## **QUASI-LIKELIHOOD FUNCTIONS**

By Peter McCullagh, 1983



### Henrique Laureano (.github.io)

LEG @ UFPR

July 23, 2021



#### QUASI-LIKELIHOOD FUNCTIONS

#### By Peter McCullagh

#### Imperial College, London and University of British Columbia

The connection between quasi-likelihood functions, exponential family models and nonlinear weighted least squares is examined. Consistency and asymptotic normality of the parameter estimates are discussed under second moment assumptions. The parameter estimates are shown to satisfy a property of asymptotic optimality similar in spirit to, but more general than, the corresponding optimal property of Gauss-Markov estimators.



- Distinguished Professor in the Department of Statistics @ University of Chicago;
- 2 Completed his PhD at Imperial College London, supervised by David Cox and Anthony Atkinson;
- 3 Also at Imperial College London, was the PhD supervisor of Gauss Cordeiro.

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- Quasi-likelihood functions
- Properties of quasi-likelihood functions
- **5** Estimation of  $\sigma^2$
- 6 Examples of quasi-likelihood functions
- A higher order theory

### Introduction



- 1 Likelihood fucntion with exponential family form
  - 4 MLE through weighted least squares
  - variance (assumed) constant: we minimize a sum of squared residuals;
  - variance not constant: estimating equations can be thought as a generalization of the scoring method.
- 2 Likelihood function without exponential family form
  - ↓ In some cases: weighted least squares
    - Jorgensen, B. (1983). Maximum likelihood estimation and large sample inference for generalized linear and non-linear regression models. *Biometrika* 70

### **Paper purposes**

- 1 Maximize the likelihood function through weighted least squares
  - ↓ In which class of problems;
- 2 Weighted least squares under 2nd moment assumptions (quasi-likelihood).

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### A class of likelihood functions



Log likelihood written in the form :  $\sigma^{-2}\{\mathbf{y}^{\top}\theta - \mathbf{b}(\theta) - \mathbf{c}(\mathbf{y}, \sigma)\}$ 

#### The first two cumulants

By differentiating it and assuming that the support does not depend on  $\boldsymbol{\theta} \mapsto \boldsymbol{E}(\mathbf{Y}) = \boldsymbol{\mu} = \mathbf{b}'(\boldsymbol{\theta})$  and  $\mathsf{Cov}(\mathbf{Y}) = \sigma^2 \mathbf{b}''(\boldsymbol{\theta}) = \sigma^2 \mathbf{V}(\boldsymbol{\mu})$ .

In fact, the *r*th order cumulants of **Y** are given by  $\kappa_r = \sigma^{2r-2} \mathbf{b}^{(r)}(\theta)$ .

- 1 The first two cumulants describe the random component of the model;
- 2 However, in applications it is usually the systematic or nonrandom variation that is of primary importance

$$\downarrow E(\mathbf{Y}) = \mu = \mu(\beta)$$
 or  $E\{(\mathbf{h}(\mathbf{Y}))\} = \psi(\beta)$  (implicitly involving  $\sigma^2$ ).

### A class of likelihood functions



If  $\sigma^2$  is known, log-likelihood is an exponential family

- 4 variance and all higher order cumulants of **Y** are functions of the mean vector alone
  - 4 exponential, Poisson, multinomial, noncentral hypergeometric and partial likelihoods (survival analysis)
    - $\downarrow$  MLE of  $\beta$  through weighted least squares
      - $\downarrow \mu$  and  $\sigma^2$  are orthogonal
        - $\downarrow \beta$  and  $\sigma^2$  also orthogonal;

If  $\sigma^2$  is unknown, log-likelihood is

Frailty-based models for (multiple) survival experiences turn out in challengeable likelihood functions with inference routines mostly done via

- Elaborated and slow expectation—maximization (EM) algorithms;
- Inefficient Markov chain Monte Carlo (MCMC) schemes.

