# Generalized Linear Mixed Models for Longitudinal Data

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Abstract The study of longitudinal data plays a significant role in medicine, epidemiology and social sciences. Typically, the interest is in the dependence of an outcome variable on the covariates. The Generalized Linear Models (GLMs) were proposed to unify the regression approach for a wide variety of discrete and continuous longitudinal data. The responses (outcomes) in longitudinal data are usually correlated. Hence, we need to use an extension of the GLMs that account for such correlation. This cam be done by inclusion of random effects in the linear predictor, that is the Generalized Linear Mixed Models (GLMMs) (also called random effects models). The maximum likelihood estimates (MLE) are obtained for the regression parameters of a logit model, when the traditional assumption of normal random effects is relaxed. In this case a more convenient distribution, such as the lognormal distribution, is used. However, adding non-normal random effects to the GLMM considerably complicates the likelihood estimation. So, the direct numerical evaluation techniques (such as Newton Raphson) become analytically and computationally tedious. To overcome such problems, we propose and develop a Monte Carlo EM (MCEM) algorithm, to obtain the maximum likelihood estimates. The proposed method is illustrated using a simulated data.

Keywords Generalized Linear Mixed Models, Logistic Regression, Longitudinal Data, Monte Carlo EM Algorithm, Random Effects Model

#### 1. Introduction

Longitudinal data consist of repeated observations, for the same subject, of an outcome variable. There may be a set of covariates for each subjects. Let  $\mathbf{y}_t = (y_{t1}, y_{t2}, ..., y_{tmt})^T$  be an  $m_t \times 1$  vector representing the observed sequence of the outcome variable  $Y_{tt}$  recorded at time  $t = 1, 2, ..., m_t$ , for the ith subject, i = 1, 2, ..., n. Also, assume that  $\mathbf{X}_{ij} = (\mathbf{x}_{ati}, \mathbf{x}_{a2}, ..., \mathbf{x}_{m_t})$  is an  $I \times p$  vector of p covariates observed at time t. Thus,  $X_t$  is an  $m_t \times p$  matrix of covariates corresponding to the ith subject taking the form:

$$\begin{pmatrix} x_{i11} & \cdots & x_{i1p} \\ \vdots & \ddots & \vdots \\ x_{omi1} & \cdots & x_{omp} \end{pmatrix}.$$

For simplicity we can assume that m = w.

The primary focus is on the dependence of the outcome on the covariates[3,5]. In other words, we are usually interested in the inference about the regression coefficients, in the usual linear models of the form

$$Y_i = X_i\beta + e_i$$
,  
where  $\beta = (\beta_1, ..., \beta_p)^T$  is a p x1 vector of the regression  
coefficients. It is usually assumed that the errors,  $e_i$ , are  
independent and identically normally distributed. Therefore,  
these models are not used in situations when response

variables have distributions other than the normal, or even when they are qualitative rather than quantitative. Examples include binary longitudinal data.

To solve this problem, Reference[18] introduce the generalized linear models (GLM) as a unified framework to model all types of longitudinal data[13, 15, 24]. These models assume that the distribution of  $Y_n$  (i = 1, ..., n, i = I, 2, ..., m) belongs to the exponential family. The exponential family distributions can be written in the form:

 $f(y_{it}, \theta_{it}) = e^{\{(y_{it}, \theta_{it} - \pi(\theta_{it}) + b(y_{it})\}\theta_{it}}$  (1) where  $\omega(.)$  and b(.) are specific functions defined according to the underlying distribution of the exponential family. Some examples of univariate distributions are given in Table (1). The parameters  $\theta_{it} = h(\eta_{it})$  and  $\eta_{it} = x_{it}\beta$ . Also  $\phi$  is the scale parameter and it is treated as a nuisance parameter when the main interest is in the regression coefficients[13]. The distribution in Equation (1) is the canonical form and  $\theta_{it}$  is called the natural parameter of the distribution.

