

Likelihood fitting and dynamic models II

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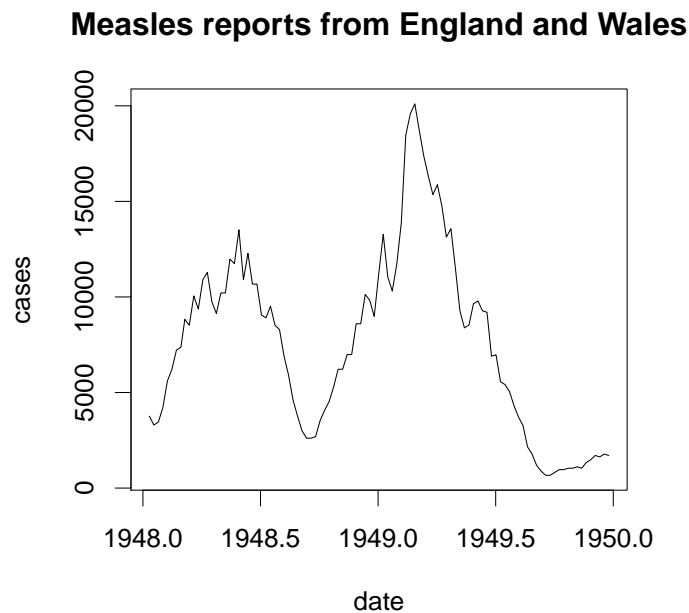
MMED 2017

Summary

- Discuss theories and approaches behind fitting models to data
- Explain relationship between least squares and likelihood
- Discuss approaches to dealing with process error and observation error

Measles data

- Reconstruct the number of susceptibles
- Divide the data into generations
- Fit \mathcal{R}_0
- Predict



Why did I get the wrong answer?

- Model structure may be wrong
- Population structure may be wrong
- Stochasticity in disease observation and recording
- Stochasticity in transmission
- Multi-parameter estimation
 - There may be different parameter combinations that work equally well
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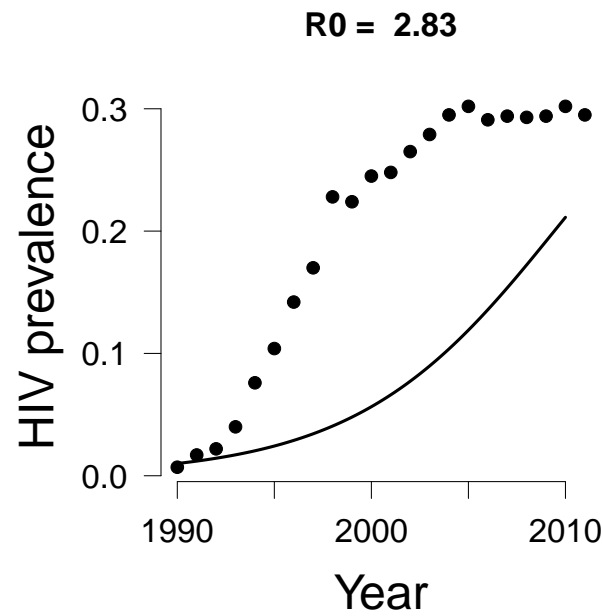
1 Conceptual framework

- How do we assume our data relate to our model world?
 - **No error:** We could attempt to model everything we see, in exact detail
 - * Impractical
 - **Observation error:** we could assume that the world is perfectly deterministic, but our *observations* are imperfect
 - * Shooting
 - **Process error:** we could assume that we observe perfectly, but that the world is stochastic
 - * Stepping
 - **Both kinds of error:** the world is stochastic, and our observations are imperfect
 - * Modern methods

Observation error only

- Point your model at the target
- Give it starting conditions and parameters
- Let it go
- Compare final results to observations