## $\textbf{Probability scale} \rightarrow \textbf{Cause-specific CIF}$



i.e.,  $CIF = \mathbb{P}[\text{ failure time } \leq t, \text{ a given cause } | \text{ features \& latent effects }].$ 

Common applications: family studies.

▶ Keywords: within-family/cluster dependence; age at disease onset; populations.

## Formally,



for a cause-specific of failure k, the cumulative incidence function (CIF) is defined as

$$F_k(t \mid \mathbf{x}) = \mathbb{P}[T \leqslant t, \ K = k \mid \mathbf{x}]$$

$$= \int_0^t f_k(z \mid \mathbf{x}) \, \mathrm{d}z \quad (f_k(t \mid \mathbf{x}) \text{ is the (sub)density for the time to a type } k \text{ failure})$$

$$= \int_0^t \underbrace{\lambda_k(z \mid \mathbf{x})}_{\text{cause-specific hazard function}} \underbrace{S(z \mid \mathbf{x})}_{\text{overall survival function}} dz, \quad t > 0, \quad k = 1, \dots, K.$$



Again, a comprehensive reference is @kalb&prentice's book.



## **SCHEIKE's CIF specification**



For two competing causes of failure, the cause-specific CIFs are specified in the following manner

$$F_k(t \mid \textbf{\textit{x}}, \ u_1, \ u_2, \ \eta_k) = \underbrace{\pi_k(\textbf{\textit{x}}, \ u_1, \ u_2)}_{\text{cluster-specific risk level}} \times \underbrace{\Phi[\textbf{\textit{w}}_k \textbf{\textit{g}}(t) - \textbf{\textit{x}} \gamma_k - \eta_k]}_{\text{cluster-specific failure time trajectory}}, \quad t > 0, \quad k = 1, \ 2, \quad (1)$$

with

$$\mathbf{1} \pi_k(\mathbf{x}, \mathbf{u}) = \exp\{\mathbf{x}\beta_k + u_k\} / \left(1 + \sum_{m=1}^{K-1} \exp\{\mathbf{x}\beta_m + u_m\}\right), \quad k = 1, 2, \quad K = 3;$$

- $\mathbf{Q} \Phi(\cdot)$  is the cumulative distribution function of a standard Gaussian distribution;
- 3  $g(t) = \operatorname{arctanh}(2t/\delta 1), \quad t \in (0, \delta), \quad g(t) \in (-\infty, \infty).$
- In @SCHEIKE, this CIF specification is modeled under a *challengeable* pairwise composite likelihood approach [@lindsay88; @varin11].

## Our contribution: a full likelihood analysis



For two competing causes of failure, a subject i, in the cluster j, in time t, we have

$$y_{ijt} \mid \underbrace{\{u_{1j}, u_{2j}, \eta_{1j}, \eta_{2j}\}}_{\text{latent effects}} \sim \text{Multinomial}(p_{1ijt}, p_{2ijt}, p_{3ijt})$$

$$\begin{bmatrix} u_1 \\ u_2 \\ \eta_1 \\ \eta_2 \end{bmatrix} \sim \begin{array}{l} \text{Multivariate} \\ \text{Normal} \end{array} \begin{pmatrix} \begin{bmatrix} 0 \\ 0 \\ 0 \\ 0 \end{bmatrix}, \begin{bmatrix} \sigma_{u_1}^2 & \text{cov}(u_1, u_2) & \text{cov}(u_1, \eta_1) & \text{cov}(u_1, \eta_2) \\ \sigma_{u_2}^2 & \text{cov}(u_2, \eta_1) & \text{cov}(u_2, \eta_2) \\ \sigma_{\eta_1}^2 & \text{cov}(\eta_1, \eta_2) \\ \sigma_{\eta_2}^2 \end{bmatrix} \end{pmatrix}$$

$$p_{kijt} = \frac{\partial}{\partial t} F_k(t \mid \boldsymbol{x}, \boldsymbol{u}, \eta_k)$$

$$= \frac{\exp\{\boldsymbol{x}_{kij}\boldsymbol{\beta}_{k} + u_{kj}\}}{1 + \sum_{m=1}^{K-1} \exp\{\boldsymbol{x}_{mij}\boldsymbol{\beta}_{m} + u_{mj}\}} \times w_{k} \frac{\delta}{2\delta t - 2t^{2}} \phi\left(w_{k}\operatorname{arctanh}\left(\frac{t - \delta/2}{\delta/2}\right) - \boldsymbol{x}_{kij}\gamma_{k} - \eta_{kj}\right), \quad k = 1, 2.$$

