimate at the  $i$ th iteration. Let

$$Q(\theta, \theta^{(t)}) = \mathbb{E} \left[ \log f(y, u|\theta, X, U) | y, \theta^{(t)} \right]. \quad (4)$$

The next value,  $\theta^{(t+1)}$ , is found by maximizing 4 with respect to  $\theta$ . The expectation in 4 is taken with respect to  $f(u|y, \theta, X, Z)$  hence if we want to obtain its closed form we need  $f(y, u|\theta, X, Z)$  and  $f_Y(y|\theta, X, Z)$ . The later function is not available for the models we are considering, therefore we need to resort to a numerical method to calculate this expectation.

The solution implemented in the `mcemGLM` package is to estimate 4 via a Markov chain Monte Carlo (MCMC) step. This works by obtaining a sample  $u_{t,1}, \dots, u_{t,m}$  from a Markov chain with stationary distribution  $f(u|y, \theta, X, Z)$  and then maximizing

$$\hat{Q}(\theta) = \sum_{j=1}^m \log f(y, u_{t,j}|\theta, X, Z) \quad (5)$$

with respect to  $\theta$  to obtain  $\theta^{(t+1)}$ .

The algorithm is run until a termination condition has been reached or the maximum number of iterations has been done.

### 3 The `mcemGLM` package

The package runs through the following steps:

1. Choose a starting value. The default method is to fit a model without random effects and using the MLEs of the fixed coefficients as starting values for  $\beta$ . For  $\sigma$  we set a predefined value of 5.
2. Obtain the sample  $u_{t,1}, \dots, u_{t,m}$ . This is done by using a Metropolis–Hastings algorithm that uses a multivariate normal random variable as its proposal. The standard deviation vector of the proposal distribution is chosen by performing an auto-tuning step before the first iteration. After each iteration the rejection rate of the chain is checked and if it is either too large ( $> 0.4$ ) or too small ( $< 0.1$ ) the package performs an auto-tuning step before the next iteration.
3. After obtaining the sample 5 is maximized with respect all the parameters using the `trust` function from the `trust` package. The maximizers are set as the current value of the estimator

of the MLEs.

4. Steps 2 and 3 are repeated until the condition

$$\max_i \left\{ \frac{|\theta_i^{(t)} - \theta_i^{(t-1)}|}{|\theta_i^{(t)}| + \delta} \right\} < \epsilon$$

for specified values of  $\delta$  and  $\epsilon$  is met three consecutive times or a maximum number of iterations have been performed.

The default values in the package are  $\delta = 0.05$  and  $\epsilon = 0.01$  but these can be easily changed by the user. The default number of iterations is 80 and this value can also be changed by the user.

5. After terminating the iterative process another sample from the random effects is obtained to estimate the information matrix of the model.

## 4 Using the mcmGLM package

```
> require(mcmGLM)
```

```
> data("simData.rdata")
```

```
> head(simData)
```

	obs	x1	x2	x3	z1	z2	z3	count	count2
1	0	9.571463	5.924451	red	D1	1	A	5	5
2	0	10.062451	5.358087	yellow	D1	1	A	4	24
3	0	8.020461	4.755584	yellow	D1	1	A	4	4
4	0	10.842312	5.610179	yellow	D1	1	A	3	9
5	0	8.457872	2.882771	yellow	D1	1	A	3	6
6	0	11.154501	4.942196	yellow	D1	1	A	3	6

```
> summary(simData)
```

	obs	x1	x2	x3	z1	z2
Min.	:0.00	Min. : 7.338	Min. :1.784	blue :65	D1:40	1:50
1st Qu.:	0.00	1st Qu.: 9.332	1st Qu.:4.405	red :67	D2:40	2:50

Median	:0.00	Median	: 9.995	Median	:5.119	yellow:68	D3:40	3:50
Mean	:0.39	Mean	: 9.990	Mean	:5.148		D4:40	4:50
3rd Qu.	:1.00	3rd Qu.	:10.629	3rd Qu.	:5.808		D5:40	
Max.	:1.00	Max.	:13.533	Max.	:7.531			
z3		count		count2				
A:100	Min.	: 0.0	Min.	: 0.00				
B: 60	1st Qu.	: 3.0	1st Qu.	: 6.00				
D: 40	Median	: 4.5	Median	: 12.00				
	Mean	: 5.1	Mean	: 16.16				
	3rd Qu.	: 7.0	3rd Qu.	: 20.00				
	Max.	:18.0	Max.	:108.00				

The data consist of three fixed effects,  $x1$ ,  $x2$ , and  $x3$ . The first two fixed effects are continuous and  $x3$  is a factor with three levels. There are three variables that we can use as variance components  $z1$  (5 levels),  $z2$  (4 levels), and  $z3$  (3 levels.) The component  $z2$  can be nested within  $z1$  and  $z3$  is crossed with these.

First we will consider a simple model based on this data using `obs` as the binary response.

## 4.1 A simple model

We will fit a model with one variance component,  $z3$  and we will consider  $z1$  as a fixed effect along with  $x1$ .

