

# Generalized linear mixed models

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- lme4: Doug Bates, Martin Mächler, Steve Walker
- Data: Josh Banta, Adrian Stier, Sea McKeon, David Julian, Jada-Simone White
- NSERC (Discovery)
- SHARCnet

# Outline

- 1 Examples and definitions
- 2 Estimation
  - Overview
  - Methods
- 3 Inference
- 4 Challenges & open questions

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# (Generalized) linear mixed models

(G)LMMs: a statistical modeling framework incorporating:

- combinations of categorical and continuous predictors, and interactions
- (some) non-Normal responses  
(e.g. binomial, Poisson, and extensions)
- (some) nonlinearity  
(e.g. logistic, exponential, hyperbolic)
- non-independent (grouped) data

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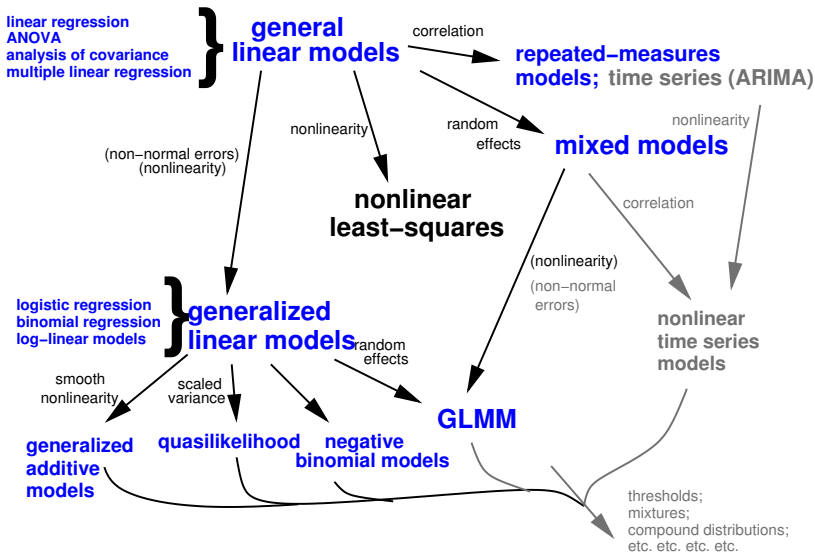
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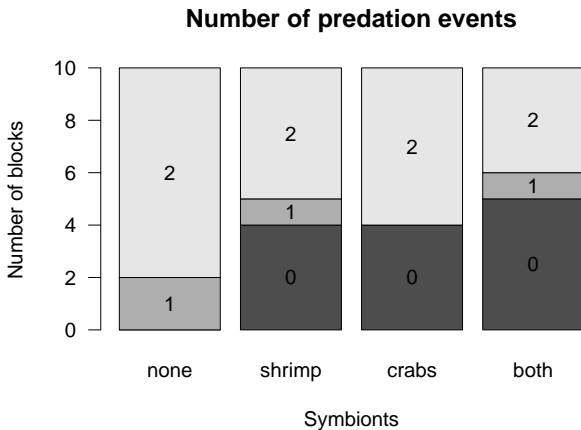
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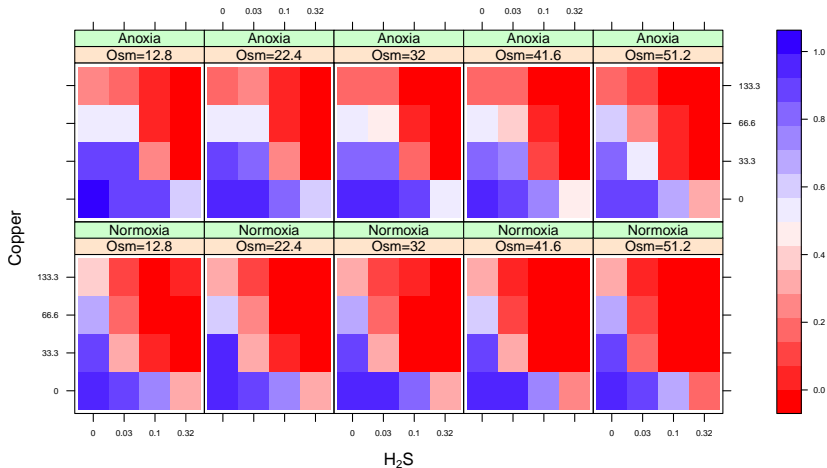