## Simulating from the model



# Marginal likelihood function for two competing causes



$$L(\theta; y) = \prod_{j=1}^{J} \int_{\mathfrak{R}^{4}} \pi(y_{j} \mid \mathbf{r}_{j}) \times \pi(\mathbf{r}_{j}) \, d\mathbf{r}_{j}$$

$$= \prod_{j=1}^{J} \int_{\mathfrak{R}^{4}} \left\{ \prod_{i=1}^{n_{j}} \prod_{t=1}^{n_{ij}} \left( \frac{\left(\sum_{k=1}^{K} y_{kijt}\right)!}{y_{1ijt}! y_{2ijt}! y_{3ijt}!} \prod_{k=1}^{K} \rho_{kijt}^{y_{kijt}} \right) \right\} \times$$
fixed effect component
$$(2\pi)^{-2} |\Sigma|^{-1/2} \exp\left\{ -\frac{1}{2} \mathbf{r}_{j}^{\top} \Sigma^{-1} \mathbf{r}_{j} \right\} d\mathbf{r}_{j}$$
latent effect component
$$= \prod_{j=1}^{J} \int \left\{ \prod_{t=1}^{n_{j}} \prod_{t=1}^{n_{ij}} \sum_{t=1}^{K} \rho_{tiit}^{y_{kijt}} \right\} (2\pi)^{-2} |\Sigma|^{-1/2} \exp\left\{ -\frac{1}{2} \mathbf{r}_{j}^{\top} \Sigma^{-1} \mathbf{r}_{i} \right\} d\mathbf{r}_{j}$$

$$= \prod_{j=1}^{J} \int_{\mathfrak{R}^4} \left\{ \underbrace{\prod_{i=1}^{H_{ij}} \prod_{t=1}^{H_{ij}} \prod_{k=1}^{K} p_{kijt}^{y_{kijt}}}_{\text{fixed effect}} \right\} \underbrace{(2\pi)^{-2} |\Sigma|^{-1/2} \exp\left\{-\frac{1}{2} \boldsymbol{r}_{j}^{\top} \Sigma^{-1} \boldsymbol{r}_{j}\right\}}_{\text{latent effect component}} \mathrm{d}\boldsymbol{r}_{j},$$

with  $p_{kijt}$  from Equation 2 and where  $\theta = [\beta \ \gamma \ \mathbf{w} \ \sigma^2 \ \rho]^{\top}$  is the parameters vector.

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# TMB: Automatic Differentiation and Laplace Approximation





An R [@R21] package for the quickly implementation of complex random effect models through simple C++ templates.

#### **Workflow**

- Write your objective function in a .cpp through a #include <TMB.hpp>;
- 2 Compile and load it in R via TMB::compile() and base::dyn.load(TMB::dynlib());
- 3 Compute your objective function derivatives with obj <- TMB::MakeADFun();</p>
- Operform the model fitting, opt <- base::nlminb(obj\$par, obj\$fn, obj\$gr);</pre>
- **5** Compute the parameters standard deviations, TMB::sdreport(obj).



For details about TMB, AD, and Laplace approximation: @laurence.

