Measurement Error and Misclassification in statistical models: Basics and applications beam Bilbao Part 3

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Methods

- ► Functional and structural
- ► Correction and method of moments and orthogonal regression
- ► Regression calibration
- ► Likelihood
- ► Quasi likelihood
- ► Bayes

Functional and structural

► Functional:

X fixed unknown constants

No assumptions about the distribution of X

► **Structural:** *X* latent random variable
Use assumptions about the distribution of *X*

Method of moments

Moments of observed data can be estimated Solve moments equations Simple linear regression:

$$\begin{array}{rcl} \mu_{X^*} & = & \mu_X \\ \mu_Y & = & \beta_0 + \beta_1 \mu_x \\ \sigma_{x^*}^2 & = & \sigma_x^2 + \sigma_u^2 \\ \sigma_y^2 & = & \beta_1^2 \sigma_x^2 + \sigma_\epsilon^2 \\ \sigma_{yx^*} & = & \beta_1 * \sigma_x^2 \end{array}$$

Orthogonal regression, total least squares

In linear Regression with classical additive measurement error:

- ► Assume $\frac{\sigma_{\epsilon}^2}{\sigma_{\mu}^2} = \eta$ is known,
 - e. g. no equation error and σ_ϵ is measurement error in Y. Minimize

$$\sum_{i=1}^{n} \left\{ (Y_i - \beta_0 - \beta_1 X_i)^2 + \eta (X_i^* - X_i)^2 \right\}$$

in
$$(\beta_0, \beta_1, X_1, X_2, \dots, X_n)$$
.

- ► Total least squares, Van Huffel (1997)
- ► Technical symmetric applications
- In other applications a problem, assumption of no equation error not realistic

Regression calibration

This simple method has been widely applied. It was suggested by different authors: Rosner et al. (1989) Carroll and Stefanski(1990)

- 1. Find a model for $E(X|X^*,Z)$ by validation data or replication
- 2. Replace the unobserved X by estimate $E(X|X^*,Z)$ in the main model
- 3. Adjust variance estimates by bootstrap or asymptotic methods
- ► Good method in many practical situations
- Calibration data can be incorporated
- ► Problems in highly nonlinear models

Regression calibration

- ▶ Berkson case: $E(X|X^*) = X^* \longrightarrow$ Naive estimation = Regression calibration
- ightharpoonup Classical : Linear regression X on X^*

$$E(X|X^*) = \frac{\sigma_X^2}{\sigma_{X^*}^2} * X^* + \mu_X * (1 - \frac{\sigma_X^2}{\sigma_{X^*}^2})$$

Correction for attenuation in linear model

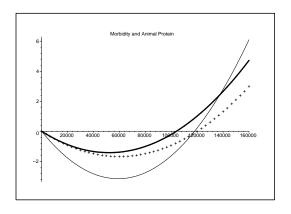
Survival

For Cox Model and rare disease assumption appropriate **Example: MONICA study, Augustin(2002)**

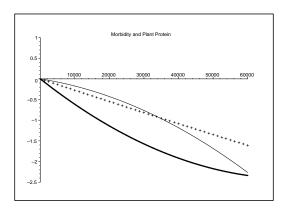
- ► CHD and fat intake
- ► Cox-Regression
- ► Quadratic model
- ► Classical additive measurement error
- ► Heteroscedastic measurement error
- ▶ Replication (7 days) for estimating measurement error variance

Results differ for assumption of homoscedastic and heteroscedastic measurement error

Results: Animal protein



Plant protein:



Likelihood methods

- Standard inference can be done with standard errors and likelihood ratio tests
- ► Efficiency
- ► Combination of different data types are possible
- ► Sometimes more accurate than approximations
- ▶ Difficult to calculate
- ► Software not available
- ▶ Parametric model for the unobserved predictor necessary
- ► Robustness to strong parametric assumptions

The classical error likelihood

$$\begin{split} [\mathbf{Y}, \mathbf{X}^* | \mathbf{Z}, \theta] &= \prod_{i=1}^n \int [y_i | x_i, z_i, \beta] [x_i^* | x_i, \eta] [x_i | z_i, \lambda] d\mu(x), \\ \text{where} \qquad \theta &= (\beta, \eta, \lambda) \end{split}$$