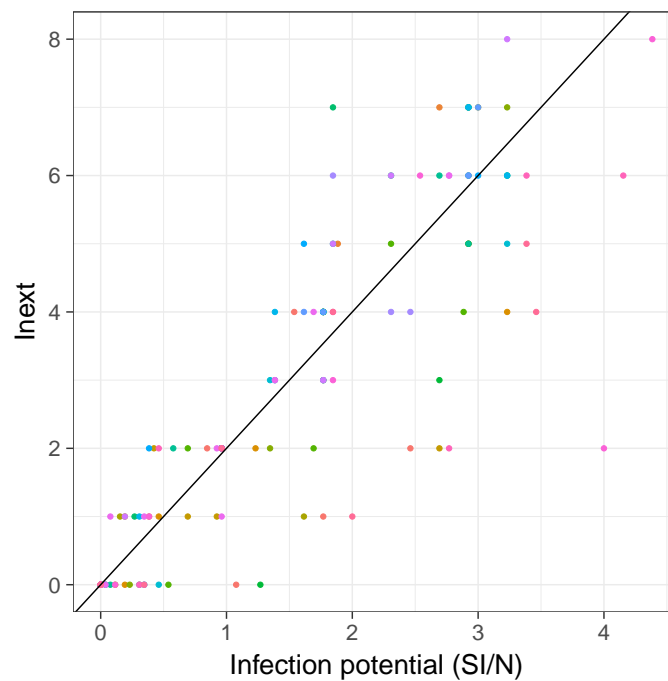
Shooting



Process error only

- Look at each step separately.
- See how the model is doing for that step.
- Reset based on observed data before taking the next step

Stepping



Modern methods

- Is it better to ignore process error, or observation error?
- What if we have a small number of cases, and good reporting (Ebola in small villages)

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- What if we have a large number of cases, and poor reporting (HIV in Harare)?

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- What if we have a new epidemic and poor reporting (West African Ebola)?

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2 Fitting

How to fit?

- Solving an equation
- By eye (fiddling with parameters)
- *Minimizing a distance function*
- Likelihood

Distance functions

$$D = \sum_i y_i - \hat{y}_i$$

Doesn't work

Distance functions

$$D = \sum_i |y_i - \hat{y}_i|$$

Not elegant

Distance functions

$$D = \sum_i (y_i - \hat{y}_i)^2$$

Simplest smooth approach

3 Likelihoods

- Assume that the difference between the estimate \hat{y}_i and the data point y_i is normally distributed. What is the log likelihood?

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$$L = \prod_i \frac{1}{\sigma\sqrt{2\pi}} \exp\left(\frac{-(\hat{y}_i - y_i)^2}{2\sigma^2}\right)$$

-

$$\ell = \sum_i -\log(\sigma\sqrt{2\pi}) - \sum_i \frac{(\hat{y}_i - y_i)^2}{2\sigma^2}$$

- *We minimize the likelihood by minimizing the sum of squares*
 - and then solving for σ

Least squares \rightarrow likelihood

- Attaching your least squares fit to a likelihood means:
 - You can *use it* for statistical inference (LRT)
 - You can *challenge* the assumptions

Mexican flu example

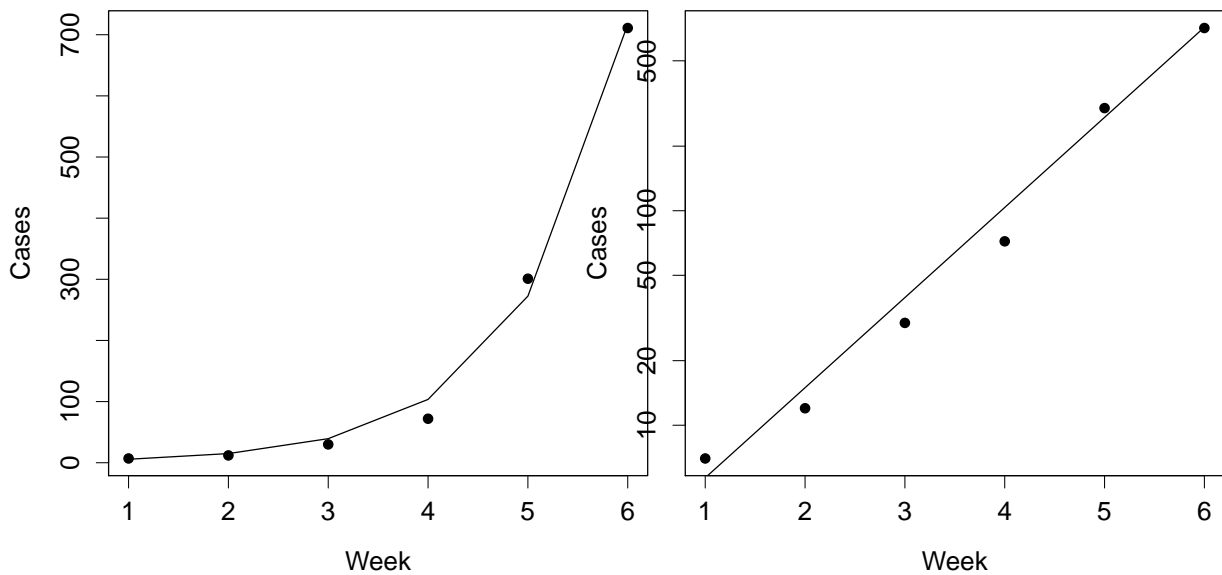
- How fast is it growing? r
- How hard will it be to control? \mathcal{R}_0