The main model arguments for the `mcemGLMM` function are `fixed` and `random`. These specify the fixed and random effects of the model. The response must be included in the `fixed` argument. In this first example we are considering  $x1$  and  $z1$  as fixed and  $z3$  as random. We can fit this model with the following command:

```
> fit0 <- mcemGLMM(fixed = obs ~ x1 + z1,
+                  random = ~0+z2,
+                  data = simData,
+                  family = "bernoulli",
+                  vcDist = "normal")
```

The rest of the used arguments are:

- **data**: argument contains the name of the data frame with the data.
- **family**: argument specifies the type of model to be fitted. The options are “bernoulli” for logistic regression, “poisson” for Poisson count regression, and “negbinom” for negative binomial count regression.
- **vcDist**: argument specifies the distribution of the random effects. The option are “normal”, and “t”. In case of  $t$  random effects an extra argument with the degrees of freedom must be supplied.

We can start by taking a look at the coefficient and variance estimates with the `summary` command:

```
> summary(fit0)
```

Call:

```
mcmGLMM(fixed = obs ~ x1 + z1, random = ~0 + z2, data = simData,  
family = "bernoulli", vcDist = "normal")
```

Two sided Wald tests for fixed effects coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	2.3683592	1.6681565	1.4197464	0.15568152
x1	-0.3368267	0.1640034	-2.0537785	0.03999714
z1D2	1.3871167	0.4908233	2.8261021	0.00471182
z1D3	-1.0466774	0.6047379	-1.7307951	0.08348832
z1D4	0.4811148	0.4944886	0.9729543	0.33057603
z1D5	1.3291029	0.4907177	2.7084876	0.00675906

One sided Wald tests for variance components:

	Estimate	Std. Error	z value	Pr(>z)
z2	0.1080782	0.1508095	0.7166536	0.2367939

We first get a print of the original call used to fit the model. The summary print out has two tables. The first table shows the estimates, standard errors and  $z$  tests for the fixed effect coefficients. While the second table contains the same information but for the variance estimates.

Now we can look at an ANOVA table based on Wald tests.

```
> anova(fit0)
```

```
Wald's Chi-squared ANOVA table
```

	Df	Wald Stat	Pr(>W)
x1	1	4.218006	0.03999714
z1	4	24.562532	0.00006160

Each line corresponds to a test on the coefficients that are related to each variable. In the case of a continuous variable or a binary this is equivalent to the  $z$  test performed with `summary`. When a categorical variable has more than two categories `anova` will test run a chi-squared test on all the coefficients that are related to that variable. In this case the chi-squared test for  $z1$  tests if the corresponding coefficients for D2, D3, D4, and D5 are both equal to zero.

We can run multiple comparison tests for the levels of  $z1$ . First we need to create a contrast matrix with each row representing a contrast that we want to test. In this case

```
> ctr0 <- rbind("D1 - D2" = c(0, 0, -1, 0, 0, 0),
+               "D1 - D3" = c(0, 0, 0, -1, 0, 0),
+               "D1 - D4" = c(0, 0, 0, 0, -1, 0),
+               "D1 - D5" = c(0, 0, 0, 0, 0, -1),
+               "D2 - D3" = c(0, 0, 1, -1, 0, 0),
+               "D2 - D4" = c(0, 0, 1, 0, -1, 0),
+               "D2 - D5" = c(0, 0, 1, 0, 0, -1),
+               "D3 - D4" = c(0, 0, 0, 1, -1, 0),
+               "D3 - D5" = c(0, 0, 0, 1, 0, -1),
+               "D4 - D5" = c(0, 0, 0, 0, 1, -1))
```

Notice that rows one and two are the contrasts that compare the baseline, D1, to the other levels, hence these will have equivalent test statistics as those obtained in `summary`. However `contrasts.mcemGLMM`



accounts for multiple comparisons by adjusting the  $p$ -values via Bonferroni correction therefore it is possible to obtain significance in `summary` and not in `contrasts.mcemGLMM` since this  $p$ -value will likely be larger.

```
> contrasts.mcemGLMM(object = fit0, ctr.mat = ctr0)
```

	Estimate	Std. Err.	Wald	Adj. p-value
D1 - D2	-1.38711666	0.4908233	7.98685283	0.0471182493
D1 - D3	1.04667741	0.6047379	2.99565159	0.8348832049
D1 - D4	-0.48111479	0.4944886	0.94664003	1.0000000000
D1 - D5	-1.32910289	0.4907177	7.33590494	0.0675906430
D2 - D3	2.43379408	0.5883939	17.10926441	0.0003528974
D2 - D4	0.90600187	0.4715595	3.69134908	0.5469535980
D2 - D5	0.05801377	0.4654850	0.01553284	1.0000000000
D3 - D4	-1.52779221	0.5909675	6.68346045	0.0973115299
D3 - D5	-2.37578031	0.5894285	16.24614167	0.0005562275
D4 - D5	-0.84798810	0.4707865	3.24437836	0.7166887291

For this simple model it is possible to plot the predicted probabilities for each level of `z1`. These estimates correspond to the population means, i.e., the random effects have been set to zero. Figure 4.1 shows the plots of the fitted probabilities as a function of `x1` for the different levels of `z1`.

We can calculate the Pearson and deviance residuals of the model with the `residuals` command. Figures 4.1 and 4.1 shows these plots.

To assess convergence we can look at trace plots of the MLE estimates and the value of the loglikelihood function across the EM iterations. These are stored in the `mcemGLMM` object returned by the function the `mcemGLMM` function on the fields `mcemEST` and `loglikeVal`. Figure 4.1 shows trace plots at each EM iteration of these quantities.

We can also take a look at the trace plots of the Markov chain used to estimate the  $Q$  function. Since this approximates an integral of dimension equal to the number of random effects it might not be practical to look at all the chains. The last MCMC step is saved on the field `randeff` as a matrix. Each column of this matrix corresponds to one random effect.