► Evaluation by numerical integration

Berkson likelihood

3 components, but the third component contains no information

$$[\mathbf{Y}, \mathbf{X}^* | \mathbf{Z}, \theta] = \prod_{i=1}^n \int [y_i | x_i, z_i, \beta] [x_i | x_i^*, \eta] [x_i | z_i, \lambda] d\mu(x)$$
$$[\mathbf{Y}, \mathbf{X}^* | \mathbf{Z}, \theta] = \prod_{i=1}^n \int [y_i | x_i, z_i, \beta] [x_i | x_i^*, \eta] d\mu(x) * const$$

The Berkson likelihood does not depend on the exposure model

Quasi likelihood

If calculation of the likelihood is too complicated use

$$E(Y|X^*, Z) = \int g(X, Z) f_{x|X^*} dx$$

$$V(Y|X^*, Z) = \int v(X, Z) f_{x|X^*} dx + Var[g(X, Z)|X^*]$$

This can be done e.g. for exponential g:

- ► Poisson regression model
- ► Parametric survival

Bayes

Richardson and Green (2002)

- ► Evaluation by MCMC techniques
- ► Conditional independence assumptions on the three models parts as seen in the likelihood approach
- ▶ The latent variable *X* is treated an unknown parameter
- ▶ Different data types can be combined
- ▶ Priori distributions for the error model
- ► Flexible handling of the exposure model

Case study: Occupational Dust and chronic bronchitis

HK/Carroll (1997) and Goessl /HK(2001)

Research question: Relationship between occupational dust and chronic bronchitis

Data form N=1246 workers:

X: log(1+average occupational dust exposure)

Y: Chronic bronchitis (CBR)

 X^* : Measurements and expert ratings

 Z_1 : Smoking

 Z_2 : Duration of exposure

No validation or replication data available!

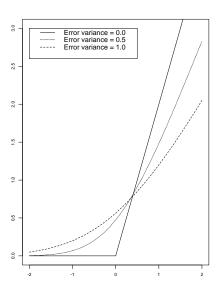
The Model

Segmented logistic regression an unknown threshold limiting value (TLV) τ

$$P(Y = 1 | X = x, Z = z) = G(z'\beta_{k-} + \beta_k(x - \tau)_+),$$

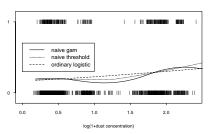
 $(x - \tau)_+ = \max(0, x - \tau).$

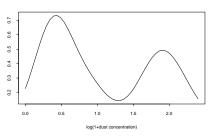
Effect of measurement error



Naive analysis

Munich Data





Likelihood

- ► Probit approximation
- ► Calculation of the integrals
- ▶ Assumption of a mixture of two normals for the exposure model
- ► Fixed additive measurement error

Regression calibration

- ▶ Assumption of a mixture of two normals for the exposure model
- ► Fixed additive measurement error

Results

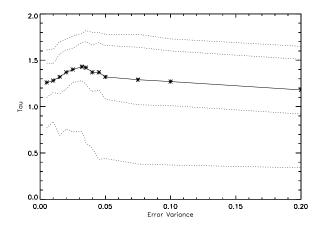
| Method | TLV- $	au_0$ | Nom s. e. | boot s.e. |
|------------------------|--------------|-----------|-----------|
| Naive | 1.27 | .41 | .24 |
| Pseudo-MLE | 1.76 | .17 | .21 |
| Regression Calibration | 1.75 | .12 | .19 |

Tabelle: Estimated TLV in the Munich data, when $\sigma_u^2 = 0.035$.

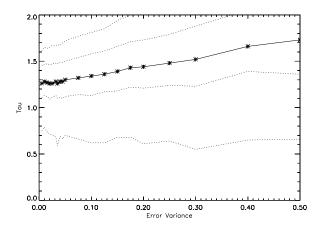
Bayes

- ► Fixed additive measurement error: Sensitivity analysis
- ▶ flat priori for measurement error :No convergence
- ► Assumption of mixture of normals for exposure model
- ▶ Both models Berkson and additive

Results: Estimation of TLV:Classical



Berkson



Conclusions

- 1. Measurement model essential: High Difference between Effect of Berkson and classical measurement error in most cases!
- 2. Additive classical non differential measurement error leads to attenuation
- 3. Many methods available
- 4. Regression calibration works in many cases
- 5. ML should be taken into account for Berkson error
- 6. Bayesian analysis is useful especially if model structure is complex