A different perspective

- Log scale shows multiplicative differences
- We could make the normal assumption on either scale
- How much does it matter?

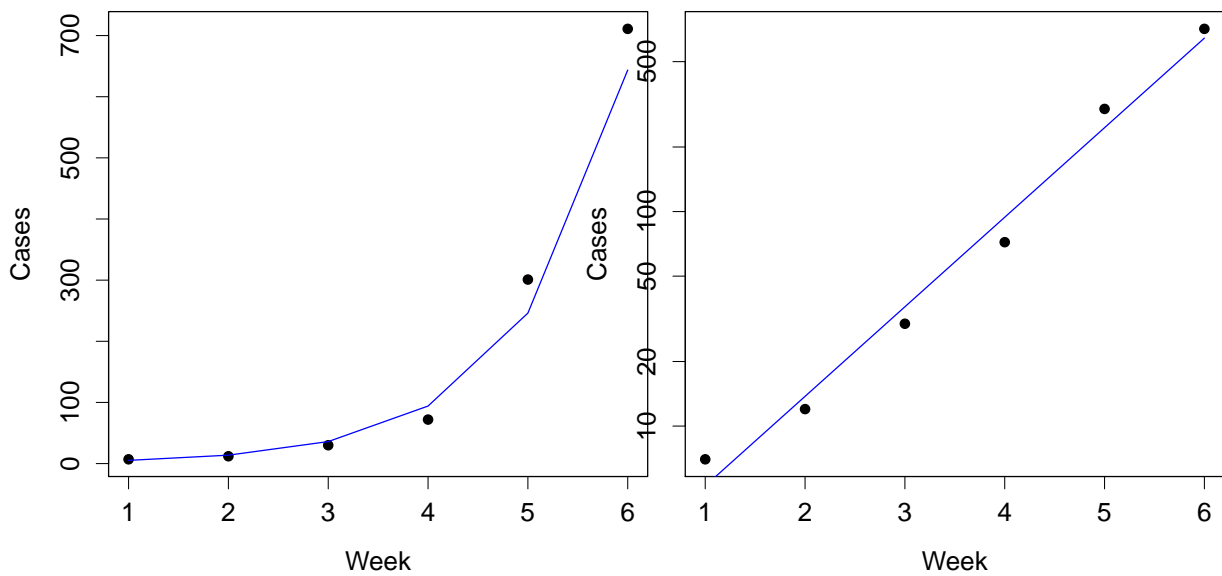
Normal assumption

- Least squares on the linear scale
- 10:50 :: 980:1020
- Gives relatively too much weight to large observations



Lognormal assumption

- Least squares on the log scale
- 3:5 :: 300:500
- Gives relatively too much weight to small observations



A more realistic error distribution

- My case counts are *individuals*

- What distributions can I use to reflect that?

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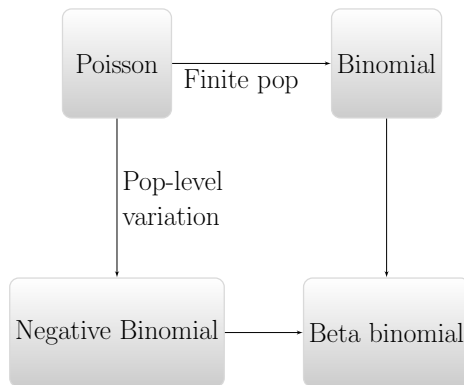
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Reality is complicated

- Poisson and binomial reflect *only* individual-level variation
 - No temporal variation
 - No clustered sampling
 - ...

