

# Likelihood fitting and dynamic models II Approaches to fitting

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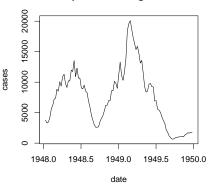
MMED 2017

# 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

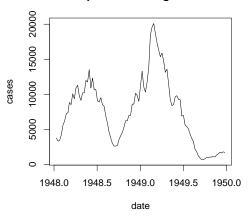
Measles reports from England and Wales



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

# Why did I get the wrong answer?

#### Measles reports from England and Wales



# 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
  - \* Parameters may not be "identifiable"

#### **Outline**

Conceptual framework

Fitting

Likelihoods

Modern approaches

Paradigms

# 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

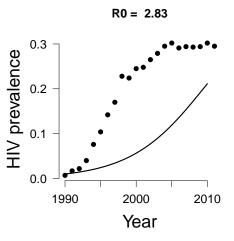
# Observation error only

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

#### **Shooting**

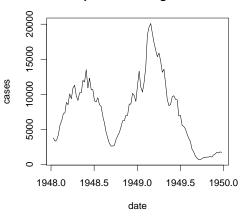


# Shooting



# Shooting

#### Measles reports from England and Wales



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# Process error only

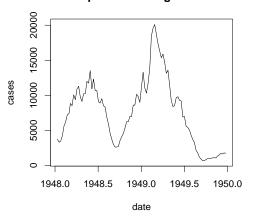
- Look at each step separately.
- See how the model is doing for that step.
- Reset based on observed data before taking the next step

**Stepping** 

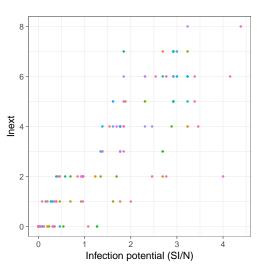


# Stepping

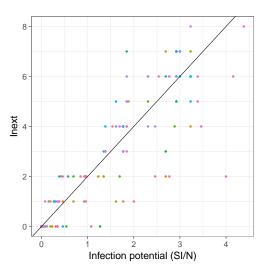
#### Measles reports from England and Wales



# Stepping



# 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)
  - \* Ignore observation error?
- What if we have a large number of cases, and poor reporting (HIV in Harare)?
  - \* Ignore process error?
- What if we have a new epidemic and poor reporting (West African Ebola)?
  - \* Can't really ignore either kind



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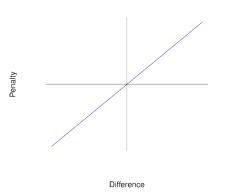
Paradigms

#### How to fit?

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

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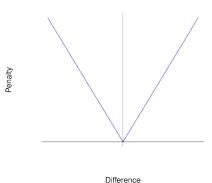
## Distance functions



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



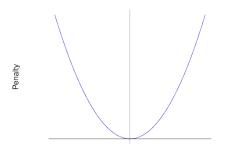
#### Distance functions



$$D=\sum_{i}|y_{i}-\hat{y}_{i}|$$



#### Distance functions



Difference

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





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#### 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?

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

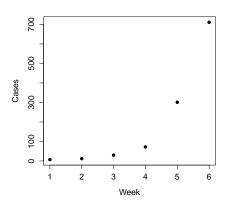
$$\ell = \sum_{i} -\log(\sigma\sqrt{2\pi}) - \sum_{i} \frac{(\hat{y}_i - y_i)^2}{2\sigma^2}$$

- We minimize the likelihood by minimizing the sum of squares
  - and then solving for  $\sigma$

## Least squares → likelihood

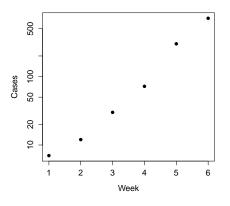
- Attaching your least squares fit to a likelihood means:
  - You can use it for statistical inference (LRT)
  - You can challenge the assumptions

# Mexican flu example



- How fast is it growing? r
- ► How hard will it be to control? R<sub>0</sub>

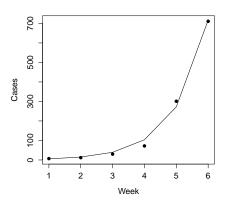
# A different perspective



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

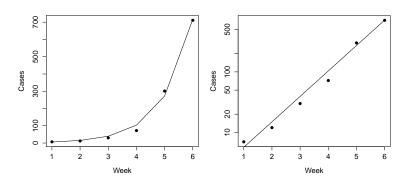


# Normal assumption

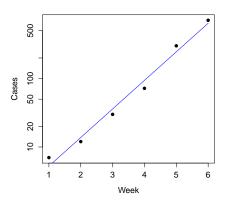


- Least squares on the linear scale
- **1**0:50 :: 980:1020
- Gives relatively too much weight to large observations

# Normal assumption

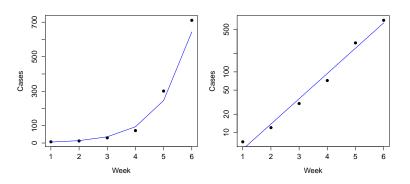


# Lognormal assumption



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

# Lognormal assumption



#### A more realistic error distribution

- My case counts are individuals
- What distributions can I use to reflect that?
- \* Poisson or binomial
  - \* WRONG!
  - **▶** \* *Sorry*:
    - \* OK, technically it's right, but you shouldn't do it.