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## Simulation study model designs



#### Risk model

Latent effects only on the risk level i.e.,

$$\Sigma = \begin{bmatrix} \sigma_{u_1}^2 & \mathsf{COV}_{u_1, u_2} \\ & \sigma_{u_2}^2 \end{bmatrix}.$$

### **Block-diag model**

Latent effects on the risk and time levels without cross-correlations i.e.,

$$\Sigma = \begin{bmatrix} \sigma_{u_1}^2 & \cos_{u_1,u_2} & 0 & 0 \\ & \sigma_{u_2}^2 & 0 & 0 \\ & & \sigma_{\eta_1}^2 & \cos_{\eta_1,\eta_2} \\ & & & \sigma_{\eta_2}^2 \end{bmatrix}$$

#### Time model

Latent effects only on the failure time trajectory level i.e.,

$$\Sigma = egin{bmatrix} \sigma_{\eta_1}^2 & \mathsf{cov}_{\eta_1,\eta_2} \ \sigma_{\eta_2}^2 \end{bmatrix}.$$

### Complete model

A complete latent effects structure i.e..

$$\Sigma = \begin{bmatrix} \sigma_{u_1}^2 & \text{cov}_{u_1,u_2} & 0 & 0 \\ & \sigma_{u_2}^2 & 0 & 0 \\ & & \sigma_{\eta_1}^2 & \text{cov}_{\eta_1,\eta_2} \\ & & & \sigma_{\eta_2}^2 \end{bmatrix}. \qquad \Sigma = \begin{bmatrix} \sigma_{u_1}^2 & \text{cov}_{u_1,u_2} & \text{cov}_{u_1,\eta_1} & \text{cov}_{u_1,\eta_2} \\ & \sigma_{u_2}^2 & \text{cov}_{u_2,\eta_1} & \text{cov}_{u_2,\eta_2} \\ & & & \sigma_{\eta_1}^2 & \text{cov}_{\eta_1,\eta_2} \\ & & & & \sigma_{\eta_2}^2 \end{bmatrix}.$$

## Simulation study setup

#### Four latent effects structures:



Risk model;

2 Time model;

3 Block-diag model;

4 Complete model.

Two CIF configurations:

**Low** max incidence  $\approx 0.15$ ;

**High** max incidence  $\approx$  0.60.

For each of those  $4 \times 2 = 8$  scenarios, we vary the sample and cluster sizes:

#### 5000 data points

- 2500 clusters of size 2;
- 1000 clusters of **size 5**;
- 500 clusters of size 10.

### 30000 data points

- 15000 clusters of size 2;
- 6000 clusters of **size 5**;
- 3000 clusters of size 10.

### 60000 data points

- 30000 clusters of size 2;
- 12000 clusters of **size 5**;
- 6000 clusters of **size 10**.

Totalizing,  $8 \times 3 \times 3 = 72$  scenarios.

For each scenario, we simulate 500 samples, totalizing  $72 \times 500 = 36000$  model fittings.

# Simulation study results



#### First of all, the **time**.

- The non-complete models (2D Laplace aprox.) are kind of fast, taking always less than 5 min.
- In the most expensive scenarios (30K 4D Laplaces),
   the complete model takes 30 min.
   In a full R implementation with 10K 4D Laplaces, it took 30hrs. TMB is fast.
- We also did a Bayesian analysis via Stan/NUTS-HMC [@RStan].
  - 1 week of parallelized processing for a 2500 size 2 clusters scenario with tuned NUTS.
     This just reinforces the MCMC impracticability for some complex models.

#### Parameters estimation.

The non-complete models fail to learn the data.
 They appear to be not structured enough to capture the data characteristics.

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### Take-home message



- The complete model works. It's not magnificent, but it works.
  - 1 It works better in the high CIF scenarios;
  - 2 As expected, as the sample size increases the results get better;
  - 3 We do not see any considerable performance difference between cluster/family sizes;
  - 4 Satisfactory full likelihood analysis under the maximum likelihood estimation framework (the estimates bias-variance could be smaller).

#### What else can we do?

- 1 Instead of a conditional approach (latent effects model), we can try a marginal approach e.g., an McGLM [@mcglm];
- 2 We can also try a copula [@copulas], on maybe two fronts:1) for a full specification; 2) to accommodate the within-cluster dependence.



For more read @laurence master thesis.

# Thanks for watching and have a great day



Special thanks to