This matrix can be used to get predictions of the observed random effects.

```
> plot(simData$x1, predict(fit0, type = "response"), col = simData$z1, xlab = "x1")
```

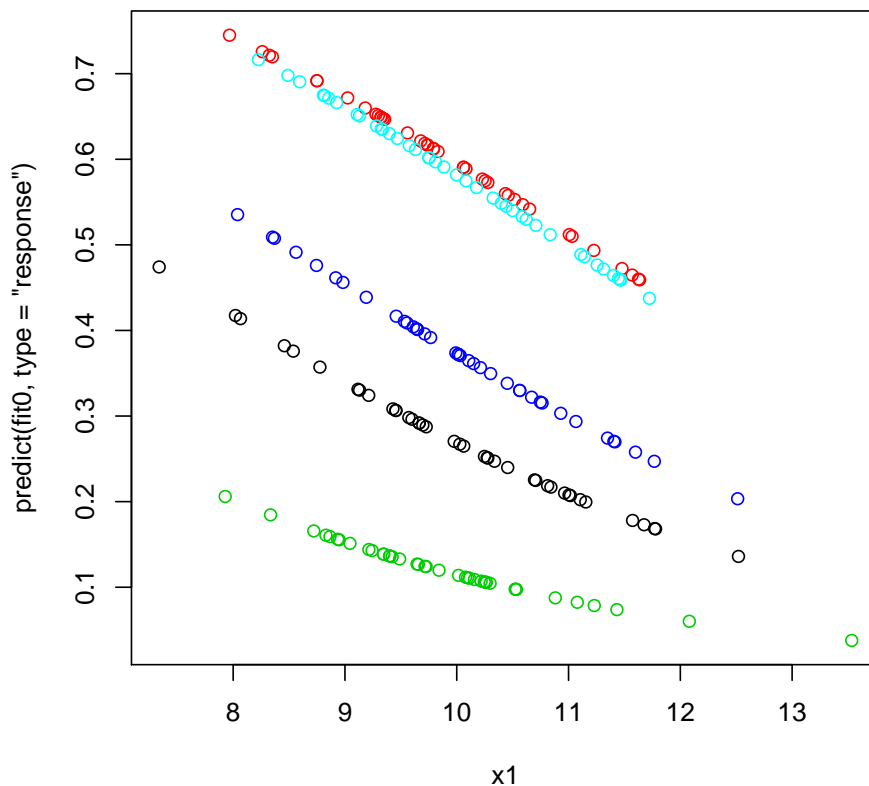


Figure 1: Fitted probabilities. Each color represent a level of  $z1$

```
> colMeans(fit0$randeff)
```

z21	z22	z23	z24
-0.0427675	-0.1262861	0.3917916	-0.2262055

To see the sampling on the loglikelihood function we can plot the values of the complete loglikelihood at each MCMC step of the last EM iteration. These values are stored in the `loglikeMCMC` field of the `mcemGLMM` object.

```
> plot(simData$x1, residuals(fit0, type = "deviance"))
```

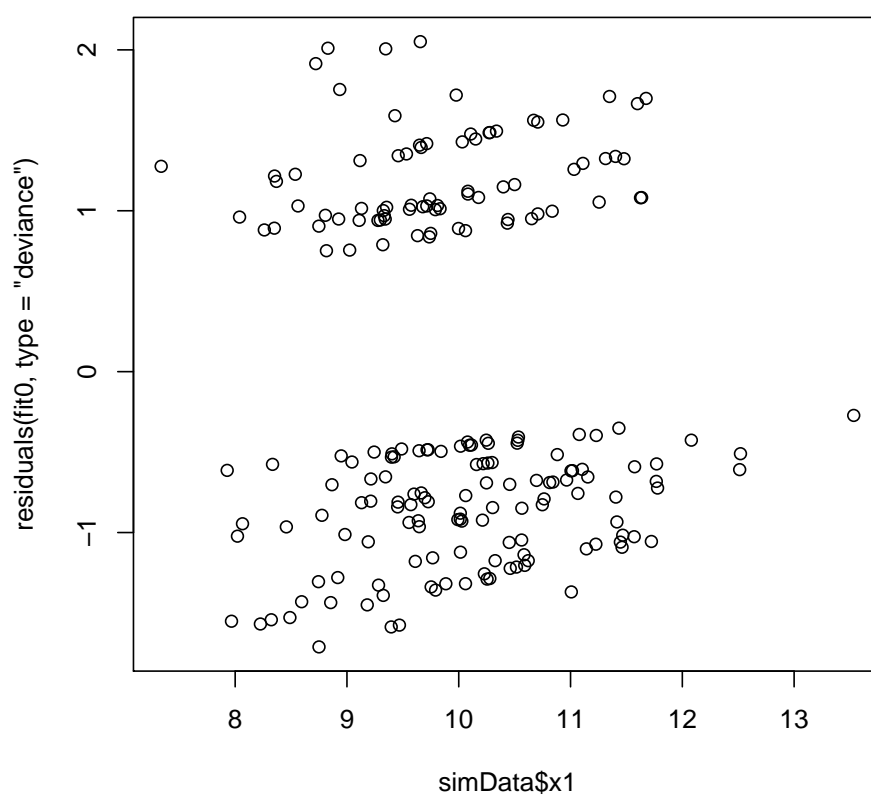


Figure 2: Deviance residuals

```
> plot(simData$x1, residuals(fit0, type = "pearson"))
```