# Coral protection from seastars (*Culcita*) by symbionts (McKeon et al., 2012)

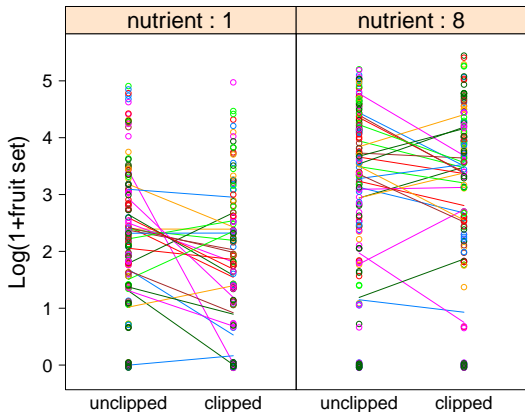


# Environmental stress: *Glycera* cell survival

(D. Julian unpubl.)

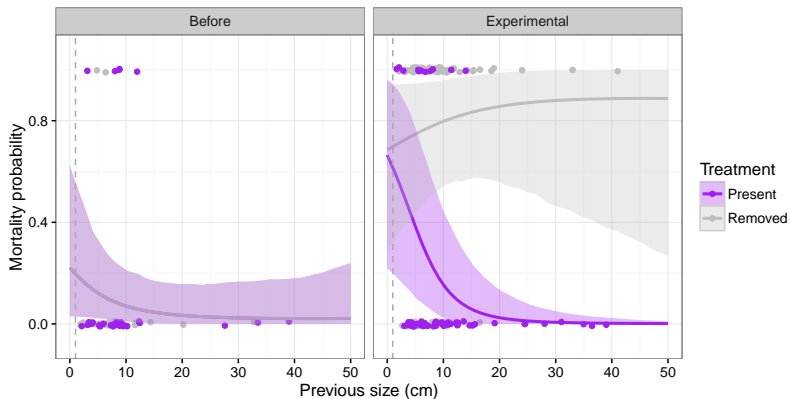


# *Arabidopsis* response to fertilization & herbivory (Banta et al., 2010)



# Coral demography

(J.-S. White unpubl.)



# Technical definition

$$\underbrace{Y_i}_{\text{response}} \sim \underbrace{\text{Distr}}_{\text{conditional distribution}} \left( \underbrace{g^{-1}(\eta_i)}_{\text{inverse link function}}, \underbrace{\phi}_{\text{scale parameter}} \right)$$

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$$\underbrace{b}_{\text{conditional modes}} \sim \text{MVN}(\mathbf{0}, \underbrace{\Sigma(\theta)}_{\text{variance-covariance matrix}})$$

# What are random effects?

A method for . . .

- accounting for among-individual, within-block correlation
- compromising between
  - complete pooling** (no among-block variance)
  - and **fixed effects** (large among-block variance)
- handling levels selected at random from a larger population
- sharing information among levels (*shrinkage estimation*)
- estimating variability among levels
- allowing predictions for unmeasured levels



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# Random-effect myths

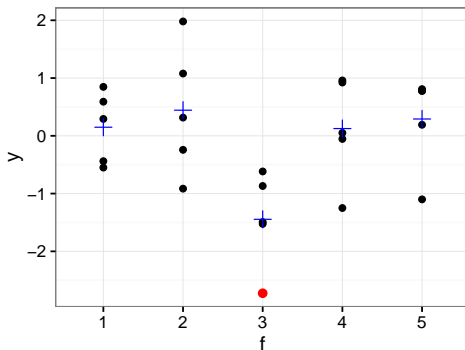
- levels of random effects must always be sampled at random
- a complete sample cannot be treated as a random effect
- random effects are always a **nuisance variable**
- nothing can be said about the predictions of a random effect
- you should always use a random effect no matter how few levels you have

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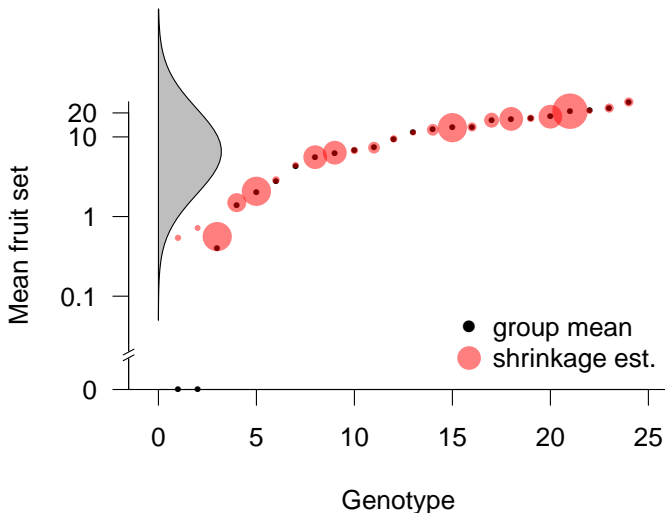
# Maximum likelihood estimation