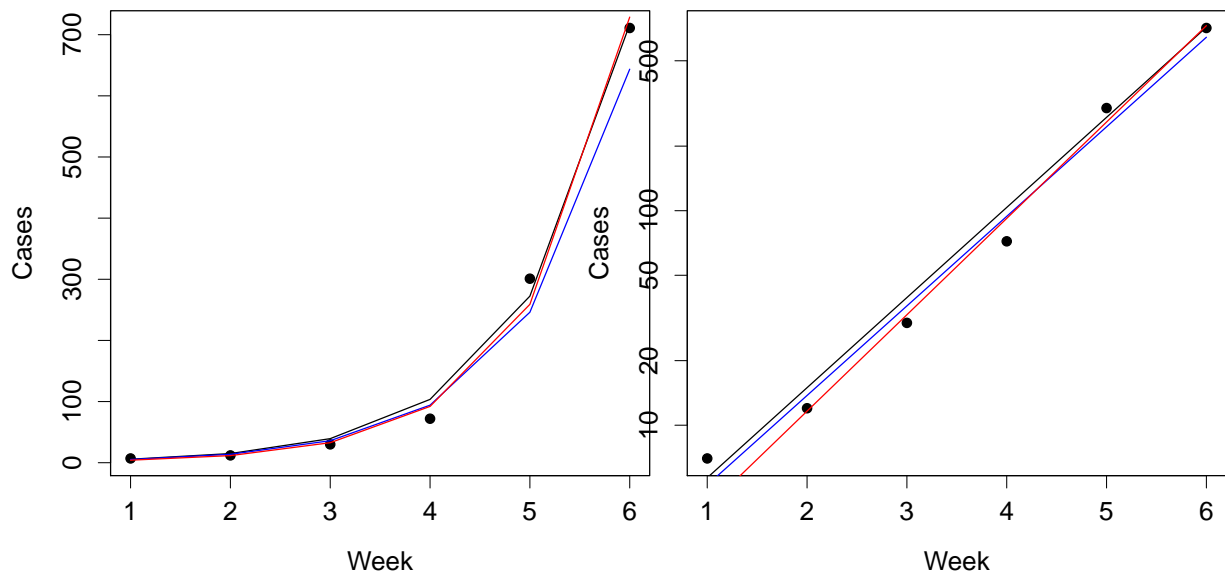
Distribution diagram



Discrete distributions

- Negative binomial is a good general-purpose discrete distribution
 - Individual- and population-level variability
- Beta binomial takes size into account
 - Good when denominator is clear and important
 - For exempling, when sampling

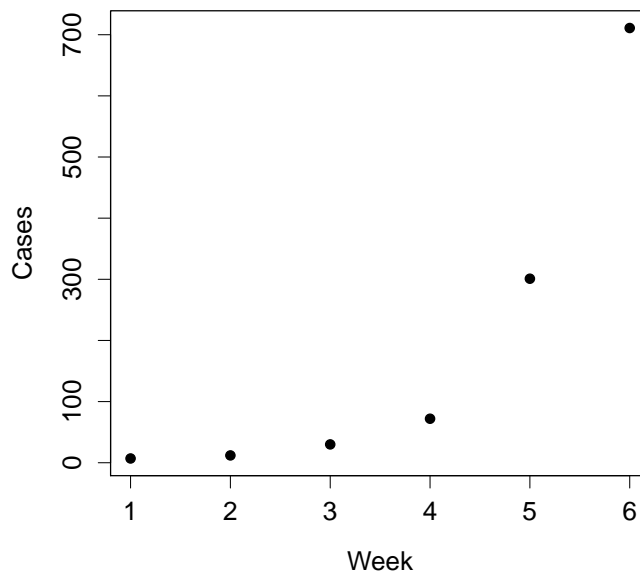
Negative binomial fits



Comparison

- Realistic error distribution provides (apparently) better fits
- Confidence intervals
 - Normal: $r = 0.96\text{--}0.97/\text{wk}$
 - Lognormal: $r = 0.64\text{--}1.29/\text{wk}$
 - Negative binomial: $r = 0.90\text{--}1.14/\text{wk}$
- How would you test these methods?
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Identifiability



- What if we tried to estimate \mathcal{R}_0 instead of r from Mexican flu data?