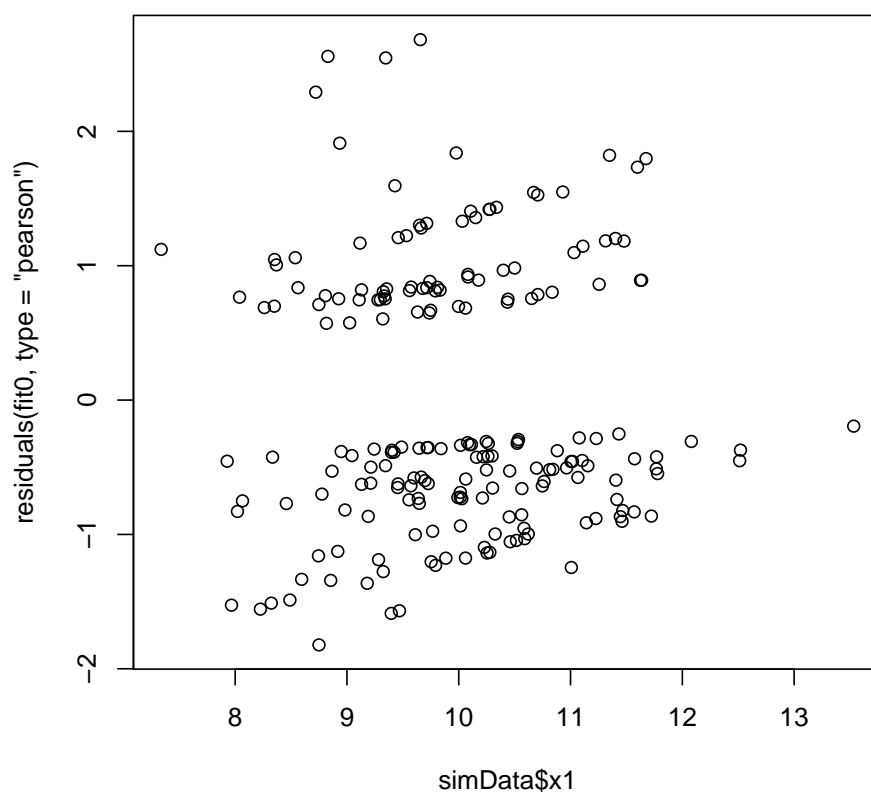


Figure 3: Pearson residuals

```

> par(mfrow = c(1, 2))
> ts.plot(fit0$mcemEST, main = "MLEs estimates",
+         xlab = "EM Iteration", ylab = "MLE value")
> ts.plot(fit0$loglikeVal, main = "Loglikelihood values",
+         xlab = "EM iteration", ylab = "Likelihood")

```

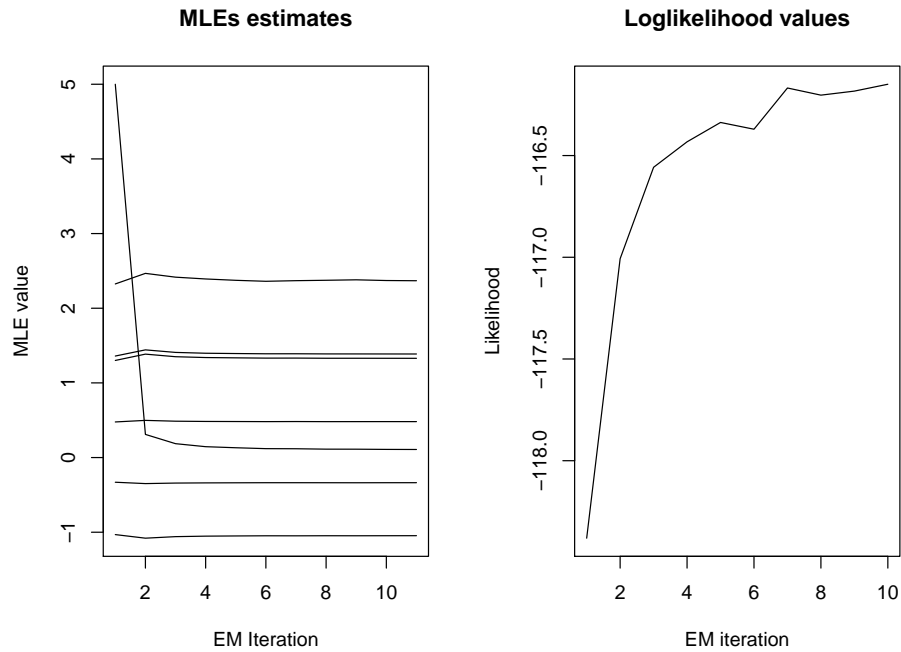


Figure 4: MLEs estimates (left) and Loglikelihood value (right) after each EM iteration.

```
> ts.plot(fit0$randeff[, 1], xlab = "MCMC iteration")
```

