# glms: a Transformative Paradigm for Statistical Practice and Education

#### John Hinde

Statistics Group, School of Mathematics, Statistics and Applied Mathematics National University of Ireland, Galway john.hinde@nuigalway.ie

Research Supported by SFI Award 07/MI/012

Imperial College, London

28 March 2015







# Summary

- The 1972 Paper
  - Software
- 2 Spreading the word
- 3 Extensions
  - Random effects
  - Overdispersion & Zero-Inflation
- 4 Examples
  - Count data
  - Multinomial
- 5 Education

#### 6 Acknowledgements

- ∢ ∃ ▶

#### The Paper — 1972

(日) (四) (三) (三) (三)

The Paper — 1972

J. R. Statist. Soc. A, (1972), 135, Part 3, p. 370 370

#### **Generalized Linear Models**

By J. A. NELDER and R. W. M. WEDDERBURN

Rothamsted Experimental Station, Harpenden, Herts

イロト イポト イヨト イヨト

The Paper — 1972

J. R. Statist. Soc. A, (1972), 135, Part 3, p. 370 370

#### **Generalized Linear Models**

By J. A. NELDER and R. W. M. WEDDERBURN

Rothamsted Experimental Station, Harpenden, Herts

• published in Series A

イロト イポト イヨト イヨト

The Paper — 1972

J. R. Statist. Soc. A, (1972), 135, Part 3, p. 370 370

#### Generalized Linear Models

By J. A. NELDER and R. W. M. WEDDERBURN

Rothamsted Experimental Station, Harpenden, Herts

- published in Series A
- 15 pages long

イロト イポト イヨト イヨト

The Paper — 1972

J. R. Statist. Soc. A, (1972), 135, Part 3, p. 370 370

#### Generalized Linear Models

By J. A. NELDER and R. W. M. WEDDERBURN

Rothamsted Experimental Station, Harpenden, Herts

- published in Series A
- 15 pages long
- many examples over half the paper

(日) (周) (三) (三)

The Paper — 1972

J. R. Statist. Soc. A, (1972), 135, Part 3, p. 370 370

#### Generalized Linear Models

By J. A. NELDER and R. W. M. WEDDERBURN

Rothamsted Experimental Station, Harpenden, Herts

- published in Series A
- 15 pages long
- many examples over half the paper
- "useful way of unifying ... unrelated statistical procedures"

#### John Nelder: 1924 — 2010

- Statistician at National Vegetable Research Station (NVRS), now Horticultural Research International, Wellesbourne — 1949-68
  - theory of general balance unifying framework for the wide range of designs in agricultural experimentation
  - initial work on GenStat

(日) (同) (三) (三)

#### John Nelder: 1924 — 2010

- Statistician at National Vegetable Research Station (NVRS), now Horticultural Research International, Wellesbourne — 1949-68
  - theory of general balance unifying framework for the wide range of designs in agricultural experimentation
  - initial work on GenStat
- Head of the Statistics Department at Rothamsted 1968-1984
  - theory of generalized linear models, with the late Robert Wedderburn
  - Applied Statistics Algorithms in Applied Statistics, JRSSC
  - further development of GenStat, with NAG
  - development of GLIM, first released in 1974

イロト イヨト イヨト

#### John Nelder: 1924 — 2010

- Statistician at National Vegetable Research Station (NVRS), now Horticultural Research International, Wellesbourne — 1949-68
  - theory of general balance unifying framework for the wide range of designs in agricultural experimentation
  - initial work on GenStat
- Head of the Statistics Department at Rothamsted 1968-1984
  - theory of generalized linear models, with the late Robert Wedderburn
  - Applied Statistics Algorithms in Applied Statistics, JRSSC
  - further development of GenStat, with NAG
  - development of GLIM, first released in 1974
- visiting Professor at Imperial College 1972-2009
  - GLIMPSE "expert system" based on GLIM
  - theory of hierarchical generalized linear models (HGLMs), with Youngjo Lee

< 由 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < 回 > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ > < □ >

#### John Nelder: 1924 — 2010

- Statistician at National Vegetable Research Station (NVRS), now Horticultural Research International, Wellesbourne — 1949-68
  - theory of general balance unifying framework for the wide range of designs in agricultural experimentation
  - initial work on GenStat
- Head of the Statistics Department at Rothamsted 1968-1984
  - theory of generalized linear models, with the late Robert Wedderburn
  - Applied Statistics Algorithms in Applied Statistics, JRSSC
  - further development of GenStat, with NAG
  - development of GLIM, first released in 1974
- visiting Professor at Imperial College 1972-2009
  - GLIMPSE "expert system" based on GLIM
  - theory of hierarchical generalized linear models (HGLMs), with Youngjo Lee

#### Robert Wedderburn: 1947 — 1975

• Died aged 28 of anaphylactic shock from an insect bite.

# John Nelder: 1924 — 2010



John Hinde (NUIG)







John at the Rothamsted Conference in 2004

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)

- 4 @ > - 4 @ > - 4 @ >

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)
- Probit analysis: Finney (1952) maximum likelihood for tolerance distribution in toxicology

< 回 > < 三 > < 三 >

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)
- Probit analysis: Finney (1952) maximum likelihood for tolerance distribution in toxicology
- Dyke & Patterson (1952): logit model for analysis of proportions in factorial experiment

(日) (周) (三) (三)

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)
- Probit analysis: Finney (1952) maximum likelihood for tolerance distribution in toxicology
- Dyke & Patterson (1952): logit model for analysis of proportions in factorial experiment
- transformations to linearity

(日) (周) (三) (三)

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)
- Probit analysis: Finney (1952) maximum likelihood for tolerance distribution in toxicology
- Dyke & Patterson (1952): logit model for analysis of proportions in factorial experiment
- transformations to linearity
- Box-Cox transformation (1964)

- 4 同 6 4 日 6 4 日 6

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)
- Probit analysis: Finney (1952) maximum likelihood for tolerance distribution in toxicology
- Dyke & Patterson (1952): logit model for analysis of proportions in factorial experiment
- transformations to linearity
- Box-Cox transformation (1964)
- Inverse polynomials, Nelder (1966)

- 4 同 6 4 日 6 4 日 6

- analysis of non-normal data variance stabilising transformation of the response
  - Poisson count data: square-root transformation,  $\sqrt{y}$
  - Binomial proportions: arc-sin-square-root,  $\sin^{-1}(\sqrt{y})$
  - Exponential times: log transformation, log(y)
- Probit analysis: Finney (1952) maximum likelihood for tolerance distribution in toxicology
- Dyke & Patterson (1952): logit model for analysis of proportions in factorial experiment
- transformations to linearity
- Box-Cox transformation (1964)
- Inverse polynomials, Nelder (1966)
- Nelder (1968): ... one transformation leads to a linear model and another to normal error.

# glms — the idea

(日) (四) (三) (三) (三)

#### glms — the idea

Gauss - You were one of the discussants of the Box-Cox 1964 paper and you also introduced the idea of inverse polynomials in 1966. How did you get the idea of Generalized Linear Models?