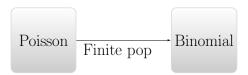
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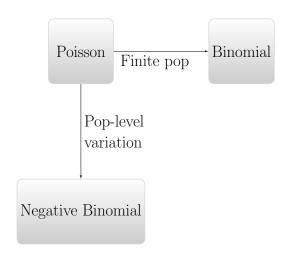
# Reality is complicated

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

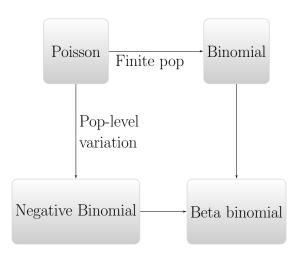


Poisson





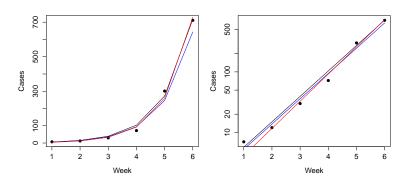
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#### Discrete distributions

- 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 exampling, when sampling

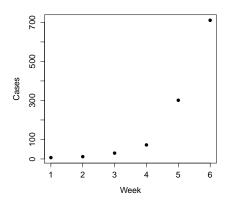
# Negative binomial fits



# Comparison

- Realistic error distribution provides (apparently) better fits
- Confidence intervals
  - Normal: r = 0.96-0.97/wk
  - Lognormal: r = 0.64-1.29/wk
  - Negative binomial: r = 0.90−1.14/wk
- How would you test these methods?
  - \* Validation: use simulated data to see if your method is reliable

# Identifiability



- What if we tried to estimate R<sub>0</sub> instead of r from Mexican flu data?
  - ▶ \* Disease could be fast with low  $\mathcal{R}_0$  or slow with high  $\mathcal{R}_0$ .

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# Modern approaches

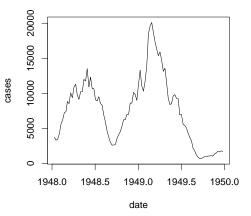
- Why are people using model worlds with no observation error?
  - or no process error?
- Sometimes they are good enough (model validation)
- Combining both is hard

# **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

# **Filtering**

#### Measles reports from England and Wales

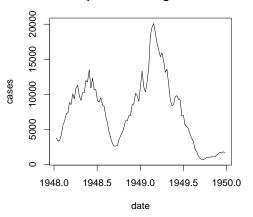


#### Latent variable methods

- 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-variable methods

#### Measles reports from England and Wales



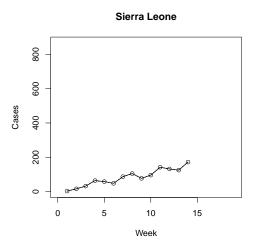
#### Latent variables

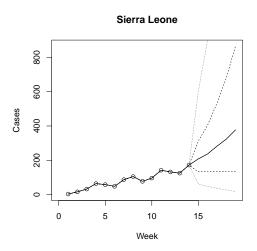
- We model observed variables:
  - e.g., reported cases, estimated prevalence
- Using unobserved latent variables
  - true cases, true prevalence, true number of susceptibles

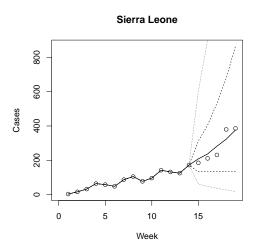
Reauires a framework that can address the fact that our latent variables have many possible values

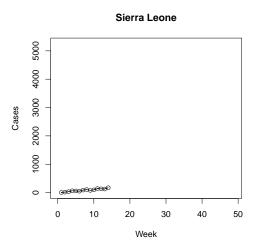
## Multi-parameter inference

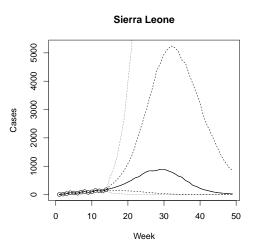
- Modern methods are already hard, and when you consider various sources of uncertainty, you're really on the bleeding edge
- Many high-profile models for Ebola, for example failed to consider process error.
- The biggest paper talking about process error neglected uncertainty in generation intervals
- 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

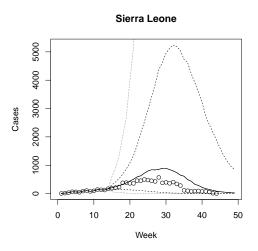












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#### Likelihood

- 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





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