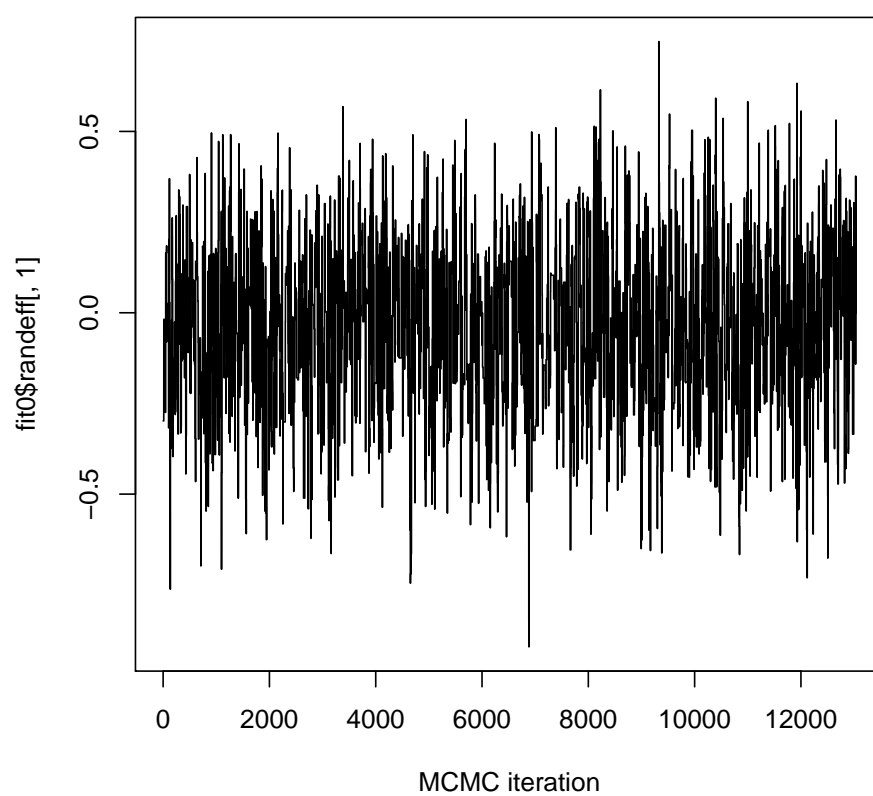


Figure 5: Trace plot for MCMC output for the first random effect.

```
> ts.plot(fit0$loglikeMCMC, xlab = "MCMC iteration")
```

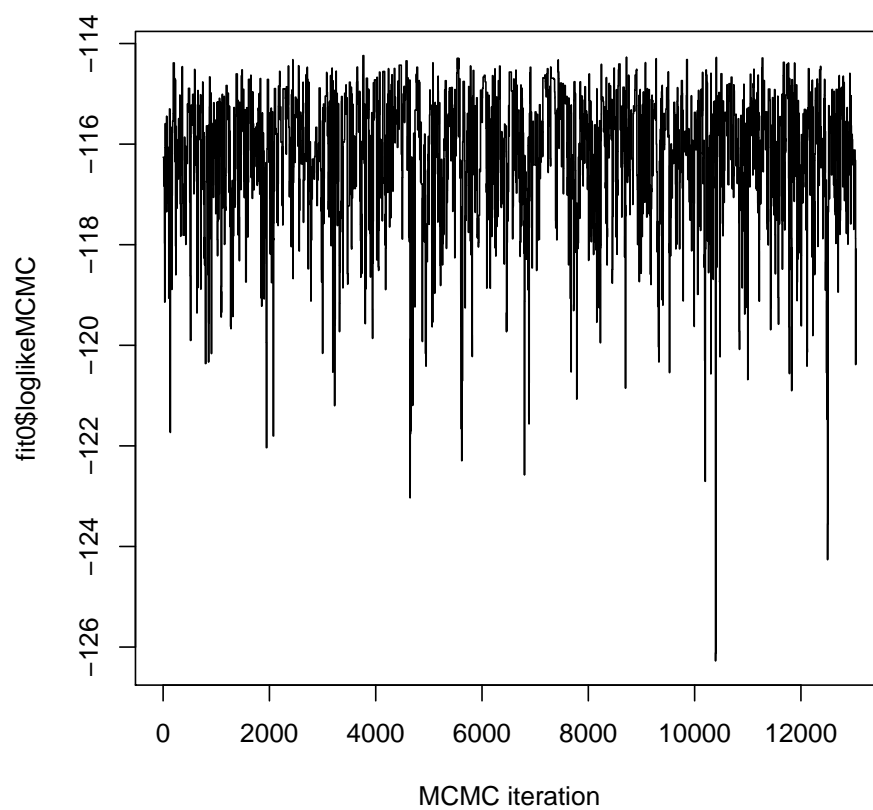


Figure 6: Trace plot for the complete loglikelihood function.

## 4.2 Fitting a more complex model

To specify more than one random effect we need to put them into a list and state that there is no intercept for that effect. In case of nested random effects if labels are repeated across it is necessary to fit the lower level by using the interaction with the upper level.

In this specific example, the labels for  $z2$ , "1", "2", "3", and "4", are used for each level of  $z1$ . If the labels of  $z2$  are unique within  $z1$  it is not necessary to use the interaction term. However it is recommended to use the interaction form for the sake of clarity in the model statement.

```
> fit1 <- mcmGLMM(fixed = obs ~ x1 + x2 + x3,
+               random = list(~0+z1, ~0+z1:z2),
+               data = simData,
+               family = "bernoulli",
+               vcDist = "t",
+               df = c(5, 5))
```

The `df` argument specifies the degrees of freedom for each variance component in `random`. If `vcDist` is "normal" this argument is not needed.