Nelder - That's an interesting question. I don't really think I know the answer to it. There is a paper that I wrote in 1970 which was published in Biometrics; in this I drew attention to the fact that there was a considerable similarity between a model with gamma errors and an inverse linear response curve and the model for Probit Analysis. I didn't understand at that time exactly what the connection was, though I could see there was one. Then in the subsequent two years somehow the idea jelled, so that Wedderburn and I could see what was common to these models. That's how it came about, but exactly how I did it I don't know. Similarly in the General Balance papers I first had the idea in a

• Intro: background (2 pages)

- Intro: background (2 pages)
  - random component: 1-parameter exponential family
  - linear predictor:  $\eta = \beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p$
  - link function:  $g(\mu) = \eta$

- Intro: background (2 pages)
  - random component: 1-parameter exponential family
  - linear predictor:  $\eta = \beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p$
  - link function:  $g(\mu) = \eta$
- Model fitting: (3 pages)
  - maximum likelihood estimation using Fisher Scoring

Iteratively (Re)-Weighted Least Squares

- Intro: background (2 pages)
  - random component: 1-parameter exponential family
  - linear predictor:  $\eta = \beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p$
  - link function:  $g(\mu) = \eta$
- Model fitting: (3 pages)
  - maximum likelihood estimation using Fisher Scoring

Iteratively (Re)-Weighted Least Squares

• sufficient statistics — canonical links

- Intro: background (2 pages)
  - random component: 1-parameter exponential family
  - linear predictor:  $\eta = \beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p$
  - link function:  $g(\mu) = \eta$
- Model fitting: (3 pages)
  - maximum likelihood estimation using Fisher Scoring

Iteratively (Re)-Weighted Least Squares

- sufficient statistics canonical links
- Analysis of Deviance

minimal  $\leftrightarrow$  complete (saturated) models

- Intro: background (2 pages)
  - random component: 1-parameter exponential family
  - linear predictor:  $\eta = \beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p$
  - link function:  $g(\mu) = \eta$
- Model fitting: (3 pages)
  - maximum likelihood estimation using Fisher Scoring

Iteratively (Re)-Weighted Least Squares

- sufficient statistics canonical links
- Analysis of Deviance

minimal  $\leftrightarrow$  complete (saturated) models

• Special distributions, examples (6 pages)

イロン 不聞と 不同と 不同と

- Intro: background (2 pages)
  - random component: 1-parameter exponential family
  - linear predictor:  $\eta = \beta_0 + \beta_1 x_1 + \cdots + \beta_p x_p$
  - link function:  $g(\mu) = \eta$
- Model fitting: (3 pages)
  - maximum likelihood estimation using Fisher Scoring

Iteratively (Re)-Weighted Least Squares

- sufficient statistics canonical links
- Analysis of Deviance

minimal  $\leftrightarrow$  complete (saturated) models

- Special distributions, examples (6 pages)
- Models in Teaching Statistics (1 page)

 Normal: observations normal on log-scale; additive effects on inverse scale

<ロト </p>

- Normal: observations normal on log-scale; additive effects on inverse scale
- Poisson: Fisher's tuberculin-test data Latin square of counts

< ロ > < 同 > < 三 > < 三

- Normal: observations normal on log-scale; additive effects on inverse scale
- Poisson: Fisher's tuberculin-test data Latin square of counts
- Poisson: multinomial distributions for contingency tables

□ ▶ ▲ □ ▶ ▲ □

- Normal: observations normal on log-scale; additive effects on inverse scale
- Poisson: Fisher's tuberculin-test data Latin square of counts
- Poisson: multinomial distributions for contingency tables
- Binomial: Probit & Logit models

- Normal: observations normal on log-scale; additive effects on inverse scale
- Poisson: Fisher's tuberculin-test data Latin square of counts
- Poisson: multinomial distributions for contingency tables
- Binomial: Probit & Logit models
- Gamma: estimation of variance components in incomplete block design

□ ▶ ▲ □ ▶ ▲ □

# John Nelder & Statistical Computing

• Anti black-box packages

3

(二)、

# John Nelder & Statistical Computing

- Anti black-box packages
- User should be in control

< A > < 3
- Anti black-box packages
- User should be in control
- Default output should be minimal

- Anti black-box packages
- User should be in control
- Default output should be minimal
- System should not allow stupid models marginality

A⊒ ▶ < ∃

- Anti black-box packages
- User should be in control
- Default output should be minimal
- System should not allow stupid models marginality
- Model specification using Wilkinson & Rogers formulæ

- Anti black-box packages
- User should be in control
- Default output should be minimal
- System should not allow stupid models marginality
- Model specification using Wilkinson & Rogers formulæ
- All structures available to the user input to other routines

#### Software

- Anti black-box packages
- User should be in control
- Default output should be minimal
- System should not allow stupid models marginality
- Model specification using Wilkinson & Rogers formulæ
- All structures available to the user input to other routines
- system should be open user extendible (GLIM, GenStat, S/R, ...)

#### Software

- Anti black-box packages
- User should be in control
- Default output should be minimal
- System should not allow stupid models marginality
- Model specification using Wilkinson & Rogers formulæ
- All structures available to the user input to other routines
- system should be open user extendible (GLIM, GenStat, S/R, ...)
- Requires user expertise/knowledge

#### Software

# John Nelder & Statistical Computing

- Anti black-box packages
- User should be in control
- Default output should be minimal
- System should not allow stupid models marginality
- Model specification using Wilkinson & Rogers formulæ
- All structures available to the user input to other routines
- system should be open user extendible (GLIM, GenStat, S/R, ...)
- Requires user expertise/knowledge

### Principles embodied in GLIM

— a system specifically for fitting glms.

- 32

- [i] ? \$yvar days \$error p \$
- [i] ? \$fit A\*S\*C\*L \$
- [o] scaled deviance = 1173.9 at cycle 4
- [o] residual df = 118

E SQA

- [i] ? \$yvar days \$error p \$
- [i] ? \$fit A\*S\*C\*L \$
- [o] scaled deviance = 1173.9 at cycle 4
- [o] residual df = 118
- Or, in John's preferred style ...

E Sac

- [i] ? \$yvar days \$error p \$
- [i] ? \$fit A\*S\*C\*L \$
- [o] scaled deviance = 1173.9 at cycle 4
- [o] residual df = 118
- Or, in John's preferred style ...
- [i] ? \$y days \$e p \$
- [i] ? \$f A\*S\*C\*L \$

• Conferences — "That's a glm!"