- Best fit is a compromise between two components  
(consistency of data with fixed effects and conditional modes;  
consistency of random effect with RE distribution)
- Goodness-of-fit *integrates* over conditional modes





# Shrinkage: *Arabidopsis* conditional modes



# Estimation methods

**deterministic** : various approximate integrals (Breslow, 2004)

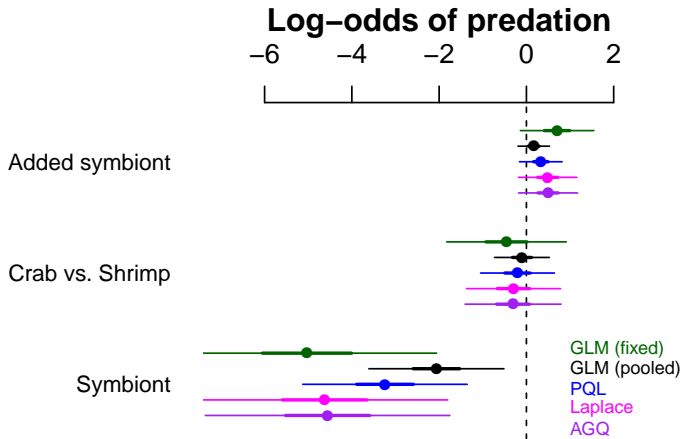
- Penalized quasi-likelihood, Laplace, Gauss-Hermite quadrature, ... (?); best methods needed for large variance, small clusters
- flexibility and speed vs. accuracy

...

**stochastic** (Monte Carlo): frequentist and Bayesian (Booth and Hobert, 1999; Ponciano et al., 2009; Sung and Geyer, 2007)

- usually slower but flexible and accurate

Estimation: *Culcita* (McKeon et al., 2012)

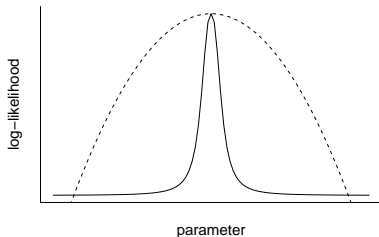


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# Wald tests

- typical results of summary
- exact for ANOVA, regression: approximation for GLM(M)s
- fast
- approximation is sometimes awful (Hauck-Donner effect)



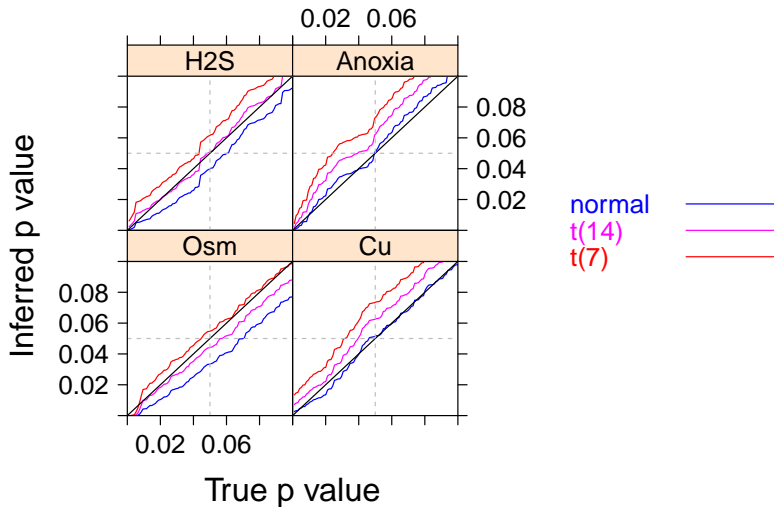
# Likelihood ratio tests

- better than Wald, but still have to two problems:
  - “denominator degrees of freedom” (when estimating scale)
  - for GLMMs, distributions are approximate anyway (Bartlett corrections)
  - Kenward-Roger correction? (Stroup, 2014)
- Profile confidence intervals: expensive/fragile

# Parametric bootstrapping

- fit null model to data
- simulate “data” from null model
- fit null and working model, compute likelihood difference
- repeat to estimate null distribution
- should be OK but ??? not well tested  
(assumes estimated parameters are “sufficiently” good)