We can look at the summary and ANOVA of the model as before

```
> summary(fit1)
```

Call:

```
mcmGLMM(fixed = obs ~ x1 + x2 + x3, random = list(~0 + z1, ~0 +
  z1:z2), data = simData, family = "bernoulli", vcDist = "t",
  df = c(5, 5))
```

Two sided Wald tests for fixed effects coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	1.0482656	2.1354187	0.4908946	0.62350096
x1	-0.5100424	0.1936848	-2.6333633	0.00845439
x2	0.5790644	0.1984313	2.9182112	0.00352046



x3red	0.2188118	0.4477685	0.4886716	0.62507423
x3yellow	0.8289974	0.4815292	1.7215932	0.08514324

One sided Wald tests for variance components:

	Estimate	Std. Error	z value	Pr(>z)
z1	0.5924914	0.9113657	0.6501138	0.2578093
z1:z2	1.3353322	1.1893710	1.1227213	0.1307779

```
> anova(fit1)
```

Wald's Chi-squared ANOVA table

	Df	Wald Stat	Pr(>W)
x1	1	6.934602	0.00845439
x2	1	8.515957	0.00352046
x3	2	3.139481	0.20809921

We can run multiple comparison tests for the levels of *x3* as before

```
> ctr1 <- rbind( "blue - red" = c(0, 0, 0,-1, 0),
+               "blue - yellow" = c(0, 0, 0, 0,-1),
+               "red - yellow" = c(0, 0, 0, 1,-1))
> contrasts.mcemGLMM(object = fit1, ctr.mat = ctr1)
```

	Estimate	Std. Err.	Wald Adj.	p-value
blue - red	-0.2188118	0.4477685	0.2387999	1.0000000
blue - yellow	-0.8289974	0.4815292	2.9638830	0.2554297
red - yellow	-0.6101857	0.4677397	1.7018277	0.5761474

Instead of performing a Wald test to test a fixed effect it is possible to run a likelihood ratio test between two nested models. First we will fit a model without *x3*:

```

> fit2 <- mcmGLMM(fixed = obs ~ x1 + x2,
+               random = list(~0+z1, ~0+z1:z2),
+               data = simData,
+               family = "bernoulli",
+               vcDist = "t",
+               df = c(5, 5),
+               controlEM = list(EMit = 3))

```

Now we can use the `anova` command to run the likelihood ratio test

```

> anova(fit1, fit2)

Test statistic value: 6.61765024168028
Degrees of freedom: 2
p value: 0.0365591

```

### 4.3 A Poisson model

To fit a Poisson model we only need to change the `family` argument in the `mcmGLMM` command. As an example we will use the `count` variable in `simData`.

```

> fit3 <- mcmGLMM(fixed = count ~ x1 + x2 + x3,
+               random = list(~0+z2),
+               data = simData,
+               family = "poisson",
+               vcDist = "normal")

```

All the previous methods are available for this type of model.

```

> summary(fit3)

```

Call:

```

mcmGLMM(fixed = count ~ x1 + x2 + x3, random = list(~0 + z2),
        data = simData, family = "poisson", vcDist = "normal")

```

Two sided Wald tests for fixed effects coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	2.429668527	0.35016284	6.93868171	0.00000000
x1	-0.100594578	0.03139176	-3.20449005	0.00135302
x2	0.042484752	0.03066740	1.38533934	0.16594873
x3red	-0.072106883	0.07945201	-0.90755269	0.36411460
x3yellow	0.002406589	0.07747179	0.03106407	0.97521845

One sided Wald tests for variance components:

	Estimate	Std. Error	z value	Pr(>z)
z2	0.003188614	0.006782275	0.4701393	0.3191277

> anova(fit3)

Wald's Chi-squared ANOVA table

	Df	Wald Stat	Pr(>W)
x1	1	10.268757	0.00135302
x2	1	1.919165	0.16594873
x3	2	1.162212	0.55927940

> contrasts.mcemGLMM(object = fit3, ctr.mat = ctr1)

	Estimate	Std. Err.	Wald	Adj. p-value
blue - red	0.072106883	0.07945201	0.8236518816	1
blue - yellow	-0.002406589	0.07747179	0.0009649761	1
red - yellow	-0.074513472	0.07727556	0.9297908976	1

```
> plot(simData$x1, predict(fit3),
+      main = "Predicted response values", xlab = "x1")
```