(日) (四) (三) (三) (三)

- Conferences "That's a glm!"
- Nelder (1984) Models for Rates with Poisson Errors: In a recent paper, Frome (1983) described the fitting of models with Poisson errors and data in the form of rates ... fitted simply by GLIM ... or the use of a program that handles iterative weighted least squares

- Conferences "That's a glm!"
- Nelder (1984) Models for Rates with Poisson Errors: In a recent paper, Frome (1983) described the fitting of models with Poisson errors and data in the form of rates ... fitted simply by GLIM ... or the use of a program that handles iterative weighted least squares
- Nelder (1991) Generalized Linear Models for Enzyme-Kinetic Data: Ruppert, Cressie, and Carroll (1989) discuss various models for fitting the Michaelis-Menten equations to data on enzyme kinetics. I find it surprising that they do not include, among the models they consider, generalized linear models (GLMs) with an inverse link

- Conferences "That's a glm!"
- Nelder (1984) Models for Rates with Poisson Errors: In a recent paper, Frome (1983) described the fitting of models with Poisson errors and data in the form of rates ... fitted simply by GLIM ... or the use of a program that handles iterative weighted least squares
- Nelder (1991) Generalized Linear Models for Enzyme-Kinetic Data: Ruppert, Cressie, and Carroll (1989) discuss various models for fitting the Michaelis-Menten equations to data on enzyme kinetics. I find it surprising that they do not include, among the models they consider, generalized linear models (GLMs) with an inverse link The data-transformation approach suffers from the disadvantage that normality of errors and linearity of systematic effects are still being sought simultaneously

John Hinde (NUIG)

### Generalized Linear Models — Monograph



# Statistical Modelling in GLIM (1989)

An applied how to text with integrated GLIM code.

3

# Statistical Modelling in GLIM (1989)

An applied how to text with integrated GLIM code.

- normal models
  - regression
  - analysis of variance
- binomial responses
- multinomial and Poisson
  - count data
  - multiway tables
- survival models
  - parametric
  - Cox PH piecewise exponential
  - discrete time



### GLIM Conferences, IWSM, Statistical Modelling

- GLIM conferences really on glms
- IWSM: International Workshop on Statistical Modelling
- Eventually led to Statistical Modelling Society



### Statistical Modelling Journal

### In 2000, founding of journal **Statistical Modelling** availability of data and code with papers $\rightarrow$ *reproducible research*

Statistical Modelling: An International Journal

http://stat.uibk.ac.at/SMIJ



Aims and Scope

For Authors

Archives

Modelling Society



### Statistical Modelling: An International Journal

#### STATISTICAL MODELLING

AN INTERNATIONAL JOURNAL





Statistical Modelling: An International Journal publishes original and high-quality articles that recognize statistical modelling as the general framework for the application of statistical ideas. Submissions must reflect important developments, extensions, and applications in statistical modelling. The journal also encourages submissions that describe scientificially interesting, complex or novel statistical modelling aspects from a wide diversity of disciplines, and submissions that embrace the diversity of applied statistical modelling.

Indexed by Science Citation Index Expanded, ISI Alerting Services, and CompuMath Citation Index, beginning with volume 3 (2003).

28 March 2015 16 / 49

### Extending the basic glm

#### • response distribution

- multivariate vector of responses
- exponential dispersion models
- generalized distributions
- quasi-distributions
- mixtures
- joint responses: longitudinal + time to event, ...

### Extending the basic glm

#### response distribution

- multivariate vector of responses
- exponential dispersion models
- generalized distributions
- quasi-distributions
- mixtures
- joint responses: longitudinal + time to event, ...

### Inear predictor

- smooth terms gams, etc
- random effects
- multiple linear predictors modelling mean and dispersion, gamlss, etc

### Extending the basic glm

### response distribution

- multivariate vector of responses
- exponential dispersion models
- generalized distributions
- quasi-distributions
- mixtures
- joint responses: longitudinal + time to event, ...

### Iinear predictor

- smooth terms gams, etc
- random effects
- multiple linear predictors modelling mean and dispersion, gamlss, etc

### Ink function

- parametric links
- composite link functions (Thompson & Baker, 1981)
- non-linear glms gnm (Turner & Firth, 2012)

イロン 不聞と 不同と 不同と

### Normal Models

$$\mathbf{y} = \boldsymbol{eta}^{\mathsf{T}} \mathbf{x} + \boldsymbol{\epsilon}$$

- single error term includes
  - individual observation/measurement error
  - experimental unit variability
  - unobserved covariates

3

イロト イヨト イヨト イヨト

### Normal Models

$$\mathbf{y} = \boldsymbol{eta}^{\mathsf{T}} \mathbf{x} + \boldsymbol{\epsilon}$$

- single error term includes
  - individual observation/measurement error
  - experimental unit variability
  - unobserved covariates

• for simplest data structures/designs use normal linear model

### Normal Models

$$\mathbf{y} = \boldsymbol{eta}^{\mathsf{T}} \mathbf{x} + \boldsymbol{\epsilon}$$

- single error term includes
  - individual observation/measurement error
  - experimental unit variability
  - unobserved covariates
- for simplest data structures/designs use normal linear model
- more complex situations
  - structure in experimental unit variability
  - repeated measures/longitudinal observations
  - ...

一日、

### Normal Mixed Model

$$\mathbf{y} = \boldsymbol{eta}^{\mathsf{T}} \mathbf{x} + \boldsymbol{\gamma}^{\mathsf{T}} \mathbf{z} + \boldsymbol{\epsilon}$$

### • z unobserved random effects

(日) (周) (日) (日)

### Normal Mixed Model

$$\mathbf{y} = oldsymbol{eta}^\mathsf{T} \mathbf{x} + oldsymbol{\gamma}^\mathsf{T} \mathbf{z} + oldsymbol{\epsilon}$$

- z unobserved random effects
- shared random effects
  - multi-level/variance components models
  - longitudinal observations
  - spatial structure

### Normal Mixed Model

$$\mathbf{y} = oldsymbol{eta}^\mathsf{T} \mathbf{x} + oldsymbol{\gamma}^\mathsf{T} \mathbf{z} + oldsymbol{\epsilon}$$

- z unobserved random effects
- shared random effects
  - multi-level/variance components models
  - longitudinal observations
  - spatial structure
- z normal
  - normal model with structured covariance matrix
- standard mixed model analyses ML, REML
- widely available in standard software

一日、

### Generalized Linear Models

Models for counts, proportions, times, ...

$$\mathbf{y} \sim \mathcal{F}(oldsymbol{\mu}) \qquad g(oldsymbol{\mu}) = oldsymbol{\eta} = oldsymbol{eta}^\mathsf{T} \mathbf{x}$$

- distributional assumption relates to the observation/measurement process
- how does this model incorporate
  - experimental/individual unit variability?
  - unobserved covariates?

## Generalized Linear Models

Models for counts, proportions, times, ...

$$\mathbf{y} \sim \mathcal{F}(oldsymbol{\mu}) \qquad g(oldsymbol{\mu}) = oldsymbol{\eta} = oldsymbol{eta}^\mathsf{T} \mathbf{x}$$

- distributional assumption relates to the observation/measurement process
- how does this model incorporate
  - experimental/individual unit variability?
  - unobserved covariates?

### It doesn't!

## Generalized Linear Models

Models for counts, proportions, times, ...

$$\mathbf{y} \sim \mathcal{F}(oldsymbol{\mu}) \qquad g(oldsymbol{\mu}) = oldsymbol{\eta} = oldsymbol{eta}^\mathsf{T} \mathbf{x}$$

- distributional assumption relates to the observation/measurement process
- how does this model incorporate
  - experimental/individual unit variability?
  - unobserved covariates?