The first two moments of  $y_{it}$  in Equation (1) are given by

$$E(y_{it}) = \mu_{it} = a'(\theta_{it}) \text{ and } Var(y_{it}) = \frac{a''(\theta_{it})}{\theta}$$
 (2)  
 $b1 = -\frac{1}{2}(y^2 + \sigma^2 \ln(2\pi\sigma^2))$ 

For the GLM, the linear combination of the parameters of interest,  $\beta_1, \beta_2, \dots, \beta_p$ , are equal to some function of the expected value of  $\gamma_{il}$  ( $\mu_{il}$ ), i. e.,  $g(\mu_{il}) = \eta_{il}$  where g is a monotone and differentiable function called the link function. From Equation (1) and Equation (2), we can write the inverse of the link function as  $g^{-1} = a$  o h. Note that a link function in which  $\theta_{il} = \eta_{il}$  (e.g., h is the identity function)

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# **Generalized Linear Mixed Models For Longitudinal Data With**

Walter W. Stroup

## **Generalized Linear Mixed Models For Longitudinal Data With:**

Generalized Linear Mixed Models Charles E. McCulloch, 2003 Wiley Series in Probability and Statistics A modern perspective on mixed models The availability of powerful computing methods in recent decades has thrust linear and nonlinear mixed models into the mainstream of statistical application. This volume offers a modern perspective on generalized linear and mixed models presenting a unified and accessible treatment of the newest statistical methods for analyzing correlated nonnormally distributed data As a follow up to Searle's classic Linear Models and Variance Components by Searle Casella and McCulloch this new work progresses from the basic one way classification to generalized linear mixed models A variety of statistical methods are explained and illustrated with an emphasis on maximum likelihood and restricted maximum likelihood An invaluable resource for applied statisticians and industrial practitioners as well as students interested in the latest results Generalized Linear and Mixed Models features A review of the basics of linear models and linear mixed models Descriptions of models for nonnormal data including generalized linear and nonlinear models Analysis and illustration of techniques for a variety of real data sets Information on the accommodation of longitudinal data using these models Coverage of the prediction of realized values of random effects A discussion of the impact of computing issues on mixed Generalized Linear Mixed Models Walter W. Stroup, 2016-04-19 With numerous examples using SAS PROC models GLIMMIX this text presents an introduction to linear modeling using the generalized linear mixed model as an overarching conceptual framework For readers new to linear models the book helps them see the big picture It shows how linear models fit with the rest of the core statistics curriculum and points out the major issues that statistical modelers must consider

Linear and Generalized Linear Mixed Models and Their Applications Jiming Jiang, 2007-05-30 Over the past decade there has been an explosion of developments in mixed e ects models and their applications. This book concentrates on two major classes of mixed e ects models linear mixed models and generalized linear mixed models with the intention of o ering an up to date account of theory and methods in the analysis of these models as well as their applications in various elds. The rst two chapters are devoted to linear mixed models. We classify I ear mixed models as Gaussian linear mixed models and non Gaussian linear mixed models. There have been extensive studies in estimation in Gaussian mixed models as well as tests and con dence intervals On the other hand the literature on non Gaussian linear mixed models is much less extensive partially because of the di culties in inference about these models. However non Gaussian linear mixed models are important because in practice one is never certain that normality holds. This book o ers a systematic approach to inference about non Gaussian linear mixed models. In particular it has included recently developed methods such as partially observed information iterative weighted least squares and jackknife in the context of mixed models. Other new methods introduced in this book include goodness of t tests p diction intervals and mixed model selection. These are of course in addition to traditional topics such as maximum likelihood and restricted maximum likelihood in Gaussian mixed models.