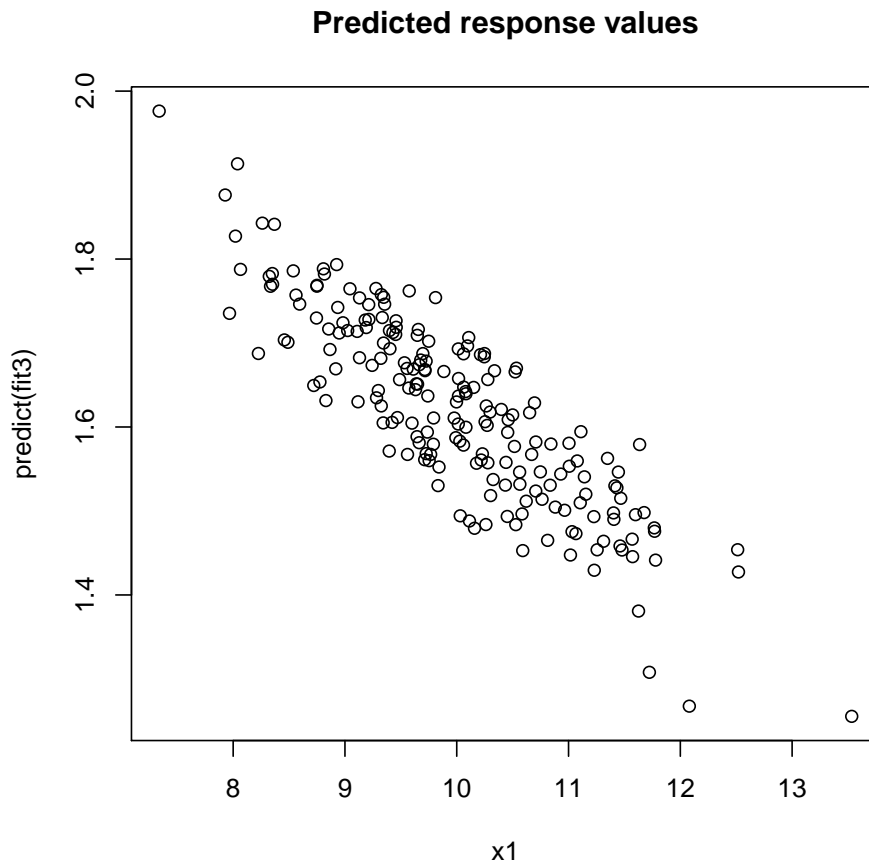


Figure 7: Predicted values as a function of  $x_1$ .

#### 4.4 A negative binomial model

To fit a negative binomial model we need to specify the `family` argument to `"negbinom"`. All the previous methods are available for this model. When we look at the summary of this model we get an estimate of the overdispersion parameter and its standard error.

```
> fit4 <- mceGLMM(fixed = count2 ~ x1 + x2 + x3,
+               random = list(~0+z1, ~0+z1:z2),
+               data = simData,
+               family = "negbinom",
```

```

> par(mfrow = c(1, 2))
> ts.plot(fit3$mcemEST, main = "MLEs estimates",
+         xlab = "EM Iteration", ylab = "MLE value")
> ts.plot(fit3$loglikeVal, main = "Loglikelihood values",
+         xlab = "EM iteration", ylab = "Likelihood")

```

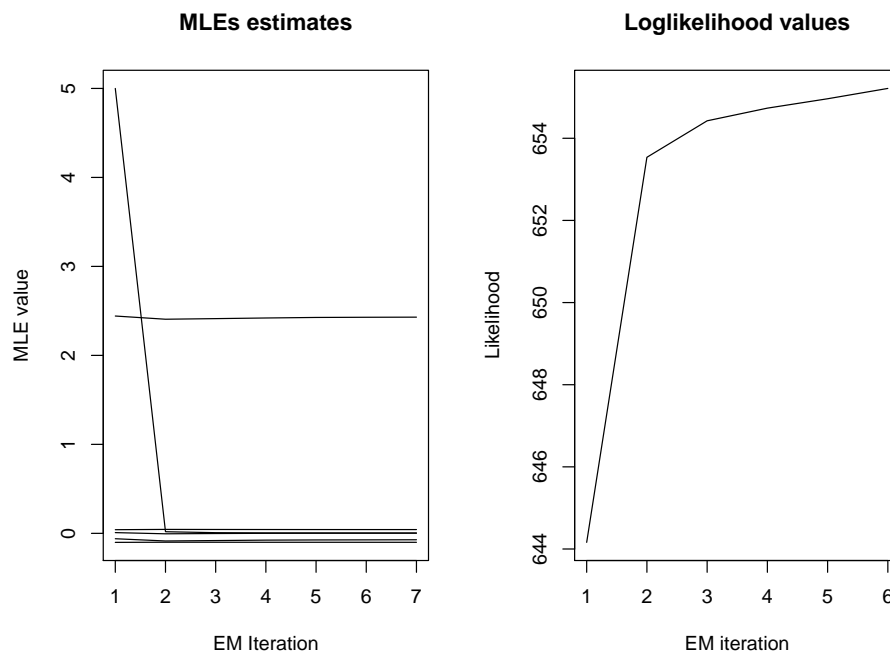


Figure 8: MLEs estimates (left) and Loglikelihood value (right) after each EM iteration.

```

+         vcDist = "normal")
> summary(fit4)

```

Call:

```

mcemGLMM(fixed = count2 ~ x1 + x2 + x3, random = list(~0 + z1,
~0 + z1:z2), data = simData, family = "negbinom", vcDist = "normal")

```

Two sided Wald tests for fixed effects coefficients:

	Estimate	Std. Error	z value	Pr(> z )
(Intercept)	4.21614618	0.57654418	7.31278944	0.00000000
x1	-0.15975518	0.04954114	-3.22469750	0.00126106

x2	0.01458830	0.04912726	0.29694920	0.76650529
x3red	-0.13490761	0.12613565	-1.06954383	0.28482469
x3yellow	-0.01146224	0.13047303	-0.08785144	0.92999476

Overdispersion parameter beta:

	Estimate	Std. Error
theta	1.388511	0.1252373

One sided Wald tests for variance components:

	Estimate	Std. Error	z value	Pr(>z)
z1	0.05705629	0.09288859	0.6142443	0.26952695
z1:z2	0.18417139	0.08842985	2.0826835	0.01864004

```
> anova(fit4)
```