### It doesn't!

hence overdispersion, etc

Include random effect(s) in the linear predictor

$$oldsymbol{\eta} = oldsymbol{eta}^{ op} \mathbf{x} + oldsymbol{\gamma}^{ op} \mathbf{z}$$

3

・ロン ・四 ・ ・ ヨン ・ ヨン

Include random effect(s) in the linear predictor

$$\boldsymbol{\eta} = \boldsymbol{eta}^{\mathsf{T}} \mathbf{x} + \boldsymbol{\gamma}^{\mathsf{T}} \mathbf{z}$$

- single conjugate random effect at individual level standard overdispersion models
  - negative binomial for count data
  - beta-binomial for proportions

Include random effect(s) in the linear predictor

$$\boldsymbol{\eta} = \boldsymbol{eta}^{\mathsf{T}} \mathbf{x} + \boldsymbol{\gamma}^{\mathsf{T}} \mathbf{z}$$

- single conjugate random effect at individual level standard overdispersion models
  - negative binomial for count data
  - beta-binomial for proportions
- $\bullet \ z \ \text{normal} \longrightarrow generalized \ linear \ mixed \ models$

・ 同 ト ・ ヨ ト ・ ヨ ト

Include random effect(s) in the linear predictor

$$oldsymbol{\eta} = oldsymbol{eta}^{\mathsf{T}} \mathsf{x} + oldsymbol{\gamma}^{\mathsf{T}} \mathsf{z}$$

- single conjugate random effect at individual level standard overdispersion models
  - negative binomial for count data
  - beta-binomial for proportions
- z normal  $\longrightarrow$  generalized linear mixed models
- ullet z unspecified  $\longrightarrow$  nonparametric maximum likelihood

・ 「 ・ ・ ・ ・ ・ ・ ・
# John's Approach (1984)

Additional random term in licen predictor  
Suppose that the linear predictor of diord be modelled  
by 
$$\eta + \varepsilon$$
, when  $\varepsilon$  is a random variable into variance  $\theta$ .  
 $\theta$  is the between-group or higher-test component of  
variance. Taking expectations over the lower terd  
error we have  
 $E_1(\eta) = \mu = g(\eta + \varepsilon) \approx g(\eta) + g'(\eta) \varepsilon$   
Hence assuming independence between the two error  
components we have  
 $var(\eta) = var_1(\eta) + g'^2 \theta$   
 $\equiv V(\mu) + \theta g'^2$  for a GLM.  
In particular for a canonical link where  $g' = V$ , we  
have  $var(\eta) \equiv V(\mu) + \theta V'(\mu)$ . (1)

< ∃→

#### Overdispersion & Zero-Inflation

# Motivating Application

- 4x2 factorial micropropagation experiment of the apple variety Trajan – a 'columnar' variety.
- Shoot tips of length 1.0-1.5 cm were placed in jars on a standard culture medium.
- 4 concentrations of cytokinin BAP added High concentrations of BAP often inhibit root formation during micropropagation of apples, but maybe not for 'columnar' varieties.
- Two growth cabinets, one with 8 hour photoperiod, the other with 16 hour.

Jars placed at random in one of the two cabinets

Response variable: number of roots after 4 weeks culture at 22°C.



## Motivating Application: Data

	Photoperiod							
	8					16		
BAP ( $\mu$ M)	2.2	4.4	8.8	17.6	2.2	4.4	8.8	17.6
No. of roots								
0	0	0	0	2	15	16	12	19
1	3	0	0	0	0	2	3	2
2	2	3	1	0	2	1	2	2
3	3	0	2	2	2	1	1	4
4	6	1	4	2	1	2	2	3
5	3	0	4	5	2	1	2	1
6	2	3	4	5	1	2	3	4
7	2	7	4	4	0	0	1	3
8	3	3	7	8	1	1	0	0
9	1	5	5	3	3	0	2	2
10	2	3	4	4	1	3	0	0
11	1	4	1	4	1	0	1	0
12	0	0	2	0	1	1	1	0
>12	13,17	13	14,14	14				
No. of shoots	30	30	40	40	30	30	30	40
Mean	5.8	7.8	7.5	7.2	3.3	2.7	3.1	2.5
Variance	14.1	7.6	8.5	8.8	16.6	14.8	13.5	8.5
Overdispersion index	1.42	-0.03	0.13	0.22	4.06	4.40	3.31	2.47

(4日) (四) (三) (三) (三)

John Hinde (NUIG)

3

#### Dispersion

Second factorial cumulant

$$S(X) = Var(X) - E[X]$$

Useful summary:

- underdispersion:
- equidispersion (Poisson):
- overdispersion:

 $-\mathsf{E}[X] \le S(X) < 0$ S(X) = 0S(X) > 0

3

- 4 週 ト - 4 三 ト - 4 三 ト

 $-\mathsf{E}[X] \le S(X) < 0$ 

S(X) = 0

S(X) > 0

#### Dispersion

Second factorial cumulant

$$S(X) = Var(X) - E[X]$$

Useful summary:

- underdispersion:
- equidispersion (Poisson):
- overdispersion:

Fisher's dispersion index

$$D(X) = \frac{\operatorname{Var}(X)}{\operatorname{E}[X]} = 1 + \frac{S(X)}{\operatorname{E}[X]}$$

3

くほと くほと くほと

Poisson (Po)

$$\operatorname{Var}(X) = \mu$$
  $S(X) = 0$ 

・ロト ・四ト ・ヨト ・ヨト

Poisson (Po)

$$Var(X) = \mu$$
  $S(X) = 0$ 

Negative binomial (NB2): Poisson-Gamma mixture

$${\sf Var}(X)=\mu+\gamma\mu^2\qquad {\cal S}(X)=\gamma\mu^2$$

Note: Poisson-lognormal mixture has same variance function

3

・ロン ・四 ・ ・ ヨン ・ ヨン

Poisson (Po)

$$Var(X) = \mu$$
  $S(X) = 0$ 

Negative binomial (NB2): Poisson-Gamma mixture

$${\sf Var}(X)=\mu+\gamma\mu^2\qquad {\cal S}(X)=\gamma\mu^2$$

Note: Poisson-lognormal mixture has same variance function **Negative binomial (NB1)**: alternative Poisson-Gamma mixture

$$Var(X) = \mu + \gamma \mu = \phi \mu$$
  $S(X) = \gamma \mu$ 

same variance function as a quasi-Poisson model

< 回 ト < 三 ト < 三 ト

Poisson (Po)

$$Var(X) = \mu$$
  $S(X) = 0$ 

Negative binomial (NB2): Poisson-Gamma mixture

$${\sf Var}(X)=\mu+\gamma\mu^2\qquad {\cal S}(X)=\gamma\mu^2$$

Note: Poisson-lognormal mixture has same variance function **Negative binomial (NB1)**: alternative Poisson-Gamma mixture

$$Var(X) = \mu + \gamma \mu = \phi \mu$$
  $S(X) = \gamma \mu$ 

same variance function as a quasi-Poisson model

#### Poisson-inverse Gaussian

$${\sf Var}(X)=\mu+\gamma\mu^3 \qquad {\cal S}(X)=\gamma\mu^3$$

< 回 ト < 三 ト < 三 ト

#### Extended variance function

An natural generalization is

$${\sf Var}(X)=\mu+\gamma\mu^{
ho}\qquad {\cal S}(X)=\gamma\mu^{
ho}$$

for some general power *p*.