with Applications in Agriculture and Biology Josafhat Salinas Ruíz, Osval Antonio Montesinos López, Gabriela Hernández Ramírez, Jose Crossa Hiriart, 2023-08-16 This open access book offers an introduction to mixed generalized linear models with applications to the biological sciences basically approached from an applications perspective without neglecting the rigor of the theory For this reason the theory that supports each of the studied methods is addressed and later through examples its application is illustrated In addition some of the assumptions and shortcomings of linear statistical models in general are also discussed An alternative to analyse non normal distributed response variables is the use of generalized linear models GLM to describe the response data with an exponential family distribution that perfectly fits the real response Extending this idea to models with random effects allows the use of Generalized Linear Mixed Models GLMMs The use of these complex models was not computationally feasible until the recent past when computational advances and improvements to statistical analysis programs allowed users to easily quickly and accurately apply GLMM to data sets GLMMs have attracted considerable attention in recent years The word Generalized refers to non normal distributions for the response variable and the word Mixed refers to random effects in addition to the fixed effects typical of analysis of variance or regression With the development of modern statistical packages such as Statistical Analysis System SAS R ASReml among others a wide variety of statistical analyzes are available to a wider audience However to be able to handle and master more sophisticated models requires proper training and great responsibility on the part of the practitioner to understand how these advanced tools work GMLM is an analysis methodology used in agriculture and biology that can accommodate complex correlation structures and types of response variables Analysis of Generalized Linear Mixed Models in the Agricultural and Natural Resources Sciences Edward E. Gbur, Walter W. Stroup, Kevin S. McCarter, Susan Durham, Linda J. Young, Mary Christman, Mark West, Matthew Kramer, 2020-01-22 Generalized Linear Mixed Models in the Agricultural and Natural Resources Sciences provides readers with an understanding and appreciation for the design and analysis of mixed models for non normally distributed data It is the only publication of its kind directed specifically toward the agricultural and natural resources sciences audience Readers will especially benefit from the numerous worked examples based on actual experimental data and the discussion of pitfalls associated with incorrect analyses Applied Regression Analysis and Generalized Linear Models John Fox, 2015-03-18 Combining a modern data analytic perspective with a focus on applications in the social sciences the Third Edition of Applied Regression Analysis and Generalized Linear Models provides in depth coverage of regression analysis generalized linear models and closely related methods such as bootstrapping and missing data Updated throughout this Third Edition includes new chapters on mixed effects models for hierarchical and longitudinal data Although the text is largely accessible to readers with a modest background in statistics and mathematics author John Fox also presents more advanced material in optional sections and chapters throughout the book Accompanying website resources containing all answers to the end of chapter exercises Answers to odd numbered questions as well as datasets and

other student resources are available on the author's website NEW Bonus chapter on Bayesian Estimation of Regression Models also available at the author's website Variable Selection Procedures for Generalized Linear Mixed Models in Longitudinal Data Analysis, 2004 Model selection is important for longitudinal data analysis But up to date little work has been done on variable selection for generalized linear mixed models GLMM In this paper we propose and study a class of variable selection methods Full likelihood FL approach is proposed for simultaneous model selection and parameter estimation Due to the intensive computation involved in FL approach Penalized Quasi Likelihood PQL procedure is developed so that model selection in GLMMs can proceed in the framework of linear mixed models Since the PQL approach will produce biased parameter estimates for sparse binary longitudinal data Two stage Penalized Quasi Likelihood approach TPQL is proposed to bias correct PQL in terms of estimation use PQL to do model selection at the first stage and existing software to do parameter estimation at the second stage Marginal approach for some special types of data is also developed A robust estimator of standard error for the fitted parameters is derived based on a sandwich formula A bias correction is proposed to improve the estimation accuracy of PQL for binary data The sampling performance of four proposed procedures is evaluated through extensive simulations and their application to real data analysis In terms of model selection all of them perform closely As for parameter estimation FL AML and TPQL yield similar results Compared with FL the other procedures greatly reduce computational load The proposed procedures can be extended to longitudinal data analysis involving missing data and the shrinkage penalty based approach allows them to work even when the number of observations n is less than the number of parameters d Generalized Linear Models Dipak K. Dey, Sujit K. Ghosh, Bani K. Mallick, 2000-05-25 This volume describes how to conceptualize perform and critique traditional generalized linear models GLMs from a Bayesian perspective and how to use modern computational methods to summarize inferences using simulation Introducing dynamic modeling for GLMs and containing over 1000 references and equations Generalized Linear Models considers parametric and semiparametric approaches to overdispersed GLMs presents methods of analyzing correlated binary data using latent variables It also proposes a semiparametric method to model link functions for binary response data and identifies areas of important future research and new applications of GLMs **Mixed Models** Eugene Demidenko, 2005-01-28 A rigorous self contained examination of mixed model theory and application Mixed modeling is one of the most promising and exciting areas of statistical analysis enabling the analysis of nontraditional clustered data that may come in the form of shapes or images This book provides in depth mathematical coverage of mixed models statistical properties and numerical algorithms as well as applications such as the analysis of tumor regrowth shape and image Paying special attention to algorithms and their implementations the book discusses Modeling of complex clustered or longitudinal data Modeling data with multiple sources of variation Modeling biological variety and heterogeneity Mixed model as a compromise between the frequentist and Bayesian approaches Mixed model for the penalized log likelihood Healthy Akaike Information Criterion HAIC How to