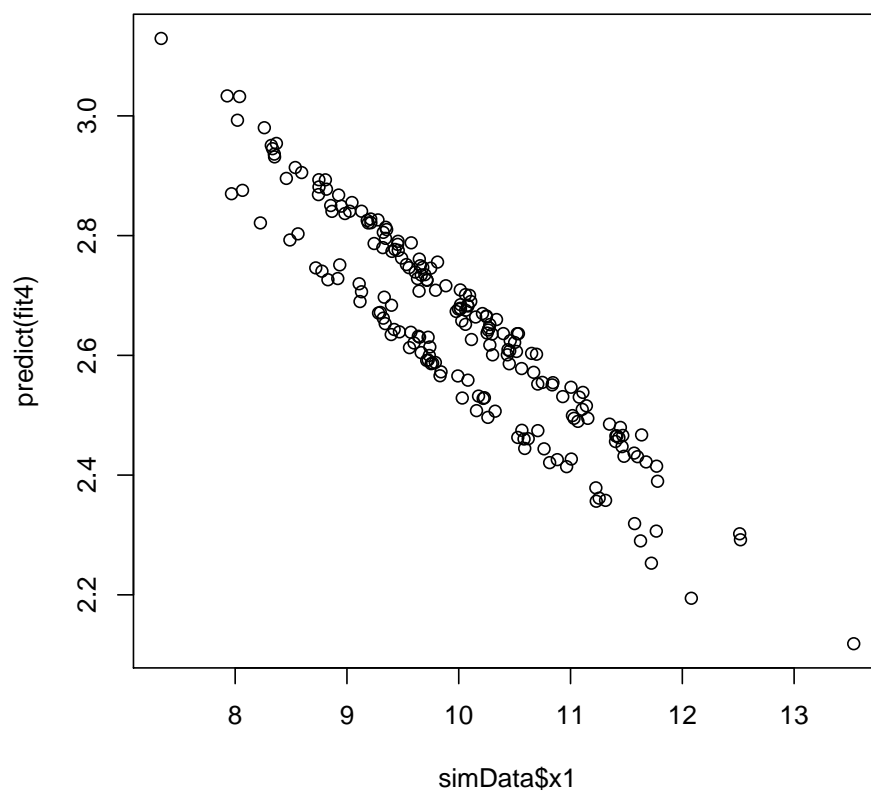
Wald's Chi-squared ANOVA table

	Df	Wald Stat	Pr(>W)
x1	1	10.39867396	0.00126106
x2	1	0.08817883	0.76650529
x3	2	1.39625335	0.49751644

```
> contrasts.mcemGLMM(object = fit4, ctr.mat = ctr1)
```

	Estimate	Std. Err.	Wald Adj.	p-value
blue - red	0.13490761	0.1261357	1.143924013	0.8544741
blue - yellow	0.01146224	0.1304730	0.007717875	1.0000000
red - yellow	-0.12344537	0.1294790	0.908972974	1.0000000

```
> plot(simData$x1, predict(fit4))
```



```

> par(mfrow = c(1, 2))
> ts.plot(fit4$mcemEST, main = "MLEs estimates",
+         xlab = "EM Iteration", ylab = "MLE value")
> ts.plot(fit4$loglikeVal, main = "Likelihood value",
+         xlab = "EM Iteration", ylab = "Likelihood")

```

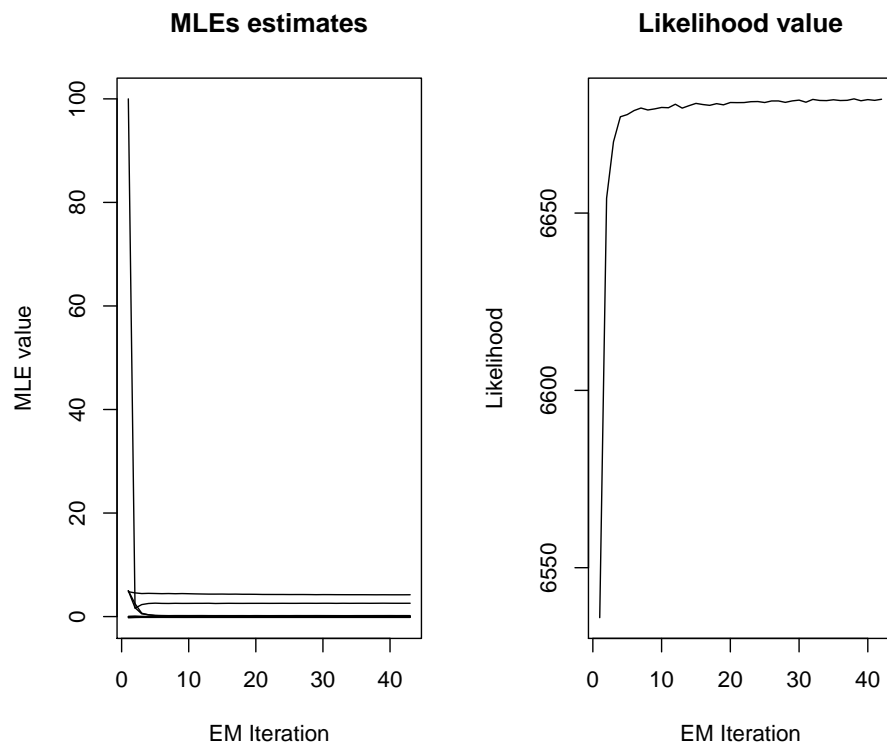


Figure 9: MLEs estimates and loglikelihood value after each EM iteration.