3

(日) (同) (三) (三)

#### Extended variance function

An natural generalization is

$${
m Var}(X)=\mu+\gamma\mu^{
ho}\qquad S(X)=\gamma\mu^{
ho}$$

for some general power *p*.

Suggested by Hinde & Demétrio (1998) and Nelder (??).

・ 同 ト ・ ヨ ト ・ ヨ ト

#### Overdispersion & Zero-Inflation

#### Extended variance function

An natural generalization is

$$\operatorname{Var}(X) = \mu + \gamma \mu^{
ho} \qquad S(X) = \gamma \mu^{
ho}$$

for some general power p.

Suggested by Hinde & Demétrio (1998) and Nelder (??).

Class of Poisson mixtures, **Poisson-Tweedie models**  $PT_p(\mu, \gamma)$ 

$$Z \sim \mathit{Tw}_{p}(\mu,\gamma), \hspace{1em} X | Z \sim \mathit{Po}(Z) \Rightarrow X \sim \mathit{PT}_{p}(\mu,\gamma)$$

has moments

$$\mathsf{E}[X] = \mathsf{E}[Z] = \mu$$
  $\mathsf{Var}(Z) = \gamma \mu^{
ho}$   $\mathsf{Var}(X) = \mu + \gamma \mu^{
ho}$ 

・ 同 ト ・ ヨ ト ・ ヨ ト

#### Tweedie Models

Family	E[ <i>Z</i> ]	Var(Z)	Туре	Support
Normal	$\mu$	$\gamma$	Continuous	R
Poisson	$\mu$	$\mu$	Discrete	N <sub>0</sub>
Non-central gamma	ntral gamma $\left  egin{array}{c c} \mu \end{array}  ight  \gamma \mu^{3/2}$		Cont. + atom	$R_0$
Gamma	$\mu$	$\gamma\mu^2$	Continuous	$R_+$
Inverse Gauss	$\mu$	$\gamma\mu^3$	Continuous	$R_+$

Only Poisson distribution is discrete.

(日) (周) (三) (三)

# Poisson-Tweedie Models

Family	E[X]	S(X)	Disp. Type	ZI(X)
Poisson	$\mu$	0	Equi	0
Hermite	$\mu$	$\gamma$	Over	+
Neyman Type A ( <i>Poisson-Poisson</i> )	$\mu$	$\gamma \mu$	Over	+
Pólya-Aeppli Type A ( <i>Poisson-compound Poisson</i> )	μ	$\gamma\mu^{3/2}$	Over	+
Negative binomial	$\mu$	$\gamma\mu^2$	Over	+
Binomial	$\mu$	$-\gamma\mu^2$	Under	+
Poisson-Inv. Gauss	$\mu$	$\gamma\mu^3$	Over	—

э

・ロト ・ 日 ・ ・ ヨ ・ ・ ヨ ・

## Motivating Application: Data

	Photoperiod							
	8					16		
BAP ( $\mu$ M)	2.2	4.4	8.8	17.6	2.2	4.4	8.8	17.6
No. of roots								
0	0	0	0	2	15	16	12	19
1	3	0	0	0	0	2	3	2
2	2	3	1	0	2	1	2	2
3	3	0	2	2	2	1	1	4
4	6	1	4	2	1	2	2	3
5	3	0	4	5	2	1	2	1
6	2	3	4	5	1	2	3	4
7	2	7	4	4	0	0	1	3
8	3	3	7	8	1	1	0	0
9	1	5	5	3	3	0	2	2
10	2	3	4	4	1	3	0	0
11	1	4	1	4	1	0	1	0
12	0	0	2	0	1	1	1	0
>12	13,17	13	14,14	14				
No. of shoots	30	30	40	40	30	30	30	40
Mean	5.8	7.8	7.5	7.2	3.3	2.7	3.1	2.5
Variance	14.1	7.6	8.5	8.8	16.6	14.8	13.5	8.5
Overdispersion index	1.42	-0.03	0.13	0.22	4.06	4.40	3.31	2.47

・ ロ ト ・ 個 ト ・ 国 ト ・ 国 ト

John Hinde (NUIG)

3

#### Zero-inflated models

If  $Y_i$  has a zero-inflated Poisson (ZIP) distribution, given by

$$\Pr(Y_i = y_i) = \begin{cases} \omega_i + (1 - \omega_i)e^{-\lambda_i} & y_i = 0\\ (1 - \omega_i)\frac{e^{-\lambda_i}\lambda_i^{y_i}}{y_i!} & y_i > 0 \end{cases}$$

(日) (同) (三) (三)

#### Zero-inflated models

If  $Y_i$  has a zero-inflated Poisson (ZIP) distribution, given by

$$\mathsf{Pr}(Y_i = y_i) = \begin{cases} \omega_i + (1 - \omega_i)e^{-\lambda_i} & y_i = 0\\ (1 - \omega_i)\frac{e^{-\lambda_i}\lambda_i^{y_i}}{y_i!} & y_i > 0 \end{cases}$$

Lambert (1992) considered models in which

$$\log(\lambda_i) = \mathbf{x}_i^T \boldsymbol{\beta}$$
 and  $\log\left(\frac{\omega_i}{1-\omega_i}\right) = \mathbf{z}_i^T \boldsymbol{\gamma}$ 

where **x** and **z** are covariate vectors and  $\beta$  and  $\gamma$  are vectors of parameters.

(日) (周) (三) (三)

#### Zero-inflated models

If  $Y_i$  has a zero-inflated Poisson (ZIP) distribution, given by

$$\mathsf{Pr}(Y_i = y_i) = \begin{cases} \omega_i + (1 - \omega_i)e^{-\lambda_i} & y_i = 0\\ (1 - \omega_i)\frac{e^{-\lambda_i}\lambda_i^{y_i}}{y_i!} & y_i > 0 \end{cases}$$

Lambert (1992) considered models in which

$$\log(\lambda_i) = \mathbf{x}_i^T \boldsymbol{eta}$$
 and  $\log\left(rac{\omega_i}{1-\omega_i}
ight) = \mathbf{z}_i^T \boldsymbol{\gamma}$ 

where **x** and **z** are covariate vectors and  $\beta$  and  $\gamma$  are vectors of parameters. Similar mixture models are available for the negative binomial distribution (ZINB), etc.