cope with parameter multidimensionality How to solve ill posed problems including image reconstruction problems Modeling of ensemble shapes and images Statistics of image processing Major results and points of discussion at the end of each chapter along with Summary Points sections make this reference not only comprehensive but also highly accessible for professionals and students alike in a broad range of fields such as cancer research computer science engineering and Generalized Linear and Nonlinear Models for Correlated Data Edward F. Vonesh, 2014-07-07 Edward Vonesh s Generalized Linear and Nonlinear Models for Correlated Data Theory and Applications Using SAS is devoted to the analysis of correlated response data using SAS with special emphasis on applications that require the use of generalized linear models or generalized nonlinear models Written in a clear easy to understand manner it provides applied statisticians with the necessary theory tools and understanding to conduct complex analyses of continuous and or discrete correlated data in a longitudinal or clustered data setting Using numerous and complex examples the book emphasizes real world applications where the underlying model requires a nonlinear rather than linear formulation and compares and contrasts the various estimation techniques for both marginal and mixed effects models The SAS procedures MIXED GENMOD GLIMMIX and NLMIXED as well as user specified macros will be used extensively in these applications In addition the book provides detailed software code with most examples so that readers can begin applying the various techniques immediately This book is part of the SAS Press program Generalized, Linear, and Mixed Models Charles E. McCulloch, Shayle R. Searle, John M. Neuhaus, 2011-09-20 An accessible and self contained introduction to statistical models now in a modernized new edition Generalized Linear and Mixed Models Second Edition provides an up to date treatment of the essential techniques for developing and applying a wide variety of statistical models. The book presents thorough and unified coverage of the theory behind generalized linear and mixed models and highlights their similarities and differences in various construction application and computational aspects A clear introduction to the basic ideas of fixed effects models random effects models and mixed models is maintained throughout and each chapter illustrates how these models are applicable in a wide array of contexts In addition a discussion of general methods for the analysis of such models is presented with an emphasis on the method of maximum likelihood for the estimation of parameters The authors also provide comprehensive coverage of the latest statistical models for correlated non normally distributed data Thoroughly updated to reflect the latest developments in the field the Second Edition features A new chapter that covers omitted covariates incorrect random effects distribution correlation of covariates and random effects and robust variance estimation A new chapter that treats shared random effects models latent class models and properties of models A revised chapter on longitudinal data which now includes a discussion of generalized linear models modern advances in longitudinal data analysis and the use between and within covariate decompositions Expanded coverage of marginal versus conditional models Numerous new and updated examples With its accessible style and wealth of illustrative exercises Generalized Linear and Mixed Models Second Edition is an ideal book for