# Parametric bootstrap results (*Glycera*)

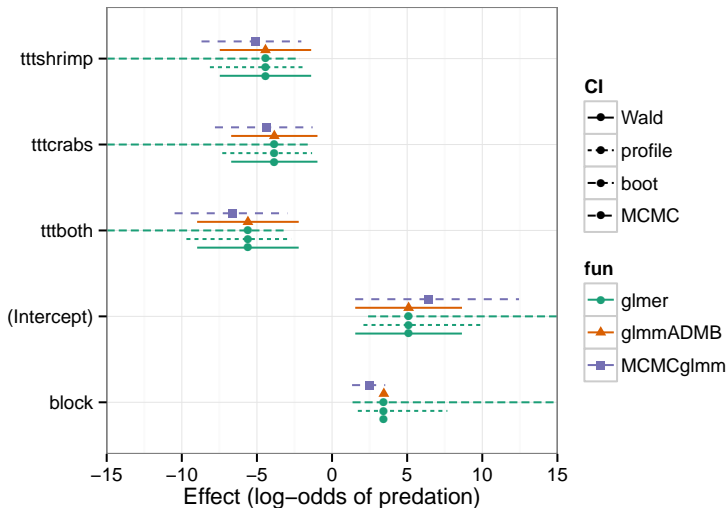




# Bayesian inference

- If we have a good sample from the posterior distribution (Markov chains have converged etc. etc.) we get most of the inferences we want for free by summarizing the marginal posteriors
- *post hoc* Bayesian methods: use deterministic/frequentist methods to find the maximum, then sample around it

# *Culcita* confidence intervals



# formula formats

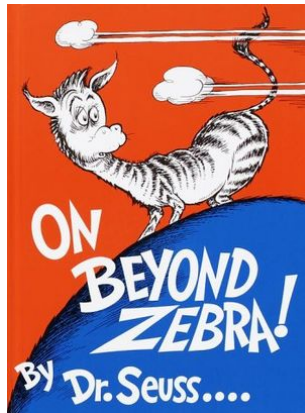
- `fixed`: fixed-effect formula
- `random`: random-effect formula (in `lme4`, combined with `fixed`)
  - generally `x|g` (term|grouping variable)
  - simplest: `1|g`, single intercept term
  - nested: `1|g1/g2`
  - random-slopes: `r|g`
  - independent terms: `(1|g)+(x+0|g)` or `(x||g)`
- `lme`: weights, correlation for heteroscedasticity and residual correlation
- `MCMCglmm`: options for variance structure

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# On beyond R

- Julia: MixedModels package
- SAS: PROC MIXED, NLMIXED
- AS-REML
- Stata (GLLAMM, xtmelogit)
- AD Model Builder; Template Model Builder
- HLM, MLWiN
- JAGS, Stan, [rethinking package](#)



# Challenges

- Small clusters: need AGQ/MCMC
- Small numbers of clusters: need finite-size corrections (KR/PB/MCMC)
- Small data sets: issues with **singular** fits  
Barr et al. (2013) vs. Bates et al. (2015)
- Big data: speed!
- Model diagnosis
- Confidence intervals accounting for uncertainty in variances

See also: <http://rpubs.com/bbolker/glmmchapter>, <https://groups.nceas.ucsb.edu/non-linear-modeling/projects>

# Spatial and temporal correlations

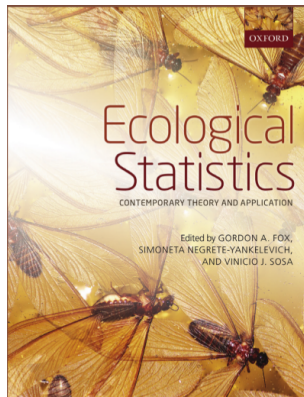
- Sometimes blocking takes care of non-independence ...
- but sometimes there is temporal or spatial correlation **within** blocks
- ... also phylogenetic ... (Ives and Zhu, 2006)
- “G-side” vs. “R-side” effects
- tricky to implement for GLMMs,  
but new possibilities on the horizon (Rousset and Ferdy, 2014;  
Rue et al., 2009);  
<https://github.com/stevencarlislewalker/lme4ord>

# Next steps

- Complex random effects:  
regularization, model selection, penalized methods  
(lasso/fence)
- Flexible correlation and variance structures
- Flexible/nonparametric random effects distributions
- hybrid & improved MCMC methods
- **Reliable** assessment of out-of-sample performance



- [http://ms.mcmaster.ca/~bolker/misc/iisc\\_private/14-Fox-Chap13.pdf](http://ms.mcmaster.ca/~bolker/misc/iisc_private/14-Fox-Chap13.pdf)
- <http://www.math.mcmaster.ca/bolker/R/misc/foxchapter>
- ?



(code ASPROMP8)

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