(日) (周) (三) (三)

### Trajan apple cultivation data: fitted frequencies

No. of		Fitted frequencies						
Roots	Observed	Poisson	Neg-bin	ZIP	ZINB	ZIGPD		
0	62	7.4	55.8	62	62	62		
1	7	21.3	19.8	1.6	5.1	4.8		
2	7	30.4	12.2	4.4	7.6	7.6		
3	8	29	8.6	7.9	8.9	9.1		
4	8	20.8	6.4	10.8	9.1	9.3		
5	6	11.9	4.9	11.8	8.4	8.5		
6	10	5.7	3.9	10.7	7.2	7.2		
7	4	2.3	3.1	8.3	5.8	5.8		
8	2	0.8	2.5	5.7	4.5	4.5		
9	7	0.3	2.1	3.4	3.4	3.4		
10	4	0.1	1.7	1.9	2.5	2.4		
11	2	0	1.4	0.9	1.8	1.7		
$\geq 12$	3	0	5.8	0.7	3.6	3.7		
-2 ×	log-lik	840.7	550.2	537.9	519.3	519.8		
	$G^2$	335.5	36.9	31.2	9.1	9.4		

イロト イヨト イヨト イヨト

▶ E の

# Trajan apple cultivation data: ZINB



Contour plot of  $2 \times \log$ -likelihood for  $\alpha$  and  $\omega$  with  $\mu$  fixed at the sample mean: maximum likelihood estimates for ZINB (\*) and negative binomial models (•).

#### Trajan Apples: model fitting results

- P is a two level factor for photoperiod
- H is a four level factor for the BAP levels
- Lin(H) is a linear trend over the levels of H

(on the log-concentration scale for BAP.)

	Mod	5					
Description	λ	ω	$\alpha$	$-2 \log L$	df	AIC	BIC
Poisson	H*P	0	0	1556.9	262	1572.9	1601.7
	Р	0	0	1571.9	268	1575.9	1583.1
Neg-Bin	H*P	0	const	1399.6	261	1417.6	1450.0
	H*P	0	Р	1264.6	260	1284.6	1320.6
	H*P	0	H*P	1254.8	254	1286.8	1344.4
	Lin(H)*P	0	Р	1270.1	264	1282.1	1303.7
	Р	0	Р	1272.4	266	1280.4	1294.8
	Р	0	const	1403.9	267	1409.9	1420.7

Image: A match a ma

## Trajan Apples: model fitting results

	Me	odels					
Description	$\lambda$	$\omega$	$\alpha$	$-2\log L$	df	AIC	BIC
ZIP	H*P	const	0	1338.0	261	1356.0	1388.4
	H*P	Р	0	1244.5	260	1264.5	1300.5
	H*P	H*P	0	1238.2	254	1270.2	1327.8
	Lin(H)*P	Р	0	1250.2	264	1262.2	1283.8
	Р	Р	0	1261.3	266	1269.3	1283.7
	Р	const	0	1355.2	267	1361.2	1372.0
ZINB	H*P	const	const	1324.8	260	1344.8	1380.8
	H*P	Р	const	1232.5	259	1254.5	1294.1
	H*P	Р	Р	1226.3	258	1250.3	1293.5
	H*P	H*P	H*P	1205.6	246	1253.6	1340.0
	Lin(H)*P	Р	Р	1231.0	262	1247.0	1275.8
	Р	Р	Р	1237.7	264	1249.7	1271.3
	Р	Р	const	1243.9	265	1253.9	1271.9
	Р	const	const	1336.5	266	1344.5	1358.9
	const	Р	const	1257.8	266	1265.8	1280.2

35 / 49

• Termite *Heterotermes tenuis*: an important pest of sugarcane in Brazil, causing damage of up to 10 metric tonnes/ha/year.

(日) (同) (三) (三)

- Termite *Heterotermes tenuis*: an important pest of sugarcane in Brazil, causing damage of up to 10 metric tonnes/ha/year.
- Fungus Beauveria bassiana: a possible microbial control.

(人間) トイヨト イヨト

- Termite *Heterotermes tenuis*: an important pest of sugarcane in Brazil, causing damage of up to 10 metric tonnes/ha/year.
- Fungus *Beauveria bassiana*: a possible microbial control.
- Experiment: on the pathogenicity and virulence of 142 different isolates of *Beauveria bassiana*.
  - Completely randomized experiment: five replicates of each of the 142 isolates.
  - Solutions of the isolates applied to groups (clusters) of *n* = 30 termites kept in plastic Petri-dishes.
  - Mortality in the groups was measured daily for eight days

(日) (周) (三) (三)

- Termite *Heterotermes tenuis*: an important pest of sugarcane in Brazil, causing damage of up to 10 metric tonnes/ha/year.
- Fungus *Beauveria bassiana*: a possible microbial control.
- Experiment: on the pathogenicity and virulence of 142 different isolates of *Beauveria bassiana*.
  - Completely randomized experiment: five replicates of each of the 142 isolates.
  - Solutions of the isolates applied to groups (clusters) of *n* = 30 termites kept in plastic Petri-dishes.
  - Mortality in the groups was measured daily for eight days
- Data: 710 ordered multinomial observations of length eight.

イロト イヨト イヨト

#### Cumulative Mortality: sample of isolates



days

John Hinde (NUIG)

3

イロト イポト イヨト イヨト

# Cumulative Mortality: spaghetti plot of all isolates



# Multinomial Model: Cumulative Proportions

Because of natural time ordering consider models for the **cumulative proportions** (isolate i, replicate k)

$$R_{ik,d}$$
 = proportion of insects dead by day  $d$ ,

 $\gamma_{ik,d} = \mathsf{E}(R_{ik,d}) = \mathsf{probability}$  an insect dies by day d,

$$\mathbf{R}_{ik} = (R_{ik,1}, R_{ik,2}, \dots, R_{ik,D})^T = \frac{1}{n} \mathbf{L} \mathbf{Y}_{ik}$$

# Multinomial Model: Cumulative Proportions

Because of natural time ordering consider models for the **cumulative proportions** (isolate i, replicate k)

$$R_{ik,d}$$
 = proportion of insects dead by day  $d$ ,

 $\gamma_{ik,d} = \mathsf{E}(R_{ik,d}) = \mathsf{probability}$  an insect dies by day d,

$$\mathbf{R}_{ik} = (R_{ik,1}, R_{ik,2}, \dots, R_{ik,D})^T = \frac{1}{n} \mathbf{L} \mathbf{Y}_{ik}$$

$$\mathsf{E}[\mathbf{R}_{ik}] = \mathbf{L}\boldsymbol{\pi}_{ik} = \boldsymbol{\gamma}_{ik}$$
$$\mathsf{Var}[\mathbf{R}_{ik}] = \frac{1}{n} \mathcal{L}[\mathsf{diag}\{\boldsymbol{\pi}_{ik}\} - \boldsymbol{\pi}_{ik}\boldsymbol{\pi}_{ik}^{\mathsf{T}}] \mathcal{L}^{\mathsf{T}} = \boldsymbol{V}(\boldsymbol{\gamma}_{ik})$$