courses on generalized linear and mixed models at the upper undergraduate and beginning graduate levels It also serves as a valuable reference for applied statisticians industrial practitioners and researchers **Models for Discrete Longitudinal Data** Geert Molenberghs, Geert Verbeke, 2006-01-28 The linear mixed model has become the main parametric tool for the analysis of continuous longitudinal data as the authors discussed in their 2000 book Without putting too much emphasis on software the book shows how the different approaches can be implemented within the SAS software package The authors received the American Statistical Association's Excellence in Continuing Education Award based on short courses on longitudinal and incomplete data at the Joint Statistical Meetings of 2002 and 2004 Generalized Linear Models and Extensions M. Ataharul Islam, Soma Chowdhury Biswas, 2025-04-30 This book presents a wide range of topics to address the needs of several groups of users of rapidly growing methods of generalized linear models Since the introduction of the idea of generalized linear models GLM in early seventies during the past four decades the modelling of statistical data have experienced a major transformation from linear models based on normality assumption to a more flexible unified approach of generalized linear models The number of readers and users of generalized linear models have increased manifold In addition the use of generalized linear models has expanded in many new fields of applications where statistical models are being employed at an increasing rate It is important to note here that the learners and users of GLM have a widely varied background in different disciplines Considering these pressing needs this book focuses on i upper undergraduate and graduate level students in need of a thorough understanding about the basic concepts of generalized linear models along with appropriate applications ii researchers and users in need of advanced generalized linear models for analysing bivariate or multivariate data stemming from longitudinal or repeated measures data and iii new challenges to analyse big data where the traditional techniques fail to provide any reasonable modelling strategy In other words this book starts with a thorough background of the generalized linear models for the new learners then provides multivariate extensions to advanced level techniques for researchers and users in various disciplines and finally some innovative modelling strategies are introduced using generalized linear models in the emerging field of big data analytics It provides materials for new learners for users researchers who are in need of more advanced techniques and also strategies for employing linear models in big data analytics Hence techniques of generalized linear models will be presented in the proposed book covering the needs of new learners users of advanced techniques researchers in need of statistical modelling of any data type and users of big data analytics wanting to increase predictive accuracy of classification and regression tree techniques **Computational and** Methodological Statistics and Biostatistics Andriëtte Bekker, (Din) Ding-Geng Chen, Johannes T. Ferreira, 2020-08-10 In the statistical domain certain topics have received considerable attention during the last decade or so necessitated by the growth and evolution of data and theoretical challenges This growth has invariably been accompanied by computational advancement which has presented end users as well as researchers with the necessary opportunities to handle data and

implement modelling solutions for statistical purposes Showcasing the interplay among a variety of disciplines this book offers pioneering theoretical and applied solutions to practice oriented problems As a carefully curated collection of prominent international thought leaders it fosters collaboration between statisticians and biostatisticians and provides an array of thought processes and tools to its readers The book thereby creates an understanding and appreciation of recent developments as well as an implementation of these contributions within the broader framework of both academia and industry Computational and Methodological Statistics and Biostatistics is composed of three main themes Recent developments in theory and applications of statistical distributions Recent developments in supervised and unsupervised modelling Recent developments in biostatistics and also features programming code and accompanying algorithms to enable readers to replicate and implement methodologies Therefore this monograph provides a concise point of reference for a variety of current trends and topics within the statistical domain With interdisciplinary appeal it will be useful to researchers graduate students and practitioners in statistics biostatistics clinical methodology geology data science and actuarial science Mixed Effects Models for Complex Data Lang Wu,2009-11-11 Although standard mixed effects models amongst others are useful in a range of studies other approaches must often be used in correlation with them when studying complex or incomplete data Mixed Effects Models for Complex Data discusses commonly used mixed effects models and presents appropriate approaches to address dropouts missing data measurement errors Grundlagen und statistische Methoden der Evaluationsforschung Heinz Holling, 2009-09-11 In nahezu allen gesellschaftlichen Bereichen wie z B Bildung Verwaltung Wirtschaft oder Gesundheit werden heutzutage unterschiedlichste Aktivit ten evaluiert Somit spielt die Evaluationsforschung nicht nur in der Psychologie sondern auch in den Nachbarwissenschaften wie der P dagogik Politologie oder konomie eine bedeutende Rolle Die Beitr ge dieses Enzyklop diebandes konzentrieren sich auf die wissenschaftlichen Grundlagen der Evaluationsforschung Im Vordergrund stehen die Definition der Evaluationsforschung und die wissenschaftstheoretische Fundierung dieses Gebietes sowie statistische und methodische Grundlagen Da Evaluationsstudien h ufig komplexer und l ngerfristiger Natur sind erfordern sie insbesondere multivariate statistische Auswertungsverfahren Ihnen wird daher besonders viel Raum gew hrt In einzelnen Kapiteln werden u a folgende Themen behandelt Facettentheorie Aufstellen und Pr fen von Hypothesen Generalisierbarkeitstheorie Metaanalyse in der Evaluationsforschung das Allgemeine Lineare Modell in der Evaluationsforschung Hierarchische Lineare Modelle HLM Generalisierte Lineare Modelle Strukturgleichungsmodelle Multidimensionale Skalierung Mischverteilungsmodelle Ver nderungsmessung Zeitreihenanalyse Resampling sowie statistische Methoden bei unvollst ndigen Daten Frontiers In Statistics Jianging Fan, Hira L Koul, 2006-07-17 During the last two decades many areas of statistical inference have experienced phenomenal growth This book presents a timely analysis and overview of some of these new developments and a contemporary outlook on the various frontiers of statistics Eminent leaders in the field have contributed 16 review articles and 6 research articles covering areas including semi

