

# **rth ORDER MARKOV MODELS BASED IN MULTIPLE LOGISTIC REGRESSION FOR COVARIATE DEPENDENCE OF SEQUENCES**

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A classical problem in many experimental investigations is to model the response variable, subject to observation, in function of the covariates that have influence in the study. The binary case has received a considerable attention (Korn and Whittemore (1979), Muenz and Rubinstein (1985), Pregibon (1982)). In this work, multiple response (or state) is considered and models based in multiple logistic regression are proposed. The basic assumption is that the sequence of states is a  $r$ th order Markov chain with stationary transition probabilities. The regression coefficients are estimated by maximum likelihood procedure, and in order to test the order of the chain, a likelihood ratio test is obtained. When  $r=1$ , these problems have been studied, considering several situations, in Molina and Requena (1986), Molina (1986a), and Molina (1986b).

## Models based in multiple logistic regression.

Suppose that a heterogeneous group of individuals is followed over time, with each individual passing through states according to a  $r$ th order Markov chain with state space  $\{1,2,\dots,m\}$  and stationary transition probabilities. Suppose that the observations are made at times  $t=0,1,\dots,T$ , where  $T$  must be at least  $r$ , and let  $Z_1,\dots,Z_k$  be the covariates.

In order to explain how the covariates relate to changes in state, we proposed to model  $p_{q,i_1\dots i_r} = P(X_t=q / X_{t-r}=i_r,\dots,X_{t-1}=i_1)$ , with  $q=1,\dots,m-1$ ,  $i_1,\dots,i_r=1,\dots,m$ ,  $t=r,\dots,T$ , in the form :

$$p_{q,i_1\dots i_r} = \frac{\exp\{f(\beta_{q,i_1\dots i_r}; Z_1\dots Z_k)\}}{1 + \sum_{q=1}^{m-1} \exp\{f(\beta_{q,i_1\dots i_r}; Z_1\dots Z_k)\}}^{-1}$$

where  $f(\beta_{q,i_1 \dots i_r}; Z_1 \dots Z_k)$  is a linear function of the covariates and the vector of regression coefficients  $\beta_{q,i_1 \dots i_r}$  whose exact form will vary according to the model under study. For simplicity let us consider the case when  $f(\beta_{q,i_1 \dots i_r}; Z_1 \dots Z_k) = \beta_{q,i_1 \dots i_r} \cdot Z'$ , where  $Z$  is the vector  $(1, z_1, \dots, z_k)$  with  $z_i$  the initial value of covariate  $Z_i$ , and  $\beta_{q,i_1 \dots i_r} = (\beta_{q,i_1 \dots i_r}^0, \dots, \beta_{q,i_1 \dots i_r}^k)$ . Such situation corresponding to model without interaction between covariates.

#### Estimating the regression coefficients.

Let the sequences observed be  $x_{0h}, x_{1h}, \dots, x_{Th}$  ( $h=1, \dots, n$ ), in  $n$  individual up to time  $T$ , and let  $Z_h = (1, z_{1h}, \dots, z_{kh})$  be the vector of covariates for the  $h$ th individual. If the  $r$  initials states of each individual are known, then the log-likelihood function can be written in the form :

$$L_T(\beta_{i_1 \dots i_r}, i_1, \dots, i_r = 1, \dots, m) = \sum_{i_1, \dots, i_r=1}^m F_{i_1 \dots i_r}(\beta_{i_1 \dots i_r}) \quad (1)$$

where

$$F_{i_1 \dots i_r}(\beta_{i_1 \dots i_r}) = \sum_{h=1}^n \left( \sum_{q=1}^{m-1} n_{q,i_1 \dots i_r}^h \beta_{q,i_1 \dots i_r} \cdot Z_h' - n_{\cdot, i_1 \dots i_r}^h \ln \left( 1 + \sum_{q=1}^{m-1} \exp(\beta_{q,i_1 \dots i_r} \cdot Z_h') \right) \right)$$

with  $\beta_{i_1 \dots i_r} = (\beta_{1,i_1 \dots i_r}, \dots, \beta_{m-1,i_1 \dots i_r})$ .

$n_{q,i_1 \dots i_r}^h$  is the number of transitions to state  $q$ , from the sequence of consecutive states  $i_r, i_{r-1}, \dots, i_1$ , observed in the  $h$ th individual

and  $n_{\cdot, i_1 \dots i_r}^h = \sum_{q=1}^m n_{q,i_1 \dots i_r}^h$

It follows from (1) that for each  $i_1, \dots, i_r \in \{1, \dots, m\}$ , the maximum likelihood estimate of  $\beta_{i_1 \dots i_r}$  can be obtained separately. For this purpose, the likelihood equation is :

$$\frac{\partial F_{i_1 \dots i_r}(\beta_{i_1 \dots i_r})}{\partial \beta_{q,i_1 \dots i_r}^s} = \sum_{h=1}^n (n_{q,i_1 \dots i_r}^h - n_{\cdot, i_1 \dots i_r}^h p_{q,i_1 \dots i_r}^h) z_{sh} = 0 \quad (2)$$

with  $s=0, \dots, k$ ,  $q=1, \dots, m-1$ , and  $z_{oh}=1$  ( $h=1, \dots, n$ )

We can maximize (2) by Newton-Raphson method as long as the second derivatives matrix has full rank, and  $\min_{i_1, \dots, i_r} \left\{ \sum_{h=1}^n n_{\cdot, i_1 \dots i_r}^h \right\} \geq k+1$

For large  $n$ , an estimate of the covariance matrix of  $\hat{\beta}_{i_1 \dots i_r}$ , is the matrix that has elements equal to  $-1$  times the inverse of the second derivatives matrix evaluated in  $\hat{\beta}_{i_1 \dots i_r}$ .

Test of the hypothesis that the chain is of a given order.

For testing the hypothesis that the process is a  $u$ th order Markov chain against the alternative it is  $r$ th but not  $u$ th order ( $u < r$ ), the statistic  $-2(L_T(\hat{\beta}_{i_1 \dots i_u}) - L_T(\hat{\beta}_{i_1 \dots i_r}))$ , that is asymptotically distributed as  $\chi^2$  with  $(m-1)m^u(m^{r-u}-1)(k+1)$  degree of freedom, can be used.

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