• Use a glm with link function:

$$g(\boldsymbol{\gamma}_{ik}) = \mathbf{X}_{ik} \boldsymbol{\beta}_i$$

3

(日) (同) (日) (日)

- Use a glm with link function:  $g(\gamma_{ik}) = \mathbf{X}_{ik} \beta_i$
- Logit link function  $\longrightarrow$  cumulative logistic model

$$g(\gamma_{_{ikj}}) = ext{logit}(\gamma_{_{ikj}}) = ext{log}\left( egin{matrix} j \ \sum \ s=1 \ rac{s=1}{D+1} \pi_{_{ik,s}} \ \sum \ s=j+1 \ rac{1}{D+1} \pi_{_{ik,s}} \end{pmatrix} = \eta_{ikj}$$

• alternative models: discrete survival models, other ordinal models

- **(() ) ) ( () ) ) () )** 

- Use a glm with link function:  $g(\gamma_{ik}) = \mathbf{X}_{ik} \beta_i$
- Logit link function  $\longrightarrow$  cumulative logistic model

$$g(\gamma_{ikj}) = ext{logit}(\gamma_{ikj}) = ext{log}\left( egin{matrix} j \ \sum {s=1} \pi_{ik,s} \ rac{j}{\sum {s=j+1}} \pi_{ik,s} \ \sum {s=j+1} \pi_{ik,s} \ \end{array} 
ight) = \eta_{ikj}$$

• Linear predictor: isolate specific factors, time dependency, ...

(日) (同) (三) (三)

- Use a glm with link function:  $g(\gamma_{ik}) = \mathbf{X}_{ik} \beta_i$
- Logit link function  $\longrightarrow$  cumulative logistic model

$$g(\gamma_{ikj}) = ext{logit}(\gamma_{ikj}) = ext{log}\left( egin{matrix} j \ \sum \ s=1 \ \pi_{ik,s} \ \hline \ \Sigma \ s=j+1 \ \pi_{ik,s} \ \end{array} 
ight) = \eta_{ikj}$$

• Linear predictor: isolate specific factors, time dependency, ... e.g. Isolate specific linear time effect, constant over replicates

$$\eta_{ikj} = \beta_{1i} + \beta_{2i} t_j,$$

- **(() ) ) ( () ) ) () )** 

# Random Effect Models

Incorporate random effects in the linear predictor:

- Add random effect for each experimental unit (groups of insects).
  - simple time shifts
  - time dependent covariates with random coefficients
  - Replicate level random effect accounts for overdispersion

・ 同 ト ・ ヨ ト ・ ヨ ト

# Random Effect Models

Incorporate random effects in the linear predictor:

- Add random effect for each experimental unit (groups of insects).
  - simple time shifts
  - time dependent covariates with random coefficients
  - Replicate level random effect accounts for overdispersion
- Model isolates as a random effect.

$$\eta_{ikj} = \mu + time_j + u_i + \epsilon_{ik}$$

Non-parametric maximum likelihood techniques give a finite mass-point distribution  $\{\omega_k; z_k\}$  for the isolate effects  $u_i$ . Using a small number of components may identify effective isolates – look at the posterior distribution of  $u_i$ .

(本語) (本語) (本語)
Additional variation across replicates  $\longrightarrow$  overdispersion

3

(日) (周) (三) (三)

Additional variation across replicates  $\longrightarrow$  overdispersion

• Allow variation in multinomial parameter  $\pi$  — two-stage model

$$\boldsymbol{Y}_{ik} \mid \boldsymbol{p}_{ik} \sim \mathsf{Multinomial}(n; \boldsymbol{p}_{ik})$$

 $oldsymbol{p}_{ik} = (p_{ik,1}, \dots, p_{ik,\mathsf{D}}, p_{ik,\mathsf{D}+1})^{\mathcal{T}}$  follows a Dirichlet distribution

(日) (周) (三) (三)

Additional variation across replicates  $\longrightarrow$  overdispersion

ullet Allow variation in multinomial parameter  $\pi$  — two-stage model

$$\begin{split} \mathbf{Y}_{ik} &| \ \mathbf{p}_{ik} \sim \text{Multinomial}(n; \mathbf{p}_{ik}) \\ \mathbf{p}_{ik} &= (p_{ik,1}, \dots, p_{ik,D}, p_{ik,D+1})^T \text{ follows a Dirichlet distribution} \end{split}$$

• Dirichlet-multinomial model for Y and R with

$$E[m{R}_{ik}] = m{\gamma}_{ik}$$

and covariance matrix given by

$$\mathsf{Var}[m{R}_{ik}] = m{V}(m{\gamma}_{ik})[1+
ho_i(n-1)]$$

where  $\rho_i$  is an (isolate specific) overdispersion parameter

Additional variation across replicates  $\longrightarrow$  overdispersion

ullet Allow variation in multinomial parameter  $\pi$  — two-stage model

$$\begin{split} \mathbf{Y}_{ik} &| \ \mathbf{p}_{ik} \sim \text{Multinomial}(n; \mathbf{p}_{ik}) \\ \mathbf{p}_{ik} &= (p_{ik,1}, \dots, p_{ik,D}, p_{ik,D+1})^T \text{ follows a Dirichlet distribution} \end{split}$$

• Dirichlet-multinomial model for Y and R with

$$E[m{R}_{ik}]=m{\gamma}_{ik}$$

and covariance matrix given by

$$\mathsf{Var}[oldsymbol{\mathcal{R}}_{ik}] = oldsymbol{\mathcal{V}}(oldsymbol{\gamma}_{ik})[1+
ho_i(n-1)]$$

where  $\rho_i$  is an (isolate specific) overdispersion parameter

• Generalization of beta-binomial model

John Hinde (NUIG)

28 March 2015 42 / 49

## Random Intercept Model

• Model additional variation by including random effects in the linear predictor

$$g(q_{ikj}) = \eta_{ikj} + \xi_{ik} = \beta_{1i} + \beta_{2i}t_j + \xi_{ik}$$

where  $\xi_{ik}$  is a random effect with  $E[\xi_{ik}] = 0$ ,  $Var[\xi_{ik}] = \sigma_i^2$ 

(日) (周) (三) (三)

## Random Intercept Model

Model additional variation by including random effects in the linear predictor

$$g(q_{ikj}) = \eta_{ikj} + \xi_{ik} = \beta_{1i} + \beta_{2i}t_j + \xi_{ik}$$

where  $\xi_{ik}$  is a random effect with  $E[\xi_{ik}] = 0$ ,  $Var[\xi_{ik}] = \sigma_i^2$ 

Taylor series approximations give

$$E[\boldsymbol{R}_{ik}] = E[E(\boldsymbol{R}_{ik}|\boldsymbol{q}_{ik})] = E[\boldsymbol{q}_{ik}] \approx \gamma_{ik}$$

and

$$\mathsf{Var}[\boldsymbol{R}_{ik}] \approx \boldsymbol{V}(\boldsymbol{\gamma}_{ik}) + \left(1 - \frac{1}{n}\right) \sigma_i^2 [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})] [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})]^T$$

where  $\boldsymbol{h}$  is inverse link function with derivative  $\boldsymbol{h}'$ 