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4 Modern approaches

- Why are people using model worlds with no observation error?
 - or no process error?
- Sometimes they are good enough (model validation)
- Combining both is *hard*

Filtering

- Filtering is a little like shooting
 - Simulate from beginning to end, but use *stochastic* simulations
- You need a lot of simulations, and often ways of selecting and refining them
- A popular, state-of-the-art method is implemented in the R package pomp

Latent variable methods

- Latent variable methods are a little like stepping
 - But we step to and from unknown values (our latent variables), so we need a way of exploring many possibilities
- Popular, state-of-the-art methods are available in the R packages `rjags` and `rstan`

Latent variables

- We model *observed* variables:
 - e.g., reported cases, estimated prevalence
- Using *unobserved* latent variables
 - true cases, true prevalence, true number of susceptibles

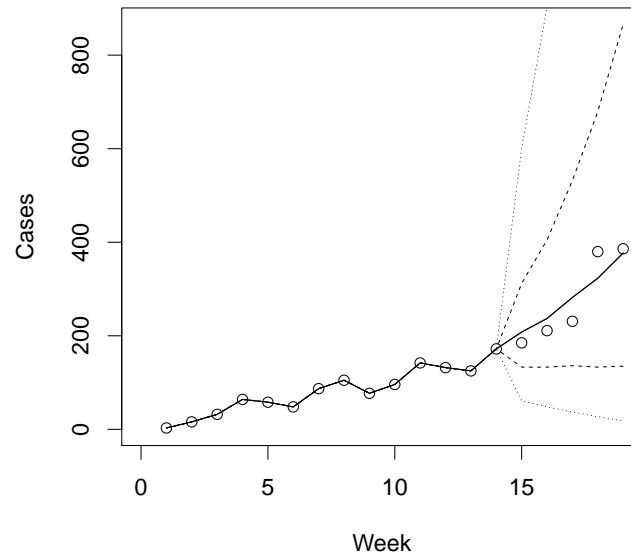
Requires a framework that can address the fact that our latent variables have many possible values

Multi-parameter inference

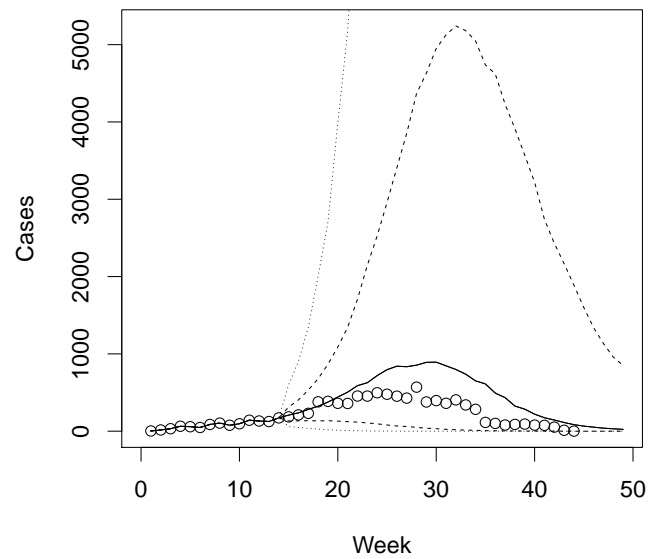
- Modern methods are already hard, and when you consider various sources of uncertainty, you're really on the bleeding edge
- Many high-profile models for Ebola, for example failed to consider process error.
- The biggest paper talking about process error neglected uncertainty in generation intervals
- Once you do multi-parameter inference, you may find that confidence intervals are very large – this may reflect the reality of knowledge, but may not make you look good

Assessing and reporting uncertainty

Sierra Leone



Sierra Leone



5 Paradigms

Likelihood

- Maximum likelihood and likelihood are not the same thing
- Bayesian approaches and frequentist approaches (including maximum likelihood) *both* depend on calculating (or approximating) likelihood

Frequentist inference

- To do frequentist inference on these complicated likelihoods, we need to:
 - estimate likelihoods
 - find the maximum likelihood
 - use the likelihood ratio test to find confidence intervals
- This is hard

Bayesian inference

- To do Bayesian inference on these complicated likelihoods, we need to:
 - construct prior distributions
 - estimate likelihoods
 - estimate the posterior
- Usually *a little* less hard
 - But still requires more assumptions

Summary

- We need **dynamics** to understand links between processes and outcomes
 - How do things work?
- We need **statistics** to understand uncertainty
 - What can we learn from *data*
- Combining these two is difficult, but progress is being made.

Summary

- Making your fit into a likelihood clarifies assumptions and creates a foundation for statistical inference
- Accounting for both process and observation error is hard
 - and not always necessary
- Stepping methods don't allow for observation error
 - filtering methods can address this
- Shooting methods don't allow for process error
 - latent variable methods can address this