parametric models data analytical nonparametric methods statistical learning network tomography longitudinal data analysis financial econometrics time series bootstrap and other re sampling methodologies statistical computing generalized nonlinear regression and mixed effects models martingale transform tests for model diagnostics robust multivariate analysis single index models and wavelets This volume is dedicated to Prof Peter J Bickel in honor of his 65th birthday The first article The SAGE Handbook of Multilevel of this volume summarizes some of Prof Bickel's distinguished contributions Modeling Marc A. Scott, Jeffrey S. Simonoff, Brian D. Marx, 2013-08-31 In this important new Handbook the editors have gathered together a range of leading contributors to introduce the theory and practice of multilevel modeling The Handbook establishes the connections in multilevel modeling bringing together leading experts from around the world to provide a roadmap for applied researchers linking theory and practice as well as a unique arsenal of state of the art tools It forges vital connections that cross traditional disciplinary divides and introduces best practice in the field Part I establishes the framework for estimation and inference including chapters dedicated to notation model selection fixed and random effects and causal inference Part II develops variations and extensions such as nonlinear semiparametric and latent class models Part III includes discussion of missing data and robust methods assessment of fit and software Part IV consists of exemplary modeling and data analyses written by methodologists working in specific disciplines Combining practical pieces with overviews of the field this Handbook is essential reading for any student or researcher looking to apply multilevel techniques Marginal Models in Analysis of Correlated Binary Data with Time Dependent Covariates in their own research Jeffrey R. Wilson, Elsa Vazquez-Arreola, (Din) Ding-Geng Chen, 2020-09-28 This monograph provides a concise point of research topics and reference for modeling correlated response data with time dependent covariates and longitudinal data for the analysis of population averaged models highlighting methods by a variety of pioneering scholars While the models presented in the volume are applied to health and health related data they can be used to analyze any kind of data that contain covariates that change over time The included data are analyzed with the use of both R and SAS and the data and computing programs are provided to readers so that they can replicate and implement covered methods It is an excellent resource for scholars of both computational and methodological statistics and biostatistics particularly in the applied areas of Innovative Statistical Methods for Public Health Data Ding-Geng (Din) Chen, Jeffrey Wilson, 2015-08-31 The health book brings together experts working in public health and multi disciplinary areas to present recent issues in statistical methodological development and their applications This timely book will impact model development and data analyses of public health research across a wide spectrum of analysis Data and software used in the studies are available for the reader to replicate the models and outcomes The fifteen chapters range in focus from techniques for dealing with missing data with Bayesian estimation health surveillance and population definition and implications in applied latent class analysis to multiple comparison and meta analysis in public health data Researchers in biomedical and public health research will find this book



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