・ロン ・四 ・ ・ ヨン ・ ヨン

# Random Intercept Model

• Model additional variation by including random effects in the linear predictor

$$g(q_{ikj}) = \eta_{ikj} + \xi_{ik} = \beta_{1i} + \beta_{2i}t_j + \xi_{ik}$$

where  $\xi_{ik}$  is a random effect with  $E[\xi_{ik}] = 0$ ,  $Var[\xi_{ik}] = \sigma_i^2$ 

Taylor series approximations give

$$E[\boldsymbol{R}_{ik}] = E[E(\boldsymbol{R}_{ik}|\boldsymbol{q}_{ik})] = E[\boldsymbol{q}_{ik}] \approx \boldsymbol{\gamma}_{ik}$$

and

$$\mathsf{Var}[\boldsymbol{R}_{ik}] \approx \boldsymbol{V}(\boldsymbol{\gamma}_{ik}) + \left(1 - \frac{1}{n}\right) \sigma_i^2 [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})] [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})]^T$$

where  $\boldsymbol{h}$  is inverse link function with derivative  $\boldsymbol{h}'$ 

• Analagous to approximate variance function for logistic-normal distribution

イロト 不得 トイヨト イヨト

## Random Intercept + Random Slope Model

• Extend to include correlated random effects for intercept and slope

$$g(q_{ikj}) = eta_{1i} + \xi_{ik} + (eta_{2i} + \zeta_{ik})t_j = \eta_{ikj} + \xi_{ik} + \zeta_{ik}t_j$$

where  $(\xi_{ik}, \zeta_{ik})^T$  has  $E[\xi_{ik}] = E[\zeta_{ik}] = 0$  and covariance matrix

$$\mathbf{\Sigma} = \begin{bmatrix} \nu_i^2 & \lambda_i \nu_i \tau_i \\ \lambda_i \nu_i \tau_i & \tau_i^2 \end{bmatrix}$$

・ロン ・四 ・ ・ ヨン ・ ヨン

## Random Intercept + Random Slope Model

• Extend to include correlated random effects for intercept and slope

$$g(q_{ikj}) = eta_{1i} + \xi_{ik} + (eta_{2i} + \zeta_{ik})t_j = \eta_{ikj} + \xi_{ik} + \zeta_{ik}t_j$$

where  $(\xi_{ik}, \zeta_{ik})^T$  has  $E[\xi_{ik}] = E[\zeta_{ik}] = 0$  and covariance matrix

$$\mathbf{\Sigma} = \begin{bmatrix} \nu_i^2 & \lambda_i \nu_i \tau_i \\ \lambda_i \nu_i \tau_i & \tau_i^2 \end{bmatrix}$$

• Approximations now give

$$E[m{R}_{ik}]pproxm{\gamma}_{ik}$$

and

$$\operatorname{Var}[\boldsymbol{R}_{ik}] \approx \boldsymbol{V}(\boldsymbol{\gamma}_{ik}) + \left(1 - \frac{1}{n}\right) \left\{ \nu_i^2 [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})] [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})]^T + \tau_i^2 [\boldsymbol{h}'(\boldsymbol{\eta}_{ik}) * \boldsymbol{t}_{ik}] [\boldsymbol{h}'(\boldsymbol{\eta}_{ik}) * \boldsymbol{t}_{ik}]^T + \lambda_i \nu_i \tau_i [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})] [\boldsymbol{h}'(\boldsymbol{\eta}_{ik})]^T * [\mathbf{1} \boldsymbol{t}_{ik}^T + \boldsymbol{t}_{ik} \mathbf{1}^T] \right\}$$

#### • Parameter estimates from all four models are identical

3

(日) (周) (三) (三)

- Parameter estimates from all four models are identical
- Robust se's from all four models are identical

(日) (同) (三) (三)

- Parameter estimates from all four models are identical
- Robust se's from all four models are identical
- Model based se's exhibit simple relationships

< 🗇 🕨 < 🖃 🕨

- Parameter estimates from all four models are identical
- Robust se's from all four models are identical
- Model based se's exhibit simple relationships
- Numerous explanations posited by various colleagues, but ....

#### All down to forms of models and matrix algebra

• Extension of general linear model

3

(a)

- Extension of general linear model
- Analysis of non-normal data

Image: A match a ma

- Extension of general linear model
- Analysis of non-normal data
- Likelihood based inference

- Extension of general linear model
- Analysis of non-normal data
- Likelihood based inference
- Model selection, comparison, validation

- Extension of general linear model
- Analysis of non-normal data
- Likelihood based inference
- Model selection, comparison, validation
- Iterative computational methods

- Extension of general linear model
- Analysis of non-normal data
- Likelihood based inference
- Model selection, comparison, validation
- Iterative computational methods
- Extending model classes

- Extension of general linear model
- Analysis of non-normal data
- Likelihood based inference
- Model selection, comparison, validation
- Iterative computational methods
- Extending model classes

#### Combination of theory & application

## Bristol: Generalised Linear Models

#### Syllabus

- Overview of data analysis, motivating examples. Review of linear models. (1 lecture)
- Generalized linear models (GLMs). Exponential family model, sufficiency issues. Link function, canonical link. (5 lectures)
- Inference for generalized linear models, based on asymptotic theory: confidence intervals, hypothesis testing, goodness of fit. Results interpretation. Diagnostics. (4 lectures)
- Binary responses, logistic regression, residuals and diagnostics. (2 lectures)
- Introduction to survival analysis. Distribution theory: standard parametric models. Proportional odds model and connection to binomial GLM's. Inference assuming a parametric form for the baseline hazard. (4 lectures)

イロト 不得下 イヨト イヨト 二日

## UCSC: Generalized Linear Models

#### Introduction to GLMs

Statistical modeling in the context of GLMs. Exponential dispersion family of distributions (definitions, properties, and examples). Components of a GLM, examples of GLMs.

#### • Likelihood inference for GLMs

Likelihood estimation (iterative weighted least squares) and inference (asymptotic interval estimates). Model diagnostics (residuals for GLMs, model comparison criteria).

• Regression models for categorical responses and count data Models for binary responses (dose-response modeling, probit and logit models). Poisson regression and log-linear models. Basic ideas for modeling of contingency tables. Multinomial response models for nominal or ordinal responses.

#### Bayesian GLMs

General setting, examples, priors for GLMs. MCMC posterior simulation methods for GLMs. Bayesian residual analysis and model choice. Hierarchical GLMs, overdispersed GLMs, generalized linear mixed models.

## Acknowledgements

- Norma Coffey
- Clarice Demétrio
- Jochen Einbeck
- Silvia de Freitas
- Emma Holian
- Naratip Jansakul
- Bent Jørgensen
- Marie-José Martinez
- Georgios Papageorgiou
- Martin Ridout
- Mariana Ragassi Urbano
- Afrânio Vieira